Clinical trial results:
A Phase 1/2 Study of CPI-0610, a Small Molecule Inhibitor of BET Proteins: Phase 1 (Dose Escalation of CPI-0610 in Patients with Hematological Malignancies) and Phase 2 (Dose Expansion of CPI-0610 with and without Ruxolitinib in Patients with Myeloproliferative Neoplasms)

Summary

EudraCT number	2018-000579-34
Trial protocol	GB NL BE PL IT
Global end of trial date	09 January 2025

Results information

Result version number	v2 (current)
This version publication date	03 July 2026
First version publication date	20 November 2025
Version creation reason	

Trial information

Trial identification

Sponsor protocol code	0610-02

Additional study identifiers

ISRCTN number	-
ClinicalTrials.gov id (NCT number)	NCT02158858
WHO universal trial number (UTN)	-
Other trial identifiers	Novartis: CDAK539A12201

Notes:

Sponsors

Sponsor organisation name	Novartis Pharma AG, on behalf of Constellation Pharmaceuticals
Sponsor organisation address	Lichtstrasse 35, Basel, Switzerland, 4056
Public contact	Clinical Disclosure Office, Novartis Pharma AG, on behalf of Constellation Pharmaceuticals, 41 613241111, Novartis.email@Novartis.com
Scientific contact	Clinical Disclosure Office, Novartis Pharma AG, on behalf of Constellation Pharmaceuticals, 41 613241111, Novartis.email@Novartis.com

Notes:

Paediatric regulatory details

Is trial part of an agreed paediatric investigation plan (PIP)	No
Does article 45 of REGULATION (EC) No 1901/2006 apply to this trial?	No
Does article 46 of REGULATION (EC) No 1901/2006 apply to this trial?	No

Notes:

Results analysis stage

Analysis stage	Final
Date of interim/final analysis	09 January 2025
Is this the analysis of the primary completion data?	Yes
Primary completion date	09 January 2025
Global end of trial reached?	Yes
Global end of trial date	09 January 2025
Was the trial ended prematurely?	No

Notes:

General information about the trial

Main objective of the trial:

The primary objective of Phase 1 (dose escalation) was to determine the dose-limiting toxicities (DLTs) and the maximum tolerated dose (MTD) of pelabresib (CPI-0610) in patients with acute leukemia, myelodysplastic syndromes (MDS), myelodysplastic/myeloproliferative neoplasms (MDS/MPN), or myelofibrosis (MF). The primary objective of Phase 2 (dose expansion of pelabresib with and without ruxolitinib in patients with MPN) was to evaluate in MF patients splenic response by imaging after 24 weeks of treatment and, for MF patients who were transfusion dependent, to assess the transfusion independence rate. In patients with essential thrombocytopenia (ET) the primary objective was to evaluate the complete hematological response rate of pelabresib monotherapy.

Protection of trial subjects:

The study was in compliance with the ethical principles derived from the Declaration of Helsinki and the International Conference on Harmonization (ICH) Good Clinical Practice (GCP) guidelines. All the local regulatory requirements pertinent to safety of trial subjects were also followed during the conduct of the trial.

Background therapy: -

Evidence for comparator: -

Actual start date of recruitment	16 July 2014
Long term follow-up planned	No
Independent data monitoring committee (IDMC) involvement?	No

Notes:

Population of trial subjects

Subjects enrolled per country

Country: Number of subjects enrolled	Belgium: 17
Country: Number of subjects enrolled	Canada: 29
Country: Number of subjects enrolled	France: 10
Country: Number of subjects enrolled	Germany: 5
Country: Number of subjects enrolled	Italy: 55
Country: Number of subjects enrolled	Netherlands: 5
Country: Number of subjects enrolled	Poland: 8
Country: Number of subjects enrolled	United Kingdom: 44
Country: Number of subjects enrolled	United States: 163
Worldwide total number of subjects	336
EEA total number of subjects	100

Notes:

Subjects enrolled per age group	
In utero	0
Preterm newborn - gestational age < 37 wk	0
Newborns (0-27 days)	0
Infants and toddlers (28 days-23 months)	0
Children (2-11 years)	0
Adolescents (12-17 years)	0
Adults (18-64 years)	108
From 65 to 84 years	222
85 years and over	6

Subject disposition

Recruitment

Recruitment details:

This study was conducted at 54 centers across 9 countries (Belgium, Canada, France, Germany, Italy, Netherlands, Poland, United Kingdom, United States)

Pre-assignment

Screening details:

Not completed = Participants who discontinued study treatment

Period 1

Period 1 title	Overall Study (overall period)
Is this the baseline period?	Yes
Allocation method	Non-randomised - controlled
Blinding used	Not blinded

Arms

Are arms mutually exclusive?	Yes
Arm title	Phase 1 (24 mg capsule PO daily)

Arm description:

Phase 1 (24 mg capsule PO daily): Eligible participants in Phase I were enrolled in sequential cohorts based on diagnosis and received escalating doses of pelabresib (CPI-0610) once daily for 14 days, followed by a 7-day break in each 21-day cycle. Treatment stopped upon disease progression, unacceptable toxicity, or pregnancy.

Arm type	Experimental
Investigational medicinal product name	Pelabresib
Investigational medicinal product code	
Other name	CPI-0610, DAK539
Pharmaceutical forms	Capsule
Routes of administration	Oral use

Dosage and administration details:

Pelabresib was administered orally once daily for 14 consecutive days, followed by a 7-day break (1 cycle = 21 days)

Arm title	Phase 1 (48 mg capsule PO daily)

Arm description:

Phase 1 (48 mg capsule PO daily): Eligible participants in Phase I were enrolled in sequential cohorts based on diagnosis and received escalating doses of pelabresib (CPI-0610) once daily for 14 days, followed by a 7-day break in each 21-day cycle. Treatment stopped upon disease progression, unacceptable toxicity, or pregnancy.

Arm type	Experimental
Investigational medicinal product name	Pelabresib
Investigational medicinal product code	
Other name	CPI-0610, DAK539
Pharmaceutical forms	Capsule
Routes of administration	Oral use

Dosage and administration details:

Pelabresib was administered orally once daily for 14 consecutive days, followed by a 7-day break (1 cycle = 21 days)

Arm title	Phase 1 (120 mg capsule PO daily)

Arm description:

Phase 1 (120 mg capsule PO daily): Eligible participants in Phase I were enrolled in sequential cohorts based on diagnosis and received escalating doses of pelabresib (CPI-0610) once daily for 14 days, followed by a 7-day break in each 21-day cycle. Treatment stopped upon disease progression, unacceptable toxicity, or pregnancy.

Arm type	Experimental
Investigational medicinal product name	Pelabresib
Investigational medicinal product code	
Other name	CPI-0610, DAK539
Pharmaceutical forms	Capsule
Routes of administration	Oral use

Dosage and administration details:

Pelabresib was administered orally once daily for 14 consecutive days, followed by a 7-day break (1 cycle = 21 days)

Arm title	Phase 1 (170 mg capsule PO daily)

Arm description:

Phase 1 (170 mg capsule PO daily): Eligible participants in Phase I were enrolled in sequential cohorts based on diagnosis and received escalating doses of pelabresib (CPI-0610) once daily for 14 days, followed by a 7-day break in each 21-day cycle. Treatment stopped upon disease progression, unacceptable toxicity, or pregnancy.

Arm type	Experimental
Investigational medicinal product name	Pelabresib
Investigational medicinal product code	
Other name	CPI-0610, DAK539
Pharmaceutical forms	Capsule
Routes of administration	Oral use

Dosage and administration details:

Pelabresib was administered orally once daily for 14 consecutive days, followed by a 7-day break (1 cycle = 21 days)

Arm title	Phase 1 (230 mg capsule PO daily)

Arm description:

Phase 1 (230 mg capsule PO daily): Eligible participants in Phase I were enrolled in sequential cohorts based on diagnosis and received escalating doses of pelabresib (CPI-0610) once daily for 14 days, followed by a 7-day break in each 21-day cycle. Treatment stopped upon disease progression, unacceptable toxicity, or pregnancy.

Arm type	Experimental
Investigational medicinal product name	Pelabresib
Investigational medicinal product code	
Other name	CPI-0610, DAK539
Pharmaceutical forms	Capsule
Routes of administration	Oral use

Dosage and administration details:

Pelabresib was administered orally once daily for 14 consecutive days, followed by a 7-day break (1 cycle = 21 days)

Arm title	Phase 1 (300 mg capsule PO daily)

Arm description:

Phase 1 (300 mg capsule PO daily): Eligible participants in Phase I were enrolled in sequential cohorts based on diagnosis and received escalating doses of pelabresib (CPI-0610) once daily for 14 days, followed by a 7-day break in each 21-day cycle. Treatment stopped upon disease progression, unacceptable toxicity, or pregnancy.

Arm type	Experimental
Investigational medicinal product name	Pelabresib
Investigational medicinal product code	
Other name	CPI-0610, DAK539
Pharmaceutical forms	Capsule
Routes of administration	Oral use

Dosage and administration details:

Pelabresib was administered orally once daily for 14 consecutive days, followed by a 7-day break (1 cycle = 21 days)

Arm title	Phase 1 (400 mg capsule PO daily)

Arm description:

Phase 1 (400 mg capsule PO daily): Eligible participants in Phase I were enrolled in sequential cohorts based on diagnosis and received escalating doses of pelabresib (CPI-0610) once daily for 14 days, followed by a 7-day break in each 21-day cycle. Treatment stopped upon disease progression, unacceptable toxicity, or pregnancy.

Arm type	Experimental
Investigational medicinal product name	Pelabresib
Investigational medicinal product code	
Other name	CPI-0610, DAK539
Pharmaceutical forms	Capsule
Routes of administration	Oral use

Dosage and administration details:

Pelabresib was administered orally once daily for 14 consecutive days, followed by a 7-day break (1 cycle = 21 days)

Arm title	Phase 1 (225 mg tablet PO daily)

Arm description:

Phase 1 (225 mg tablet PO daily): Eligible participants in Phase I were enrolled in sequential cohorts based on diagnosis and received escalating doses of pelabresib (CPI-0610) once daily for 14 days, followed by a 7-day break in each 21-day cycle. Treatment stopped upon disease progression, unacceptable toxicity, or pregnancy.

Arm type	Experimental
Investigational medicinal product name	Pelabresib
Investigational medicinal product code	
Other name	CPI-0610, DAK539
Pharmaceutical forms	Tablet
Routes of administration	Oral use

Dosage and administration details:

Pelabresib was administered orally once daily for 14 consecutive days, followed by a 7-day break (1 cycle = 21 days)

Arm title	Phase 1 (275 mg tablet PO daily)

Arm description:

Phase 1 (275 mg tablet PO daily): Eligible participants in Phase I were enrolled in sequential cohorts based on diagnosis and received escalating doses of pelabresib (CPI-0610) once daily for 14 days, followed by a 7-day break in each 21-day cycle. Treatment stopped upon disease progression, unacceptable toxicity, or pregnancy.

Arm type	Experimental
Investigational medicinal product name	Pelabresib
Investigational medicinal product code	
Other name	CPI-0610, DAK539
Pharmaceutical forms	Tablet
Routes of administration	Oral use

Dosage and administration details:

Pelabresib was administered orally once daily for 14 consecutive days, followed by a 7-day break (1 cycle = 21 days)

Arm title	Ph2 Arm1 MF w/ prior JAKi – Cohort 1A

Arm description:

In Phase 2 (Arm 1) - Cohort 1A, eligible transfusion-dependent (TD) participants received Pelabresib 125 mg QD (tablet) for 14 days, then 7-day break (21-day cycle). Upward titration allowed up to 225 mg QD based on platelet count, hemoglobin, and safety. Treatment stopped upon disease progression, unacceptable toxicity, or pregnancy.

Arm type	Experimental

Investigational medicinal product name	Pelabresib
Investigational medicinal product code	
Other name	CPI-0610, DAK539
Pharmaceutical forms	Tablet
Routes of administration	Oral use

Dosage and administration details:

Pelabresib was administered orally once daily for 14 consecutive days, followed by a 7-day break (1 cycle = 21 days)

Arm title	Ph2 Arm1 MF w/ prior JAKi – Cohort 1B

Arm description:

In Phase 2 (Arm 1) - Cohort 1B, eligible non-transfusion-dependent (non-TD) participants received Pelabresib 125 mg QD (tablet) for 14 days, then 7-day break (21-day cycle). Upward titration allowed up to 225 mg QD based on platelet count, hemoglobin, and safety. Treatment stopped upon disease progression, unacceptable toxicity, or pregnancy.

Arm type	Experimental
Investigational medicinal product name	Pelabresib
Investigational medicinal product code	
Other name	CPI-0610, DAK539
Pharmaceutical forms	Tablet
Routes of administration	Oral use

Dosage and administration details:

Pelabresib was administered orally once daily for 14 consecutive days, followed by a 7-day break (1 cycle = 21 days)

Arm title	Ph2 Arm2 MF JAKi combo – Cohort 2A

Arm description:

In Phase 2 (Arm 2) – Cohort 2A, eligible transfusion-dependent (TD) participants already on ruxolitinib received Pelabresib 125 mg once daily for 14 days followed by a 7-day break (21-day cycle), alongside their stable dose of ruxolitinib. Pelabresib could be titrated up to 225 mg daily. Treatment was discontinued upon disease progression, unacceptable toxicity, or pregnancy.

Arm type	Experimental
Investigational medicinal product name	Ruxolitinib
Investigational medicinal product code	
Other name	
Pharmaceutical forms	Tablet
Routes of administration	Oral use

Dosage and administration details:

Ruxolitinib was given orally, twice daily (BID), on a continuous basis for 21 consecutive days of each 21-day cycle.

Investigational medicinal product name	Pelabresib
Investigational medicinal product code	
Other name	CPI-0610, DAK539
Pharmaceutical forms	Tablet
Routes of administration	Oral use

Dosage and administration details:

Pelabresib was administered orally once daily for 14 consecutive days, followed by a 7-day break (1 cycle = 21 days)

Arm title	Ph2 Arm2 MF JAKi combo – Cohort 2B

Arm description:

In Phase 2 (Arm 2) – Cohort 2B, eligible non-transfusion-dependent (non-TD) participants already receiving ruxolitinib were treated with Pelabresib 125 mg once daily for 14 days, followed by a 7-day break (21-day cycle), alongside their stable ruxolitinib dose. Pelabresib could be titrated up to 225 mg daily. Treatment was discontinued upon disease progression, unacceptable toxicity, or pregnancy.

Arm type	Experimental

Investigational medicinal product name	Ruxolitinib
Investigational medicinal product code	
Other name	
Pharmaceutical forms	Tablet
Routes of administration	Oral use

Dosage and administration details:

Ruxolitinib was given orally, twice daily (BID), on a continuous basis for 21 consecutive days of each 21-day cycle.

Investigational medicinal product name	Pelabresib
Investigational medicinal product code	
Other name	CPI-0610, DAK539
Pharmaceutical forms	Tablet
Routes of administration	Oral use

Dosage and administration details:

Pelabresib was administered orally once daily for 14 consecutive days, followed by a 7-day break (1 cycle = 21 days)

Arm title	Ph2 Arm3 MF JAKi-naïve – Combo Tx

Arm description:

Eligible participants in Phase 2 (Arm 3) received Pelabresib 125 mg once daily for 14 days followed by a 7-day break (21-day cycle), combined with Ruxolitinib, initiated at one dose level below the recommended amount based on baseline platelet count. Dose escalation: a) Ruxolitinib: Required increase of 5 mg twice daily at Cycle 3 Day 1 if criteria were met, up to 25 mg twice daily; b) Pelabresib: Optional increase from Cycle 5 Day 1 in 25 mg steps, no more than once every two cycles, up to 175 mg once daily. Treatment was discontinued in cases of disease progression, unacceptable toxicity, or pregnancy.

Arm type	Experimental
Investigational medicinal product name	Ruxolitinib
Investigational medicinal product code	
Other name	
Pharmaceutical forms	Tablet
Routes of administration	Oral use

Dosage and administration details:

Ruxolitinib was given orally, twice daily (BID), on a continuous basis for 21 consecutive days of each 21-day cycle.

Investigational medicinal product name	Pelabresib
Investigational medicinal product code	
Other name	CPI-0610, DAK539
Pharmaceutical forms	Tablet
Routes of administration	Oral use

Dosage and administration details:

Pelabresib was administered orally once daily for 14 consecutive days, followed by a 7-day break (1 cycle = 21 days)

Arm title	Ph2 Arm4 ET – Monotherapy

Arm description:

Eligible participants in Phase 2 (Arm 4) received Pelabresib 225 mg once daily (tablet) was administered for 14 days followed by a 7-day break (21-day cycle). No dose increases beyond 225 mg once daily were permitted. Treatment was discontinued in cases of disease progression, unacceptable toxicity, or pregnancy.

Arm type	Experimental
Investigational medicinal product name	Pelabresib
Investigational medicinal product code	
Other name	CPI-0610, DAK539
Pharmaceutical forms	Tablet
Routes of administration	Oral use

Dosage and administration details:

Pelabresib was administered orally once daily for 14 consecutive days, followed by a 7-day break (1 cycle = 21 days)

Number of subjects in period 1	Phase 1 (24 mg capsule PO daily)	Phase 1 (48 mg capsule PO daily)	Phase 1 (120 mg capsule PO daily)
Started	3	5	5
Completed	0	0	0
Not completed	3	5	5
Adverse event, serious fatal	-	-	-
Consent withdrawn by subject	1	1	1
Physician decision	-	-	-
Disease progression	1	3	3
Adverse event, non-fatal	-	1	1
Transitioned to pelabresib extension study	-	-	-
Other protocol defined stopping criteria	1	-	-
Cell transplant	-	-	-

Number of subjects in period 1	Phase 1 (170 mg capsule PO daily)	Phase 1 (230 mg capsule PO daily)	Phase 1 (300 mg capsule PO daily)
Started	3	3	4
Completed	0	0	0
Not completed	3	3	4
Adverse event, serious fatal	-	-	-
Consent withdrawn by subject	-	-	-
Physician decision	-	-	-
Disease progression	2	3	2
Adverse event, non-fatal	1	-	2
Transitioned to pelabresib extension study	-	-	-
Other protocol defined stopping criteria	-	-	-
Cell transplant	-	-	-

Number of subjects in period 1	Phase 1 (400 mg capsule PO daily)	Phase 1 (225 mg tablet PO daily)	Phase 1 (275 mg tablet PO daily)
Started	7	8	6
Completed	0	0	0
Not completed	7	8	6
Adverse event, serious fatal	-	1	1
Consent withdrawn by subject	1	3	2

Physician decision	1	-	2
Disease progression	3	1	-
Adverse event, non-fatal	1	3	1
Transitioned to pelabresib extension study	-	-	-
Other protocol defined stopping criteria	1	-	-
Cell transplant	-	-	-

Number of subjects in period 1	Ph2 Arm1 MF w/ prior JAKi – Cohort 1A	Ph2 Arm1 MF w/ prior JAKi – Cohort 1B	Ph2 Arm2 MF JAKi combo – Cohort 2A
Started	48	52	59
Completed	0	0	0
Not completed	48	52	59
Adverse event, serious fatal	4	3	4
Consent withdrawn by subject	5	8	6
Physician decision	20	13	14
Disease progression	5	9	13
Adverse event, non-fatal	10	7	17
Transitioned to pelabresib extension study	2	7	3
Other protocol defined stopping criteria	2	1	-
Cell transplant	-	4	2

Number of subjects in period 1	Ph2 Arm2 MF JAKi combo – Cohort 2B	Ph2 Arm3 MF JAKi-naïve – Combo Tx	Ph2 Arm4 ET – Monotherapy
Started	28	84	21
Completed	0	0	0
Not completed	28	84	21
Adverse event, serious fatal	1	8	-
Consent withdrawn by subject	3	10	4
Physician decision	6	10	3
Disease progression	7	12	1
Adverse event, non-fatal	6	14	4
Transitioned to pelabresib extension study	-	14	9
Other protocol defined stopping criteria	2	3	-
Cell transplant	3	13	-

Baseline characteristics

Reporting groups

Reporting group title	Phase 1 (24 mg capsule PO daily)

Reporting group description:

Phase 1 (24 mg capsule PO daily): Eligible participants in Phase I were enrolled in sequential cohorts based on diagnosis and received escalating doses of pelabresib (CPI-0610) once daily for 14 days, followed by a 7-day break in each 21-day cycle. Treatment stopped upon disease progression, unacceptable toxicity, or pregnancy.

Reporting group title	Phase 1 (48 mg capsule PO daily)

Reporting group description:

Phase 1 (48 mg capsule PO daily): Eligible participants in Phase I were enrolled in sequential cohorts based on diagnosis and received escalating doses of pelabresib (CPI-0610) once daily for 14 days, followed by a 7-day break in each 21-day cycle. Treatment stopped upon disease progression, unacceptable toxicity, or pregnancy.

Reporting group title	Phase 1 (120 mg capsule PO daily)

Reporting group description:

Phase 1 (120 mg capsule PO daily): Eligible participants in Phase I were enrolled in sequential cohorts based on diagnosis and received escalating doses of pelabresib (CPI-0610) once daily for 14 days, followed by a 7-day break in each 21-day cycle. Treatment stopped upon disease progression, unacceptable toxicity, or pregnancy.

Reporting group title	Phase 1 (170 mg capsule PO daily)

Reporting group description:

Phase 1 (170 mg capsule PO daily): Eligible participants in Phase I were enrolled in sequential cohorts based on diagnosis and received escalating doses of pelabresib (CPI-0610) once daily for 14 days, followed by a 7-day break in each 21-day cycle. Treatment stopped upon disease progression, unacceptable toxicity, or pregnancy.

Reporting group title	Phase 1 (230 mg capsule PO daily)

Reporting group description:

Phase 1 (230 mg capsule PO daily): Eligible participants in Phase I were enrolled in sequential cohorts based on diagnosis and received escalating doses of pelabresib (CPI-0610) once daily for 14 days, followed by a 7-day break in each 21-day cycle. Treatment stopped upon disease progression, unacceptable toxicity, or pregnancy.

Reporting group title	Phase 1 (300 mg capsule PO daily)

Reporting group description:

Phase 1 (300 mg capsule PO daily): Eligible participants in Phase I were enrolled in sequential cohorts based on diagnosis and received escalating doses of pelabresib (CPI-0610) once daily for 14 days, followed by a 7-day break in each 21-day cycle. Treatment stopped upon disease progression, unacceptable toxicity, or pregnancy.

Reporting group title	Phase 1 (400 mg capsule PO daily)

Reporting group description:

Phase 1 (400 mg capsule PO daily): Eligible participants in Phase I were enrolled in sequential cohorts based on diagnosis and received escalating doses of pelabresib (CPI-0610) once daily for 14 days, followed by a 7-day break in each 21-day cycle. Treatment stopped upon disease progression, unacceptable toxicity, or pregnancy.

Reporting group title	Phase 1 (225 mg tablet PO daily)

Reporting group description:

Phase 1 (225 mg tablet PO daily): Eligible participants in Phase I were enrolled in sequential cohorts based on diagnosis and received escalating doses of pelabresib (CPI-0610) once daily for 14 days, followed by a 7-day break in each 21-day cycle. Treatment stopped upon disease progression, unacceptable toxicity, or pregnancy.

Reporting group title	Phase 1 (275 mg tablet PO daily)

Reporting group description:

Phase 1 (275 mg tablet PO daily): Eligible participants in Phase I were enrolled in sequential cohorts based on diagnosis and received escalating doses of pelabresib (CPI-0610) once daily for 14 days, followed by a 7-day break in each 21-day cycle. Treatment stopped upon disease progression, unacceptable toxicity, or pregnancy.

Reporting group title	Ph2 Arm1 MF w/ prior JAKi – Cohort 1A

Reporting group description:

In Phase 2 (Arm 1) - Cohort 1A, eligible transfusion-dependent (TD) participants received Pelabresib 125 mg QD (tablet) for 14 days, then 7-day break (21-day cycle). Upward titration allowed up to 225 mg QD based on platelet count, hemoglobin, and safety. Treatment stopped upon disease progression, unacceptable toxicity, or pregnancy.

Reporting group title	Ph2 Arm1 MF w/ prior JAKi – Cohort 1B

Reporting group description:

In Phase 2 (Arm 1) - Cohort 1B, eligible non-transfusion-dependent (non-TD) participants received Pelabresib 125 mg QD (tablet) for 14 days, then 7-day break (21-day cycle). Upward titration allowed up to 225 mg QD based on platelet count, hemoglobin, and safety. Treatment stopped upon disease progression, unacceptable toxicity, or pregnancy.

Reporting group title	Ph2 Arm2 MF JAKi combo – Cohort 2A

Reporting group description:

In Phase 2 (Arm 2) – Cohort 2A, eligible transfusion-dependent (TD) participants already on ruxolitinib received Pelabresib 125 mg once daily for 14 days followed by a 7-day break (21-day cycle), alongside their stable dose of ruxolitinib. Pelabresib could be titrated up to 225 mg daily. Treatment was discontinued upon disease progression, unacceptable toxicity, or pregnancy.

Reporting group title	Ph2 Arm2 MF JAKi combo – Cohort 2B

Reporting group description:

In Phase 2 (Arm 2) – Cohort 2B, eligible non-transfusion-dependent (non-TD) participants already receiving ruxolitinib were treated with Pelabresib 125 mg once daily for 14 days, followed by a 7-day break (21-day cycle), alongside their stable ruxolitinib dose. Pelabresib could be titrated up to 225 mg daily. Treatment was discontinued upon disease progression, unacceptable toxicity, or pregnancy.

Reporting group title	Ph2 Arm3 MF JAKi-naïve – Combo Tx

Reporting group description:

Eligible participants in Phase 2 (Arm 3) received Pelabresib 125 mg once daily for 14 days followed by a 7-day break (21-day cycle), combined with Ruxolitinib, initiated at one dose level below the recommended amount based on baseline platelet count. Dose escalation: a) Ruxolitinib: Required increase of 5 mg twice daily at Cycle 3 Day 1 if criteria were met, up to 25 mg twice daily; b) Pelabresib: Optional increase from Cycle 5 Day 1 in 25 mg steps, no more than once every two cycles, up to 175 mg once daily. Treatment was discontinued in cases of disease progression, unacceptable toxicity, or pregnancy.

Reporting group title	Ph2 Arm4 ET – Monotherapy

Reporting group description:

Eligible participants in Phase 2 (Arm 4) received Pelabresib 225 mg once daily (tablet) was administered for 14 days followed by a 7-day break (21-day cycle). No dose increases beyond 225 mg once daily were permitted. Treatment was discontinued in cases of disease progression, unacceptable toxicity, or pregnancy.

Reporting group values	Phase 1 (24 mg capsule PO daily)	Phase 1 (48 mg capsule PO daily)	Phase 1 (120 mg capsule PO daily)
Number of subjects	3	5	5
Age categorical Units: Subjects			
In utero	0	0	0
Preterm newborn infants (gestational age < 37 wks)	0	0	0
Newborns (0-27 days)	0	0	0
Infants and toddlers (28 days-23 months)	0	0	0
Children (2-11 years)	0	0	0
Adolescents (12-17 years)	0	0	0
Adults (18-64 years)	1	2	2
From 65-84 years	2	3	3
85 years and over	0	0	0

Age Continuous Units: Years			
median	79.0	65.0	69.0
full range (min-max)	37 to 79	55 to 76	34 to 80
Sex: Female, Male Units: Participants			
Female	3	1	1
Male	0	4	4
Race (NIH/OMB) Units: Subjects			
American Indian or Alaska Native	0	0	0
Asian	0	1	0
Native Hawaiian or Other Pacific Islander	0	0	0
Black or African American	0	0	1
White	3	4	4
More than one race	0	0	0
Unknown or Not Reported	0	0	0

Reporting group values	Phase 1 (170 mg capsule PO daily)	Phase 1 (230 mg capsule PO daily)	Phase 1 (300 mg capsule PO daily)
Number of subjects	3	3	4
Age categorical Units: Subjects			
In utero	0	0	0
Preterm newborn infants (gestational age < 37 wks)	0	0	0
Newborns (0-27 days)	0	0	0
Infants and toddlers (28 days-23 months)	0	0	0
Children (2-11 years)	0	0	0
Adolescents (12-17 years)	0	0	0
Adults (18-64 years)	0	1	2
From 65-84 years	3	2	2
85 years and over	0	0	0
Age Continuous Units: Years			
median	76.0	80.0	66.5
full range (min-max)	69 to 79	57 to 82	27 to 76
Sex: Female, Male Units: Participants			
Female	3	1	0
Male	0	2	4
Race (NIH/OMB) Units: Subjects			
American Indian or Alaska Native	0	0	0
Asian	0	0	0
Native Hawaiian or Other Pacific Islander	0	0	0
Black or African American	0	0	1
White	3	3	1
More than one race	0	0	2
Unknown or Not Reported	0	0	0

Reporting group values	Phase 1 (400 mg capsule PO daily)	Phase 1 (225 mg tablet PO daily)	Phase 1 (275 mg tablet PO daily)
Number of subjects	7	8	6
Age categorical Units: Subjects			
In utero	0	0	0
Preterm newborn infants (gestational age < 37 wks)	0	0	0
Newborns (0-27 days)	0	0	0
Infants and toddlers (28 days-23 months)	0	0	0
Children (2-11 years)	0	0	0
Adolescents (12-17 years)	0	0	0
Adults (18-64 years)	4	4	3
From 65-84 years	3	4	3
85 years and over	0	0	0
Age Continuous Units: Years			
median	63.0	65.5	61.0
full range (min-max)	59 to 77	54 to 82	18 to 80
Sex: Female, Male Units: Participants			
Female	2	1	2
Male	5	7	4
Race (NIH/OMB) Units: Subjects			
American Indian or Alaska Native	0	0	0
Asian	0	0	0
Native Hawaiian or Other Pacific Islander	0	0	0
Black or African American	1	0	2
White	6	8	4
More than one race	0	0	0
Unknown or Not Reported	0	0	0

Reporting group values	Ph2 Arm1 MF w/ prior JAKi – Cohort 1A	Ph2 Arm1 MF w/ prior JAKi – Cohort 1B	Ph2 Arm2 MF JAKi combo – Cohort 2A
Number of subjects	48	52	59
Age categorical Units: Subjects			
In utero	0	0	0
Preterm newborn infants (gestational age < 37 wks)	0	0	0
Newborns (0-27 days)	0	0	0
Infants and toddlers (28 days-23 months)	0	0	0
Children (2-11 years)	0	0	0
Adolescents (12-17 years)	0	0	0
Adults (18-64 years)	8	15	11
From 65-84 years	37	35	48
85 years and over	3	2	0

Age Continuous Units: Years			
median	73.0	69.5	72.0
full range (min-max)	60 to 88	43 to 85	41 to 83
Sex: Female, Male Units: Participants			
Female	16	29	18
Male	32	23	41
Race (NIH/OMB) Units: Subjects			
American Indian or Alaska Native	0	0	0
Asian	1	1	1
Native Hawaiian or Other Pacific Islander	0	0	0
Black or African American	4	1	2
White	42	47	54
More than one race	1	2	1
Unknown or Not Reported	0	1	1

Reporting group values	Ph2 Arm2 MF JAKi combo – Cohort 2B	Ph2 Arm3 MF JAKi-naïve – Combo Tx	Ph2 Arm4 ET – Monotherapy
Number of subjects	28	84	21
Age categorical Units: Subjects			
In utero	0	0	0
Preterm newborn infants (gestational age < 37 wks)	0	0	0
Newborns (0-27 days)	0	0	0
Infants and toddlers (28 days-23 months)	0	0	0
Children (2-11 years)	0	0	0
Adolescents (12-17 years)	0	0	0
Adults (18-64 years)	13	31	11
From 65-84 years	15	52	10
85 years and over	0	1	0
Age Continuous Units: Years			
median	65.5	68.0	64.0
full range (min-max)	49 to 78	37 to 85	42 to 83
Sex: Female, Male Units: Participants			
Female	13	25	13
Male	15	59	8
Race (NIH/OMB) Units: Subjects			
American Indian or Alaska Native	0	0	0
Asian	1	3	0
Native Hawaiian or Other Pacific Islander	0	0	0
Black or African American	4	3	0
White	22	75	19
More than one race	1	3	2
Unknown or Not Reported	0	0	0

Reporting group values	Total		
Number of subjects	336		
Age categorical Units: Subjects			
In utero	0		
Preterm newborn infants (gestational age < 37 wks)	0		
Newborns (0-27 days)	0		
Infants and toddlers (28 days-23 months)	0		
Children (2-11 years)	0		
Adolescents (12-17 years)	0		
Adults (18-64 years)	108		
From 65-84 years	222		
85 years and over	6		
Age Continuous Units: Years			
median			
full range (min-max)	-		
Sex: Female, Male Units: Participants			
Female	128		
Male	208		
Race (NIH/OMB) Units: Subjects			
American Indian or Alaska Native	0		
Asian	8		
Native Hawaiian or Other Pacific Islander	0		
Black or African American	19		
White	295		
More than one race	12		
Unknown or Not Reported	2		

End points reporting groups

Reporting group title	Phase 1 (24 mg capsule PO daily)

Reporting group description:

Phase 1 (24 mg capsule PO daily): Eligible participants in Phase I were enrolled in sequential cohorts based on diagnosis and received escalating doses of pelabresib (CPI-0610) once daily for 14 days, followed by a 7-day break in each 21-day cycle. Treatment stopped upon disease progression, unacceptable toxicity, or pregnancy.

Reporting group title	Phase 1 (48 mg capsule PO daily)

Reporting group description:

Phase 1 (48 mg capsule PO daily): Eligible participants in Phase I were enrolled in sequential cohorts based on diagnosis and received escalating doses of pelabresib (CPI-0610) once daily for 14 days, followed by a 7-day break in each 21-day cycle. Treatment stopped upon disease progression, unacceptable toxicity, or pregnancy.

Reporting group title	Phase 1 (120 mg capsule PO daily)

Reporting group description:

Phase 1 (120 mg capsule PO daily): Eligible participants in Phase I were enrolled in sequential cohorts based on diagnosis and received escalating doses of pelabresib (CPI-0610) once daily for 14 days, followed by a 7-day break in each 21-day cycle. Treatment stopped upon disease progression, unacceptable toxicity, or pregnancy.

Reporting group title	Phase 1 (170 mg capsule PO daily)

Reporting group description:

Phase 1 (170 mg capsule PO daily): Eligible participants in Phase I were enrolled in sequential cohorts based on diagnosis and received escalating doses of pelabresib (CPI-0610) once daily for 14 days, followed by a 7-day break in each 21-day cycle. Treatment stopped upon disease progression, unacceptable toxicity, or pregnancy.

Reporting group title	Phase 1 (230 mg capsule PO daily)

Reporting group description:

Phase 1 (230 mg capsule PO daily): Eligible participants in Phase I were enrolled in sequential cohorts based on diagnosis and received escalating doses of pelabresib (CPI-0610) once daily for 14 days, followed by a 7-day break in each 21-day cycle. Treatment stopped upon disease progression, unacceptable toxicity, or pregnancy.

Reporting group title	Phase 1 (300 mg capsule PO daily)

Reporting group description:

Phase 1 (300 mg capsule PO daily): Eligible participants in Phase I were enrolled in sequential cohorts based on diagnosis and received escalating doses of pelabresib (CPI-0610) once daily for 14 days, followed by a 7-day break in each 21-day cycle. Treatment stopped upon disease progression, unacceptable toxicity, or pregnancy.

Reporting group title	Phase 1 (400 mg capsule PO daily)

Reporting group description:

Phase 1 (400 mg capsule PO daily): Eligible participants in Phase I were enrolled in sequential cohorts based on diagnosis and received escalating doses of pelabresib (CPI-0610) once daily for 14 days, followed by a 7-day break in each 21-day cycle. Treatment stopped upon disease progression, unacceptable toxicity, or pregnancy.

Reporting group title	Phase 1 (225 mg tablet PO daily)

Reporting group description:

Phase 1 (225 mg tablet PO daily): Eligible participants in Phase I were enrolled in sequential cohorts based on diagnosis and received escalating doses of pelabresib (CPI-0610) once daily for 14 days, followed by a 7-day break in each 21-day cycle. Treatment stopped upon disease progression, unacceptable toxicity, or pregnancy.

Reporting group title	Phase 1 (275 mg tablet PO daily)

Reporting group description:

Phase 1 (275 mg tablet PO daily): Eligible participants in Phase I were enrolled in sequential cohorts based on diagnosis and received escalating doses of pelabresib (CPI-0610) once daily for 14 days, followed by a 7-day break in each 21-day cycle. Treatment stopped upon disease progression, unacceptable toxicity, or pregnancy.

Reporting group title	Ph2 Arm1 MF w/ prior JAKi – Cohort 1A

Reporting group description:

In Phase 2 (Arm 1) - Cohort 1A, eligible transfusion-dependent (TD) participants received Pelabresib 125 mg QD (tablet) for 14 days, then 7-day break (21-day cycle). Upward titration allowed up to 225 mg QD based on platelet count, hemoglobin, and safety. Treatment stopped upon disease progression, unacceptable toxicity, or pregnancy.

Reporting group title	Ph2 Arm1 MF w/ prior JAKi – Cohort 1B

Reporting group description:

In Phase 2 (Arm 1) - Cohort 1B, eligible non-transfusion-dependent (non-TD) participants received Pelabresib 125 mg QD (tablet) for 14 days, then 7-day break (21-day cycle). Upward titration allowed up to 225 mg QD based on platelet count, hemoglobin, and safety. Treatment stopped upon disease progression, unacceptable toxicity, or pregnancy.

Reporting group title	Ph2 Arm2 MF JAKi combo – Cohort 2A

Reporting group description:

In Phase 2 (Arm 2) – Cohort 2A, eligible transfusion-dependent (TD) participants already on ruxolitinib received Pelabresib 125 mg once daily for 14 days followed by a 7-day break (21-day cycle), alongside their stable dose of ruxolitinib. Pelabresib could be titrated up to 225 mg daily. Treatment was discontinued upon disease progression, unacceptable toxicity, or pregnancy.

Reporting group title	Ph2 Arm2 MF JAKi combo – Cohort 2B

Reporting group description:

In Phase 2 (Arm 2) – Cohort 2B, eligible non-transfusion-dependent (non-TD) participants already receiving ruxolitinib were treated with Pelabresib 125 mg once daily for 14 days, followed by a 7-day break (21-day cycle), alongside their stable ruxolitinib dose. Pelabresib could be titrated up to 225 mg daily. Treatment was discontinued upon disease progression, unacceptable toxicity, or pregnancy.

Reporting group title	Ph2 Arm3 MF JAKi-naïve – Combo Tx

Reporting group description:

Eligible participants in Phase 2 (Arm 3) received Pelabresib 125 mg once daily for 14 days followed by a 7-day break (21-day cycle), combined with Ruxolitinib, initiated at one dose level below the recommended amount based on baseline platelet count. Dose escalation: a) Ruxolitinib: Required increase of 5 mg twice daily at Cycle 3 Day 1 if criteria were met, up to 25 mg twice daily; b) Pelabresib: Optional increase from Cycle 5 Day 1 in 25 mg steps, no more than once every two cycles, up to 175 mg once daily. Treatment was discontinued in cases of disease progression, unacceptable toxicity, or pregnancy.

Reporting group title	Ph2 Arm4 ET – Monotherapy

Reporting group description:

Eligible participants in Phase 2 (Arm 4) received Pelabresib 225 mg once daily (tablet) was administered for 14 days followed by a 7-day break (21-day cycle). No dose increases beyond 225 mg once daily were permitted. Treatment was discontinued in cases of disease progression, unacceptable toxicity, or pregnancy.

Primary: Phase 1: Frequency of Dose-limiting toxicities (DLTs)

End point title	Phase 1: Frequency of Dose-limiting toxicities (DLTs)[1][2]

End point description:

A dose-limiting toxicity (DLT) was defined as an adverse event or abnormal laboratory value assessed by the Investigator as unrelated to disease progression, intercurrent illness, or concomitant medications that occurred within the first cycle of treatment (21-day cycle) with pelabresib (CPI-0610), and that met any of the criteria specified in the protocol.

End point type	Primary

End point timeframe:

Up to 21 days

Notes:

[1] - No statistical analyses have been specified for this primary end point. It is expected there is at least one statistical analysis for each primary end point.
Justification: Only descriptive statistics performed

[2] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: Endpoint only applicable to Phase 1

End point values	Phase 1 (24 mg capsule PO daily)	Phase 1 (48 mg capsule PO daily)	Phase 1 (120 mg capsule PO daily)	Phase 1 (170 mg capsule PO daily)
Subject group type	Reporting group	Reporting group	Reporting group	Reporting group
Number of subjects analysed	3	4	5	3
Units: Participants	0	0	0	0

End point values	Phase 1 (230 mg capsule PO daily)	Phase 1 (300 mg capsule PO daily)	Phase 1 (400 mg capsule PO daily)	Phase 1 (225 mg tablet PO daily)
Subject group type	Reporting group	Reporting group	Reporting group	Reporting group
Number of subjects analysed	3	3	6	6
Units: Participants	0	0	1	2

End point values	Phase 1 (275 mg tablet PO daily)			
Subject group type	Reporting group			
Number of subjects analysed	5			
Units: Participants	4			

Statistical analyses

No statistical analyses for this end point

Primary: Phase 2 (Cohorts 1B, 2B, and Arm 3): Number of Participants with Splenic Response Rate (SVR35) at week 24

End point title	Phase 2 (Cohorts 1B, 2B, and Arm 3): Number of Participants with Splenic Response Rate (SVR35) at week 24[3][4]

End point description:

Splenic Response Rate (SVR35) at Week 24 was defined as the proportion of participants who demonstrated a reduction of at least 35% in spleen size from baseline, as measured by imaging techniques (MRI or CT), following 24 weeks of treatment.

End point type	Primary

End point timeframe:

Week 24 (Cycle 9 Day 1)

Notes:

[3] - No statistical analyses have been specified for this primary end point. It is expected there is at least one statistical analysis for each primary end point.
Justification: Only descriptive statistics performed

[4] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: Endpoint only applicable to Phase 2 (Cohorts 1B, 2B, and Arm 3)

End point values	Ph2 Arm1 MF w/ prior JAKi – Cohort 1B	Ph2 Arm2 MF JAKi combo – Cohort 2B	Ph2 Arm3 MF JAKi-naïve – Combo Tx	
Subject group type	Reporting group	Reporting group	Reporting group	
Number of subjects analysed	52	28	84	
Units: Percentage of Participants				
number (confidence interval 95%)	19.2 (9.6 to 32.5)	21.4 (8.3 to 41.0)	67.9 (56.8 to 77.6)	

Statistical analyses

No statistical analyses for this end point

Primary: Phase 2 (Cohorts 1A and 2A): Number of Participants enrolled as Transfusion Dependent (TD) with Conversion rate from Red Blood Cell (RBC) transfusion dependence (TD) to transfusion independence (TI)

End point title	Phase 2 (Cohorts 1A and 2A): Number of Participants enrolled as Transfusion Dependent (TD) with Conversion rate from Red Blood Cell (RBC) transfusion dependence (TD) to transfusion independence (TI)[5][6]

End point description:

Conversion rate was defined as the proportion of participants who converted from transfusion dependence (TD) to transfusion independence (TI). TD was characterized by receiving an average of at least 2 units of red blood cell (RBC) transfusions per month—amounting to a minimum of 6 units over the 12 weeks prior to enrollment—while TI was defined as the absence of RBC transfusions during any consecutive 12-week period.

End point type	Primary

End point timeframe:

Any 12 consecutive weeks (rolling window) during active treatment with pelabresib (CPI0610), from first dose through treatment discontinuation (up to approximately 8 years for Phase II)

Notes:

[5] - No statistical analyses have been specified for this primary end point. It is expected there is at least one statistical analysis for each primary end point.
Justification: Only descriptive statistics performed

[6] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: Endpoint only applicable to Phase 2 (Cohorts 1A and 2A)

End point values	Ph2 Arm1 MF w/ prior JAKi – Cohort 1A	Ph2 Arm2 MF JAKi combo – Cohort 2A		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	48	59		
Units: Percentage of Participants				
number (confidence interval 95%)	25.6 (13.5 to 41.2)	26.1 (14.3 to 41.1)		

Statistical analyses

No statistical analyses for this end point

Primary: Phase 2 (Arm 4): Number of Participants with Complete Hematological

Response (CHR) Rate

End point title	Phase 2 (Arm 4): Number of Participants with Complete Hematological Response (CHR) Rate[7][8]

End point description:

Complete Hematological Response (CHR) Rate was defined as the proportion of participants who fulfilled the criteria for CHR, based on the modified European LeukemiaNet (ELN) guidelines (Barosi et al 2009): platelet count ≤400 × 10/L, white blood cell (WBC) count ≤10 × 10/L, confirmation of laboratory values after one treatment cycle, and normal spleen size determined by palpation or imaging.

End point type	Primary

End point timeframe:

Over 2 consecutive cycles (rolling window) (1 cycle = 21 days)

Notes:

[7] - No statistical analyses have been specified for this primary end point. It is expected there is at least one statistical analysis for each primary end point.
Justification: Only descriptive statistics performed

[8] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: Endpoint only applicable to Phase 2 (Arm 4)

End point values	Ph2 Arm4 ET – Monotherapy			
Subject group type	Reporting group			
Number of subjects analysed	21			
Units: Percentage of Participants				
number (confidence interval 95%)	57.1 (34.0 to 78.2)			

Statistical analyses

No statistical analyses for this end point

Secondary: Phase 1: Number of adverse events and serious adverse events as assessed by CTCAE criteria

End point title	Phase 1: Number of adverse events and serious adverse events as assessed by CTCAE criteria[9]

End point description:

The distribution of adverse events was performed through the analysis of frequencies for treatment-emergent adverse events (TEAEs) and serious adverse events (TESAEs), based on the monitoring of relevant clinical and laboratory safety parameters. Treatment-emergent adverse events (TEAEs) in this study were defined as events that began after the first dose of study treatment and continued until 30 days after the last dose, or events that were present prior to the first dose and increased in severity based on preferred term within 30 days following the last dose of pelabresib (CPI-0610).

End point type	Secondary

End point timeframe:

Up to approximately 6 months

Notes:

[9] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: Endpoint only applicable to Phase 1

End point values	Phase 1 (24 mg capsule PO daily)	Phase 1 (48 mg capsule PO daily)	Phase 1 (120 mg capsule PO daily)	Phase 1 (170 mg capsule PO daily)
Subject group type	Reporting group	Reporting group	Reporting group	Reporting group
Number of subjects analysed	3	5	5	3
Units: Participants				
Participants with at least 1 TEAE	3	5	5	3
Participants with ≥ Grade 3 TEAEs	2	4	3	3
Pts w/ any drug-related TEAE	3	3	5	3
Pts w/ drug-related TEAEs ≥ Gr3 (CTCAE)	1	1	1	0
Participants with any serious TEAEs	1	4	3	1
Participants with any drug-related serious TEAEs	0	0	1	0
Pts w/ TEAEs leading to drug discontinuation	0	2	2	1

End point values	Phase 1 (230 mg capsule PO daily)	Phase 1 (300 mg capsule PO daily)	Phase 1 (400 mg capsule PO daily)	Phase 1 (225 mg tablet PO daily)
Subject group type	Reporting group	Reporting group	Reporting group	Reporting group
Number of subjects analysed	3	4	7	8
Units: Participants				
Participants with at least 1 TEAE	3	4	7	8
Participants with ≥ Grade 3 TEAEs	2	4	7	8
Pts w/ any drug-related TEAE	3	3	6	7
Pts w/ drug-related TEAEs ≥ Gr3 (CTCAE)	0	0	5	3
Participants with any serious TEAEs	2	4	6	6
Participants with any drug-related serious TEAEs	0	0	1	0
Pts w/ TEAEs leading to drug discontinuation	0	2	2	4

End point values	Phase 1 (275 mg tablet PO daily)			
Subject group type	Reporting group			
Number of subjects analysed	6			
Units: Participants				
Participants with at least 1 TEAE	6			
Participants with ≥ Grade 3 TEAEs	6			
Pts w/ any drug-related TEAE	5			
Pts w/ drug-related TEAEs ≥ Gr3 (CTCAE)	4			
Participants with any serious TEAEs	6			
Participants with any drug-related serious TEAEs	3			
Pts w/ TEAEs leading to drug discontinuation	3			

Statistical analyses

No statistical analyses for this end point

Secondary: Phase 2 (All Arms): Number of adverse events and serious adverse events as assessed by CTCAE criteria

End point title	Phase 2 (All Arms): Number of adverse events and serious adverse events as assessed by CTCAE criteria[10]

End point description:

The distribution of adverse events was performed through the analysis of frequencies for treatment-emergent adverse events (TEAEs) and serious adverse events (TESAEs), based on the monitoring of relevant clinical and laboratory safety parameters. Treatment-emergent adverse events (TEAEs) in this study were defined as events that began after the first dose of study treatment and continued until 30 days after the last dose, or events that were present prior to the first dose and increased in severity based on preferred term within 30 days following the last dose of pelabresib (CPI-0610).

End point type	Secondary

End point timeframe:

Up to approximately 387 weeks

Notes:

[10] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: Endpoint only applicable to Phase 2 (All Arms)

End point values	Ph2 Arm1 MF w/ prior JAKi – Cohort 1A	Ph2 Arm1 MF w/ prior JAKi – Cohort 1B	Ph2 Arm2 MF JAKi combo – Cohort 2A	Ph2 Arm2 MF JAKi combo – Cohort 2B
Subject group type	Reporting group	Reporting group	Reporting group	Reporting group
Number of subjects analysed	48	52	59	28
Units: Participants				
Participants with at least 1 TEAE	48	51	59	28
Participants with ≥ Grade 3 TEAEs	38	40	49	21
Pts w/ ≥1 TEAE related to pelabresib	45	44	53	27
Pts w/ ≥Gr3 TEAEs related to pelabresib	27	21	33	14
Participants with any serious TEAEs	18	23	29	18
Pts w/ serious TEAEs related to pelabresib	4	3	2	6
Pts w/ TEAEs causing pelabresib interruption	20	16	32	13
Pts w/ TEAEs causing pelabresib dose reduction	16	10	15	7
Pts w/ TEAEs causing pelabresib discontinuation	12	12	17	7
Pts w/ TEAEs causing study discontinuation	11	11	18	9

End point values	Ph2 Arm3 MF JAKi-naïve –	Ph2 Arm4 ET – Monotherapy		

	Combo Tx			
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	84	21		
Units: Participants				
Participants with at least 1 TEAE	84	21		
Participants with ≥ Grade 3 TEAEs	66	10		
Pts w/ ≥1 TEAE related to pelabresib	74	20		
Pts w/ ≥Gr3 TEAEs related to pelabresib	41	6		
Participants with any serious TEAEs	40	5		
Pts w/ serious TEAEs related to pelabresib	16	2		
Pts w/ TEAEs causing pelabresib interruption	39	9		
Pts w/ TEAEs causing pelabresib dose reduction	38	6		
Pts w/ TEAEs causing pelabresib discontinuation	22	3		
Pts w/ TEAEs causing study discontinuation	22	5		

Statistical analyses

No statistical analyses for this end point

Secondary: Phase 2 (All Arms): Symptom improvement from the Patient Global Impression of Change (PGIC) at 12 and 24 weeks

End point title	Phase 2 (All Arms): Symptom improvement from the Patient Global Impression of Change (PGIC) at 12 and 24 weeks[11]

End point description:

The Patient Global Impression of Change (PGIC) was a single-question, patient-reported assessment that asked individuals to rate their overall change in myeloproliferative neoplasm (MPN) symptoms since starting study treatment. The participants selected one of seven options, ranging from 'Very much improved' to 'Very much worse'. 'No Change from Baseline' indicated stable symptoms (no improvement or worsening), negative change from Baseline indicated a reduction in symptom severity (improvement), and positive change from Baseline indicated an increase in symptom severity (worsening).

End point type	Secondary

End point timeframe:

Week 12 (Cycle 5 Day 1), Week 24 weeks (Cycle 9 Day 1)

Notes:

[11] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: Endpoint only applicable to Phase 2 (All Arms)

End point values	Ph2 Arm1 MF w/ prior JAKi – Cohort 1A	Ph2 Arm1 MF w/ prior JAKi – Cohort 1B	Ph2 Arm2 MF JAKi combo – Cohort 2A	Ph2 Arm2 MF JAKi combo – Cohort 2B
Subject group type	Reporting group	Reporting group	Reporting group	Reporting group
Number of subjects analysed	39	43	47	25
Units: Participants				
PGIC status at Week 12|Improvement	28	36	22	18
PGIC status at Week 24|Improvement	20	29	27	16
PGIC status at Week 12|No change	7	2	15	4

PGIC status at Week 24|No change	10	3	12	6
PGIC status at Week 12|Worsening	4	5	10	3
PGIC status at Week 24|Worsening	2	5	4	0

End point values	Ph2 Arm3 MF JAKi-naïve – Combo Tx	Ph2 Arm4 ET – Monotherapy		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	80	19		
Units: Participants				
PGIC status at Week 12|Improvement	66	13		
PGIC status at Week 24|Improvement	58	8		
PGIC status at Week 12|No change	9	4		
PGIC status at Week 24|No change	15	3		
PGIC status at Week 12|Worsening	5	2		
PGIC status at Week 24|Worsening	6	4		

Statistical analyses

No statistical analyses for this end point

Secondary: Phase 2 (Arms 1, 2 and 3): Percent Change from Baseline in Total Symptom Score (TSS) from the Myelofibrosis Symptom Assessment Form (MFSAF v4.0) at 12 and 24 weeks

End point title	Phase 2 (Arms 1, 2 and 3): Percent Change from Baseline in Total Symptom Score (TSS) from the Myelofibrosis Symptom Assessment Form (MFSAF v4.0) at 12 and 24 weeks[12]

End point description:

The MFSAF (Myelofibrosis Symptom Assessment Form) was completed by participants every day for 7 days before Day 1 of each treatment cycle, including the 7 days before starting Cycle 1. It used a 24-hour recall format, asking participants to rate the worst severity of seven symptoms (fatigue, night sweats, pruritus, abdominal discomfort, pain under the ribs on the left side, early satiety, and bone pain) during the past 24 hours. Each symptom was rated on a scale from 0 (Absent) to 10 (Worst Imaginable). The Total Symptom Score (TSS) was the sum of 7 symptoms (range: 0–70). 'No Change from Baseline' indicated stable symptoms (no improvement or worsening), negative change from Baseline indicated a reduction in symptom severity (improvement), and positive change from Baseline indicated an increase in symptom severity (worsening).

End point type	Secondary

End point timeframe:

Baseline, Week 12 (Cycle 5 Day 1), Week 24 (Cycle 9 Day 1)

Notes:

[12] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: Endpoint only applicable to Phase 2 (Arms 1, 2 and 3)

End point values	Ph2 Arm1 MF w/ prior JAKi – Cohort 1A	Ph2 Arm1 MF w/ prior JAKi – Cohort 1B	Ph2 Arm2 MF JAKi combo – Cohort 2A	Ph2 Arm2 MF JAKi combo – Cohort 2B
Subject group type	Reporting group	Reporting group	Reporting group	Reporting group
Number of subjects analysed	38	43	48	25
Units: Score on a scale				
arithmetic mean (standard deviation)				
Percent Change from baseline to Week 12	-38.77 (± 38.637)	-30.98 (± 35.394)	-27.18 (± 75.391)	-29.70 (± 36.106)
Percent Change from baseline to Week 24	-32.69 (± 36.474)	-39.82 (± 42.228)	-38.93 (± 61.818)	-44.73 (± 35.224)

End point values	Ph2 Arm3 MF JAKi-naïve – Combo Tx			
Subject group type	Reporting group			
Number of subjects analysed	82			
Units: Score on a scale				
arithmetic mean (standard deviation)				
Percent Change from baseline to Week 12	-39.91 (± 69.251)			
Percent Change from baseline to Week 24	-47.43 (± 52.460)			

Statistical analyses

No statistical analyses for this end point

Secondary: Phase 2 (Arms 1, 2 and 3): Number of Participants who achieved a ≥ 50% reduction in Total Symptom Score (TSS) at 12 and 24 weeks

End point title	Phase 2 (Arms 1, 2 and 3): Number of Participants who achieved a ≥ 50% reduction in Total Symptom Score (TSS) at 12 and 24 weeks[13]

End point description:

The proportion of study participants who experienced a reduction of at least 50% in their Total Symptom Score (TSS), as assessed using the Myelofibrosis Symptom Assessment Form (MFSAF v4.0), was evaluated at both 12 and 24 weeks relative to their baseline score.

End point type	Secondary

End point timeframe:

Week 12 (Cycle 5 Day 1), Week 24 (Cycle 9 Day 1)

Notes:

[13] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: Endpoint only applicable to Phase 2 (Arms 1, 2 and 3)

End point values	Ph2 Arm1 MF w/ prior JAKi – Cohort 1A	Ph2 Arm1 MF w/ prior JAKi – Cohort 1B	Ph2 Arm2 MF JAKi combo – Cohort 2A	Ph2 Arm2 MF JAKi combo – Cohort 2B
Subject group type	Reporting group	Reporting group	Reporting group	Reporting group
Number of subjects analysed	48	52	59	28
Units: Percentage of Participants				
number (confidence interval 95%)				
MFSAF TSS50 response at week 12	33.3 (20.0 to 49.0)	31.3 (18.7 to 46.3)	36.2 (24.0 to 49.9)	33.3 (16.5 to 54.0)
MFSAF TSS50 response at week 24	15.6 (6.5 to 29.5)	34.7 (21.7 to 49.6)	36.2 (24.0 to 49.9)	40.7 (22.4 to 61.2)

End point values	Ph2 Arm3 MF JAKi-naïve – Combo Tx			
Subject group type	Reporting group			
Number of subjects analysed	84			
Units: Percentage of Participants				
number (confidence interval 95%)				
MFSAF TSS50 response at week 12	52.4 (41.1 to 63.6)			
MFSAF TSS50 response at week 24	56.1 (44.7 to 67.0)			

Statistical analyses

No statistical analyses for this end point

Secondary: Phase 2 (Arms 1, 2 and 3): Number of Participants with Overall Splenic Response Rate (overall SVR35)

End point title	Phase 2 (Arms 1, 2 and 3): Number of Participants with Overall Splenic Response Rate (overall SVR35)[14]

End point description:

Overall Splenic Response Rate (SVR35) was defined as the proportion of participants who achieved a reduction of at least 35% in spleen size from baseline, as measured by imaging (MRI or CT), at any point between Cycle 1 Day 1 and the End of Study Visit, whichever occurred first.

End point type	Secondary

End point timeframe:

Through Phase II completion, an average of 6 years

Notes:

[14] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: Endpoint only applicable to Phase 2 (Arms 1, 2 and 3)

End point values	Ph2 Arm1 MF w/ prior JAKi – Cohort 1A	Ph2 Arm1 MF w/ prior JAKi – Cohort 1B	Ph2 Arm2 MF JAKi combo – Cohort 2A	Ph2 Arm2 MF JAKi combo – Cohort 2B
Subject group type	Reporting group	Reporting group	Reporting group	Reporting group
Number of subjects analysed	48	52	59	28
Units: Percentage of Participants				
number (confidence interval 95%)	16.7 (7.0 to 31.4)	28.8 (17.1 to 43.1)	30.8 (18.7 to 45.1)	25.0 (10.7 to 44.9)

End point values	Ph2 Arm3 MF JAKi-naïve – Combo Tx			
Subject group type	Reporting group			
Number of subjects analysed	84			
Units: Percentage of Participants				
number (confidence interval 95%)	79.8 (69.6 to 87.7)			

Statistical analyses

No statistical analyses for this end point

Secondary: Phase 2 (Arms 1, 2 and 3): Duration of Overall Splenic Response (overall SVR35)

End point title	Phase 2 (Arms 1, 2 and 3): Duration of Overall Splenic Response (overall SVR35)[15]

End point description:

Duration of Overall Splenic Response (overall SVR35) was defined as the time from the first occurrence of a at least 35% reduction in spleen volume from baseline until the earliest of the following: a reduction of less than 35% from baseline combined with an increase of more than 25% from the nadir in spleen volume (as measured by MRI or CT), or death. The nadir was defined as the lowest spleen volume recorded after baseline and up to the evaluation point at which the initial splenic response was achieved.

End point type	Secondary

End point timeframe:

From first onset of splenic response until loss of response, assessed up to approximately 6 years

Notes:

[15] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: Endpoint only applicable to Phase 2 (Arms 1, 2 and 3)

End point values	Ph2 Arm1 MF w/ prior JAKi – Cohort 1A	Ph2 Arm1 MF w/ prior JAKi – Cohort 1B	Ph2 Arm2 MF JAKi combo – Cohort 2A	Ph2 Arm2 MF JAKi combo – Cohort 2B
Subject group type	Reporting group	Reporting group	Reporting group	Reporting group
Number of subjects analysed	48	52	59	28
Units: Weeks				
median (confidence interval 95%)	24.1 (12.4 to 999)	73.1 (27.1 to 106.4)	180.9 (73.0 to 999)	228.4 (36.1 to 999)

End point values	Ph2 Arm3 MF JAKi-naïve – Combo Tx			
Subject group type	Reporting group			
Number of subjects analysed	84			
Units: Weeks				
median (confidence interval 95%)	197.6 (95.7 to 999)			

Statistical analyses

No statistical analyses for this end point

Secondary: Phase 2 (Cohorts 1A and 2A): Number of Participants with Splenic Response Rate (SVR35) at 12 and 24 weeks

End point title	Phase 2 (Cohorts 1A and 2A): Number of Participants with Splenic Response Rate (SVR35) at 12 and 24 weeks[16]

End point description:

Splenic Response Rate (SVR35) at 12 and 24 weeks was defined as the proportion of participants who achieved a reduction of at least 35% in spleen size from baseline, as determined by imaging (MRI or CT), following 12 and 24 weeks of treatment, respectively.

End point type	Secondary

End point timeframe:

Week 12 (Cycle 5 Day 1), Week 24 weeks (Cycle 9 Day 1)

Notes:

[16] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: Endpoint only applicable to Phase 2 (Cohorts 1A and 2A)

End point values	Ph2 Arm1 MF w/ prior JAKi – Cohort 1A	Ph2 Arm2 MF JAKi combo – Cohort 2A		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	48	59		
Units: Percentage of Participants				
number (confidence interval 95%)				
SVR35 at week 12	9.3 (2.6 to 22.1)	9.4 (3.1 to 20.7)		
SVR35 at week 24	0 (0 to 999)	16.4 (7.8 to 28.8)		

Statistical analyses

No statistical analyses for this end point

Secondary: Phase 2 (Cohorts 1A and 2A): Duration of Red Blood Cell (RBC)

Transfusion Independence (TI) in participants who enroll as Transfusion Dependent (TD)

End point title	Phase 2 (Cohorts 1A and 2A): Duration of Red Blood Cell (RBC) Transfusion Independence (TI) in participants who enroll as Transfusion Dependent (TD)[17]

End point description:

Duration of Red Blood Cell (RBC) Transfusion Independence (TI) was defined as the longest continuous period during which participants, having achieved at least 12 weeks of transfusion independence, remained free from RBC transfusions.

End point type	Secondary

End point timeframe:

From first onset of TI to earliest onset of loss of TI, assessed up to approximately 6 years

Notes:

[17] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: Endpoint only applicable to Phase 2 (Cohorts 1A and 2A)

End point values	Ph2 Arm1 MF w/ prior JAKi – Cohort 1A	Ph2 Arm2 MF JAKi combo – Cohort 2A		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	48	59		
Units: Weeks				
median (full range (min-max))	35.8 (15.3 to 999)	68.0 (34.7 to 999)		

Statistical analyses

No statistical analyses for this end point

Secondary: Phase 2 (Cohorts 1A and 2A): Early anemic response rate in participants who enroll as Transfusion Dependent (TD)

End point title	Phase 2 (Cohorts 1A and 2A): Early anemic response rate in participants who enroll as Transfusion Dependent (TD)[18]

End point description:

Early anemic response rate was defined as the proportion of participants who achieved an average increase of at least 1 g/dL in hemoglobin concentration over any rolling 8-week (56-day) period following baseline. This calculation was performed after applying the 14/3 day rule, which stipulates that hemoglobin measurements must be spaced at least 14 days apart, and that at least 3 such measurements are required within the 8-week window to ensure a reliable average. Importantly, this increase had to occur without any red blood cell (RBC) transfusions during the treatment period and up to the time of the latest hemoglobin assessment for each patient.

End point type	Secondary

End point timeframe:

Through Phase II completion, an average of 6 years

Notes:

[18] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: Endpoint only applicable to Phase 2 (Cohorts 1A and 2A)

End point values	Ph2 Arm1 MF w/ prior JAKi – Cohort 1A	Ph2 Arm2 MF JAKi combo – Cohort 2A		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	48	59		
Units: Percentage of Participants				
number (confidence interval 95%)	18.6 (8.4 to 33.4)	21.7 (10.9 to 36.4)		

Statistical analyses

No statistical analyses for this end point

Secondary: Phase 2 (Cohorts 1B, 2B, and Arm 3): Anemic response rate in participants who enroll as non-transfusion-dependent (non-TD)

End point title	Phase 2 (Cohorts 1B, 2B, and Arm 3): Anemic response rate in participants who enroll as non-transfusion-dependent (non-TD)[19]

End point description:

Anemic response was defined as a sustained average increase in hemoglobin concentration of at least 1.5 g/dL over any rolling 12-week (84-day) period following baseline. This calculation excluded periods involving red blood cell (RBC) transfusions and was performed after applying the 14/3-day rule for valid hemoglobin assessments. The response had to be maintained through to the most recent available hemoglobin measurement for each patient.

End point type	Secondary

End point timeframe:

Through Phase II completion, an average of 6 years

Notes:

[19] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: Endpoint only applicable to Phase 2 (Cohorts 1B, 2B, and Arm 3)

End point values	Ph2 Arm1 MF w/ prior JAKi – Cohort 1B	Ph2 Arm2 MF JAKi combo – Cohort 2B	Ph2 Arm3 MF JAKi-naïve – Combo Tx	
Subject group type	Reporting group	Reporting group	Reporting group	
Number of subjects analysed	52	28	84	
Units: Percentage of Participants				
number (confidence interval 95%)	47.1 (32.9 to 61.5)	21.4 (8.3 to 41.0)	35.4 (25.0 to 47.0)	

Statistical analyses

No statistical analyses for this end point

Secondary: Phase 2 (Cohorts 1B, 2B, and Arm 3): Number of Participants with Splenic Response Rate (SVR35) at 12 weeks

End point title	Phase 2 (Cohorts 1B, 2B, and Arm 3): Number of Participants with Splenic Response Rate (SVR35) at 12 weeks[20]

End point description:

Splenic Response Rate (SVR35) at Week 12 was defined as the proportion of participants who achieved

a reduction of at least 35% in spleen size from baseline, as measured by imaging (MRI or CT), following 12 weeks of treatment.

End point type	Secondary

End point timeframe:

Week 12 (Cycle 5 Day 1)

Notes:

[20] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: Endpoint only applicable to Phase 2 (Cohorts 1B, 2B, and Arm 3)

End point values	Ph2 Arm1 MF w/ prior JAKi – Cohort 1B	Ph2 Arm2 MF JAKi combo – Cohort 2B	Ph2 Arm3 MF JAKi-naïve – Combo Tx	
Subject group type	Reporting group	Reporting group	Reporting group	
Number of subjects analysed	52	28	84	
Units: Percentage of Participants				
number (confidence interval 95%)	15.4 (6.9 to 28.1)	10.7 (2.3 to 28.2)	66.7 (55.5 to 76.6)	

Statistical analyses

No statistical analyses for this end point

Secondary: Phase 2 (Arm 4): Partial Hematological Response Rate (PHR)

End point title	Phase 2 (Arm 4): Partial Hematological Response Rate (PHR)[21]

End point description:

Partial Hematological Response Rate (PHR) was defined as the proportion of participants who met the following criteria over two consecutive cycles up to the last administration of study treatment:
• Platelet count > 400-600 x 10^9/L
• WBC count within normal range (i.e., ≤ 10 x 10^9/L)
• Laboratory results confirmed after 1 cycle (after 3weeks)

End point type	Secondary

End point timeframe:

Over 2 consecutive cycles (rolling window) (1 cycle = 21 days)

Notes:

[21] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: Endpoint only applicable to Phase 2 (Arm 4)

End point values	Ph2 Arm4 ET – Monotherapy			
Subject group type	Reporting group			
Number of subjects analysed	21			
Units: Percentage of Participants				
number (confidence interval 95%)	38.1 (18.1 to 61.6)			

Statistical analyses

No statistical analyses for this end point

Secondary: Phase 2 (Arm 4): Percentage of participants who achieve a ≥50% reduction from baseline in the Myeloproliferative Neoplasm Symptom Assessment Form (MPN-SAF) total score

End point title	Phase 2 (Arm 4): Percentage of participants who achieve a ≥50% reduction from baseline in the Myeloproliferative Neoplasm Symptom Assessment Form (MPN-SAF) total score[22]

End point description:

The Myeloproliferative Neoplasm Symptom Assessment Form (MPN-SAF) is a patient-reported questionnaire designed to measure symptom burden in participants with myeloproliferative neoplasms (MPNs). Participants rate the severity of several symptoms (such as fatigue, night sweats, itching, abdominal discomfort, bone pain, early satiety, and others) over the past 24 hours. Each symptom is scored on a scale from 0 (absent/as good as it can be) to 10 (worst imaginable/as bad as it can be). The total score is the sum of all individual symptom scores, providing an overall measure of symptom burden. A 50% reduction in the MPN-SAF total score at 12 or 24 weeks means the patient's overall symptom burden has improved by half compared to their baseline (pre-treatment) score.

End point type	Secondary

End point timeframe:

Baseline, 12 weeks (Cycle 5, Day 1), 24 weeks (Cycle 9, Day 1)

Notes:

[22] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: Endpoint only applicable to Phase 2 (Arm 4)

End point values	Ph2 Arm4 ET – Monotherapy			
Subject group type	Reporting group			
Number of subjects analysed	21			
Units: Percentage of Participants				
number (confidence interval 95%)				
MPN-SAF TSS50 response at week 12	25.0 (8.7 to 49.1)			
MPN-SAF TSS50 response at week 24	20.0 (5.7 to 43.7)			

Statistical analyses

No statistical analyses for this end point

Secondary: Phase 2 (Arm 4): Duration of Overall Hematological Response Rate (OHR)

End point title	Phase 2 (Arm 4): Duration of Overall Hematological Response Rate (OHR)[23]

End point description:

Duration of Overall Hematological Response (OHR) Rate was defined as the time from when the overall hematological response was first met until the time at which the overall hematological response was lost, i.e., the criteria for an overall hematological response (CHR or PHR) were not observed or death occurred, whichever came first.

End point type	Secondary

End point timeframe:

Through Phase II completion, an average of 6 years

Notes:

[23] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.

Justification: Endpoint only applicable to Phase 2 (Arm 4)

End point values	Ph2 Arm4 ET – Monotherapy			
Subject group type	Reporting group			
Number of subjects analysed	21			
Units: Weeks				
median (confidence interval 95%)	21.1 (9.1 to 33.1)			

Statistical analyses

No statistical analyses for this end point

Secondary: Phase 2 (Arm 4): Overall Hematological Response Rate (OHR)

End point title	Phase 2 (Arm 4): Overall Hematological Response Rate

End point description:

Confirmed Overall Hematological Response (OHR) Rate was defined as the proportion of participants with either a confirmed complete or a partial hematological response at any time.

End point type	Secondary

End point timeframe:

Through Phase II completion, an average of 6 years

Notes:

[24] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.

Justification: Endpoint only applicable to Phase 2 (Arm 4)

End point values	Ph2 Arm4 ET – Monotherapy			
Subject group type	Reporting group			
Number of subjects analysed	21			
Units: Percentage of Participants				
number (confidence interval 95%)	66.7 (43.0 to 85.4)			

Statistical analyses

No statistical analyses for this end point

Secondary: Phase 2 (Arm 4): Rate of hemorrhagic and thromboembolic (TE) events

End point title	Phase 2 (Arm 4): Rate of hemorrhagic and thromboembolic (TE) events[25]

End point description:

Rate of hemorrhagic and thromboembolic (TE) events was defined as the proportion of participants with hemorrhagic or thromboembolic events throughout the study.

End point type	Secondary

End point timeframe:

Through Phase II completion, an average of 6 years

Notes:

[25] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: Endpoint only applicable to Phase 2 (Arm 4)

End point values	Ph2 Arm4 ET – Monotherapy			
Subject group type	Reporting group			
Number of subjects analysed	21			
Units: Percentage of Participants				
number (confidence interval 95%)	61.9 (38.4 to 81.9)			

Statistical analyses

No statistical analyses for this end point

Secondary: Phase 2 (All Arms): Maximum observed plasma concentration (Cmax) of pelabresib

End point title	Phase 2 (All Arms): Maximum observed plasma concentration (Cmax) of pelabresib[26]

End point description:

Pharmacokinetic (PK) parameters were calculated based on Pelabresib plasma concentrations and actual sampling time points.
Cmax was listed and summarized using descriptive statistics.

End point type	Secondary

End point timeframe:

Cycle 1 Day 14 (30 minutes, 1 hour, 2 hours, 4 hours, 6 hours, and 8 hours post-dose). 1 cycle = 21 days.

Notes:

[26] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: Endpoint only applicable to Phase 2 (All Arms)

End point values	Ph2 Arm1 MF w/ prior JAKi – Cohort 1A	Ph2 Arm1 MF w/ prior JAKi – Cohort 1B	Ph2 Arm2 MF JAKi combo – Cohort 2A	Ph2 Arm2 MF JAKi combo – Cohort 2B
Subject group type	Reporting group	Reporting group	Reporting group	Reporting group
Number of subjects analysed	32	39	38	16
Units: ng/mL				
geometric mean (geometric coefficient of variation)	1280 (± 29.1)	1350 (± 37.0)	1210 (± 40.0)	1190 (± 29.0)

End point values	Ph2 Arm3 MF JAKi-naïve – Combo Tx	Ph2 Arm4 ET – Monotherapy		

Subject group type	Reporting group	Reporting group		
Number of subjects analysed	64	14		
Units: ng/mL				
geometric mean (geometric coefficient of variation)	1070 (± 26.3)	2130 (± 24.7)		

Statistical analyses

No statistical analyses for this end point

Secondary: Phase 2 (All Arms): Time to reach maximum concentration (Tmax) of pelabresib

End point title	Phase 2 (All Arms): Time to reach maximum concentration (Tmax) of pelabresib[27]

End point description:

Pharmacokinetic (PK) parameters were calculated based on Pelabresib plasma concentrations and actual sampling time points.
Tmax was listed and summarized using descriptive statistics.

End point type	Secondary

End point timeframe:

Cycle 1 Day 14 (predose, 30 minutes, 1 hour, 2 hours, 4 hours, 6 hours, and 8 hours post-dose). 1 cycle = 21 days.

Notes:

[27] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: Endpoint only applicable to Phase 2 (All Arms)

End point values	Ph2 Arm1 MF w/ prior JAKi – Cohort 1A	Ph2 Arm1 MF w/ prior JAKi – Cohort 1B	Ph2 Arm2 MF JAKi combo – Cohort 2A	Ph2 Arm2 MF JAKi combo – Cohort 2B
Subject group type	Reporting group	Reporting group	Reporting group	Reporting group
Number of subjects analysed	32	39	38	16
Units: Hour				
median (full range (min-max))	1.94 (0.85 to 4.02)	1.58 (0.83 to 3.63)	1.90 (0.50 to 4.00)	1.81 (0.92 to 3.13)

End point values	Ph2 Arm3 MF JAKi-naïve – Combo Tx	Ph2 Arm4 ET – Monotherapy		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	64	14		
Units: Hour				
median (full range (min-max))	1.71 (0.50 to 3.87)	2.08 (1.07 to 6.00)		

Statistical analyses

No statistical analyses for this end point

Secondary: Phase 2 (All Arms): Predose (trough) concentration at the end of a dosing interval (Ctrough) of pelabresib

End point title	Phase 2 (All Arms): Predose (trough) concentration at the end of a dosing interval (Ctrough) of pelabresib[28]

End point description:

Pharmacokinetic (PK) parameters were calculated based on Pelabresib plasma concentrations and actual sampling time points.
Ctrough was listed and summarized using descriptive statistics.

End point type	Secondary

End point timeframe:

Cycle 1 Day 14 (predose, 30 minutes, 1 hour, 2 hours, 4 hours, 6 hours, and 8 hours post-dose). 1 cycle = 21 days.

Notes:

[28] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: Endpoint only applicable to Phase 2 (All Arms)

End point values	Ph2 Arm1 MF w/ prior JAKi – Cohort 1A	Ph2 Arm1 MF w/ prior JAKi – Cohort 1B	Ph2 Arm2 MF JAKi combo – Cohort 2A	Ph2 Arm2 MF JAKi combo – Cohort 2B
Subject group type	Reporting group	Reporting group	Reporting group	Reporting group
Number of subjects analysed	31	37	36	16
Units: ng/mL				
geometric mean (geometric coefficient of variation)	120 (± 489.0)	201 (± 107.0)	244 (± 151.0)	162 (± 84.0)

End point values	Ph2 Arm3 MF JAKi-naïve – Combo Tx	Ph2 Arm4 ET – Monotherapy		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	63	12		
Units: ng/mL				
geometric mean (geometric coefficient of variation)	240 (± 79.1)	174 (± 112.0)		

Statistical analyses

No statistical analyses for this end point

Secondary: Phase 2 (All Arms): Area under the concentration–time curve from time zero to the last observed concentration (AUClast) of pelabresib

End point title	Phase 2 (All Arms): Area under the concentration–time curve from time zero to the last observed concentration (AUClast) of pelabresib[29]

End point description:

Pharmacokinetic (PK) parameters were calculated based on Pelabresib plasma concentrations and actual sampling time points.
AUClast was listed and summarized using descriptive statistics.

End point type	Secondary

End point timeframe:

Cycle 1 Day 14 (30 minutes, 1 hour, 2 hours, 4 hours, 6 hours, and 8 hours post-dose). 1 cycle = 21 days.

Notes:

[29] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: Endpoint only applicable to Phase 2 (All Arms)

End point values	Ph2 Arm1 MF w/ prior JAKi – Cohort 1A	Ph2 Arm1 MF w/ prior JAKi – Cohort 1B	Ph2 Arm2 MF JAKi combo – Cohort 2A	Ph2 Arm2 MF JAKi combo – Cohort 2B
Subject group type	Reporting group	Reporting group	Reporting group	Reporting group
Number of subjects analysed	32	39	38	16
Units: h*ng/mL				
geometric mean (geometric coefficient of variation)	5660 (± 36.7)	5610 (± 35.5)	5190 (± 40.9)	4760 (± 26.7)

End point values	Ph2 Arm3 MF JAKi-naïve – Combo Tx	Ph2 Arm4 ET – Monotherapy		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	64	14		
Units: h*ng/mL				
geometric mean (geometric coefficient of variation)	4380 (± 26.8)	9590 (± 28.7)		

Statistical analyses

No statistical analyses for this end point

Secondary: Phase 2 (All Arms): Area under the concentration–time curve from time zero to 8 hours post-dose at steady state (AUC0-8,ss) of pelabresib

End point title	Phase 2 (All Arms): Area under the concentration–time curve from time zero to 8 hours post-dose at steady state (AUC0-8,ss) of pelabresib[30]

End point description:

Pharmacokinetic (PK) parameters were calculated based on Pelabresib plasma concentrations and actual sampling time points.
AUC0-8,ss was listed and summarized using descriptive statistics.

End point type	Secondary

End point timeframe:

Cycle 1 Day 14 (30 minutes, 1 hour, 2 hours, 4 hours, 6 hours, and 8 hours post-dose). 1 cycle = 21 days.

Notes:

[30] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: Endpoint only applicable to Phase 2 (All Arms)

End point values	Ph2 Arm1 MF w/ prior JAKi – Cohort 1A	Ph2 Arm1 MF w/ prior JAKi – Cohort 1B	Ph2 Arm2 MF JAKi combo – Cohort 2A	Ph2 Arm2 MF JAKi combo – Cohort 2B
Subject group type	Reporting group	Reporting group	Reporting group	Reporting group
Number of subjects analysed	24	31	34	14
Units: h*ng/mL				
geometric mean (geometric coefficient of variation)	6200 (± 35.3)	6470 (± 31.7)	5320 (± 38.1)	4930 (± 28.8)

End point values	Ph2 Arm3 MF JAKi-naïve – Combo Tx	Ph2 Arm4 ET – Monotherapy		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	57	10		
Units: h*ng/mL				
geometric mean (geometric coefficient of variation)	4540 (± 25.3)	10800 (± 29.0)		

Statistical analyses

No statistical analyses for this end point

Secondary: Phase 2 (All Arms): Time of Last Measurable Concentration (Tlast) of pelabresib

End point title	Phase 2 (All Arms): Time of Last Measurable Concentration (Tlast) of pelabresib[31]

End point description:

Pharmacokinetic (PK) parameters were calculated based on Pelabresib plasma concentrations and actual sampling time points.
Tlast was listed and summarized using descriptive statistics.

End point type	Secondary

End point timeframe:

Cycle 1 Day 14 (30 minutes, 1 hour, 2 hours, 4 hours, 6 hours, and 8 hours post-dose). 1 cycle = 21 days.

Notes:

[31] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: Endpoint only applicable to Phase 2 (All Arms)

End point values	Ph2 Arm1 MF w/ prior JAKi – Cohort 1A	Ph2 Arm1 MF w/ prior JAKi – Cohort 1B	Ph2 Arm2 MF JAKi combo – Cohort 2A	Ph2 Arm2 MF JAKi combo – Cohort 2B
Subject group type	Reporting group	Reporting group	Reporting group	Reporting group
Number of subjects analysed	32	39	38	16
Units: Hour				
median (full range (min-max))	7.08 (4.50 to 9.00)	7.07 (2.58 to 8.08)	7.19 (6.10 to 8.05)	7.12 (7.00 to 8.17)

End point values	Ph2 Arm3 MF JAKi-naïve – Combo Tx	Ph2 Arm4 ET – Monotherapy		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	64	14		
Units: Hour				
median (full range (min-max))	7.08 (5.07 to 8.58)	7.08 (5.88 to 10.08)		

Statistical analyses

No statistical analyses for this end point

Secondary: Phase 2 (Arm 2 - Cohort 2A): Maximum observed plasma concentration (Cmax) of ruxolitinib

End point title	Phase 2 (Arm 2 - Cohort 2A): Maximum observed plasma concentration (Cmax) of ruxolitinib[32]

End point description:

Pharmacokinetic (PK) parameters were calculated based on ruxolitinib plasma concentrations and actual sampling time points.
Cmax was listed and summarized using descriptive statistics.

End point type	Secondary

End point timeframe:

Cycle 1 Day 14 (30 minutes, 1 hour, 2 hours, 4 hours, 6 hours, and 8 hours post-dose). 1 cycle = 21 days.

Notes:

[32] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: Endpoint only applicable to Phase 2 (Arm 2 - Cohort 2A)

End point values	Ph2 Arm2 MF JAKi combo – Cohort 2A			
Subject group type	Reporting group			
Number of subjects analysed	30			
Units: ng/mL				
geometric mean (geometric coefficient of variation)				
Cohort 2A, 5 mg	65.9 (± 56.5)			
Cohort 2A, 7.5 mg	142.0 (± 31.0)			
Cohort 2A, 10 mg	108.0 (± 29.3)			
Cohort 2A, 15 mg	310.0 (± 88.9)			
Cohort 2A, 20 mg	325.0 (± 37.7)			

Statistical analyses

No statistical analyses for this end point

Secondary: Phase 2 (Arm 2 - Cohort 2A): Time to reach maximum concentration (Tmax) of ruxolitinib

End point title	Phase 2 (Arm 2 - Cohort 2A): Time to reach maximum concentration (Tmax) of ruxolitinib[33]

End point description:

Pharmacokinetic (PK) parameters were calculated based on ruxolitinib plasma concentrations and actual sampling time points.
Tmax was listed and summarized using descriptive statistics.

End point type	Secondary

End point timeframe:

Cycle 1 Day 14 (30 minutes, 1 hour, 2 hours, 4 hours, 6 hours, and 8 hours post-dose). 1 cycle = 21 days.

Notes:

[33] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: Endpoint only applicable to Phase 2 (Arm 2 - Cohort 2A)

End point values	Ph2 Arm2 MF JAKi combo – Cohort 2A			
Subject group type	Reporting group			
Number of subjects analysed	30			
Units: Hour				
median (full range (min-max))				
Cohort 2A, 5 mg	0.92 (0.50 to 2.00)			
Cohort 2A, 7.5 mg	1.00 (0.50 to 2.02)			
Cohort 2A, 10 mg	0.76 (0.50 to 1.50)			
Cohort 2A, 15 mg	1.55 (0.67 to 3.50)			
Cohort 2A, 20 mg	0.53 (0.33 to 1.58)			

Statistical analyses

No statistical analyses for this end point

Secondary: Phase 2 (Arm 2 - Cohort 2A): Predose (trough) concentration at the end of a dosing interval (Ctrough) of ruxolitinib

End point title	Phase 2 (Arm 2 - Cohort 2A): Predose (trough) concentration at the end of a dosing interval (Ctrough) of ruxolitinib[34]

End point description:

Pharmacokinetic (PK) parameters were calculated based on ruxolitinib plasma concentrations and actual sampling time points.
Ctrough was listed and summarized using descriptive statistics.

End point type	Secondary

End point timeframe:

Cycle 1 Day 14 (predose, 30 minutes, 1 hour, 2 hours, 4 hours, 6 hours, and 8 hours post-dose). 1 cycle = 21 days.

Notes:

[34] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: Endpoint only applicable to Phase 2 (Arm 2 - Cohort 2A)

End point values	Ph2 Arm2 MF JAKi combo – Cohort 2A			
Subject group type	Reporting group			
Number of subjects analysed	28			
Units: ng/mL				
geometric mean (geometric coefficient of variation)				
Cohort 2A, 5 mg	999 (± 999)			
Cohort 2A, 7.5 mg	14.7 (± 74.9)			
Cohort 2A, 10 mg	13.4 (± 81.6)			
Cohort 2A, 15 mg	74.6 (± 94.7)			
Cohort 2A, 20 mg	19.8 (± 54.2)			

Statistical analyses

No statistical analyses for this end point

Secondary: Phase 2 (Arm 2 - Cohort 2A): Area under the concentration–time curve from time zero to the last observed concentration (AUClast) of ruxolitinib

End point title	Phase 2 (Arm 2 - Cohort 2A): Area under the concentration–time curve from time zero to the last observed concentration (AUClast) of ruxolitinib[35]

End point description:

Pharmacokinetic (PK) parameters were calculated based on ruxolitinib plasma concentrations and actual sampling time points.
AUClast was listed and summarized using descriptive statistics.

End point type	Secondary

End point timeframe:

Cycle 1 Day 14 (30 minutes, 1 hour, 2 hours, 4 hours, 6 hours, and 8 hours post-dose). 1 cycle = 21 days.

Notes:

[35] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: Endpoint only applicable to Phase 2 (Arm 2 - Cohort 2A)

End point values	Ph2 Arm2 MF JAKi combo – Cohort 2A			
Subject group type	Reporting group			
Number of subjects analysed	30			
Units: h*ng/mL				
geometric mean (geometric coefficient of variation)				
Cohort 2A, 5 mg	230 (± 63.3)			
Cohort 2A, 7.5 mg	390 (± 34.0)			
Cohort 2A, 10 mg	347 (± 44.7)			
Cohort 2A, 15 mg	1240 (± 64.7)			
Cohort 2A, 20 mg	829 (± 42.3)			

Statistical analyses

No statistical analyses for this end point

Secondary: Phase 2 (Arm 2 - Cohort 2A): Area under the concentration–time curve from time zero to 8 hours post-dose at steady state (AUC0-8,ss) of ruxolitinib

End point title	Phase 2 (Arm 2 - Cohort 2A): Area under the concentration–time curve from time zero to 8 hours post-dose at steady state (AUC0-8,ss) of ruxolitinib[36]

End point description:

Pharmacokinetic (PK) parameters were calculated based on ruxolitinib plasma concentrations and actual sampling time points.
AUC0-8,ss was listed and summarized using descriptive statistics.

End point type	Secondary

End point timeframe:

Cycle 1 Day 14 (30 minutes, 1 hour, 2 hours, 4 hours, 6 hours, and 8 hours post-dose). 1 cycle = 21 days.

Notes:

[36] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: Endpoint only applicable to Phase 2 (Arm 2 - Cohort 2A)

End point values	Ph2 Arm2 MF JAKi combo – Cohort 2A			
Subject group type	Reporting group			
Number of subjects analysed	26			
Units: h*ng/mL				
geometric mean (geometric coefficient of variation)				
Cohort 2A, 5 mg	239 (± 64.5)			
Cohort 2A, 7.5 mg	369 (± 35.9)			
Cohort 2A, 10 mg	375 (± 47.2)			
Cohort 2A, 15 mg	999 (± 999)			
Cohort 2A, 20 mg	887 (± 42.3)			

Statistical analyses

No statistical analyses for this end point

Secondary: Phase 2 (Arm 2 - Cohort 2B): Maximum observed plasma concentration (Cmax) of ruxolitinib

End point title	Phase 2 (Arm 2 - Cohort 2B): Maximum observed plasma concentration (Cmax) of ruxolitinib[37]

End point description:

Pharmacokinetic (PK) parameters were calculated based on ruxolitinib plasma concentrations and actual sampling time points.
Cmax was listed and summarized using descriptive statistics.

End point type	Secondary

End point timeframe:

Cycle 1 Day 14 (30 minutes, 1 hour, 2 hours, 4 hours, 6 hours, and 8 hours post-dose). 1 cycle = 21 days.

Notes:

[37] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: Endpoint only applicable to Phase 2 (Arm 2 - Cohort 2B)

End point values	Ph2 Arm2 MF JAKi combo – Cohort 2B			
Subject group type	Reporting group			
Number of subjects analysed	7			
Units: ng/mL				
geometric mean (geometric coefficient of variation)				
Cohort 2B, 5 mg	69.5 (± 27.8)			
Cohort 2B, 15 mg	999 (± 999)			
Cohort 2B, 20 mg	999 (± 999)			
Cohort 2B, 25 mg	999 (± 999)			

Statistical analyses

No statistical analyses for this end point

Secondary: Phase 2 (Arm 2 - Cohort 2B): Time to reach maximum concentration (Tmax) of ruxolitinib

End point title	Phase 2 (Arm 2 - Cohort 2B): Time to reach maximum concentration (Tmax) of ruxolitinib[38]

End point description:

Pharmacokinetic (PK) parameters were calculated based on ruxolitinib plasma concentrations and actual sampling time points.
Tmax was listed and summarized using descriptive statistics.

End point type	Secondary

End point timeframe:

Cycle 1 Day 14 (30 minutes, 1 hour, 2 hours, 4 hours, 6 hours, and 8 hours post-dose). 1 cycle = 21 days.

Notes:

[38] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: Endpoint only applicable to Phase 2 (Arm 2 - Cohort 2B)

End point values	Ph2 Arm2 MF JAKi combo – Cohort 2B			
Subject group type	Reporting group			
Number of subjects analysed	7			
Units: Hour				
median (full range (min-max))				
Cohort 2B, 5 mg	1.57 (0.47 to 1.60)			
Cohort 2B, 15 mg	0.48 (0 to 999)			
Cohort 2B, 20 mg	0.53 (0 to 999)			
Cohort 2B, 25 mg	0.40 (0 to 999)			

Statistical analyses

No statistical analyses for this end point

Secondary: Phase 2 (Arm 2 - Cohort 2B): Predose (trough) concentration at the end of a dosing interval (Ctrough) of ruxolitinib

End point title	Phase 2 (Arm 2 - Cohort 2B): Predose (trough) concentration at the end of a dosing interval (Ctrough) of ruxolitinib[39]

End point description:

Pharmacokinetic (PK) parameters were calculated based on ruxolitinib plasma concentrations and actual sampling time points.
Ctrough was listed and summarized using descriptive statistics.

End point type	Secondary

End point timeframe:

Cycle 1 Day 14 (30 minutes, 1 hour, 2 hours, 4 hours, 6 hours, and 8 hours post-dose). 1 cycle = 21 days.

Notes:

[39] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: Endpoint only applicable to Phase 2 (Cohort 2B)

End point values	Ph2 Arm2 MF JAKi combo – Cohort 2B			
Subject group type	Reporting group			
Number of subjects analysed	7			
Units: ng/mL				
geometric mean (geometric coefficient of variation)				
Cohort 2B, 5 mg	8.49 (± 28.4)			
Cohort 2B, 15 mg	999 (± 999)			
Cohort 2B, 20 mg	999 (± 999)			
Cohort 2B, 25 mg	999 (± 999)			

Statistical analyses

No statistical analyses for this end point

Secondary: Phase 2 (Arm 2 - Cohort 2B): Area under the concentration–time curve from time zero to the last observed concentration (AUClast) of ruxolitinib

End point title	Phase 2 (Arm 2 - Cohort 2B): Area under the concentration–time curve from time zero to the last observed concentration (AUClast) of ruxolitinib[40]

End point description:

Pharmacokinetic (PK) parameters were calculated based on ruxolitinib plasma concentrations and actual sampling time points.

AUClast was listed and summarized using descriptive statistics.

End point type	Secondary

End point timeframe:

Cycle 1 Day 14 (30 minutes, 1 hour, 2 hours, 4 hours, 6 hours, and 8 hours post-dose). 1 cycle = 21 days.

Notes:

[40] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: Endpoint only applicable to Phase 2 (Arm 2 - Cohort 2B)

End point values	Ph2 Arm2 MF JAKi combo – Cohort 2B			
Subject group type	Reporting group			
Number of subjects analysed	7			
Units: h*ng/mL				
geometric mean (geometric coefficient of variation)				
Cohort 2B, 5 mg	207 (± 3.08)			
Cohort 2B, 15 mg	999 (± 999)			
Cohort 2B, 20 mg	999 (± 999)			
Cohort 2B, 25 mg	999 (± 999)			

Statistical analyses

No statistical analyses for this end point

Secondary: Phase 2 (Arm 2 - Cohort 2B): Area under the concentration–time curve from time zero to 8 hours post-dose at steady state (AUC0-8,ss) of ruxolitinib

End point title	Phase 2 (Arm 2 - Cohort 2B): Area under the concentration–time curve from time zero to 8 hours post-dose at steady state (AUC0-8,ss) of ruxolitinib[41]

End point description:

Pharmacokinetic (PK) parameters were calculated based on ruxolitinib plasma concentrations and actual sampling time points.
AUC0-8,ss was listed and summarized using descriptive statistics.

End point type	Secondary

End point timeframe:

Cycle 1 Day 14 (30 minutes, 1 hour, 2 hours, 4 hours, 6 hours, and 8 hours post-dose). 1 cycle = 21 days.

Notes:

[41] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: Endpoint only applicable to Phase 2 (Arm 2 - Cohort 2B)

End point values	Ph2 Arm2 MF JAKi combo – Cohort 2B			
Subject group type	Reporting group			
Number of subjects analysed	7			
Units: ng/mL				
geometric mean (geometric coefficient				

of variation)				
Cohort 2B, 5 mg	214 (± 3.67)			
Cohort 2B, 15 mg	999 (± 999)			
Cohort 2B, 20 mg	999 (± 999)			
Cohort 2B, 25 mg	999 (± 999)			

Statistical analyses

No statistical analyses for this end point

Secondary: Phase 2 (Arm 3): Maximum observed plasma concentration (Cmax) of ruxolitinib

End point title	Phase 2 (Arm 3): Maximum observed plasma concentration (Cmax) of ruxolitinib[42]

End point description:

Pharmacokinetic (PK) parameters were calculated based on ruxolitinib plasma concentrations and actual sampling time points.
Cmax was listed and summarized using descriptive statistics.

End point type	Secondary

End point timeframe:

Cycle 1 Day 14 (30 minutes, 1 hour, 2 hours, 4 hours, 6 hours, and 8 hours post-dose). 1 cycle = 21 days.

Notes:

[42] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: Endpoint only applicable to Phase 2 (Arm 3)

End point values	Ph2 Arm3 MF JAKi-naïve – Combo Tx			
Subject group type	Reporting group			
Number of subjects analysed	44			
Units: ng/mL				
geometric mean (geometric coefficient of variation)				
Cohort 3, 5 mg	999 (± 999)			
Cohort 3, 10 mg	190 (± 28.6)			
Cohort 3, 15 mg	213 (± 34.7)			
Cohort 3, 20 mg	999 (± 999)			

Statistical analyses

No statistical analyses for this end point

Secondary: Phase 2 (Arm 3): Time to reach maximum concentration (Tmax) of ruxolitinib

End point title	Phase 2 (Arm 3): Time to reach maximum concentration (Tmax) of ruxolitinib[43]

End point description:

Pharmacokinetic (PK) parameters were calculated based on ruxolitinib plasma concentrations and actual sampling time points.
Tmax was listed and summarized using descriptive statistics.

End point type	Secondary

End point timeframe:

Cycle 1 Day 14 (30 minutes, 1 hour, 2 hours, 4 hours, 6 hours, and 8 hours post-dose). 1 cycle = 21 days.

Notes:

[43] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: Endpoint only applicable to Phase 2 (Arm 3)

End point values	Ph2 Arm3 MF JAKi-naïve – Combo Tx			
Subject group type	Reporting group			
Number of subjects analysed	44			
Units: Hour				
median (full range (min-max))				
Cohort 3, 5 mg	0.72 (0 to 999)			
Cohort 3, 10 mg	0.62 (0.28 to 7.00)			
Cohort 3, 15 mg	0.95 (0.22 to 3.87)			
Cohort 3, 20 mg	1.25 (0 to 999)			

Statistical analyses

No statistical analyses for this end point

Secondary: Phase 2 (Arm 3): Predose (trough) concentration at the end of a dosing interval (Ctrough) of ruxolitinib

End point title	Phase 2 (Arm 3): Predose (trough) concentration at the end of a dosing interval (Ctrough) of ruxolitinib[44]

End point description:

Pharmacokinetic (PK) parameters were calculated based on ruxolitinib plasma concentrations and actual sampling time points.
Ctrough was listed and summarized using descriptive statistics.

End point type	Secondary

End point timeframe:

Cycle 1 Day 14 (predose, 30 minutes, 1 hour, 2 hours, 4 hours, 6 hours, and 8 hours post-dose). 1 cycle = 21 days.

Notes:

[44] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: Endpoint only applicable to Phase 2 (Arm 3)

End point values	Ph2 Arm3 MF JAKi-naïve – Combo Tx			
Subject group type	Reporting group			
Number of subjects analysed	43			
Units: ng/mL				
geometric mean (geometric coefficient of variation)				
Cohort 3, 5 mg	999 (± 999)			
Cohort 3, 10 mg	25.3 (± 99.2)			
Cohort 3, 15 mg	31.4 (± 58.1)			
Cohort 3, 20 mg	999 (± 999)			

Statistical analyses

No statistical analyses for this end point

Secondary: Phase 2 (Arm 3): Area under the concentration–time curve from time zero to the last observed concentration (AUClast) of ruxolitinib

End point title	Phase 2 (Arm 3): Area under the concentration–time curve from time zero to the last observed concentration (AUClast) of ruxolitinib[45]

End point description:

Pharmacokinetic (PK) parameters were calculated based on ruxolitinib plasma concentrations and actual sampling time points.
AUClast was listed and summarized using descriptive statistics.

End point type	Secondary

End point timeframe:

Cycle 1 Day 14 (30 minutes, 1 hour, 2 hours, 4 hours, 6 hours, and 8 hours post-dose). 1 cycle = 21 days.

Notes:

[45] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: Endpoint only applicable to Phase 2 (Arm 3)

End point values	Ph2 Arm3 MF JAKi-naïve – Combo Tx			
Subject group type	Reporting group			
Number of subjects analysed	44			
Units: h*ng/mL				
geometric mean (geometric coefficient of variation)				
Cohort 3, 5 mg	999 (± 999)			
Cohort 3, 10 mg	519 (± 54.8)			
Cohort 3, 15 mg	677 (± 33.9)			
Cohort 3, 20 mg	999 (± 999)			

Statistical analyses

No statistical analyses for this end point

Secondary: Phase 2 (Arm 3): Area under the concentration–time curve from time zero to 8 hours post-dose at steady state (AUC0-8,ss) of ruxolitinib

End point title	Phase 2 (Arm 3): Area under the concentration–time curve from time zero to 8 hours post-dose at steady state (AUC0-8,ss) of ruxolitinib[46]

End point description:

Pharmacokinetic (PK) parameters were calculated based on ruxolitinib plasma concentrations and actual sampling time points.
AUC0-8,ss was listed and summarized using descriptive statistics.

End point type	Secondary

End point timeframe:

Cycle 1 Day 14 (30 minutes, 1 hour, 2 hours, 4 hours, 6 hours, and 8 hours post-dose). 1 cycle = 21 days.

Notes:

[46] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: Endpoint only applicable to Phase 2 (Arm 3)

End point values	Ph2 Arm3 MF JAKi-naïve – Combo Tx			
Subject group type	Reporting group			
Number of subjects analysed	39			
Units: h*ng/mL				
geometric mean (geometric coefficient of variation)				
Cohort 3, 5 mg	999 (± 999)			
Cohort 3, 10 mg	587 (± 45.5)			
Cohort 3, 15 mg	696 (± 34.7)			
Cohort 3, 20 mg	999 (± 999)			

Statistical analyses

No statistical analyses for this end point

Adverse events

Adverse events information

Timeframe for reporting adverse events:

Adverse events were collected from first study dose through 30 days after the last pelabresib (CPI0610) dose. Duration varied by participant and phase, with follow up up to about two years in Phase I and about eight years in Phase II.

Adverse event reporting additional description:

Consistent with EudraCT disclosure specifications, Novartis has reported under the Serious adverse events field "number of deaths resulting from adverse events" all those deaths, resulting from serious adverse events that are deemed to be causally related to treatment by the investigator.

Assessment type	Systematic

Dictionary used

Dictionary name	MedDRA
Dictionary version	26.0

Reporting groups

Reporting group title	Phase 1 225 mg tablet PO daily
Reporting group description:	
Phase 1 225 mg tablet PO daily	
Reporting group title	Phase 1 400 mg capsule PO daily
Reporting group description:	
Phase 1 400 mg capsule PO daily	
Reporting group title	Phase 1 300 mg capsule PO daily
Reporting group description:	
Phase 1 300 mg capsule PO daily	
Reporting group title	Phase 1 230 mg capsule PO daily
Reporting group description:	
Phase 1 230 mg capsule PO daily	
Reporting group title	Phase 1 170 mg capsule PO daily
Reporting group description:	
Phase 1 170 mg capsule PO daily	
Reporting group title	Phase 1 120 mg capsule PO daily
Reporting group description:	
Phase 1 120 mg capsule PO daily	
Reporting group title	Phase 1 48 mg capsule PO daily
Reporting group description:	
Phase 1 48 mg capsule PO daily	
Reporting group title	Phase 1 24 mg capsule PO daily
Reporting group description:	
Phase 1 24 mg capsule PO daily	
Reporting group title	Phase 2 Arm 4
Reporting group description:	
Phase 2 Arm 4	
Reporting group title	Phase 2 Arm 3
Reporting group description:	
Phase 2 Arm 3	
Reporting group title	Phase 2 Arm 2 2B
Reporting group description:	
Phase 2 Arm 2 2B	
Reporting group title	Phase 2 Arm 2 2A

Reporting group description:	
Phase 2 Arm 2 2A	
Reporting group title	Phase 2 Arm 1 1B
Reporting group description:	
Phase 2 Arm 1 1B	
Reporting group title	Phase 2 Arm 1 1A
Reporting group description:	
Phase 2 Arm 1 1A	
Reporting group title	Phase 1 overall
Reporting group description:	
Phase 1 overall	
Reporting group title	Phase 2 Overall
Reporting group description:	
Phase 2 Overall	
Reporting group title	Phase 1 275 mg tablet PO daily
Reporting group description:	
Phase 1 275 mg tablet PO daily	

Serious adverse events	Phase 1 225 mg tablet PO daily	Phase 1 400 mg capsule PO daily	Phase 1 300 mg capsule PO daily
Total subjects affected by serious adverse events			
subjects affected / exposed	6 / 8 (75.00%)	6 / 7 (85.71%)	4 / 4 (100.00%)
number of deaths (all causes)	1	1	1
number of deaths resulting from adverse events	0	0	0
Neoplasms benign, malignant and unspecified (incl cysts and polyps)			
Basal cell carcinoma			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Acute myeloid leukaemia			
subjects affected / exposed	1 / 8 (12.50%)	1 / 7 (14.29%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 1	0 / 0	0 / 0
Breast cancer			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Bowen's disease			

subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Colon cancer metastatic			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Glioblastoma			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Invasive breast carcinoma			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Haematopoietic neoplasm			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Keratoacanthoma			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Lung neoplasm malignant			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Malignant melanoma			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Metastatic squamous cell carcinoma			

subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Neuroendocrine carcinoma of the skin			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Oropharyngeal squamous cell carcinoma			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Ovarian neoplasm			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Squamous cell carcinoma of head and neck			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Squamous cell carcinoma of skin			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Transformation to acute myeloid leukaemia			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Vascular disorders			
Aortic dissection			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0

Haematoma			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hypertension			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hypotension			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Peripheral ischaemia			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
General disorders and administration site conditions			
Asthenia			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Chills			
subjects affected / exposed	1 / 8 (12.50%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Chest pain			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Drug withdrawal syndrome			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0

Disease progression			
subjects affected / exposed	0 / 8 (0.00%)	1 / 7 (14.29%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 1	0 / 0
Fatigue			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Malaise			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Multiple organ dysfunction syndrome			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Non-cardiac chest pain			
subjects affected / exposed	1 / 8 (12.50%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Oedema			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pyrexia			
subjects affected / exposed	0 / 8 (0.00%)	1 / 7 (14.29%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Immune system disorders			
Serum sickness			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Respiratory, thoracic and mediastinal			

disorders			
Acute respiratory failure			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Bronchospasm			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Dyspnoea			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Epistaxis			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Organising pneumonia			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pleuritic pain			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pneumonia aspiration			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pneumonitis			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pneumothorax			

subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pulmonary embolism			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pulmonary haemorrhage			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pulmonary oedema			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pulmonary hypertension			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Respiratory failure			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Psychiatric disorders			
Confusional state			
subjects affected / exposed	0 / 8 (0.00%)	2 / 7 (28.57%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 2	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Delirium			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Depression			

subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Suicide attempt			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Investigations			
Blast cell count increased			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Blood bilirubin increased			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Ejection fraction decreased			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Blood creatinine increased			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Platelet count decreased			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Injury, poisoning and procedural complications			
Extradural haematoma			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0

Fall			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pelvic fracture			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hip fracture			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Post procedural haemorrhage			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Subdural haematoma			
subjects affected / exposed	1 / 8 (12.50%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cardiac disorders			
Acute left ventricular failure			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Acute myocardial infarction			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Atrial fibrillation			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Angina pectoris			

subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cardiac arrest			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cardiac failure			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cardiac failure congestive			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Chronic coronary syndrome			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cardiac iron overload			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Dilated cardiomyopathy			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Left ventricular dysfunction			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Left ventricular failure			

subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pericarditis			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Sinus tachycardia			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Right ventricular failure			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Nervous system disorders			
Brain stem haemorrhage			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cerebral infarction			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cerebrovascular accident			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Haemorrhage intracranial			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Metabolic encephalopathy			

subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Spinal cord compression			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Paraesthesia			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Subarachnoid haemorrhage			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Superior sagittal sinus thrombosis			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Syncope			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Blood and lymphatic system disorders			
Anaemia			
subjects affected / exposed	0 / 8 (0.00%)	1 / 7 (14.29%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Febrile neutropenia			
subjects affected / exposed	1 / 8 (12.50%)	2 / 7 (28.57%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 2	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Leukocytosis			

subjects affected / exposed	1 / 8 (12.50%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Leukostasis syndrome			
subjects affected / exposed	1 / 8 (12.50%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Splenic infarction			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Splenomegaly			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Thrombocytopenia			
subjects affected / exposed	0 / 8 (0.00%)	1 / 7 (14.29%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	1 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Ear and labyrinth disorders			
Vertigo			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Gastrointestinal disorders			
Abdominal pain			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Colitis			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Abdominal wall haematoma			

subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Constipation			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Diarrhoea			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Duodenal ulcer haemorrhage			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Gastric ulcer			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Gastrointestinal disorder			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Gastrointestinal haemorrhage			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Haemorrhoids			
subjects affected / exposed	1 / 8 (12.50%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Intestinal haematoma			

subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Intestinal obstruction			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Lower gastrointestinal haemorrhage			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Rectal haemorrhage			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Nausea			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Small intestinal obstruction			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	1 / 4 (25.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Vomiting			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hepatobiliary disorders			
Biliary dilatation			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hepatocellular injury			

subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hypertransaminasaemia			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Skin and subcutaneous tissue disorders			
Cutaneous vasculitis			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Drug eruption			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Renal and urinary disorders			
Acute kidney injury			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Haematuria			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Renal cyst ruptured			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Renal haemorrhage			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Endocrine disorders			

Adrenal haematoma			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Inappropriate antidiuretic hormone secretion			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Musculoskeletal and connective tissue disorders			
Back pain			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Bone pain			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Joint effusion			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Muscular weakness			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Musculoskeletal pain			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Neck pain			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0

Infections and infestations			
Abscess limb			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Aspergillus infection			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Bacteraemia			
subjects affected / exposed	0 / 8 (0.00%)	1 / 7 (14.29%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Bronchiolitis			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
COVID-19			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Bronchopulmonary aspergillosis			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
COVID-19 pneumonia			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Clostridium difficile infection			
subjects affected / exposed	1 / 8 (12.50%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cellulitis			

subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Dermo-hypodermitis			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Device related sepsis			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	1 / 4 (25.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Device related infection			
subjects affected / exposed	1 / 8 (12.50%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Diverticulitis			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Empyema			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Endocarditis bacterial			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Enterococcal bacteraemia			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	1 / 4 (25.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 1
Escherichia bacteraemia			

subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Escherichia infection			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Escherichia urinary tract infection			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Fournier's gangrene			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Herpes zoster			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Herpes simplex reactivation			
subjects affected / exposed	1 / 8 (12.50%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Gastroenteritis viral			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Influenza			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Localised infection			

subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Nocardiosis			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pneumonia			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	1 / 4 (25.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pneumonia acinetobacter			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pneumonia bacterial			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pneumonia pneumococcal			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pneumonia staphylococcal			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Post procedural infection			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pyelonephritis			

subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pyelonephritis acute			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Respiratory tract infection			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Septic shock			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Sepsis			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Streptococcal sepsis			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Tooth infection			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Urinary tract infection			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	1 / 4 (25.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Urosepsis			

subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Viral infection			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Metabolism and nutrition disorders			
Decreased appetite			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Dehydration			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Failure to thrive			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Gout			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hypercalcaemia			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hyperglycaemia			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hyperkalaemia			

subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hyponatraemia			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0

Serious adverse events	Phase 1 230 mg capsule PO daily	Phase 1 170 mg capsule PO daily	Phase 1 120 mg capsule PO daily
Total subjects affected by serious adverse events			
subjects affected / exposed	2 / 3 (66.67%)	1 / 3 (33.33%)	3 / 5 (60.00%)
number of deaths (all causes)	1	0	2
number of deaths resulting from adverse events	0	0	0
Neoplasms benign, malignant and unspecified (incl cysts and polyps)			
Basal cell carcinoma			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Acute myeloid leukaemia			
subjects affected / exposed	1 / 3 (33.33%)	0 / 3 (0.00%)	1 / 5 (20.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 1	0 / 0	0 / 1
Breast cancer			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Bowen's disease			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Colon cancer metastatic			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0

Glioblastoma			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Invasive breast carcinoma			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Haematopoietic neoplasm			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Keratoacanthoma			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Lung neoplasm malignant			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Malignant melanoma			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Metastatic squamous cell carcinoma			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Neuroendocrine carcinoma of the skin			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Oropharyngeal squamous cell carcinoma			

subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Ovarian neoplasm			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Squamous cell carcinoma of head and neck			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Squamous cell carcinoma of skin			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Transformation to acute myeloid leukaemia			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Vascular disorders			
Aortic dissection			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Haematoma			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hypertension			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hypotension			

subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Peripheral ischaemia			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	1 / 5 (20.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
General disorders and administration site conditions			
Asthenia			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Chills			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Chest pain			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Drug withdrawal syndrome			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Disease progression			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Fatigue			
subjects affected / exposed	1 / 3 (33.33%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 2	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Malaise			

subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Multiple organ dysfunction syndrome			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Non-cardiac chest pain			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Oedema			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pyrexia			
subjects affected / exposed	0 / 3 (0.00%)	1 / 3 (33.33%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Immune system disorders			
Serum sickness			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Respiratory, thoracic and mediastinal disorders			
Acute respiratory failure			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Bronchospasm			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0

Dyspnoea			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Epistaxis			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Organising pneumonia			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pleuritic pain			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pneumonia aspiration			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pneumonitis			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pneumothorax			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pulmonary embolism			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pulmonary haemorrhage			

subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	1 / 5 (20.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pulmonary oedema			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pulmonary hypertension			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Respiratory failure			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	1 / 5 (20.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 1
Psychiatric disorders			
Confusional state			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Delirium			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Depression			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Suicide attempt			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Investigations			

Blast cell count increased			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Blood bilirubin increased			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Ejection fraction decreased			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Blood creatinine increased			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Platelet count decreased			
subjects affected / exposed	0 / 3 (0.00%)	1 / 3 (33.33%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Injury, poisoning and procedural complications			
Extradural haematoma			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Fall			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pelvic fracture			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0

Hip fracture			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	1 / 5 (20.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Post procedural haemorrhage			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Subdural haematoma			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cardiac disorders			
Acute left ventricular failure			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Acute myocardial infarction			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Atrial fibrillation			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Angina pectoris			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cardiac arrest			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cardiac failure			

subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cardiac failure congestive			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Chronic coronary syndrome			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cardiac iron overload			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Dilated cardiomyopathy			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Left ventricular dysfunction			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Left ventricular failure			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pericarditis			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Sinus tachycardia			

subjects affected / exposed	0 / 3 (0.00%)	1 / 3 (33.33%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Right ventricular failure			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Nervous system disorders			
Brain stem haemorrhage			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cerebral infarction			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cerebrovascular accident			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Haemorrhage intracranial			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Metabolic encephalopathy			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Spinal cord compression			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Paraesthesia			

subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Subarachnoid haemorrhage			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Superior sagittal sinus thrombosis			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Syncope			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Blood and lymphatic system disorders			
Anaemia			
subjects affected / exposed	0 / 3 (0.00%)	1 / 3 (33.33%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Febrile neutropenia			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	1 / 5 (20.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Leukocytosis			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Leukostasis syndrome			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Splenic infarction			

subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Splenomegaly			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Thrombocytopenia			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Ear and labyrinth disorders			
Vertigo			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Gastrointestinal disorders			
Abdominal pain			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	1 / 5 (20.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Colitis			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Abdominal wall haematoma			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Constipation			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	1 / 5 (20.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Diarrhoea			

subjects affected / exposed	0 / 3 (0.00%)	1 / 3 (33.33%)	1 / 5 (20.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	1 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Duodenal ulcer haemorrhage			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Gastric ulcer			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Gastrointestinal disorder			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Gastrointestinal haemorrhage			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Haemorrhoids			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Intestinal haematoma			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Intestinal obstruction			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Lower gastrointestinal haemorrhage			

subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Rectal haemorrhage			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Nausea			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Small intestinal obstruction			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Vomiting			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hepatobiliary disorders			
Biliary dilatation			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hepatocellular injury			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hypertransaminasaemia			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Skin and subcutaneous tissue disorders			

Cutaneous vasculitis			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Drug eruption			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Renal and urinary disorders			
Acute kidney injury			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Haematuria			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Renal cyst ruptured			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Renal haemorrhage			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Endocrine disorders			
Adrenal haematoma			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Inappropriate antidiuretic hormone secretion			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0

Musculoskeletal and connective tissue disorders			
Back pain			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Bone pain			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Joint effusion			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Muscular weakness			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Musculoskeletal pain			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Neck pain			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Infections and infestations			
Abscess limb			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Aspergillus infection			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0

Bacteraemia			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Bronchiolitis			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
COVID-19			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Bronchopulmonary aspergillosis			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
COVID-19 pneumonia			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Clostridium difficile infection			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cellulitis			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Dermo-hypodermitis			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Device related sepsis			

subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Device related infection			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Diverticulitis			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Empyema			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Endocarditis bacterial			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Enterococcal bacteraemia			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Escherichia bacteraemia			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Escherichia infection			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Escherichia urinary tract infection			

subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Fournier's gangrene			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Herpes zoster			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Herpes simplex reactivation			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Gastroenteritis viral			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Influenza			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Localised infection			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Nocardiosis			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pneumonia			

subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	1 / 5 (20.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pneumonia acinetobacter			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pneumonia bacterial			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pneumonia pneumococcal			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pneumonia staphylococcal			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Post procedural infection			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pyelonephritis			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pyelonephritis acute			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Respiratory tract infection			

subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Septic shock			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Sepsis			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	1 / 5 (20.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Streptococcal sepsis			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Tooth infection			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Urinary tract infection			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Urosepsis			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Viral infection			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Metabolism and nutrition disorders			
Decreased appetite			

subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Dehydration			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Failure to thrive			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Gout			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hypercalcaemia			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hyperglycaemia			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hyperkalaemia			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hyponatraemia			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0

Serious adverse events	Phase 1 48 mg capsule PO daily	Phase 1 24 mg capsule PO daily	Phase 2 Arm 4

Total subjects affected by serious adverse events			
subjects affected / exposed	4 / 5 (80.00%)	1 / 3 (33.33%)	5 / 21 (23.81%)
number of deaths (all causes)	1	0	0
number of deaths resulting from adverse events	0	0	0
Neoplasms benign, malignant and unspecified (incl cysts and polyps)			
Basal cell carcinoma			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Acute myeloid leukaemia			
subjects affected / exposed	1 / 5 (20.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 1	0 / 0	0 / 0
Breast cancer			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	1 / 21 (4.76%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Bowen's disease			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Colon cancer metastatic			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Glioblastoma			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Invasive breast carcinoma			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Haematopoietic neoplasm			

subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Keratoacanthoma			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Lung neoplasm malignant			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Malignant melanoma			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Metastatic squamous cell carcinoma			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Neuroendocrine carcinoma of the skin			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Oropharyngeal squamous cell carcinoma			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Ovarian neoplasm			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Squamous cell carcinoma of head and neck			

subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Squamous cell carcinoma of skin			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Transformation to acute myeloid leukaemia			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Vascular disorders			
Aortic dissection			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Haematoma			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hypertension			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hypotension			
subjects affected / exposed	1 / 5 (20.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 1	0 / 0	0 / 0
Peripheral ischaemia			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
General disorders and administration site conditions			

Asthenia			
subjects affected / exposed	1 / 5 (20.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Chills			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Chest pain			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Drug withdrawal syndrome			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Disease progression			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Fatigue			
subjects affected / exposed	2 / 5 (40.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 2	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Malaise			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Multiple organ dysfunction syndrome			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Non-cardiac chest pain			

subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Oedema			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pyrexia			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Immune system disorders			
Serum sickness			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Respiratory, thoracic and mediastinal disorders			
Acute respiratory failure			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Bronchospasm			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Dyspnoea			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	1 / 21 (4.76%)
occurrences causally related to treatment / all	0 / 0	0 / 0	1 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Epistaxis			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0

Organising pneumonia			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pleuritic pain			
subjects affected / exposed	0 / 5 (0.00%)	1 / 3 (33.33%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pneumonia aspiration			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pneumonitis			
subjects affected / exposed	0 / 5 (0.00%)	1 / 3 (33.33%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pneumothorax			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pulmonary embolism			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	1 / 21 (4.76%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pulmonary haemorrhage			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pulmonary oedema			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pulmonary hypertension			

subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Respiratory failure			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Psychiatric disorders			
Confusional state			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Delirium			
subjects affected / exposed	1 / 5 (20.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Depression			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Suicide attempt			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Investigations			
Blast cell count increased			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Blood bilirubin increased			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Ejection fraction decreased			

subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Blood creatinine increased			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Platelet count decreased			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Injury, poisoning and procedural complications			
Extradural haematoma			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Fall			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pelvic fracture			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hip fracture			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Post procedural haemorrhage			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Subdural haematoma			

subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cardiac disorders			
Acute left ventricular failure			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Acute myocardial infarction			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Atrial fibrillation			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Angina pectoris			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cardiac arrest			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cardiac failure			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cardiac failure congestive			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Chronic coronary syndrome			

subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cardiac iron overload			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Dilated cardiomyopathy			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Left ventricular dysfunction			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Left ventricular failure			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pericarditis			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Sinus tachycardia			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Right ventricular failure			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Nervous system disorders			
Brain stem haemorrhage			

subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cerebral infarction			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cerebrovascular accident			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Haemorrhage intracranial			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Metabolic encephalopathy			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Spinal cord compression			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Paraesthesia			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Subarachnoid haemorrhage			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Superior sagittal sinus thrombosis			

subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Syncope			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Blood and lymphatic system disorders			
Anaemia			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Febrile neutropenia			
subjects affected / exposed	2 / 5 (40.00%)	1 / 3 (33.33%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 2	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 1	0 / 0	0 / 0
Leukocytosis			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Leukostasis syndrome			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Splenic infarction			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Splenomegaly			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Thrombocytopenia			

subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Ear and labyrinth disorders			
Vertigo			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Gastrointestinal disorders			
Abdominal pain			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Colitis			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Abdominal wall haematoma			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Constipation			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Diarrhoea			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Duodenal ulcer haemorrhage			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Gastric ulcer			

subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Gastrointestinal disorder			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Gastrointestinal haemorrhage			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	1 / 21 (4.76%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Haemorrhoids			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Intestinal haematoma			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Intestinal obstruction			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Lower gastrointestinal haemorrhage			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Rectal haemorrhage			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Nausea			

subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Small intestinal obstruction			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Vomiting			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hepatobiliary disorders			
Biliary dilatation			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hepatocellular injury			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hypertransaminasaemia			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Skin and subcutaneous tissue disorders			
Cutaneous vasculitis			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Drug eruption			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Renal and urinary disorders			

Acute kidney injury			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	1 / 21 (4.76%)
occurrences causally related to treatment / all	0 / 0	0 / 0	1 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Haematuria			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Renal cyst ruptured			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Renal haemorrhage			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Endocrine disorders			
Adrenal haematoma			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Inappropriate antidiuretic hormone secretion			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Musculoskeletal and connective tissue disorders			
Back pain			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Bone pain			

subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Joint effusion			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Muscular weakness			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Musculoskeletal pain			
subjects affected / exposed	1 / 5 (20.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Neck pain			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Infections and infestations			
Abscess limb			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Aspergillus infection			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Bacteraemia			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Bronchiolitis			

subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
COVID-19			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Bronchopulmonary aspergillosis			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
COVID-19 pneumonia			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Clostridium difficile infection			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cellulitis			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Dermo-hypodermitis			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Device related sepsis			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Device related infection			

subjects affected / exposed	0 / 5 (0.00%)	1 / 3 (33.33%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Diverticulitis			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Empyema			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Endocarditis bacterial			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Enterococcal bacteraemia			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Escherichia bacteraemia			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Escherichia infection			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Escherichia urinary tract infection			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Fournier's gangrene			

subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Herpes zoster			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Herpes simplex reactivation			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Gastroenteritis viral			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Influenza			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Localised infection			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Nocardiosis			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pneumonia			
subjects affected / exposed	1 / 5 (20.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pneumonia acinetobacter			

subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pneumonia bacterial			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pneumonia pneumococcal			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pneumonia staphylococcal			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Post procedural infection			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pyelonephritis			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pyelonephritis acute			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Respiratory tract infection			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Septic shock			

subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Sepsis			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Streptococcal sepsis			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Tooth infection			
subjects affected / exposed	0 / 5 (0.00%)	1 / 3 (33.33%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Urinary tract infection			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Urosepsis			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Viral infection			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	1 / 21 (4.76%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Metabolism and nutrition disorders			
Decreased appetite			
subjects affected / exposed	1 / 5 (20.00%)	0 / 3 (0.00%)	1 / 21 (4.76%)
occurrences causally related to treatment / all	0 / 1	0 / 0	1 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Dehydration			

subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Failure to thrive			
subjects affected / exposed	1 / 5 (20.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Gout			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hypercalcaemia			
subjects affected / exposed	1 / 5 (20.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hyperglycaemia			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hyperkalaemia			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hyponatraemia			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0

Serious adverse events	Phase 2 Arm 3	Phase 2 Arm 2 2B	Phase 2 Arm 2 2A
Total subjects affected by serious adverse events			
subjects affected / exposed	40 / 84 (47.62%)	18 / 28 (64.29%)	29 / 59 (49.15%)
number of deaths (all causes)	7	3	3
number of deaths resulting from adverse events	2	1	0

Neoplasms benign, malignant and unspecified (incl cysts and polyps)			
Basal cell carcinoma			
subjects affected / exposed	1 / 84 (1.19%)	1 / 28 (3.57%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 3	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Acute myeloid leukaemia			
subjects affected / exposed	0 / 84 (0.00%)	1 / 28 (3.57%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 1	0 / 0
Breast cancer			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Bowen's disease			
subjects affected / exposed	0 / 84 (0.00%)	1 / 28 (3.57%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Colon cancer metastatic			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences causally related to treatment / all	0 / 0	0 / 0	1 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Glioblastoma			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Invasive breast carcinoma			
subjects affected / exposed	0 / 84 (0.00%)	1 / 28 (3.57%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Haematopoietic neoplasm			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0

Keratoacanthoma			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Lung neoplasm malignant			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Malignant melanoma			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	1 / 2	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Metastatic squamous cell carcinoma			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Neuroendocrine carcinoma of the skin			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Oropharyngeal squamous cell carcinoma			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Ovarian neoplasm			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Squamous cell carcinoma of head and neck			
subjects affected / exposed	1 / 84 (1.19%)	1 / 28 (3.57%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0

Squamous cell carcinoma of skin			
subjects affected / exposed	1 / 84 (1.19%)	1 / 28 (3.57%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	4 / 5	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Transformation to acute myeloid leukaemia			
subjects affected / exposed	6 / 84 (7.14%)	1 / 28 (3.57%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 6	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 1	0 / 0	0 / 0
Vascular disorders			
Aortic dissection			
subjects affected / exposed	0 / 84 (0.00%)	1 / 28 (3.57%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Haematoma			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hypertension			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hypotension			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Peripheral ischaemia			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	2 / 59 (3.39%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
General disorders and administration site conditions			
Asthenia			

subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Chills			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Chest pain			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Drug withdrawal syndrome			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 1	0 / 0	0 / 0
Disease progression			
subjects affected / exposed	0 / 84 (0.00%)	1 / 28 (3.57%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 1	0 / 0
Fatigue			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Malaise			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Multiple organ dysfunction syndrome			
subjects affected / exposed	2 / 84 (2.38%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	1 / 2	0 / 0	0 / 0
deaths causally related to treatment / all	1 / 2	0 / 0	0 / 0
Non-cardiac chest pain			

subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	2 / 59 (3.39%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Oedema			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pyrexia			
subjects affected / exposed	3 / 84 (3.57%)	1 / 28 (3.57%)	1 / 59 (1.69%)
occurrences causally related to treatment / all	1 / 3	0 / 2	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Immune system disorders			
Serum sickness			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Respiratory, thoracic and mediastinal disorders			
Acute respiratory failure			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Bronchospasm			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Dyspnoea			
subjects affected / exposed	1 / 84 (1.19%)	1 / 28 (3.57%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 1	1 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Epistaxis			
subjects affected / exposed	1 / 84 (1.19%)	1 / 28 (3.57%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	1 / 2	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0

Organising pneumonia			
subjects affected / exposed	0 / 84 (0.00%)	1 / 28 (3.57%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pleuritic pain			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pneumonia aspiration			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pneumonitis			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pneumothorax			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pulmonary embolism			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pulmonary haemorrhage			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pulmonary oedema			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pulmonary hypertension			

subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Respiratory failure			
subjects affected / exposed	1 / 84 (1.19%)	2 / 28 (7.14%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 2	0 / 0
deaths causally related to treatment / all	0 / 1	0 / 1	0 / 0
Psychiatric disorders			
Confusional state			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Delirium			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Depression			
subjects affected / exposed	0 / 84 (0.00%)	1 / 28 (3.57%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Suicide attempt			
subjects affected / exposed	0 / 84 (0.00%)	1 / 28 (3.57%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Investigations			
Blast cell count increased			
subjects affected / exposed	5 / 84 (5.95%)	3 / 28 (10.71%)	1 / 59 (1.69%)
occurrences causally related to treatment / all	2 / 5	0 / 3	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Blood bilirubin increased			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Ejection fraction decreased			

subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Blood creatinine increased			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Platelet count decreased			
subjects affected / exposed	0 / 84 (0.00%)	1 / 28 (3.57%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Injury, poisoning and procedural complications			
Extradural haematoma			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Fall			
subjects affected / exposed	2 / 84 (2.38%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 2	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pelvic fracture			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hip fracture			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Post procedural haemorrhage			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Subdural haematoma			

subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cardiac disorders			
Acute left ventricular failure			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Acute myocardial infarction			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Atrial fibrillation			
subjects affected / exposed	2 / 84 (2.38%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 2	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Angina pectoris			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cardiac arrest			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cardiac failure			
subjects affected / exposed	2 / 84 (2.38%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	1 / 2	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cardiac failure congestive			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 1
Chronic coronary syndrome			

subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cardiac iron overload			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Dilated cardiomyopathy			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Left ventricular dysfunction			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	1 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Left ventricular failure			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pericarditis			
subjects affected / exposed	0 / 84 (0.00%)	1 / 28 (3.57%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Sinus tachycardia			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Right ventricular failure			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Nervous system disorders			
Brain stem haemorrhage			

subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 1
Cerebral infarction			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 1
Cerebrovascular accident			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Haemorrhage intracranial			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 1
Metabolic encephalopathy			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Spinal cord compression			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Paraesthesia			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Subarachnoid haemorrhage			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Superior sagittal sinus thrombosis			

subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Syncope			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Blood and lymphatic system disorders			
Anaemia			
subjects affected / exposed	1 / 84 (1.19%)	5 / 28 (17.86%)	2 / 59 (3.39%)
occurrences causally related to treatment / all	0 / 1	2 / 5	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Febrile neutropenia			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Leukocytosis			
subjects affected / exposed	0 / 84 (0.00%)	1 / 28 (3.57%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Leukostasis syndrome			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Splenic infarction			
subjects affected / exposed	0 / 84 (0.00%)	1 / 28 (3.57%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Splenomegaly			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Thrombocytopenia			

subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Ear and labyrinth disorders			
Vertigo			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Gastrointestinal disorders			
Abdominal pain			
subjects affected / exposed	0 / 84 (0.00%)	1 / 28 (3.57%)	1 / 59 (1.69%)
occurrences causally related to treatment / all	0 / 0	0 / 1	1 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Colitis			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Abdominal wall haematoma			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	1 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Constipation			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Diarrhoea			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Duodenal ulcer haemorrhage			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	1 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Gastric ulcer			

subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Gastrointestinal disorder			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Gastrointestinal haemorrhage			
subjects affected / exposed	2 / 84 (2.38%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 2	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Haemorrhoids			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Intestinal haematoma			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Intestinal obstruction			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Lower gastrointestinal haemorrhage			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Rectal haemorrhage			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Nausea			

subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Small intestinal obstruction			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Vomiting			
subjects affected / exposed	0 / 84 (0.00%)	1 / 28 (3.57%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	1 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hepatobiliary disorders			
Biliary dilatation			
subjects affected / exposed	0 / 84 (0.00%)	1 / 28 (3.57%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hepatocellular injury			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hypertransaminasaemia			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	1 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Skin and subcutaneous tissue disorders			
Cutaneous vasculitis			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Drug eruption			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Renal and urinary disorders			

Acute kidney injury			
subjects affected / exposed	1 / 84 (1.19%)	1 / 28 (3.57%)	1 / 59 (1.69%)
occurrences causally related to treatment / all	0 / 1	1 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	1 / 1	0 / 0
Haematuria			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Renal cyst ruptured			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Renal haemorrhage			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Endocrine disorders			
Adrenal haematoma			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Inappropriate antidiuretic hormone secretion			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Musculoskeletal and connective tissue disorders			
Back pain			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Bone pain			

subjects affected / exposed	0 / 84 (0.00%)	1 / 28 (3.57%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Joint effusion			
subjects affected / exposed	0 / 84 (0.00%)	1 / 28 (3.57%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Muscular weakness			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Musculoskeletal pain			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Neck pain			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Infections and infestations			
Abscess limb			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Aspergillus infection			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Bacteraemia			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Bronchiolitis			

subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	1 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
COVID-19			
subjects affected / exposed	3 / 84 (3.57%)	1 / 28 (3.57%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	1 / 3	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 1	0 / 0	0 / 0
Bronchopulmonary aspergillosis			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
COVID-19 pneumonia			
subjects affected / exposed	4 / 84 (4.76%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences causally related to treatment / all	2 / 4	0 / 0	0 / 1
deaths causally related to treatment / all	1 / 1	0 / 0	0 / 0
Clostridium difficile infection			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cellulitis			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Dermo-hypodermitis			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Device related sepsis			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Device related infection			

subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Diverticulitis			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Empyema			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Endocarditis bacterial			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Enterococcal bacteraemia			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Escherichia bacteraemia			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Escherichia infection			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Escherichia urinary tract infection			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Fournier's gangrene			

subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Herpes zoster			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Herpes simplex reactivation			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Gastroenteritis viral			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Influenza			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Localised infection			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Nocardiosis			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pneumonia			
subjects affected / exposed	5 / 84 (5.95%)	2 / 28 (7.14%)	4 / 59 (6.78%)
occurrences causally related to treatment / all	2 / 5	1 / 2	0 / 4
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pneumonia acinetobacter			

subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pneumonia bacterial			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 1	0 / 0	0 / 0
Pneumonia pneumococcal			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pneumonia staphylococcal			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	1 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	1 / 1	0 / 0	0 / 0
Post procedural infection			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pyelonephritis			
subjects affected / exposed	0 / 84 (0.00%)	2 / 28 (7.14%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	1 / 2	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pyelonephritis acute			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Respiratory tract infection			
subjects affected / exposed	2 / 84 (2.38%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 2	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Septic shock			

subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Sepsis			
subjects affected / exposed	2 / 84 (2.38%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	1 / 2	0 / 0	0 / 0
deaths causally related to treatment / all	1 / 2	0 / 0	0 / 0
Streptococcal sepsis			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	1 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Tooth infection			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Urinary tract infection			
subjects affected / exposed	2 / 84 (2.38%)	1 / 28 (3.57%)	1 / 59 (1.69%)
occurrences causally related to treatment / all	0 / 2	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 1	0 / 0	0 / 0
Urosepsis			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Viral infection			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Metabolism and nutrition disorders			
Decreased appetite			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Dehydration			

subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Failure to thrive			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Gout			
subjects affected / exposed	0 / 84 (0.00%)	1 / 28 (3.57%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hypercalcaemia			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hyperglycaemia			
subjects affected / exposed	0 / 84 (0.00%)	1 / 28 (3.57%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	1 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hyperkalaemia			
subjects affected / exposed	0 / 84 (0.00%)	1 / 28 (3.57%)	0 / 59 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 2	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hyponatraemia			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences causally related to treatment / all	1 / 1	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0

Serious adverse events	Phase 2 Arm 1 1B	Phase 2 Arm 1 1A	Phase 1 overall
Total subjects affected by serious adverse events			
subjects affected / exposed	23 / 52 (44.23%)	18 / 48 (37.50%)	33 / 44 (75.00%)
number of deaths (all causes)	3	6	8
number of deaths resulting from adverse events	0	0	0

Neoplasms benign, malignant and unspecified (incl cysts and polyps)			
Basal cell carcinoma			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Acute myeloid leukaemia			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	5 / 44 (11.36%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 5
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 4
Breast cancer			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Bowen's disease			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Colon cancer metastatic			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Glioblastoma			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Invasive breast carcinoma			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Haematopoietic neoplasm			
subjects affected / exposed	1 / 52 (1.92%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0

Keratoacanthoma			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Lung neoplasm malignant			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Malignant melanoma			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Metastatic squamous cell carcinoma			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Neuroendocrine carcinoma of the skin			
subjects affected / exposed	0 / 52 (0.00%)	1 / 48 (2.08%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 1	0 / 0
Oropharyngeal squamous cell carcinoma			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Ovarian neoplasm			
subjects affected / exposed	0 / 52 (0.00%)	1 / 48 (2.08%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Squamous cell carcinoma of head and neck			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0

Squamous cell carcinoma of skin			
subjects affected / exposed	0 / 52 (0.00%)	1 / 48 (2.08%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Transformation to acute myeloid leukaemia			
subjects affected / exposed	2 / 52 (3.85%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 2	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Vascular disorders			
Aortic dissection			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Haematoma			
subjects affected / exposed	1 / 52 (1.92%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hypertension			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences causally related to treatment / all	0 / 0	0 / 0	1 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hypotension			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	2 / 44 (4.55%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 1
Peripheral ischaemia			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
General disorders and administration site conditions			
Asthenia			

subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Chills			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Chest pain			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Drug withdrawal syndrome			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Disease progression			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 1
Fatigue			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	4 / 44 (9.09%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 5
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Malaise			
subjects affected / exposed	1 / 52 (1.92%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Multiple organ dysfunction syndrome			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Non-cardiac chest pain			

subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Oedema			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pyrexia			
subjects affected / exposed	0 / 52 (0.00%)	1 / 48 (2.08%)	3 / 44 (6.82%)
occurrences causally related to treatment / all	0 / 0	1 / 1	0 / 3
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Immune system disorders			
Serum sickness			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences causally related to treatment / all	0 / 0	0 / 0	1 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Respiratory, thoracic and mediastinal disorders			
Acute respiratory failure			
subjects affected / exposed	1 / 52 (1.92%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 1	0 / 0	0 / 0
Bronchospasm			
subjects affected / exposed	0 / 52 (0.00%)	1 / 48 (2.08%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Dyspnoea			
subjects affected / exposed	2 / 52 (3.85%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 3	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Epistaxis			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0

Organising pneumonia			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pleuritic pain			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pneumonia aspiration			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pneumonitis			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pneumothorax			
subjects affected / exposed	1 / 52 (1.92%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pulmonary embolism			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pulmonary haemorrhage			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pulmonary oedema			
subjects affected / exposed	0 / 52 (0.00%)	1 / 48 (2.08%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pulmonary hypertension			

subjects affected / exposed	0 / 52 (0.00%)	1 / 48 (2.08%)	1 / 44 (2.27%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Respiratory failure			
subjects affected / exposed	0 / 52 (0.00%)	1 / 48 (2.08%)	2 / 44 (4.55%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 1	0 / 1
Psychiatric disorders			
Confusional state			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	2 / 44 (4.55%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Delirium			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Depression			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Suicide attempt			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Investigations			
Blast cell count increased			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Blood bilirubin increased			
subjects affected / exposed	1 / 52 (1.92%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Ejection fraction decreased			

subjects affected / exposed	0 / 52 (0.00%)	1 / 48 (2.08%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Blood creatinine increased			
subjects affected / exposed	0 / 52 (0.00%)	1 / 48 (2.08%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	1 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Platelet count decreased			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Injury, poisoning and procedural complications			
Extradural haematoma			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Fall			
subjects affected / exposed	0 / 52 (0.00%)	1 / 48 (2.08%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pelvic fracture			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hip fracture			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Post procedural haemorrhage			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Subdural haematoma			

subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cardiac disorders			
Acute left ventricular failure			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Acute myocardial infarction			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Atrial fibrillation			
subjects affected / exposed	0 / 52 (0.00%)	1 / 48 (2.08%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 3	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Angina pectoris			
subjects affected / exposed	1 / 52 (1.92%)	1 / 48 (2.08%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cardiac arrest			
subjects affected / exposed	1 / 52 (1.92%)	1 / 48 (2.08%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 1	0 / 1	0 / 0
Cardiac failure			
subjects affected / exposed	1 / 52 (1.92%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cardiac failure congestive			
subjects affected / exposed	1 / 52 (1.92%)	1 / 48 (2.08%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Chronic coronary syndrome			

subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cardiac iron overload			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Dilated cardiomyopathy			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Left ventricular dysfunction			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Left ventricular failure			
subjects affected / exposed	1 / 52 (1.92%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pericarditis			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Sinus tachycardia			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Right ventricular failure			
subjects affected / exposed	0 / 52 (0.00%)	1 / 48 (2.08%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Nervous system disorders			
Brain stem haemorrhage			

subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cerebral infarction			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cerebrovascular accident			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Haemorrhage intracranial			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Metabolic encephalopathy			
subjects affected / exposed	0 / 52 (0.00%)	1 / 48 (2.08%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Spinal cord compression			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Paraesthesia			
subjects affected / exposed	1 / 52 (1.92%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Subarachnoid haemorrhage			
subjects affected / exposed	0 / 52 (0.00%)	1 / 48 (2.08%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Superior sagittal sinus thrombosis			

subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Syncope			
subjects affected / exposed	1 / 52 (1.92%)	1 / 48 (2.08%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Blood and lymphatic system disorders			
Anaemia			
subjects affected / exposed	2 / 52 (3.85%)	1 / 48 (2.08%)	2 / 44 (4.55%)
occurrences causally related to treatment / all	0 / 2	0 / 4	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Febrile neutropenia			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	9 / 44 (20.45%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 9
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 1
Leukocytosis			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Leukostasis syndrome			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Splenic infarction			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Splenomegaly			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Thrombocytopenia			

subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences causally related to treatment / all	0 / 0	0 / 0	1 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Ear and labyrinth disorders			
Vertigo			
subjects affected / exposed	0 / 52 (0.00%)	1 / 48 (2.08%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Gastrointestinal disorders			
Abdominal pain			
subjects affected / exposed	1 / 52 (1.92%)	0 / 48 (0.00%)	2 / 44 (4.55%)
occurrences causally related to treatment / all	0 / 1	0 / 0	1 / 2
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Colitis			
subjects affected / exposed	1 / 52 (1.92%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences causally related to treatment / all	2 / 3	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Abdominal wall haematoma			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Constipation			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	2 / 44 (4.55%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Diarrhoea			
subjects affected / exposed	1 / 52 (1.92%)	0 / 48 (0.00%)	3 / 44 (6.82%)
occurrences causally related to treatment / all	1 / 1	0 / 0	2 / 3
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Duodenal ulcer haemorrhage			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Gastric ulcer			

subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Gastrointestinal disorder			
subjects affected / exposed	1 / 52 (1.92%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Gastrointestinal haemorrhage			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Haemorrhoids			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Intestinal haematoma			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Intestinal obstruction			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Lower gastrointestinal haemorrhage			
subjects affected / exposed	1 / 52 (1.92%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Rectal haemorrhage			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Nausea			

subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	2 / 44 (4.55%)
occurrences causally related to treatment / all	0 / 0	0 / 0	2 / 2
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Small intestinal obstruction			
subjects affected / exposed	1 / 52 (1.92%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Vomiting			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	3 / 44 (6.82%)
occurrences causally related to treatment / all	0 / 0	0 / 0	2 / 3
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hepatobiliary disorders			
Biliary dilatation			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hepatocellular injury			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hypertransaminasaemia			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Skin and subcutaneous tissue disorders			
Cutaneous vasculitis			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Drug eruption			
subjects affected / exposed	1 / 52 (1.92%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	1 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Renal and urinary disorders			

Acute kidney injury			
subjects affected / exposed	1 / 52 (1.92%)	1 / 48 (2.08%)	1 / 44 (2.27%)
occurrences causally related to treatment / all	0 / 1	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Haematuria			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Renal cyst ruptured			
subjects affected / exposed	1 / 52 (1.92%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Renal haemorrhage			
subjects affected / exposed	0 / 52 (0.00%)	1 / 48 (2.08%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Endocrine disorders			
Adrenal haematoma			
subjects affected / exposed	1 / 52 (1.92%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Inappropriate antidiuretic hormone secretion			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Musculoskeletal and connective tissue disorders			
Back pain			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Bone pain			

subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Joint effusion			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Muscular weakness			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Musculoskeletal pain			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Neck pain			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Infections and infestations			
Abscess limb			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Aspergillus infection			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 1
Bacteraemia			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Bronchiolitis			

subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
COVID-19			
subjects affected / exposed	1 / 52 (1.92%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 1	0 / 0	0 / 0
Bronchopulmonary aspergillosis			
subjects affected / exposed	0 / 52 (0.00%)	1 / 48 (2.08%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
COVID-19 pneumonia			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Clostridium difficile infection			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cellulitis			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Dermo-hypodermitis			
subjects affected / exposed	0 / 52 (0.00%)	1 / 48 (2.08%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Device related sepsis			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Device related infection			

subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	2 / 44 (4.55%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Diverticulitis			
subjects affected / exposed	0 / 52 (0.00%)	1 / 48 (2.08%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Empyema			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Endocarditis bacterial			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Enterococcal bacteraemia			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 1
Escherichia bacteraemia			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 1
Escherichia infection			
subjects affected / exposed	0 / 52 (0.00%)	1 / 48 (2.08%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Escherichia urinary tract infection			
subjects affected / exposed	0 / 52 (0.00%)	1 / 48 (2.08%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Fournier's gangrene			

subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Herpes zoster			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Herpes simplex reactivation			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Gastroenteritis viral			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Influenza			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Localised infection			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Nocardiosis			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pneumonia			
subjects affected / exposed	3 / 52 (5.77%)	4 / 48 (8.33%)	4 / 44 (9.09%)
occurrences causally related to treatment / all	0 / 3	1 / 4	0 / 5
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 1
Pneumonia acinetobacter			

subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pneumonia bacterial			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pneumonia pneumococcal			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pneumonia staphylococcal			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Post procedural infection			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pyelonephritis			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pyelonephritis acute			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Respiratory tract infection			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Septic shock			

subjects affected / exposed	0 / 52 (0.00%)	2 / 48 (4.17%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	1 / 2	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 1	0 / 0
Sepsis			
subjects affected / exposed	1 / 52 (1.92%)	1 / 48 (2.08%)	3 / 44 (6.82%)
occurrences causally related to treatment / all	0 / 1	0 / 1	0 / 3
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Streptococcal sepsis			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Tooth infection			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Urinary tract infection			
subjects affected / exposed	0 / 52 (0.00%)	2 / 48 (4.17%)	1 / 44 (2.27%)
occurrences causally related to treatment / all	0 / 0	0 / 2	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Urosepsis			
subjects affected / exposed	1 / 52 (1.92%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Viral infection			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Metabolism and nutrition disorders			
Decreased appetite			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Dehydration			

subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Failure to thrive			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Gout			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hypercalcaemia			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hyperglycaemia			
subjects affected / exposed	0 / 52 (0.00%)	1 / 48 (2.08%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 0	1 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hyperkalaemia			
subjects affected / exposed	1 / 52 (1.92%)	2 / 48 (4.17%)	1 / 44 (2.27%)
occurrences causally related to treatment / all	0 / 1	2 / 2	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hyponatraemia			
subjects affected / exposed	2 / 52 (3.85%)	1 / 48 (2.08%)	0 / 44 (0.00%)
occurrences causally related to treatment / all	0 / 2	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0

Serious adverse events	Phase 2 Overall	Phase 1 275 mg tablet PO daily	
Total subjects affected by serious adverse events			
subjects affected / exposed	133 / 292 (45.55%)	6 / 6 (100.00%)	
number of deaths (all causes)	22	1	
number of deaths resulting from adverse events	3	0	

Neoplasms benign, malignant and unspecified (incl cysts and polyps)			
Basal cell carcinoma			
subjects affected / exposed	2 / 292 (0.68%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 4	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Acute myeloid leukaemia			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 1	0 / 0	
Breast cancer			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Bowen's disease			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Colon cancer metastatic			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	1 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Glioblastoma			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Invasive breast carcinoma			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Haematopoietic neoplasm			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	

Keratoacanthoma				
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)		
occurrences causally related to treatment / all	0 / 1	0 / 0		
deaths causally related to treatment / all	0 / 0	0 / 0		
Lung neoplasm malignant				
subjects affected / exposed	2 / 292 (0.68%)	0 / 6 (0.00%)		
occurrences causally related to treatment / all	0 / 2	0 / 0		
deaths causally related to treatment / all	0 / 0	0 / 0		
Malignant melanoma				
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)		
occurrences causally related to treatment / all	1 / 2	0 / 0		
deaths causally related to treatment / all	0 / 0	0 / 0		
Metastatic squamous cell carcinoma				
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)		
occurrences causally related to treatment / all	0 / 1	0 / 0		
deaths causally related to treatment / all	0 / 0	0 / 0		
Neuroendocrine carcinoma of the skin				
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)		
occurrences causally related to treatment / all	0 / 1	0 / 0		
deaths causally related to treatment / all	0 / 1	0 / 0		
Oropharyngeal squamous cell carcinoma				
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)		
occurrences causally related to treatment / all	0 / 1	0 / 0		
deaths causally related to treatment / all	0 / 0	0 / 0		
Ovarian neoplasm				
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)		
occurrences causally related to treatment / all	0 / 1	0 / 0		
deaths causally related to treatment / all	0 / 0	0 / 0		
Squamous cell carcinoma of head and neck				
subjects affected / exposed	2 / 292 (0.68%)	0 / 6 (0.00%)		
occurrences causally related to treatment / all	0 / 2	0 / 0		
deaths causally related to treatment / all	0 / 0	0 / 0		

Squamous cell carcinoma of skin			
subjects affected / exposed	3 / 292 (1.03%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	4 / 7	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Transformation to acute myeloid leukaemia			
subjects affected / exposed	9 / 292 (3.08%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 9	0 / 0	
deaths causally related to treatment / all	0 / 1	0 / 0	
Vascular disorders			
Aortic dissection			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Haematoma			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hypertension			
subjects affected / exposed	0 / 292 (0.00%)	1 / 6 (16.67%)	
occurrences causally related to treatment / all	0 / 0	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hypotension			
subjects affected / exposed	0 / 292 (0.00%)	1 / 6 (16.67%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Peripheral ischaemia			
subjects affected / exposed	3 / 292 (1.03%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 3	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
General disorders and administration site conditions			
Asthenia			

subjects affected / exposed	0 / 292 (0.00%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Chills			
subjects affected / exposed	0 / 292 (0.00%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Chest pain			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Drug withdrawal syndrome			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 1	0 / 0	
Disease progression			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 1	0 / 0	
Fatigue			
subjects affected / exposed	0 / 292 (0.00%)	1 / 6 (16.67%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Malaise			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Multiple organ dysfunction syndrome			
subjects affected / exposed	2 / 292 (0.68%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	1 / 2	0 / 0	
deaths causally related to treatment / all	1 / 2	0 / 0	
Non-cardiac chest pain			

subjects affected / exposed	2 / 292 (0.68%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Oedema			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pyrexia			
subjects affected / exposed	6 / 292 (2.05%)	1 / 6 (16.67%)	
occurrences causally related to treatment / all	2 / 7	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Immune system disorders			
Serum sickness			
subjects affected / exposed	0 / 292 (0.00%)	1 / 6 (16.67%)	
occurrences causally related to treatment / all	0 / 0	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Respiratory, thoracic and mediastinal disorders			
Acute respiratory failure			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 1	0 / 0	
Bronchospasm			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Dyspnoea			
subjects affected / exposed	5 / 292 (1.71%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	2 / 6	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Epistaxis			
subjects affected / exposed	2 / 292 (0.68%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	1 / 3	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	

Organising pneumonia			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pleuritic pain			
subjects affected / exposed	0 / 292 (0.00%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pneumonia aspiration			
subjects affected / exposed	0 / 292 (0.00%)	1 / 6 (16.67%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pneumonitis			
subjects affected / exposed	0 / 292 (0.00%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pneumothorax			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pulmonary embolism			
subjects affected / exposed	2 / 292 (0.68%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pulmonary haemorrhage			
subjects affected / exposed	0 / 292 (0.00%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pulmonary oedema			
subjects affected / exposed	2 / 292 (0.68%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pulmonary hypertension			

subjects affected / exposed	1 / 292 (0.34%)	1 / 6 (16.67%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Respiratory failure			
subjects affected / exposed	4 / 292 (1.37%)	1 / 6 (16.67%)	
occurrences causally related to treatment / all	0 / 4	0 / 1	
deaths causally related to treatment / all	0 / 3	0 / 0	
Psychiatric disorders			
Confusional state			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Delirium			
subjects affected / exposed	0 / 292 (0.00%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Depression			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Suicide attempt			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Investigations			
Blast cell count increased			
subjects affected / exposed	9 / 292 (3.08%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	2 / 9	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Blood bilirubin increased			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Ejection fraction decreased			

subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Blood creatinine increased			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	1 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Platelet count decreased			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Injury, poisoning and procedural complications			
Extradural haematoma			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Fall			
subjects affected / exposed	3 / 292 (1.03%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 3	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pelvic fracture			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hip fracture			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Post procedural haemorrhage			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Subdural haematoma			

subjects affected / exposed	0 / 292 (0.00%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cardiac disorders			
Acute left ventricular failure			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Acute myocardial infarction			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Atrial fibrillation			
subjects affected / exposed	3 / 292 (1.03%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 5	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Angina pectoris			
subjects affected / exposed	2 / 292 (0.68%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cardiac arrest			
subjects affected / exposed	2 / 292 (0.68%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 2	0 / 0	
Cardiac failure			
subjects affected / exposed	3 / 292 (1.03%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	1 / 3	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cardiac failure congestive			
subjects affected / exposed	3 / 292 (1.03%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 3	0 / 0	
deaths causally related to treatment / all	0 / 1	0 / 0	
Chronic coronary syndrome			

subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cardiac iron overload			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Dilated cardiomyopathy			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Left ventricular dysfunction			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	1 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Left ventricular failure			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pericarditis			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Sinus tachycardia			
subjects affected / exposed	0 / 292 (0.00%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Right ventricular failure			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Nervous system disorders			
Brain stem haemorrhage			

subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 1	0 / 0	
Cerebral infarction			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 1	0 / 0	
Cerebrovascular accident			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Haemorrhage intracranial			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 1	0 / 0	
Metabolic encephalopathy			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Spinal cord compression			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Paraesthesia			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Subarachnoid haemorrhage			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Superior sagittal sinus thrombosis			

subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Syncope			
subjects affected / exposed	3 / 292 (1.03%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 3	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Blood and lymphatic system disorders			
Anaemia			
subjects affected / exposed	11 / 292 (3.77%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	2 / 14	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Febrile neutropenia			
subjects affected / exposed	1 / 292 (0.34%)	2 / 6 (33.33%)	
occurrences causally related to treatment / all	0 / 1	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Leukocytosis			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Leukostasis syndrome			
subjects affected / exposed	0 / 292 (0.00%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Splenic infarction			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Splenomegaly			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Thrombocytopenia			

subjects affected / exposed	0 / 292 (0.00%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Ear and labyrinth disorders			
Vertigo			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Gastrointestinal disorders			
Abdominal pain			
subjects affected / exposed	3 / 292 (1.03%)	1 / 6 (16.67%)	
occurrences causally related to treatment / all	1 / 3	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Colitis			
subjects affected / exposed	1 / 292 (0.34%)	1 / 6 (16.67%)	
occurrences causally related to treatment / all	2 / 3	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Abdominal wall haematoma			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	1 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Constipation			
subjects affected / exposed	0 / 292 (0.00%)	1 / 6 (16.67%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Diarrhoea			
subjects affected / exposed	1 / 292 (0.34%)	1 / 6 (16.67%)	
occurrences causally related to treatment / all	1 / 1	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Duodenal ulcer haemorrhage			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	1 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Gastric ulcer			

subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Gastrointestinal disorder			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Gastrointestinal haemorrhage			
subjects affected / exposed	3 / 292 (1.03%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 3	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Haemorrhoids			
subjects affected / exposed	0 / 292 (0.00%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Intestinal haematoma			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Intestinal obstruction			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Lower gastrointestinal haemorrhage			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Rectal haemorrhage			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Nausea			

subjects affected / exposed	0 / 292 (0.00%)	2 / 6 (33.33%)	
occurrences causally related to treatment / all	0 / 0	2 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Small intestinal obstruction			
subjects affected / exposed	2 / 292 (0.68%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Vomiting			
subjects affected / exposed	1 / 292 (0.34%)	3 / 6 (50.00%)	
occurrences causally related to treatment / all	1 / 1	2 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hepatobiliary disorders			
Biliary dilatation			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hepatocellular injury			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hypertransaminasaemia			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	1 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Skin and subcutaneous tissue disorders			
Cutaneous vasculitis			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Drug eruption			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	1 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Renal and urinary disorders			

Acute kidney injury			
subjects affected / exposed	6 / 292 (2.05%)	1 / 6 (16.67%)	
occurrences causally related to treatment / all	2 / 6	0 / 1	
deaths causally related to treatment / all	1 / 1	0 / 0	
Haematuria			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Renal cyst ruptured			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Renal haemorrhage			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Endocrine disorders			
Adrenal haematoma			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Inappropriate antidiuretic hormone secretion			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Musculoskeletal and connective tissue disorders			
Back pain			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Bone pain			

subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Joint effusion			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Muscular weakness			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Musculoskeletal pain			
subjects affected / exposed	0 / 292 (0.00%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Neck pain			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Infections and infestations			
Abscess limb			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Aspergillus infection			
subjects affected / exposed	0 / 292 (0.00%)	1 / 6 (16.67%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 1	
Bacteraemia			
subjects affected / exposed	0 / 292 (0.00%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Bronchiolitis			

subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	1 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
COVID-19			
subjects affected / exposed	5 / 292 (1.71%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	1 / 5	0 / 0	
deaths causally related to treatment / all	0 / 2	0 / 0	
Bronchopulmonary aspergillosis			
subjects affected / exposed	2 / 292 (0.68%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
COVID-19 pneumonia			
subjects affected / exposed	5 / 292 (1.71%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	2 / 5	0 / 0	
deaths causally related to treatment / all	1 / 1	0 / 0	
Clostridium difficile infection			
subjects affected / exposed	0 / 292 (0.00%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cellulitis			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Dermo-hypodermitis			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Device related sepsis			
subjects affected / exposed	0 / 292 (0.00%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Device related infection			

subjects affected / exposed	0 / 292 (0.00%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Diverticulitis			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Empyema			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Endocarditis bacterial			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Enterococcal bacteraemia			
subjects affected / exposed	0 / 292 (0.00%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Escherichia bacteraemia			
subjects affected / exposed	0 / 292 (0.00%)	1 / 6 (16.67%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 1	
Escherichia infection			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Escherichia urinary tract infection			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Fournier's gangrene			

subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Herpes zoster			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Herpes simplex reactivation			
subjects affected / exposed	0 / 292 (0.00%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Gastroenteritis viral			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Influenza			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Localised infection			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Nocardiosis			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pneumonia			
subjects affected / exposed	18 / 292 (6.16%)	1 / 6 (16.67%)	
occurrences causally related to treatment / all	4 / 18	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 1	
Pneumonia acinetobacter			

subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pneumonia bacterial			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 1	0 / 0	
Pneumonia pneumococcal			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pneumonia staphylococcal			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	1 / 1	0 / 0	
deaths causally related to treatment / all	1 / 1	0 / 0	
Post procedural infection			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pyelonephritis			
subjects affected / exposed	2 / 292 (0.68%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	1 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pyelonephritis acute			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Respiratory tract infection			
subjects affected / exposed	2 / 292 (0.68%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Septic shock			

subjects affected / exposed	3 / 292 (1.03%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	1 / 3	0 / 0	
deaths causally related to treatment / all	0 / 1	0 / 0	
Sepsis			
subjects affected / exposed	4 / 292 (1.37%)	2 / 6 (33.33%)	
occurrences causally related to treatment / all	1 / 4	0 / 2	
deaths causally related to treatment / all	1 / 2	0 / 0	
Streptococcal sepsis			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	1 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Tooth infection			
subjects affected / exposed	0 / 292 (0.00%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Urinary tract infection			
subjects affected / exposed	6 / 292 (2.05%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 6	0 / 0	
deaths causally related to treatment / all	0 / 1	0 / 0	
Urosepsis			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Viral infection			
subjects affected / exposed	2 / 292 (0.68%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Metabolism and nutrition disorders			
Decreased appetite			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	1 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Dehydration			

subjects affected / exposed	0 / 292 (0.00%)	1 / 6 (16.67%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Failure to thrive			
subjects affected / exposed	0 / 292 (0.00%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Gout			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hypercalcaemia			
subjects affected / exposed	0 / 292 (0.00%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hyperglycaemia			
subjects affected / exposed	2 / 292 (0.68%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	2 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hyperkalaemia			
subjects affected / exposed	4 / 292 (1.37%)	1 / 6 (16.67%)	
occurrences causally related to treatment / all	2 / 5	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hyponatraemia			
subjects affected / exposed	5 / 292 (1.71%)	0 / 6 (0.00%)	
occurrences causally related to treatment / all	1 / 5	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	

Frequency threshold for reporting non-serious adverse events: 5 %

Non-serious adverse events	Phase 1 225 mg tablet PO daily	Phase 1 400 mg capsule PO daily	Phase 1 300 mg capsule PO daily
Total subjects affected by non-serious adverse events			
subjects affected / exposed	8 / 8 (100.00%)	7 / 7 (100.00%)	4 / 4 (100.00%)
Neoplasms benign, malignant and unspecified (incl cysts and polyps)			
Squamous cell carcinoma			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Basal cell carcinoma			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Vascular disorders			
Flushing			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Haematoma			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Hot flush			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Hypotension			
subjects affected / exposed	1 / 8 (12.50%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	1	0	0
Hypertension			
subjects affected / exposed	0 / 8 (0.00%)	1 / 7 (14.29%)	0 / 4 (0.00%)
occurrences (all)	0	1	0
General disorders and administration site conditions			
Chest pain			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	1 / 4 (25.00%)
occurrences (all)	0	0	1
Asthenia			
subjects affected / exposed	0 / 8 (0.00%)	2 / 7 (28.57%)	0 / 4 (0.00%)
occurrences (all)	0	2	0
Early satiety			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0

Chills			
subjects affected / exposed	1 / 8 (12.50%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	1	0	0
Fatigue			
subjects affected / exposed	3 / 8 (37.50%)	5 / 7 (71.43%)	0 / 4 (0.00%)
occurrences (all)	3	6	0
Gait disturbance			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Influenza like illness			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Malaise			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Mucosal inflammation			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Nodule			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Non-cardiac chest pain			
subjects affected / exposed	1 / 8 (12.50%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	1	0	0
Oedema peripheral			
subjects affected / exposed	2 / 8 (25.00%)	1 / 7 (14.29%)	1 / 4 (25.00%)
occurrences (all)	2	1	1
Pain			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Pyrexia			
subjects affected / exposed	0 / 8 (0.00%)	2 / 7 (28.57%)	1 / 4 (25.00%)
occurrences (all)	0	2	1
Peripheral swelling			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0

Reproductive system and breast disorders			
Genital lesion			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Vaginal haemorrhage			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Respiratory, thoracic and mediastinal disorders			
Dyspnoea			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Cough			
subjects affected / exposed	1 / 8 (12.50%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	1	0	0
Dyspnoea exertional			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Nasal congestion			
subjects affected / exposed	1 / 8 (12.50%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	1	0	0
Hypoxia			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Epistaxis			
subjects affected / exposed	3 / 8 (37.50%)	1 / 7 (14.29%)	1 / 4 (25.00%)
occurrences (all)	6	1	1
Oropharyngeal pain			
subjects affected / exposed	1 / 8 (12.50%)	0 / 7 (0.00%)	1 / 4 (25.00%)
occurrences (all)	1	0	1
Pleural effusion			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Productive cough			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Pulmonary oedema			

subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Rhinorrhoea			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Sinus pain			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Upper-airway cough syndrome			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Wheezing			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Psychiatric disorders			
Anxiety			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Confusional state			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Delirium			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Depression			
subjects affected / exposed	0 / 8 (0.00%)	1 / 7 (14.29%)	0 / 4 (0.00%)
occurrences (all)	0	1	0
Insomnia			
subjects affected / exposed	0 / 8 (0.00%)	1 / 7 (14.29%)	0 / 4 (0.00%)
occurrences (all)	0	1	0
Investigations			
Bilirubin conjugated increased			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Aspartate aminotransferase increased			

subjects affected / exposed	1 / 8 (12.50%)	1 / 7 (14.29%)	0 / 4 (0.00%)
occurrences (all)	1	2	0
Alanine aminotransferase increased			
subjects affected / exposed	1 / 8 (12.50%)	2 / 7 (28.57%)	0 / 4 (0.00%)
occurrences (all)	1	2	0
Activated partial thromboplastin time prolonged			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Blast cell count increased			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Blood alkaline phosphatase increased			
subjects affected / exposed	1 / 8 (12.50%)	1 / 7 (14.29%)	0 / 4 (0.00%)
occurrences (all)	1	2	0
Blood bilirubin increased			
subjects affected / exposed	1 / 8 (12.50%)	1 / 7 (14.29%)	0 / 4 (0.00%)
occurrences (all)	1	1	0
Cardiac murmur			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Blood phosphorus increased			
subjects affected / exposed	1 / 8 (12.50%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	1	0	0
Blood uric acid increased			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Blood creatinine increased			
subjects affected / exposed	0 / 8 (0.00%)	1 / 7 (14.29%)	0 / 4 (0.00%)
occurrences (all)	0	2	0
International normalised ratio increased			
subjects affected / exposed	1 / 8 (12.50%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	1	0	0
Ejection fraction decreased			

subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Electrocardiogram QRS complex abnormal			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Electrocardiogram QT prolonged			
subjects affected / exposed	1 / 8 (12.50%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	1	0	0
Haematocrit increased			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Lymphocyte count decreased			
subjects affected / exposed	0 / 8 (0.00%)	1 / 7 (14.29%)	0 / 4 (0.00%)
occurrences (all)	0	4	0
Neutrophil count decreased			
subjects affected / exposed	1 / 8 (12.50%)	2 / 7 (28.57%)	0 / 4 (0.00%)
occurrences (all)	2	5	0
Platelet count decreased			
subjects affected / exposed	2 / 8 (25.00%)	2 / 7 (28.57%)	0 / 4 (0.00%)
occurrences (all)	2	2	0
Transaminases increased			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Troponin T increased			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Weight decreased			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	1 / 4 (25.00%)
occurrences (all)	0	0	1
Weight increased			
subjects affected / exposed	1 / 8 (12.50%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	1	0	0
White blood cell count decreased			
subjects affected / exposed	0 / 8 (0.00%)	2 / 7 (28.57%)	0 / 4 (0.00%)
occurrences (all)	0	7	0

Injury, poisoning and procedural complications			
Contusion			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Fall			
subjects affected / exposed	1 / 8 (12.50%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	1	0	0
Vascular access site complication			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	1 / 4 (25.00%)
occurrences (all)	0	0	1
Skin laceration			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Procedural site reaction			
subjects affected / exposed	0 / 8 (0.00%)	1 / 7 (14.29%)	0 / 4 (0.00%)
occurrences (all)	0	2	0
Procedural pain			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Nail injury			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Wound			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Cardiac disorders			
Angina pectoris			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Atrial fibrillation			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	1 / 4 (25.00%)
occurrences (all)	0	0	1
Atrioventricular block			
subjects affected / exposed	1 / 8 (12.50%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	1	0	0
Hyperdynamic left ventricle			

subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Palpitations			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Sinus tachycardia			
subjects affected / exposed	0 / 8 (0.00%)	1 / 7 (14.29%)	0 / 4 (0.00%)
occurrences (all)	0	1	0
Tachycardia			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Nervous system disorders			
Ageusia			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Dysarthria			
subjects affected / exposed	1 / 8 (12.50%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	1	0	0
Dizziness			
subjects affected / exposed	0 / 8 (0.00%)	1 / 7 (14.29%)	0 / 4 (0.00%)
occurrences (all)	0	1	0
Cerebrovascular accident			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Memory impairment			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Lethargy			
subjects affected / exposed	0 / 8 (0.00%)	1 / 7 (14.29%)	0 / 4 (0.00%)
occurrences (all)	0	1	0
Hypoaesthesia			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Headache			
subjects affected / exposed	1 / 8 (12.50%)	2 / 7 (28.57%)	0 / 4 (0.00%)
occurrences (all)	1	2	0

Dysgeusia			
subjects affected / exposed	3 / 8 (37.50%)	1 / 7 (14.29%)	1 / 4 (25.00%)
occurrences (all)	3	1	1
Presyncope			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Paraesthesia			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Neuropathy peripheral			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Myoclonus			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Sciatica			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Migraine			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Taste disorder			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Syncope			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Blood and lymphatic system disorders			
Anaemia			
subjects affected / exposed	1 / 8 (12.50%)	4 / 7 (57.14%)	1 / 4 (25.00%)
occurrences (all)	1	7	1
Leukocytosis			
subjects affected / exposed	1 / 8 (12.50%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	1	0	0
Lymphadenopathy			

subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Neutropenia			
subjects affected / exposed	0 / 8 (0.00%)	1 / 7 (14.29%)	1 / 4 (25.00%)
occurrences (all)	0	1	1
Spleen disorder			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Thrombocytosis			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Thrombocytopenia			
subjects affected / exposed	0 / 8 (0.00%)	1 / 7 (14.29%)	1 / 4 (25.00%)
occurrences (all)	0	1	1
Ear and labyrinth disorders			
Ear discomfort			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Ear pain			
subjects affected / exposed	1 / 8 (12.50%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	1	0	0
Ear haemorrhage			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Hypoacusis			
subjects affected / exposed	1 / 8 (12.50%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	1	0	0
Tinnitus			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Eye disorders			
Eye haemorrhage			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Vision blurred			

subjects affected / exposed	1 / 8 (12.50%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	1	0	0
Gastrointestinal disorders			
Abdominal discomfort			
subjects affected / exposed	1 / 8 (12.50%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	1	0	0
Abdominal distension			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Abdominal pain			
subjects affected / exposed	1 / 8 (12.50%)	1 / 7 (14.29%)	0 / 4 (0.00%)
occurrences (all)	1	1	0
Abdominal pain upper			
subjects affected / exposed	0 / 8 (0.00%)	1 / 7 (14.29%)	0 / 4 (0.00%)
occurrences (all)	0	1	0
Anal incontinence			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Constipation			
subjects affected / exposed	1 / 8 (12.50%)	1 / 7 (14.29%)	0 / 4 (0.00%)
occurrences (all)	1	1	0
Diarrhoea			
subjects affected / exposed	5 / 8 (62.50%)	3 / 7 (42.86%)	4 / 4 (100.00%)
occurrences (all)	8	3	4
Dry mouth			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Enterocolitis			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Dysphagia			
subjects affected / exposed	0 / 8 (0.00%)	1 / 7 (14.29%)	0 / 4 (0.00%)
occurrences (all)	0	1	0
Dyspepsia			
subjects affected / exposed	1 / 8 (12.50%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	1	0	0

Eructation			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Flatulence			
subjects affected / exposed	1 / 8 (12.50%)	1 / 7 (14.29%)	0 / 4 (0.00%)
occurrences (all)	1	1	0
Gastrooesophageal reflux disease			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Gingival bleeding			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Gingival pain			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Haemorrhoidal haemorrhage			
subjects affected / exposed	0 / 8 (0.00%)	1 / 7 (14.29%)	0 / 4 (0.00%)
occurrences (all)	0	1	0
Haemorrhoids			
subjects affected / exposed	1 / 8 (12.50%)	1 / 7 (14.29%)	0 / 4 (0.00%)
occurrences (all)	1	1	0
Mouth ulceration			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Mouth haemorrhage			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Melaena			
subjects affected / exposed	0 / 8 (0.00%)	1 / 7 (14.29%)	0 / 4 (0.00%)
occurrences (all)	0	1	0
Nausea			
subjects affected / exposed	5 / 8 (62.50%)	2 / 7 (28.57%)	2 / 4 (50.00%)
occurrences (all)	5	2	2
Proctalgia			
subjects affected / exposed	1 / 8 (12.50%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	1	0	0

Oral disorder			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Stomatitis			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Vomiting			
subjects affected / exposed	1 / 8 (12.50%)	3 / 7 (42.86%)	1 / 4 (25.00%)
occurrences (all)	2	3	1
Hepatobiliary disorders			
Hyperbilirubinaemia			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Skin and subcutaneous tissue disorders			
Dermatitis acneiform			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Dry skin			
subjects affected / exposed	0 / 8 (0.00%)	1 / 7 (14.29%)	0 / 4 (0.00%)
occurrences (all)	0	1	0
Erythema			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Hyperhidrosis			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Night sweats			
subjects affected / exposed	1 / 8 (12.50%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	1	0	0
Petechiae			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Pruritus			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Skin discolouration			

subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Rash maculo-papular			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Rash macular			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Rash			
subjects affected / exposed	0 / 8 (0.00%)	1 / 7 (14.29%)	0 / 4 (0.00%)
occurrences (all)	0	1	0
Skin ulcer			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Skin lesion			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Urticaria			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Skin induration			
subjects affected / exposed	0 / 8 (0.00%)	1 / 7 (14.29%)	0 / 4 (0.00%)
occurrences (all)	0	1	0
Renal and urinary disorders			
Acute kidney injury			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	1 / 4 (25.00%)
occurrences (all)	0	0	1
Dysuria			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Haematuria			
subjects affected / exposed	1 / 8 (12.50%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	1	0	0
Micturition urgency			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0

Nocturia			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Urinary incontinence			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Pollakiuria			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Musculoskeletal and connective tissue disorders			
Arthralgia			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Bone cyst			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Back pain			
subjects affected / exposed	0 / 8 (0.00%)	1 / 7 (14.29%)	1 / 4 (25.00%)
occurrences (all)	0	1	1
Bone pain			
subjects affected / exposed	1 / 8 (12.50%)	1 / 7 (14.29%)	0 / 4 (0.00%)
occurrences (all)	2	1	0
Groin pain			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Gouty arthritis			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Bursitis			
subjects affected / exposed	1 / 8 (12.50%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	1	0	0
Muscle spasms			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Joint swelling			

subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Muscular weakness			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Musculoskeletal chest pain			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Musculoskeletal disorder			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Musculoskeletal pain			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	1 / 4 (25.00%)
occurrences (all)	0	0	1
Myalgia			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Neck pain			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Pain in extremity			
subjects affected / exposed	1 / 8 (12.50%)	0 / 7 (0.00%)	1 / 4 (25.00%)
occurrences (all)	1	0	1
Pain in jaw			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Rhabdomyolysis			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Infections and infestations			
Cellulitis			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
COVID-19			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0

Bronchitis			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Cystitis			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Eye infection			
subjects affected / exposed	0 / 8 (0.00%)	1 / 7 (14.29%)	0 / 4 (0.00%)
occurrences (all)	0	1	0
Folliculitis			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Fungal skin infection			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Gastroenteritis			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Herpes simplex reactivation			
subjects affected / exposed	0 / 8 (0.00%)	1 / 7 (14.29%)	0 / 4 (0.00%)
occurrences (all)	0	1	0
Herpes zoster			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Hordeolum			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Influenza			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Localised infection			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Lower respiratory tract infection			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0

Orchitis			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	1 / 4 (25.00%)
occurrences (all)	0	0	1
Otitis media			
subjects affected / exposed	1 / 8 (12.50%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	1	0	0
Pharyngitis			
subjects affected / exposed	1 / 8 (12.50%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	1	0	0
Pneumonia			
subjects affected / exposed	1 / 8 (12.50%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	1	0	0
Nasopharyngitis			
subjects affected / exposed	1 / 8 (12.50%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	1	0	0
Oral herpes			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Skin infection			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Sinusitis			
subjects affected / exposed	0 / 8 (0.00%)	1 / 7 (14.29%)	0 / 4 (0.00%)
occurrences (all)	0	1	0
Septic shock			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Rhinitis			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Respiratory tract infection			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Staphylococcal infection			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	1 / 4 (25.00%)
occurrences (all)	0	0	1

Urinary tract infection bacterial			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Urinary tract infection			
subjects affected / exposed	0 / 8 (0.00%)	1 / 7 (14.29%)	0 / 4 (0.00%)
occurrences (all)	0	1	0
Upper respiratory tract infection			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Tooth infection			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Metabolism and nutrition disorders			
Decreased appetite			
subjects affected / exposed	5 / 8 (62.50%)	2 / 7 (28.57%)	1 / 4 (25.00%)
occurrences (all)	5	2	1
Dehydration			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	1 / 4 (25.00%)
occurrences (all)	0	0	1
Folate deficiency			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Gout			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Hyperglycaemia			
subjects affected / exposed	2 / 8 (25.00%)	0 / 7 (0.00%)	1 / 4 (25.00%)
occurrences (all)	2	0	1
Hypercholesterolaemia			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Hyperkalaemia			
subjects affected / exposed	1 / 8 (12.50%)	1 / 7 (14.29%)	0 / 4 (0.00%)
occurrences (all)	1	2	0
Hyperphosphataemia			

subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	1 / 4 (25.00%)
occurrences (all)	0	0	1
Hypermagnesaemia			
subjects affected / exposed	0 / 8 (0.00%)	1 / 7 (14.29%)	0 / 4 (0.00%)
occurrences (all)	0	2	0
Hypertriglyceridaemia			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Hyperuricaemia			
subjects affected / exposed	2 / 8 (25.00%)	2 / 7 (28.57%)	0 / 4 (0.00%)
occurrences (all)	2	3	0
Hypocalcaemia			
subjects affected / exposed	0 / 8 (0.00%)	1 / 7 (14.29%)	0 / 4 (0.00%)
occurrences (all)	0	1	0
Hypokalaemia			
subjects affected / exposed	1 / 8 (12.50%)	0 / 7 (0.00%)	1 / 4 (25.00%)
occurrences (all)	1	0	1
Hypomagnesaemia			
subjects affected / exposed	2 / 8 (25.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	2	0	0
Hyponatraemia			
subjects affected / exposed	0 / 8 (0.00%)	1 / 7 (14.29%)	0 / 4 (0.00%)
occurrences (all)	0	3	0
Hypophosphataemia			
subjects affected / exposed	0 / 8 (0.00%)	1 / 7 (14.29%)	0 / 4 (0.00%)
occurrences (all)	0	1	0
Iron overload			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Tumour lysis syndrome			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Vitamin D deficiency			
subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0
Type 2 diabetes mellitus			

subjects affected / exposed	0 / 8 (0.00%)	0 / 7 (0.00%)	0 / 4 (0.00%)
occurrences (all)	0	0	0

Non-serious adverse events	Phase 1 230 mg capsule PO daily	Phase 1 170 mg capsule PO daily	Phase 1 120 mg capsule PO daily
Total subjects affected by non-serious adverse events			
subjects affected / exposed	3 / 3 (100.00%)	3 / 3 (100.00%)	5 / 5 (100.00%)
Neoplasms benign, malignant and unspecified (incl cysts and polyps)			
Squamous cell carcinoma			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Basal cell carcinoma			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Vascular disorders			
Flushing			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Haematoma			
subjects affected / exposed	2 / 3 (66.67%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	2	0	0
Hot flush			
subjects affected / exposed	1 / 3 (33.33%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	1	0	0
Hypotension			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	1 / 5 (20.00%)
occurrences (all)	0	0	1
Hypertension			
subjects affected / exposed	1 / 3 (33.33%)	0 / 3 (0.00%)	1 / 5 (20.00%)
occurrences (all)	1	0	1
General disorders and administration site conditions			
Chest pain			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Asthenia			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	2 / 5 (40.00%)
occurrences (all)	0	0	2

Early satiety			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Chills			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Fatigue			
subjects affected / exposed	0 / 3 (0.00%)	2 / 3 (66.67%)	4 / 5 (80.00%)
occurrences (all)	0	2	4
Gait disturbance			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Influenza like illness			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Malaise			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Mucosal inflammation			
subjects affected / exposed	0 / 3 (0.00%)	1 / 3 (33.33%)	0 / 5 (0.00%)
occurrences (all)	0	1	0
Nodule			
subjects affected / exposed	0 / 3 (0.00%)	1 / 3 (33.33%)	0 / 5 (0.00%)
occurrences (all)	0	1	0
Non-cardiac chest pain			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Oedema peripheral			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Pain			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Pyrexia			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	1 / 5 (20.00%)
occurrences (all)	0	0	1

Peripheral swelling			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Reproductive system and breast disorders			
Genital lesion			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	1 / 5 (20.00%)
occurrences (all)	0	0	1
Vaginal haemorrhage			
subjects affected / exposed	0 / 3 (0.00%)	1 / 3 (33.33%)	0 / 5 (0.00%)
occurrences (all)	0	1	0
Respiratory, thoracic and mediastinal disorders			
Dyspnoea			
subjects affected / exposed	1 / 3 (33.33%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	1	0	0
Cough			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	1 / 5 (20.00%)
occurrences (all)	0	0	1
Dyspnoea exertional			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Nasal congestion			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Hypoxia			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Epistaxis			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	1 / 5 (20.00%)
occurrences (all)	0	0	2
Oropharyngeal pain			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Pleural effusion			
subjects affected / exposed	1 / 3 (33.33%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	1	0	0
Productive cough			

subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Pulmonary oedema			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Rhinorrhoea			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Sinus pain			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Upper-airway cough syndrome			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Wheezing			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Psychiatric disorders			
Anxiety			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Confusional state			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Delirium			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	1 / 5 (20.00%)
occurrences (all)	0	0	1
Depression			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Insomnia			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Investigations			
Bilirubin conjugated increased			

subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Aspartate aminotransferase increased			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Alanine aminotransferase increased			
subjects affected / exposed	0 / 3 (0.00%)	1 / 3 (33.33%)	0 / 5 (0.00%)
occurrences (all)	0	1	0
Activated partial thromboplastin time prolonged			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	1 / 5 (20.00%)
occurrences (all)	0	0	1
Blast cell count increased			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Blood alkaline phosphatase increased			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Blood bilirubin increased			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Cardiac murmur			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Blood phosphorus increased			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Blood uric acid increased			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Blood creatinine increased			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	1 / 5 (20.00%)
occurrences (all)	0	0	1
International normalised ratio increased			

subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Ejection fraction decreased			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Electrocardiogram QRS complex abnormal			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Electrocardiogram QT prolonged			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Haematocrit increased			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Lymphocyte count decreased			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Neutrophil count decreased			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Platelet count decreased			
subjects affected / exposed	0 / 3 (0.00%)	1 / 3 (33.33%)	0 / 5 (0.00%)
occurrences (all)	0	1	0
Transaminases increased			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	1 / 5 (20.00%)
occurrences (all)	0	0	1
Troponin T increased			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Weight decreased			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Weight increased			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0

White blood cell count decreased			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Injury, poisoning and procedural complications			
Contusion			
subjects affected / exposed	0 / 3 (0.00%)	1 / 3 (33.33%)	0 / 5 (0.00%)
occurrences (all)	0	1	0
Fall			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Vascular access site complication			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Skin laceration			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	1 / 5 (20.00%)
occurrences (all)	0	0	1
Procedural site reaction			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Procedural pain			
subjects affected / exposed	1 / 3 (33.33%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	1	0	0
Nail injury			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Wound			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Cardiac disorders			
Angina pectoris			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Atrial fibrillation			
subjects affected / exposed	1 / 3 (33.33%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	1	0	0
Atrioventricular block			

subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Hyperdynamic left ventricle			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	1 / 5 (20.00%)
occurrences (all)	0	0	1
Palpitations			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Sinus tachycardia			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Tachycardia			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Nervous system disorders			
Ageusia			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Dysarthria			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Dizziness			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	1 / 5 (20.00%)
occurrences (all)	0	0	1
Cerebrovascular accident			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Memory impairment			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Lethargy			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Hypoaesthesia			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0

Headache			
subjects affected / exposed	0 / 3 (0.00%)	1 / 3 (33.33%)	2 / 5 (40.00%)
occurrences (all)	0	1	2
Dysgeusia			
subjects affected / exposed	0 / 3 (0.00%)	1 / 3 (33.33%)	1 / 5 (20.00%)
occurrences (all)	0	1	1
Presyncope			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	1 / 5 (20.00%)
occurrences (all)	0	0	1
Paraesthesia			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Neuropathy peripheral			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Myoclonus			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	1 / 5 (20.00%)
occurrences (all)	0	0	1
Sciatica			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Migraine			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Taste disorder			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Syncope			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Blood and lymphatic system disorders			
Anaemia			
subjects affected / exposed	1 / 3 (33.33%)	0 / 3 (0.00%)	1 / 5 (20.00%)
occurrences (all)	1	0	2
Leukocytosis			

subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Lymphadenopathy			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Neutropenia			
subjects affected / exposed	0 / 3 (0.00%)	1 / 3 (33.33%)	0 / 5 (0.00%)
occurrences (all)	0	1	0
Spleen disorder			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Thrombocytosis			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Thrombocytopenia			
subjects affected / exposed	0 / 3 (0.00%)	1 / 3 (33.33%)	0 / 5 (0.00%)
occurrences (all)	0	1	0
Ear and labyrinth disorders			
Ear discomfort			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Ear pain			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Ear haemorrhage			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Hypoacusis			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Tinnitus			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Eye disorders			
Eye haemorrhage			

subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	1 / 5 (20.00%)
occurrences (all)	0	0	1
Vision blurred			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Gastrointestinal disorders			
Abdominal discomfort			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Abdominal distension			
subjects affected / exposed	1 / 3 (33.33%)	1 / 3 (33.33%)	0 / 5 (0.00%)
occurrences (all)	1	1	0
Abdominal pain			
subjects affected / exposed	1 / 3 (33.33%)	0 / 3 (0.00%)	1 / 5 (20.00%)
occurrences (all)	1	0	1
Abdominal pain upper			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Anal incontinence			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	1 / 5 (20.00%)
occurrences (all)	0	0	1
Constipation			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	1 / 5 (20.00%)
occurrences (all)	0	0	1
Diarrhoea			
subjects affected / exposed	0 / 3 (0.00%)	1 / 3 (33.33%)	1 / 5 (20.00%)
occurrences (all)	0	1	2
Dry mouth			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	1 / 5 (20.00%)
occurrences (all)	0	0	1
Enterocolitis			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Dysphagia			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0

Dyspepsia			
subjects affected / exposed	0 / 3 (0.00%)	1 / 3 (33.33%)	0 / 5 (0.00%)
occurrences (all)	0	1	0
Eructation			
subjects affected / exposed	1 / 3 (33.33%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	1	0	0
Flatulence			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Gastrooesophageal reflux disease			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Gingival bleeding			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Gingival pain			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Haemorrhoidal haemorrhage			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Haemorrhoids			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Mouth ulceration			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Mouth haemorrhage			
subjects affected / exposed	1 / 3 (33.33%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	1	0	0
Melaena			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Nausea			
subjects affected / exposed	0 / 3 (0.00%)	3 / 3 (100.00%)	4 / 5 (80.00%)
occurrences (all)	0	4	5

Proctalgia			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Oral disorder			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Stomatitis			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	1 / 5 (20.00%)
occurrences (all)	0	0	1
Vomiting			
subjects affected / exposed	0 / 3 (0.00%)	2 / 3 (66.67%)	1 / 5 (20.00%)
occurrences (all)	0	3	1
Hepatobiliary disorders			
Hyperbilirubinaemia			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Skin and subcutaneous tissue disorders			
Dermatitis acneiform			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Dry skin			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Erythema			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Hyperhidrosis			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Night sweats			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Petechiae			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Pruritus			

subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Skin discolouration			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Rash maculo-papular			
subjects affected / exposed	0 / 3 (0.00%)	1 / 3 (33.33%)	0 / 5 (0.00%)
occurrences (all)	0	1	0
Rash macular			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Rash			
subjects affected / exposed	0 / 3 (0.00%)	2 / 3 (66.67%)	0 / 5 (0.00%)
occurrences (all)	0	2	0
Skin ulcer			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Skin lesion			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Urticaria			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Skin induration			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Renal and urinary disorders			
Acute kidney injury			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Dysuria			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Haematuria			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0

Micturition urgency			
subjects affected / exposed	0 / 3 (0.00%)	1 / 3 (33.33%)	0 / 5 (0.00%)
occurrences (all)	0	1	0
Nocturia			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Urinary incontinence			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	1 / 5 (20.00%)
occurrences (all)	0	0	1
Pollakiuria			
subjects affected / exposed	0 / 3 (0.00%)	1 / 3 (33.33%)	0 / 5 (0.00%)
occurrences (all)	0	2	0
Musculoskeletal and connective tissue disorders			
Arthralgia			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	1 / 5 (20.00%)
occurrences (all)	0	0	1
Bone cyst			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Back pain			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Bone pain			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Groin pain			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Gouty arthritis			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	1 / 5 (20.00%)
occurrences (all)	0	0	1
Bursitis			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Muscle spasms			

subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Joint swelling			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Muscular weakness			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Musculoskeletal chest pain			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Musculoskeletal disorder			
subjects affected / exposed	0 / 3 (0.00%)	1 / 3 (33.33%)	0 / 5 (0.00%)
occurrences (all)	0	1	0
Musculoskeletal pain			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Myalgia			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Neck pain			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Pain in extremity			
subjects affected / exposed	1 / 3 (33.33%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	2	0	0
Pain in jaw			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Rhabdomyolysis			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Infections and infestations			
Cellulitis			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	1 / 5 (20.00%)
occurrences (all)	0	0	1

COVID-19			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Bronchitis			
subjects affected / exposed	1 / 3 (33.33%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	1	0	0
Cystitis			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Eye infection			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Folliculitis			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Fungal skin infection			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	1 / 5 (20.00%)
occurrences (all)	0	0	1
Gastroenteritis			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Herpes simplex reactivation			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Herpes zoster			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Hordeolum			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Influenza			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Localised infection			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0

Lower respiratory tract infection			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Orchitis			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Otitis media			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Pharyngitis			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Pneumonia			
subjects affected / exposed	1 / 3 (33.33%)	0 / 3 (0.00%)	1 / 5 (20.00%)
occurrences (all)	1	0	1
Nasopharyngitis			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Oral herpes			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Skin infection			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Sinusitis			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Septic shock			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Rhinitis			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Respiratory tract infection			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0

Staphylococcal infection			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Urinary tract infection bacterial			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	1 / 5 (20.00%)
occurrences (all)	0	0	1
Urinary tract infection			
subjects affected / exposed	0 / 3 (0.00%)	1 / 3 (33.33%)	1 / 5 (20.00%)
occurrences (all)	0	1	1
Upper respiratory tract infection			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Tooth infection			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Metabolism and nutrition disorders			
Decreased appetite			
subjects affected / exposed	2 / 3 (66.67%)	2 / 3 (66.67%)	3 / 5 (60.00%)
occurrences (all)	2	2	3
Dehydration			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Folate deficiency			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Gout			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Hyperglycaemia			
subjects affected / exposed	1 / 3 (33.33%)	0 / 3 (0.00%)	1 / 5 (20.00%)
occurrences (all)	1	0	1
Hypercholesterolaemia			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Hyperkalaemia			

subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Hyperphosphataemia			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Hypermagnesaemia			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Hypertriglyceridaemia			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Hyperuricaemia			
subjects affected / exposed	1 / 3 (33.33%)	0 / 3 (0.00%)	1 / 5 (20.00%)
occurrences (all)	1	0	1
Hypocalcaemia			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	1 / 5 (20.00%)
occurrences (all)	0	0	1
Hypokalaemia			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Hypomagnesaemia			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	1 / 5 (20.00%)
occurrences (all)	0	0	1
Hyponatraemia			
subjects affected / exposed	1 / 3 (33.33%)	0 / 3 (0.00%)	1 / 5 (20.00%)
occurrences (all)	1	0	1
Hypophosphataemia			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Iron overload			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Tumour lysis syndrome			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Vitamin D deficiency			

subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0
Type 2 diabetes mellitus			
subjects affected / exposed	0 / 3 (0.00%)	0 / 3 (0.00%)	0 / 5 (0.00%)
occurrences (all)	0	0	0

Non-serious adverse events	Phase 1 48 mg capsule PO daily	Phase 1 24 mg capsule PO daily	Phase 2 Arm 4
Total subjects affected by non-serious adverse events			
subjects affected / exposed	5 / 5 (100.00%)	3 / 3 (100.00%)	21 / 21 (100.00%)
Neoplasms benign, malignant and unspecified (incl cysts and polyps)			
Squamous cell carcinoma			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	1 / 21 (4.76%)
occurrences (all)	0	0	2
Basal cell carcinoma			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Vascular disorders			
Flushing			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	1 / 21 (4.76%)
occurrences (all)	0	0	1
Haematoma			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	1 / 21 (4.76%)
occurrences (all)	0	0	1
Hot flush			
subjects affected / exposed	0 / 5 (0.00%)	1 / 3 (33.33%)	1 / 21 (4.76%)
occurrences (all)	0	1	1
Hypotension			
subjects affected / exposed	1 / 5 (20.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	1	0	0
Hypertension			
subjects affected / exposed	1 / 5 (20.00%)	0 / 3 (0.00%)	2 / 21 (9.52%)
occurrences (all)	1	0	3
General disorders and administration site conditions			
Chest pain			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	2 / 21 (9.52%)
occurrences (all)	0	0	2

Asthenia			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Early satiety			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Chills			
subjects affected / exposed	0 / 5 (0.00%)	1 / 3 (33.33%)	0 / 21 (0.00%)
occurrences (all)	0	1	0
Fatigue			
subjects affected / exposed	3 / 5 (60.00%)	2 / 3 (66.67%)	5 / 21 (23.81%)
occurrences (all)	4	2	11
Gait disturbance			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Influenza like illness			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	1 / 21 (4.76%)
occurrences (all)	0	0	2
Malaise			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Mucosal inflammation			
subjects affected / exposed	1 / 5 (20.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	1	0	0
Nodule			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Non-cardiac chest pain			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	2 / 21 (9.52%)
occurrences (all)	0	0	3
Oedema peripheral			
subjects affected / exposed	3 / 5 (60.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	3	0	0
Pain			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0

Pyrexia			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	2 / 21 (9.52%)
occurrences (all)	0	0	3
Peripheral swelling			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Reproductive system and breast disorders			
Genital lesion			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Vaginal haemorrhage			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	1 / 21 (4.76%)
occurrences (all)	0	0	1
Respiratory, thoracic and mediastinal disorders			
Dyspnoea			
subjects affected / exposed	1 / 5 (20.00%)	1 / 3 (33.33%)	1 / 21 (4.76%)
occurrences (all)	1	1	1
Cough			
subjects affected / exposed	0 / 5 (0.00%)	2 / 3 (66.67%)	4 / 21 (19.05%)
occurrences (all)	0	2	7
Dyspnoea exertional			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Nasal congestion			
subjects affected / exposed	0 / 5 (0.00%)	1 / 3 (33.33%)	1 / 21 (4.76%)
occurrences (all)	0	1	1
Hypoxia			
subjects affected / exposed	1 / 5 (20.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	1	0	0
Epistaxis			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	2 / 21 (9.52%)
occurrences (all)	0	0	3
Oropharyngeal pain			
subjects affected / exposed	0 / 5 (0.00%)	1 / 3 (33.33%)	2 / 21 (9.52%)
occurrences (all)	0	1	2
Pleural effusion			

subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Productive cough			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Pulmonary oedema			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Rhinorrhoea			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	3 / 21 (14.29%)
occurrences (all)	0	0	3
Sinus pain			
subjects affected / exposed	1 / 5 (20.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	1	0	0
Upper-airway cough syndrome			
subjects affected / exposed	0 / 5 (0.00%)	1 / 3 (33.33%)	1 / 21 (4.76%)
occurrences (all)	0	1	1
Wheezing			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Psychiatric disorders			
Anxiety			
subjects affected / exposed	1 / 5 (20.00%)	2 / 3 (66.67%)	0 / 21 (0.00%)
occurrences (all)	1	2	0
Confusional state			
subjects affected / exposed	0 / 5 (0.00%)	1 / 3 (33.33%)	1 / 21 (4.76%)
occurrences (all)	0	1	1
Delirium			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Depression			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Insomnia			
subjects affected / exposed	1 / 5 (20.00%)	0 / 3 (0.00%)	2 / 21 (9.52%)
occurrences (all)	1	0	2

Investigations			
Bilirubin conjugated increased			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Aspartate aminotransferase increased			
subjects affected / exposed	1 / 5 (20.00%)	0 / 3 (0.00%)	1 / 21 (4.76%)
occurrences (all)	1	0	1
Alanine aminotransferase increased			
subjects affected / exposed	1 / 5 (20.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	1	0	0
Activated partial thromboplastin time prolonged			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Blast cell count increased			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Blood alkaline phosphatase increased			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Blood bilirubin increased			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	2 / 21 (9.52%)
occurrences (all)	0	0	4
Cardiac murmur			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	1 / 21 (4.76%)
occurrences (all)	0	0	1
Blood phosphorus increased			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Blood uric acid increased			
subjects affected / exposed	1 / 5 (20.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	1	0	0
Blood creatinine increased			
subjects affected / exposed	1 / 5 (20.00%)	0 / 3 (0.00%)	1 / 21 (4.76%)
occurrences (all)	1	0	1
International normalised ratio increased			

subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Ejection fraction decreased			
subjects affected / exposed	1 / 5 (20.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	1	0	0
Electrocardiogram QRS complex abnormal			
subjects affected / exposed	0 / 5 (0.00%)	1 / 3 (33.33%)	0 / 21 (0.00%)
occurrences (all)	0	1	0
Electrocardiogram QT prolonged			
subjects affected / exposed	1 / 5 (20.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	1	0	0
Haematocrit increased			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	2 / 21 (9.52%)
occurrences (all)	0	0	8
Lymphocyte count decreased			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Neutrophil count decreased			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Platelet count decreased			
subjects affected / exposed	0 / 5 (0.00%)	2 / 3 (66.67%)	2 / 21 (9.52%)
occurrences (all)	0	3	2
Transaminases increased			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Troponin T increased			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Weight decreased			
subjects affected / exposed	0 / 5 (0.00%)	1 / 3 (33.33%)	4 / 21 (19.05%)
occurrences (all)	0	1	4
Weight increased			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0

White blood cell count decreased			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Injury, poisoning and procedural complications			
Contusion			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	2 / 21 (9.52%)
occurrences (all)	0	0	4
Fall			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Vascular access site complication			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Skin laceration			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Procedural site reaction			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Procedural pain			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	1 / 21 (4.76%)
occurrences (all)	0	0	1
Nail injury			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	2 / 21 (9.52%)
occurrences (all)	0	0	3
Wound			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Cardiac disorders			
Angina pectoris			
subjects affected / exposed	0 / 5 (0.00%)	1 / 3 (33.33%)	0 / 21 (0.00%)
occurrences (all)	0	1	0
Atrial fibrillation			
subjects affected / exposed	1 / 5 (20.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	1	0	0
Atrioventricular block			

subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Hyperdynamic left ventricle			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Palpitations			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	3 / 21 (14.29%)
occurrences (all)	0	0	5
Sinus tachycardia			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Tachycardia			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	1 / 21 (4.76%)
occurrences (all)	0	0	1
Nervous system disorders			
Ageusia			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	6 / 21 (28.57%)
occurrences (all)	0	0	7
Dysarthria			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Dizziness			
subjects affected / exposed	0 / 5 (0.00%)	1 / 3 (33.33%)	4 / 21 (19.05%)
occurrences (all)	0	1	4
Cerebrovascular accident			
subjects affected / exposed	1 / 5 (20.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	1	0	0
Memory impairment			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	1 / 21 (4.76%)
occurrences (all)	0	0	1
Lethargy			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Hypoaesthesia			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0

Headache			
subjects affected / exposed	2 / 5 (40.00%)	0 / 3 (0.00%)	6 / 21 (28.57%)
occurrences (all)	2	0	11
Dysgeusia			
subjects affected / exposed	1 / 5 (20.00%)	0 / 3 (0.00%)	9 / 21 (42.86%)
occurrences (all)	1	0	12
Presyncope			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	3 / 21 (14.29%)
occurrences (all)	0	0	3
Paraesthesia			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	1 / 21 (4.76%)
occurrences (all)	0	0	1
Neuropathy peripheral			
subjects affected / exposed	1 / 5 (20.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	1	0	0
Myoclonus			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Sciatica			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Migraine			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	1 / 21 (4.76%)
occurrences (all)	0	0	1
Taste disorder			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	2 / 21 (9.52%)
occurrences (all)	0	0	3
Syncope			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	1 / 21 (4.76%)
occurrences (all)	0	0	1
Blood and lymphatic system disorders			
Anaemia			
subjects affected / exposed	1 / 5 (20.00%)	1 / 3 (33.33%)	1 / 21 (4.76%)
occurrences (all)	1	1	1
Leukocytosis			

subjects affected / exposed	1 / 5 (20.00%)	1 / 3 (33.33%)	1 / 21 (4.76%)
occurrences (all)	1	1	3
Lymphadenopathy			
subjects affected / exposed	1 / 5 (20.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	1	0	0
Neutropenia			
subjects affected / exposed	1 / 5 (20.00%)	0 / 3 (0.00%)	1 / 21 (4.76%)
occurrences (all)	1	0	1
Spleen disorder			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Thrombocytosis			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	1 / 21 (4.76%)
occurrences (all)	0	0	8
Thrombocytopenia			
subjects affected / exposed	1 / 5 (20.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	1	0	0
Ear and labyrinth disorders			
Ear discomfort			
subjects affected / exposed	0 / 5 (0.00%)	1 / 3 (33.33%)	0 / 21 (0.00%)
occurrences (all)	0	1	0
Ear pain			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Ear haemorrhage			
subjects affected / exposed	0 / 5 (0.00%)	1 / 3 (33.33%)	0 / 21 (0.00%)
occurrences (all)	0	1	0
Hypoacusis			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Tinnitus			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	1 / 21 (4.76%)
occurrences (all)	0	0	1
Eye disorders			
Eye haemorrhage			

subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Vision blurred			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	1 / 21 (4.76%)
occurrences (all)	0	0	1
Gastrointestinal disorders			
Abdominal discomfort			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Abdominal distension			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	1 / 21 (4.76%)
occurrences (all)	0	0	1
Abdominal pain			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	6 / 21 (28.57%)
occurrences (all)	0	0	7
Abdominal pain upper			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	3 / 21 (14.29%)
occurrences (all)	0	0	3
Anal incontinence			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Constipation			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	10 / 21 (47.62%)
occurrences (all)	0	0	13
Diarrhoea			
subjects affected / exposed	0 / 5 (0.00%)	1 / 3 (33.33%)	10 / 21 (47.62%)
occurrences (all)	0	1	18
Dry mouth			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Enterocolitis			
subjects affected / exposed	1 / 5 (20.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	1	0	0
Dysphagia			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	1 / 21 (4.76%)
occurrences (all)	0	0	1

Dyspepsia			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	1 / 21 (4.76%)
occurrences (all)	0	0	1
Eructation			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Flatulence			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	1 / 21 (4.76%)
occurrences (all)	0	0	1
Gastrooesophageal reflux disease			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	1 / 21 (4.76%)
occurrences (all)	0	0	1
Gingival bleeding			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Gingival pain			
subjects affected / exposed	2 / 5 (40.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	3	0	0
Haemorrhoidal haemorrhage			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Haemorrhoids			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Mouth ulceration			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	1 / 21 (4.76%)
occurrences (all)	0	0	1
Mouth haemorrhage			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Melaena			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Nausea			
subjects affected / exposed	3 / 5 (60.00%)	2 / 3 (66.67%)	16 / 21 (76.19%)
occurrences (all)	4	2	28

Proctalgia			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Oral disorder			
subjects affected / exposed	1 / 5 (20.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	1	0	0
Stomatitis			
subjects affected / exposed	1 / 5 (20.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	1	0	0
Vomiting			
subjects affected / exposed	2 / 5 (40.00%)	0 / 3 (0.00%)	9 / 21 (42.86%)
occurrences (all)	2	0	15
Hepatobiliary disorders			
Hyperbilirubinaemia			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Skin and subcutaneous tissue disorders			
Dermatitis acneiform			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	1 / 21 (4.76%)
occurrences (all)	0	0	1
Dry skin			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Erythema			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Hyperhidrosis			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	1 / 21 (4.76%)
occurrences (all)	0	0	1
Night sweats			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Petechiae			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	1 / 21 (4.76%)
occurrences (all)	0	0	1
Pruritus			

subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	3 / 21 (14.29%)
occurrences (all)	0	0	3
Skin discolouration			
subjects affected / exposed	0 / 5 (0.00%)	1 / 3 (33.33%)	0 / 21 (0.00%)
occurrences (all)	0	1	0
Rash maculo-papular			
subjects affected / exposed	0 / 5 (0.00%)	1 / 3 (33.33%)	0 / 21 (0.00%)
occurrences (all)	0	1	0
Rash macular			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	1 / 21 (4.76%)
occurrences (all)	0	0	1
Rash			
subjects affected / exposed	1 / 5 (20.00%)	0 / 3 (0.00%)	4 / 21 (19.05%)
occurrences (all)	1	0	5
Skin ulcer			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Skin lesion			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	4 / 21 (19.05%)
occurrences (all)	0	0	5
Urticaria			
subjects affected / exposed	0 / 5 (0.00%)	1 / 3 (33.33%)	0 / 21 (0.00%)
occurrences (all)	0	1	0
Skin induration			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Renal and urinary disorders			
Acute kidney injury			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Dysuria			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	1 / 21 (4.76%)
occurrences (all)	0	0	1
Haematuria			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	2 / 21 (9.52%)
occurrences (all)	0	0	2

Micturition urgency			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Nocturia			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Urinary incontinence			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Pollakiuria			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	1 / 21 (4.76%)
occurrences (all)	0	0	1
Musculoskeletal and connective tissue disorders			
Arthralgia			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	3 / 21 (14.29%)
occurrences (all)	0	0	6
Bone cyst			
subjects affected / exposed	1 / 5 (20.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	1	0	0
Back pain			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	3 / 21 (14.29%)
occurrences (all)	0	0	3
Bone pain			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	2 / 21 (9.52%)
occurrences (all)	0	0	2
Groin pain			
subjects affected / exposed	2 / 5 (40.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	3	0	0
Gouty arthritis			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Bursitis			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Muscle spasms			

subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	6 / 21 (28.57%)
occurrences (all)	0	0	9
Joint swelling			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	2 / 21 (9.52%)
occurrences (all)	0	0	2
Muscular weakness			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	1 / 21 (4.76%)
occurrences (all)	0	0	1
Musculoskeletal chest pain			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	1 / 21 (4.76%)
occurrences (all)	0	0	1
Musculoskeletal disorder			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Musculoskeletal pain			
subjects affected / exposed	0 / 5 (0.00%)	1 / 3 (33.33%)	0 / 21 (0.00%)
occurrences (all)	0	1	0
Myalgia			
subjects affected / exposed	1 / 5 (20.00%)	0 / 3 (0.00%)	3 / 21 (14.29%)
occurrences (all)	1	0	3
Neck pain			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Pain in extremity			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Pain in jaw			
subjects affected / exposed	1 / 5 (20.00%)	0 / 3 (0.00%)	1 / 21 (4.76%)
occurrences (all)	1	0	1
Rhabdomyolysis			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Infections and infestations			
Cellulitis			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	1 / 21 (4.76%)
occurrences (all)	0	0	1

COVID-19			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	4 / 21 (19.05%)
occurrences (all)	0	0	4
Bronchitis			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	1 / 21 (4.76%)
occurrences (all)	0	0	1
Cystitis			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	2 / 21 (9.52%)
occurrences (all)	0	0	4
Eye infection			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Folliculitis			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	2 / 21 (9.52%)
occurrences (all)	0	0	4
Fungal skin infection			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Gastroenteritis			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	4 / 21 (19.05%)
occurrences (all)	0	0	5
Herpes simplex reactivation			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Herpes zoster			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	1 / 21 (4.76%)
occurrences (all)	0	0	1
Hordeolum			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Influenza			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Localised infection			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0

Lower respiratory tract infection			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	1 / 21 (4.76%)
occurrences (all)	0	0	1
Orchitis			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Otitis media			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Pharyngitis			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	1 / 21 (4.76%)
occurrences (all)	0	0	1
Pneumonia			
subjects affected / exposed	1 / 5 (20.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	1	0	0
Nasopharyngitis			
subjects affected / exposed	0 / 5 (0.00%)	1 / 3 (33.33%)	2 / 21 (9.52%)
occurrences (all)	0	2	4
Oral herpes			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	1 / 21 (4.76%)
occurrences (all)	0	0	1
Skin infection			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Sinusitis			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Septic shock			
subjects affected / exposed	1 / 5 (20.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	1	0	0
Rhinitis			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	2 / 21 (9.52%)
occurrences (all)	0	0	2
Respiratory tract infection			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0

Staphylococcal infection			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Urinary tract infection bacterial			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Urinary tract infection			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	4 / 21 (19.05%)
occurrences (all)	0	0	6
Upper respiratory tract infection			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	2 / 21 (9.52%)
occurrences (all)	0	0	3
Tooth infection			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Metabolism and nutrition disorders			
Decreased appetite			
subjects affected / exposed	1 / 5 (20.00%)	0 / 3 (0.00%)	4 / 21 (19.05%)
occurrences (all)	2	0	4
Dehydration			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Folate deficiency			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Gout			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Hyperglycaemia			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	2 / 21 (9.52%)
occurrences (all)	0	0	2
Hypercholesterolaemia			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Hyperkalaemia			

subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	1 / 21 (4.76%)
occurrences (all)	0	0	1
Hyperphosphataemia			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Hypermagnesaemia			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Hypertriglyceridaemia			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Hyperuricaemia			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Hypocalcaemia			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Hypokalaemia			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Hypomagnesaemia			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Hyponatraemia			
subjects affected / exposed	1 / 5 (20.00%)	0 / 3 (0.00%)	1 / 21 (4.76%)
occurrences (all)	1	0	1
Hypophosphataemia			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	1 / 21 (4.76%)
occurrences (all)	0	0	2
Iron overload			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	0	0	0
Tumour lysis syndrome			
subjects affected / exposed	1 / 5 (20.00%)	0 / 3 (0.00%)	0 / 21 (0.00%)
occurrences (all)	1	0	0
Vitamin D deficiency			

subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	1 / 21 (4.76%)
occurrences (all)	0	0	1
Type 2 diabetes mellitus			
subjects affected / exposed	0 / 5 (0.00%)	0 / 3 (0.00%)	2 / 21 (9.52%)
occurrences (all)	0	0	2

Non-serious adverse events	Phase 2 Arm 3	Phase 2 Arm 2 2B	Phase 2 Arm 2 2A
Total subjects affected by non-serious adverse events			
subjects affected / exposed	84 / 84 (100.00%)	28 / 28 (100.00%)	58 / 59 (98.31%)
Neoplasms benign, malignant and unspecified (incl cysts and polyps)			
Squamous cell carcinoma			
subjects affected / exposed	2 / 84 (2.38%)	1 / 28 (3.57%)	3 / 59 (5.08%)
occurrences (all)	2	1	3
Basal cell carcinoma			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	3 / 59 (5.08%)
occurrences (all)	3	0	3
Vascular disorders			
Flushing			
subjects affected / exposed	0 / 84 (0.00%)	2 / 28 (7.14%)	0 / 59 (0.00%)
occurrences (all)	0	2	0
Haematoma			
subjects affected / exposed	4 / 84 (4.76%)	3 / 28 (10.71%)	1 / 59 (1.69%)
occurrences (all)	5	3	1
Hot flush			
subjects affected / exposed	1 / 84 (1.19%)	2 / 28 (7.14%)	1 / 59 (1.69%)
occurrences (all)	1	2	1
Hypotension			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	3 / 59 (5.08%)
occurrences (all)	1	0	3
Hypertension			
subjects affected / exposed	6 / 84 (7.14%)	0 / 28 (0.00%)	3 / 59 (5.08%)
occurrences (all)	10	0	8
General disorders and administration site conditions			
Chest pain			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences (all)	0	0	1

Asthenia			
subjects affected / exposed	7 / 84 (8.33%)	3 / 28 (10.71%)	8 / 59 (13.56%)
occurrences (all)	11	4	15
Early satiety			
subjects affected / exposed	5 / 84 (5.95%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences (all)	5	0	0
Chills			
subjects affected / exposed	10 / 84 (11.90%)	6 / 28 (21.43%)	2 / 59 (3.39%)
occurrences (all)	11	9	2
Fatigue			
subjects affected / exposed	27 / 84 (32.14%)	13 / 28 (46.43%)	13 / 59 (22.03%)
occurrences (all)	44	15	17
Gait disturbance			
subjects affected / exposed	1 / 84 (1.19%)	2 / 28 (7.14%)	0 / 59 (0.00%)
occurrences (all)	1	2	0
Influenza like illness			
subjects affected / exposed	4 / 84 (4.76%)	3 / 28 (10.71%)	0 / 59 (0.00%)
occurrences (all)	6	7	0
Malaise			
subjects affected / exposed	3 / 84 (3.57%)	2 / 28 (7.14%)	2 / 59 (3.39%)
occurrences (all)	3	2	2
Mucosal inflammation			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences (all)	2	0	0
Nodule			
subjects affected / exposed	1 / 84 (1.19%)	1 / 28 (3.57%)	0 / 59 (0.00%)
occurrences (all)	1	1	0
Non-cardiac chest pain			
subjects affected / exposed	5 / 84 (5.95%)	5 / 28 (17.86%)	1 / 59 (1.69%)
occurrences (all)	8	5	2
Oedema peripheral			
subjects affected / exposed	12 / 84 (14.29%)	6 / 28 (21.43%)	12 / 59 (20.34%)
occurrences (all)	12	7	15
Pain			
subjects affected / exposed	2 / 84 (2.38%)	0 / 28 (0.00%)	2 / 59 (3.39%)
occurrences (all)	3	0	2

Pyrexia			
subjects affected / exposed	14 / 84 (16.67%)	6 / 28 (21.43%)	8 / 59 (13.56%)
occurrences (all)	17	9	9
Peripheral swelling			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences (all)	1	0	0
Reproductive system and breast disorders			
Genital lesion			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences (all)	0	0	0
Vaginal haemorrhage			
subjects affected / exposed	1 / 84 (1.19%)	1 / 28 (3.57%)	0 / 59 (0.00%)
occurrences (all)	1	1	0
Respiratory, thoracic and mediastinal disorders			
Dyspnoea			
subjects affected / exposed	20 / 84 (23.81%)	7 / 28 (25.00%)	9 / 59 (15.25%)
occurrences (all)	21	7	11
Cough			
subjects affected / exposed	22 / 84 (26.19%)	14 / 28 (50.00%)	13 / 59 (22.03%)
occurrences (all)	25	21	19
Dyspnoea exertional			
subjects affected / exposed	5 / 84 (5.95%)	0 / 28 (0.00%)	2 / 59 (3.39%)
occurrences (all)	5	0	2
Nasal congestion			
subjects affected / exposed	3 / 84 (3.57%)	2 / 28 (7.14%)	2 / 59 (3.39%)
occurrences (all)	4	2	2
Hypoxia			
subjects affected / exposed	0 / 84 (0.00%)	1 / 28 (3.57%)	0 / 59 (0.00%)
occurrences (all)	0	1	0
Epistaxis			
subjects affected / exposed	13 / 84 (15.48%)	9 / 28 (32.14%)	10 / 59 (16.95%)
occurrences (all)	37	12	33
Oropharyngeal pain			
subjects affected / exposed	9 / 84 (10.71%)	5 / 28 (17.86%)	6 / 59 (10.17%)
occurrences (all)	12	6	8
Pleural effusion			

subjects affected / exposed	2 / 84 (2.38%)	0 / 28 (0.00%)	2 / 59 (3.39%)
occurrences (all)	2	0	2
Productive cough			
subjects affected / exposed	3 / 84 (3.57%)	1 / 28 (3.57%)	3 / 59 (5.08%)
occurrences (all)	3	1	4
Pulmonary oedema			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences (all)	0	0	1
Rhinorrhoea			
subjects affected / exposed	4 / 84 (4.76%)	2 / 28 (7.14%)	0 / 59 (0.00%)
occurrences (all)	4	2	0
Sinus pain			
subjects affected / exposed	0 / 84 (0.00%)	1 / 28 (3.57%)	0 / 59 (0.00%)
occurrences (all)	0	1	0
Upper-airway cough syndrome			
subjects affected / exposed	3 / 84 (3.57%)	1 / 28 (3.57%)	6 / 59 (10.17%)
occurrences (all)	3	1	6
Wheezing			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences (all)	1	0	1
Psychiatric disorders			
Anxiety			
subjects affected / exposed	4 / 84 (4.76%)	1 / 28 (3.57%)	3 / 59 (5.08%)
occurrences (all)	4	1	3
Confusional state			
subjects affected / exposed	2 / 84 (2.38%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences (all)	2	0	1
Delirium			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences (all)	1	0	1
Depression			
subjects affected / exposed	2 / 84 (2.38%)	3 / 28 (10.71%)	1 / 59 (1.69%)
occurrences (all)	2	3	2
Insomnia			
subjects affected / exposed	7 / 84 (8.33%)	3 / 28 (10.71%)	4 / 59 (6.78%)
occurrences (all)	7	3	5

Investigations			
Bilirubin conjugated increased			
subjects affected / exposed	1 / 84 (1.19%)	2 / 28 (7.14%)	2 / 59 (3.39%)
occurrences (all)	1	3	2
Aspartate aminotransferase increased			
subjects affected / exposed	6 / 84 (7.14%)	5 / 28 (17.86%)	5 / 59 (8.47%)
occurrences (all)	7	5	10
Alanine aminotransferase increased			
subjects affected / exposed	8 / 84 (9.52%)	6 / 28 (21.43%)	5 / 59 (8.47%)
occurrences (all)	10	6	10
Activated partial thromboplastin time prolonged			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences (all)	0	0	0
Blast cell count increased			
subjects affected / exposed	2 / 84 (2.38%)	2 / 28 (7.14%)	1 / 59 (1.69%)
occurrences (all)	2	2	1
Blood alkaline phosphatase increased			
subjects affected / exposed	5 / 84 (5.95%)	3 / 28 (10.71%)	0 / 59 (0.00%)
occurrences (all)	5	3	0
Blood bilirubin increased			
subjects affected / exposed	5 / 84 (5.95%)	4 / 28 (14.29%)	5 / 59 (8.47%)
occurrences (all)	8	4	5
Cardiac murmur			
subjects affected / exposed	0 / 84 (0.00%)	2 / 28 (7.14%)	0 / 59 (0.00%)
occurrences (all)	0	3	0
Blood phosphorus increased			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences (all)	1	0	0
Blood uric acid increased			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	2 / 59 (3.39%)
occurrences (all)	0	0	2
Blood creatinine increased			
subjects affected / exposed	11 / 84 (13.10%)	1 / 28 (3.57%)	5 / 59 (8.47%)
occurrences (all)	15	1	6
International normalised ratio increased			

subjects affected / exposed	2 / 84 (2.38%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences (all)	2	0	0
Ejection fraction decreased			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences (all)	0	0	1
Electrocardiogram QRS complex abnormal			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences (all)	0	0	0
Electrocardiogram QT prolonged			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences (all)	1	0	0
Haematocrit increased			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences (all)	0	0	0
Lymphocyte count decreased			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	2 / 59 (3.39%)
occurrences (all)	1	0	6
Neutrophil count decreased			
subjects affected / exposed	4 / 84 (4.76%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences (all)	12	0	1
Platelet count decreased			
subjects affected / exposed	22 / 84 (26.19%)	8 / 28 (28.57%)	11 / 59 (18.64%)
occurrences (all)	37	13	22
Transaminases increased			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences (all)	0	0	0
Troponin T increased			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences (all)	0	0	1
Weight decreased			
subjects affected / exposed	2 / 84 (2.38%)	2 / 28 (7.14%)	8 / 59 (13.56%)
occurrences (all)	2	2	8
Weight increased			
subjects affected / exposed	4 / 84 (4.76%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences (all)	4	0	0

White blood cell count decreased			
subjects affected / exposed	4 / 84 (4.76%)	0 / 28 (0.00%)	5 / 59 (8.47%)
occurrences (all)	10	0	5
Injury, poisoning and procedural complications			
Contusion			
subjects affected / exposed	16 / 84 (19.05%)	8 / 28 (28.57%)	9 / 59 (15.25%)
occurrences (all)	21	11	11
Fall			
subjects affected / exposed	9 / 84 (10.71%)	7 / 28 (25.00%)	4 / 59 (6.78%)
occurrences (all)	15	10	4
Vascular access site complication			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences (all)	0	0	0
Skin laceration			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences (all)	1	0	2
Procedural site reaction			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences (all)	0	0	0
Procedural pain			
subjects affected / exposed	1 / 84 (1.19%)	2 / 28 (7.14%)	1 / 59 (1.69%)
occurrences (all)	1	2	1
Nail injury			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences (all)	0	0	0
Wound			
subjects affected / exposed	1 / 84 (1.19%)	1 / 28 (3.57%)	2 / 59 (3.39%)
occurrences (all)	1	1	2
Cardiac disorders			
Angina pectoris			
subjects affected / exposed	1 / 84 (1.19%)	1 / 28 (3.57%)	0 / 59 (0.00%)
occurrences (all)	1	1	0
Atrial fibrillation			
subjects affected / exposed	3 / 84 (3.57%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences (all)	3	0	0
Atrioventricular block			

subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences (all)	0	0	0
Hyperdynamic left ventricle			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences (all)	0	0	0
Palpitations			
subjects affected / exposed	3 / 84 (3.57%)	1 / 28 (3.57%)	2 / 59 (3.39%)
occurrences (all)	9	2	2
Sinus tachycardia			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences (all)	0	0	1
Tachycardia			
subjects affected / exposed	2 / 84 (2.38%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences (all)	2	0	0
Nervous system disorders			
Ageusia			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences (all)	0	0	1
Dysarthria			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences (all)	1	0	2
Dizziness			
subjects affected / exposed	21 / 84 (25.00%)	5 / 28 (17.86%)	13 / 59 (22.03%)
occurrences (all)	30	6	16
Cerebrovascular accident			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences (all)	0	0	0
Memory impairment			
subjects affected / exposed	0 / 84 (0.00%)	2 / 28 (7.14%)	0 / 59 (0.00%)
occurrences (all)	0	2	0
Lethargy			
subjects affected / exposed	1 / 84 (1.19%)	1 / 28 (3.57%)	1 / 59 (1.69%)
occurrences (all)	2	2	1
Hypoaesthesia			
subjects affected / exposed	2 / 84 (2.38%)	2 / 28 (7.14%)	0 / 59 (0.00%)
occurrences (all)	5	2	0

Headache			
subjects affected / exposed	19 / 84 (22.62%)	3 / 28 (10.71%)	6 / 59 (10.17%)
occurrences (all)	24	6	6
Dysgeusia			
subjects affected / exposed	21 / 84 (25.00%)	7 / 28 (25.00%)	16 / 59 (27.12%)
occurrences (all)	23	7	23
Presyncope			
subjects affected / exposed	4 / 84 (4.76%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences (all)	4	0	0
Paraesthesia			
subjects affected / exposed	4 / 84 (4.76%)	4 / 28 (14.29%)	5 / 59 (8.47%)
occurrences (all)	7	4	7
Neuropathy peripheral			
subjects affected / exposed	2 / 84 (2.38%)	1 / 28 (3.57%)	1 / 59 (1.69%)
occurrences (all)	2	1	1
Myoclonus			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences (all)	0	0	0
Sciatica			
subjects affected / exposed	4 / 84 (4.76%)	1 / 28 (3.57%)	3 / 59 (5.08%)
occurrences (all)	6	1	7
Migraine			
subjects affected / exposed	4 / 84 (4.76%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences (all)	4	0	0
Taste disorder			
subjects affected / exposed	2 / 84 (2.38%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences (all)	2	0	0
Syncope			
subjects affected / exposed	3 / 84 (3.57%)	1 / 28 (3.57%)	0 / 59 (0.00%)
occurrences (all)	3	2	0
Blood and lymphatic system disorders			
Anaemia			
subjects affected / exposed	41 / 84 (48.81%)	7 / 28 (25.00%)	18 / 59 (30.51%)
occurrences (all)	76	16	23
Leukocytosis			

subjects affected / exposed	4 / 84 (4.76%)	2 / 28 (7.14%)	1 / 59 (1.69%)
occurrences (all)	4	2	1
Lymphadenopathy			
subjects affected / exposed	2 / 84 (2.38%)	0 / 28 (0.00%)	2 / 59 (3.39%)
occurrences (all)	3	0	2
Neutropenia			
subjects affected / exposed	2 / 84 (2.38%)	1 / 28 (3.57%)	10 / 59 (16.95%)
occurrences (all)	3	2	17
Spleen disorder			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	2 / 59 (3.39%)
occurrences (all)	0	0	2
Thrombocytosis			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences (all)	0	0	0
Thrombocytopenia			
subjects affected / exposed	32 / 84 (38.10%)	9 / 28 (32.14%)	25 / 59 (42.37%)
occurrences (all)	53	21	55
Ear and labyrinth disorders			
Ear discomfort			
subjects affected / exposed	2 / 84 (2.38%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences (all)	2	0	0
Ear pain			
subjects affected / exposed	2 / 84 (2.38%)	1 / 28 (3.57%)	1 / 59 (1.69%)
occurrences (all)	2	1	1
Ear haemorrhage			
subjects affected / exposed	0 / 84 (0.00%)	1 / 28 (3.57%)	0 / 59 (0.00%)
occurrences (all)	0	1	0
Hypoacusis			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences (all)	1	0	0
Tinnitus			
subjects affected / exposed	6 / 84 (7.14%)	0 / 28 (0.00%)	2 / 59 (3.39%)
occurrences (all)	8	0	2
Eye disorders			
Eye haemorrhage			

subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences (all)	0	0	1
Vision blurred			
subjects affected / exposed	3 / 84 (3.57%)	1 / 28 (3.57%)	2 / 59 (3.39%)
occurrences (all)	3	1	3
Gastrointestinal disorders			
Abdominal discomfort			
subjects affected / exposed	4 / 84 (4.76%)	1 / 28 (3.57%)	5 / 59 (8.47%)
occurrences (all)	4	1	6
Abdominal distension			
subjects affected / exposed	13 / 84 (15.48%)	8 / 28 (28.57%)	6 / 59 (10.17%)
occurrences (all)	14	12	7
Abdominal pain			
subjects affected / exposed	13 / 84 (15.48%)	7 / 28 (25.00%)	4 / 59 (6.78%)
occurrences (all)	17	9	6
Abdominal pain upper			
subjects affected / exposed	11 / 84 (13.10%)	4 / 28 (14.29%)	4 / 59 (6.78%)
occurrences (all)	18	5	5
Anal incontinence			
subjects affected / exposed	0 / 84 (0.00%)	1 / 28 (3.57%)	0 / 59 (0.00%)
occurrences (all)	0	1	0
Constipation			
subjects affected / exposed	27 / 84 (32.14%)	6 / 28 (21.43%)	9 / 59 (15.25%)
occurrences (all)	29	7	11
Diarrhoea			
subjects affected / exposed	40 / 84 (47.62%)	18 / 28 (64.29%)	34 / 59 (57.63%)
occurrences (all)	71	34	65
Dry mouth			
subjects affected / exposed	3 / 84 (3.57%)	1 / 28 (3.57%)	0 / 59 (0.00%)
occurrences (all)	3	1	0
Enterocolitis			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences (all)	0	0	0
Dysphagia			
subjects affected / exposed	1 / 84 (1.19%)	1 / 28 (3.57%)	1 / 59 (1.69%)
occurrences (all)	1	1	1

Dyspepsia			
subjects affected / exposed	4 / 84 (4.76%)	1 / 28 (3.57%)	2 / 59 (3.39%)
occurrences (all)	5	1	3
Eructation			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences (all)	0	0	0
Flatulence			
subjects affected / exposed	7 / 84 (8.33%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences (all)	7	0	1
Gastrooesophageal reflux disease			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences (all)	1	0	0
Gingival bleeding			
subjects affected / exposed	2 / 84 (2.38%)	0 / 28 (0.00%)	3 / 59 (5.08%)
occurrences (all)	2	0	4
Gingival pain			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences (all)	1	0	0
Haemorrhoidal haemorrhage			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences (all)	2	0	0
Haemorrhoids			
subjects affected / exposed	5 / 84 (5.95%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences (all)	6	0	0
Mouth ulceration			
subjects affected / exposed	5 / 84 (5.95%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences (all)	6	0	1
Mouth haemorrhage			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences (all)	1	0	1
Melaena			
subjects affected / exposed	1 / 84 (1.19%)	1 / 28 (3.57%)	1 / 59 (1.69%)
occurrences (all)	1	1	1
Nausea			
subjects affected / exposed	26 / 84 (30.95%)	12 / 28 (42.86%)	21 / 59 (35.59%)
occurrences (all)	39	18	31

Proctalgia			
subjects affected / exposed	0 / 84 (0.00%)	1 / 28 (3.57%)	0 / 59 (0.00%)
occurrences (all)	0	1	0
Oral disorder			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences (all)	1	0	0
Stomatitis			
subjects affected / exposed	3 / 84 (3.57%)	3 / 28 (10.71%)	5 / 59 (8.47%)
occurrences (all)	3	3	5
Vomiting			
subjects affected / exposed	15 / 84 (17.86%)	8 / 28 (28.57%)	10 / 59 (16.95%)
occurrences (all)	18	9	14
Hepatobiliary disorders			
Hyperbilirubinaemia			
subjects affected / exposed	2 / 84 (2.38%)	0 / 28 (0.00%)	2 / 59 (3.39%)
occurrences (all)	6	0	4
Skin and subcutaneous tissue disorders			
Dermatitis acneiform			
subjects affected / exposed	3 / 84 (3.57%)	0 / 28 (0.00%)	3 / 59 (5.08%)
occurrences (all)	3	0	5
Dry skin			
subjects affected / exposed	4 / 84 (4.76%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences (all)	4	0	1
Erythema			
subjects affected / exposed	1 / 84 (1.19%)	1 / 28 (3.57%)	0 / 59 (0.00%)
occurrences (all)	1	1	0
Hyperhidrosis			
subjects affected / exposed	2 / 84 (2.38%)	0 / 28 (0.00%)	2 / 59 (3.39%)
occurrences (all)	3	0	2
Night sweats			
subjects affected / exposed	12 / 84 (14.29%)	2 / 28 (7.14%)	6 / 59 (10.17%)
occurrences (all)	18	3	8
Petechiae			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences (all)	1	0	1
Pruritus			

subjects affected / exposed	18 / 84 (21.43%)	7 / 28 (25.00%)	2 / 59 (3.39%)
occurrences (all)	21	9	2
Skin discolouration			
subjects affected / exposed	0 / 84 (0.00%)	1 / 28 (3.57%)	0 / 59 (0.00%)
occurrences (all)	0	1	0
Rash maculo-papular			
subjects affected / exposed	4 / 84 (4.76%)	1 / 28 (3.57%)	0 / 59 (0.00%)
occurrences (all)	7	2	0
Rash macular			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences (all)	0	0	0
Rash			
subjects affected / exposed	6 / 84 (7.14%)	4 / 28 (14.29%)	3 / 59 (5.08%)
occurrences (all)	8	5	3
Skin ulcer			
subjects affected / exposed	2 / 84 (2.38%)	0 / 28 (0.00%)	2 / 59 (3.39%)
occurrences (all)	2	0	2
Skin lesion			
subjects affected / exposed	5 / 84 (5.95%)	2 / 28 (7.14%)	2 / 59 (3.39%)
occurrences (all)	5	2	2
Urticaria			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences (all)	0	0	0
Skin induration			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences (all)	0	0	0
Renal and urinary disorders			
Acute kidney injury			
subjects affected / exposed	0 / 84 (0.00%)	1 / 28 (3.57%)	1 / 59 (1.69%)
occurrences (all)	0	1	1
Dysuria			
subjects affected / exposed	2 / 84 (2.38%)	0 / 28 (0.00%)	5 / 59 (8.47%)
occurrences (all)	3	0	6
Haematuria			
subjects affected / exposed	3 / 84 (3.57%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences (all)	3	0	2

Micturition urgency			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences (all)	0	0	1
Nocturia			
subjects affected / exposed	2 / 84 (2.38%)	2 / 28 (7.14%)	0 / 59 (0.00%)
occurrences (all)	2	2	0
Urinary incontinence			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences (all)	0	0	0
Pollakiuria			
subjects affected / exposed	4 / 84 (4.76%)	2 / 28 (7.14%)	1 / 59 (1.69%)
occurrences (all)	4	2	1
Musculoskeletal and connective tissue disorders			
Arthralgia			
subjects affected / exposed	21 / 84 (25.00%)	7 / 28 (25.00%)	11 / 59 (18.64%)
occurrences (all)	26	8	12
Bone cyst			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences (all)	0	0	0
Back pain			
subjects affected / exposed	17 / 84 (20.24%)	4 / 28 (14.29%)	8 / 59 (13.56%)
occurrences (all)	23	4	9
Bone pain			
subjects affected / exposed	8 / 84 (9.52%)	2 / 28 (7.14%)	8 / 59 (13.56%)
occurrences (all)	10	2	10
Groin pain			
subjects affected / exposed	2 / 84 (2.38%)	1 / 28 (3.57%)	1 / 59 (1.69%)
occurrences (all)	2	1	1
Gouty arthritis			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences (all)	0	0	0
Bursitis			
subjects affected / exposed	1 / 84 (1.19%)	1 / 28 (3.57%)	0 / 59 (0.00%)
occurrences (all)	1	1	0
Muscle spasms			

subjects affected / exposed	18 / 84 (21.43%)	10 / 28 (35.71%)	7 / 59 (11.86%)
occurrences (all)	24	11	7
Joint swelling			
subjects affected / exposed	1 / 84 (1.19%)	2 / 28 (7.14%)	2 / 59 (3.39%)
occurrences (all)	1	2	3
Muscular weakness			
subjects affected / exposed	5 / 84 (5.95%)	1 / 28 (3.57%)	7 / 59 (11.86%)
occurrences (all)	7	1	9
Musculoskeletal chest pain			
subjects affected / exposed	6 / 84 (7.14%)	0 / 28 (0.00%)	3 / 59 (5.08%)
occurrences (all)	7	0	3
Musculoskeletal disorder			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences (all)	0	0	0
Musculoskeletal pain			
subjects affected / exposed	4 / 84 (4.76%)	0 / 28 (0.00%)	2 / 59 (3.39%)
occurrences (all)	4	0	2
Myalgia			
subjects affected / exposed	10 / 84 (11.90%)	3 / 28 (10.71%)	5 / 59 (8.47%)
occurrences (all)	12	5	7
Neck pain			
subjects affected / exposed	6 / 84 (7.14%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences (all)	6	0	1
Pain in extremity			
subjects affected / exposed	15 / 84 (17.86%)	9 / 28 (32.14%)	8 / 59 (13.56%)
occurrences (all)	20	10	11
Pain in jaw			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences (all)	1	0	0
Rhabdomyolysis			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences (all)	0	0	0
Infections and infestations			
Cellulitis			
subjects affected / exposed	2 / 84 (2.38%)	2 / 28 (7.14%)	2 / 59 (3.39%)
occurrences (all)	3	2	2

COVID-19			
subjects affected / exposed	24 / 84 (28.57%)	3 / 28 (10.71%)	9 / 59 (15.25%)
occurrences (all)	29	3	10
Bronchitis			
subjects affected / exposed	3 / 84 (3.57%)	3 / 28 (10.71%)	0 / 59 (0.00%)
occurrences (all)	3	3	0
Cystitis			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	2 / 59 (3.39%)
occurrences (all)	1	0	2
Eye infection			
subjects affected / exposed	2 / 84 (2.38%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences (all)	2	0	0
Folliculitis			
subjects affected / exposed	2 / 84 (2.38%)	0 / 28 (0.00%)	2 / 59 (3.39%)
occurrences (all)	3	0	4
Fungal skin infection			
subjects affected / exposed	0 / 84 (0.00%)	1 / 28 (3.57%)	0 / 59 (0.00%)
occurrences (all)	0	1	0
Gastroenteritis			
subjects affected / exposed	2 / 84 (2.38%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences (all)	2	0	1
Herpes simplex reactivation			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences (all)	1	0	0
Herpes zoster			
subjects affected / exposed	14 / 84 (16.67%)	4 / 28 (14.29%)	4 / 59 (6.78%)
occurrences (all)	15	4	6
Hordeolum			
subjects affected / exposed	4 / 84 (4.76%)	0 / 28 (0.00%)	2 / 59 (3.39%)
occurrences (all)	8	0	2
Influenza			
subjects affected / exposed	5 / 84 (5.95%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences (all)	7	0	0
Localised infection			
subjects affected / exposed	1 / 84 (1.19%)	0 / 28 (0.00%)	3 / 59 (5.08%)
occurrences (all)	1	0	3

Lower respiratory tract infection			
subjects affected / exposed	2 / 84 (2.38%)	3 / 28 (10.71%)	2 / 59 (3.39%)
occurrences (all)	2	5	3
Orchitis			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences (all)	0	0	0
Otitis media			
subjects affected / exposed	0 / 84 (0.00%)	1 / 28 (3.57%)	2 / 59 (3.39%)
occurrences (all)	0	1	2
Pharyngitis			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences (all)	0	0	0
Pneumonia			
subjects affected / exposed	8 / 84 (9.52%)	3 / 28 (10.71%)	5 / 59 (8.47%)
occurrences (all)	8	5	6
Nasopharyngitis			
subjects affected / exposed	10 / 84 (11.90%)	2 / 28 (7.14%)	0 / 59 (0.00%)
occurrences (all)	17	2	0
Oral herpes			
subjects affected / exposed	2 / 84 (2.38%)	4 / 28 (14.29%)	5 / 59 (8.47%)
occurrences (all)	3	8	6
Skin infection			
subjects affected / exposed	2 / 84 (2.38%)	2 / 28 (7.14%)	0 / 59 (0.00%)
occurrences (all)	2	2	0
Sinusitis			
subjects affected / exposed	4 / 84 (4.76%)	1 / 28 (3.57%)	2 / 59 (3.39%)
occurrences (all)	7	1	4
Septic shock			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences (all)	0	0	0
Rhinitis			
subjects affected / exposed	3 / 84 (3.57%)	1 / 28 (3.57%)	1 / 59 (1.69%)
occurrences (all)	3	1	1
Respiratory tract infection			
subjects affected / exposed	5 / 84 (5.95%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences (all)	6	0	1

Staphylococcal infection			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences (all)	0	0	0
Urinary tract infection bacterial			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences (all)	0	0	0
Urinary tract infection			
subjects affected / exposed	14 / 84 (16.67%)	3 / 28 (10.71%)	6 / 59 (10.17%)
occurrences (all)	26	11	10
Upper respiratory tract infection			
subjects affected / exposed	14 / 84 (16.67%)	8 / 28 (28.57%)	11 / 59 (18.64%)
occurrences (all)	17	14	19
Tooth infection			
subjects affected / exposed	4 / 84 (4.76%)	2 / 28 (7.14%)	0 / 59 (0.00%)
occurrences (all)	5	2	0
Metabolism and nutrition disorders			
Decreased appetite			
subjects affected / exposed	15 / 84 (17.86%)	7 / 28 (25.00%)	13 / 59 (22.03%)
occurrences (all)	20	7	13
Dehydration			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	3 / 59 (5.08%)
occurrences (all)	0	0	3
Folate deficiency			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences (all)	0	0	0
Gout			
subjects affected / exposed	2 / 84 (2.38%)	1 / 28 (3.57%)	2 / 59 (3.39%)
occurrences (all)	2	1	2
Hyperglycaemia			
subjects affected / exposed	1 / 84 (1.19%)	2 / 28 (7.14%)	4 / 59 (6.78%)
occurrences (all)	1	2	5
Hypercholesterolaemia			
subjects affected / exposed	3 / 84 (3.57%)	2 / 28 (7.14%)	0 / 59 (0.00%)
occurrences (all)	3	2	0
Hyperkalaemia			

subjects affected / exposed	6 / 84 (7.14%)	2 / 28 (7.14%)	4 / 59 (6.78%)
occurrences (all)	9	9	7
Hyperphosphataemia			
subjects affected / exposed	2 / 84 (2.38%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences (all)	7	0	1
Hypermagnesaemia			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences (all)	0	0	0
Hypertriglyceridaemia			
subjects affected / exposed	5 / 84 (5.95%)	0 / 28 (0.00%)	1 / 59 (1.69%)
occurrences (all)	12	0	1
Hyperuricaemia			
subjects affected / exposed	5 / 84 (5.95%)	4 / 28 (14.29%)	5 / 59 (8.47%)
occurrences (all)	6	4	5
Hypocalcaemia			
subjects affected / exposed	1 / 84 (1.19%)	1 / 28 (3.57%)	3 / 59 (5.08%)
occurrences (all)	1	1	6
Hypokalaemia			
subjects affected / exposed	4 / 84 (4.76%)	1 / 28 (3.57%)	4 / 59 (6.78%)
occurrences (all)	4	1	4
Hypomagnesaemia			
subjects affected / exposed	1 / 84 (1.19%)	1 / 28 (3.57%)	2 / 59 (3.39%)
occurrences (all)	1	2	4
Hyponatraemia			
subjects affected / exposed	11 / 84 (13.10%)	2 / 28 (7.14%)	3 / 59 (5.08%)
occurrences (all)	13	2	3
Hypophosphataemia			
subjects affected / exposed	1 / 84 (1.19%)	1 / 28 (3.57%)	0 / 59 (0.00%)
occurrences (all)	1	3	0
Iron overload			
subjects affected / exposed	2 / 84 (2.38%)	0 / 28 (0.00%)	4 / 59 (6.78%)
occurrences (all)	2	0	4
Tumour lysis syndrome			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences (all)	0	0	0
Vitamin D deficiency			

subjects affected / exposed	1 / 84 (1.19%)	1 / 28 (3.57%)	0 / 59 (0.00%)
occurrences (all)	1	1	0
Type 2 diabetes mellitus			
subjects affected / exposed	0 / 84 (0.00%)	0 / 28 (0.00%)	0 / 59 (0.00%)
occurrences (all)	0	0	0

Non-serious adverse events	Phase 2 Arm 1 1B	Phase 2 Arm 1 1A	Phase 1 overall
Total subjects affected by non-serious adverse events			
subjects affected / exposed	50 / 52 (96.15%)	48 / 48 (100.00%)	43 / 44 (97.73%)
Neoplasms benign, malignant and unspecified (incl cysts and polyps)			
Squamous cell carcinoma			
subjects affected / exposed	1 / 52 (1.92%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences (all)	1	0	0
Basal cell carcinoma			
subjects affected / exposed	1 / 52 (1.92%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences (all)	3	0	0
Vascular disorders			
Flushing			
subjects affected / exposed	1 / 52 (1.92%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences (all)	1	0	0
Haematoma			
subjects affected / exposed	3 / 52 (5.77%)	0 / 48 (0.00%)	2 / 44 (4.55%)
occurrences (all)	3	0	2
Hot flush			
subjects affected / exposed	1 / 52 (1.92%)	0 / 48 (0.00%)	2 / 44 (4.55%)
occurrences (all)	1	0	2
Hypotension			
subjects affected / exposed	3 / 52 (5.77%)	0 / 48 (0.00%)	4 / 44 (9.09%)
occurrences (all)	3	0	4
Hypertension			
subjects affected / exposed	4 / 52 (7.69%)	1 / 48 (2.08%)	4 / 44 (9.09%)
occurrences (all)	5	1	4
General disorders and administration site conditions			
Chest pain			
subjects affected / exposed	1 / 52 (1.92%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	1	0	1

Asthenia			
subjects affected / exposed	3 / 52 (5.77%)	9 / 48 (18.75%)	4 / 44 (9.09%)
occurrences (all)	3	9	4
Early satiety			
subjects affected / exposed	4 / 52 (7.69%)	2 / 48 (4.17%)	0 / 44 (0.00%)
occurrences (all)	5	3	0
Chills			
subjects affected / exposed	2 / 52 (3.85%)	1 / 48 (2.08%)	2 / 44 (4.55%)
occurrences (all)	2	1	2
Fatigue			
subjects affected / exposed	17 / 52 (32.69%)	14 / 48 (29.17%)	20 / 44 (45.45%)
occurrences (all)	19	18	23
Gait disturbance			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences (all)	0	0	0
Influenza like illness			
subjects affected / exposed	2 / 52 (3.85%)	3 / 48 (6.25%)	0 / 44 (0.00%)
occurrences (all)	2	5	0
Malaise			
subjects affected / exposed	2 / 52 (3.85%)	1 / 48 (2.08%)	1 / 44 (2.27%)
occurrences (all)	2	1	1
Mucosal inflammation			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	2 / 44 (4.55%)
occurrences (all)	0	0	2
Nodule			
subjects affected / exposed	1 / 52 (1.92%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	1	0	1
Non-cardiac chest pain			
subjects affected / exposed	1 / 52 (1.92%)	2 / 48 (4.17%)	1 / 44 (2.27%)
occurrences (all)	1	3	1
Oedema peripheral			
subjects affected / exposed	8 / 52 (15.38%)	13 / 48 (27.08%)	9 / 44 (20.45%)
occurrences (all)	11	13	9
Pain			
subjects affected / exposed	0 / 52 (0.00%)	1 / 48 (2.08%)	1 / 44 (2.27%)
occurrences (all)	0	1	1

Pyrexia			
subjects affected / exposed	6 / 52 (11.54%)	8 / 48 (16.67%)	5 / 44 (11.36%)
occurrences (all)	8	10	5
Peripheral swelling			
subjects affected / exposed	0 / 52 (0.00%)	3 / 48 (6.25%)	0 / 44 (0.00%)
occurrences (all)	0	4	0
Reproductive system and breast disorders			
Genital lesion			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	0	0	1
Vaginal haemorrhage			
subjects affected / exposed	1 / 52 (1.92%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	1	0	1
Respiratory, thoracic and mediastinal disorders			
Dyspnoea			
subjects affected / exposed	10 / 52 (19.23%)	11 / 48 (22.92%)	4 / 44 (9.09%)
occurrences (all)	12	14	4
Cough			
subjects affected / exposed	11 / 52 (21.15%)	9 / 48 (18.75%)	6 / 44 (13.64%)
occurrences (all)	17	10	6
Dyspnoea exertional			
subjects affected / exposed	0 / 52 (0.00%)	1 / 48 (2.08%)	0 / 44 (0.00%)
occurrences (all)	0	1	0
Nasal congestion			
subjects affected / exposed	1 / 52 (1.92%)	1 / 48 (2.08%)	2 / 44 (4.55%)
occurrences (all)	1	1	2
Hypoxia			
subjects affected / exposed	0 / 52 (0.00%)	2 / 48 (4.17%)	2 / 44 (4.55%)
occurrences (all)	0	2	2
Epistaxis			
subjects affected / exposed	7 / 52 (13.46%)	6 / 48 (12.50%)	6 / 44 (13.64%)
occurrences (all)	17	7	10
Oropharyngeal pain			
subjects affected / exposed	2 / 52 (3.85%)	2 / 48 (4.17%)	3 / 44 (6.82%)
occurrences (all)	2	2	3
Pleural effusion			

subjects affected / exposed	2 / 52 (3.85%)	1 / 48 (2.08%)	1 / 44 (2.27%)
occurrences (all)	2	1	1
Productive cough			
subjects affected / exposed	2 / 52 (3.85%)	3 / 48 (6.25%)	0 / 44 (0.00%)
occurrences (all)	2	3	0
Pulmonary oedema			
subjects affected / exposed	1 / 52 (1.92%)	1 / 48 (2.08%)	1 / 44 (2.27%)
occurrences (all)	1	1	1
Rhinorrhoea			
subjects affected / exposed	2 / 52 (3.85%)	2 / 48 (4.17%)	0 / 44 (0.00%)
occurrences (all)	2	2	0
Sinus pain			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	0	0	1
Upper-airway cough syndrome			
subjects affected / exposed	3 / 52 (5.77%)	1 / 48 (2.08%)	1 / 44 (2.27%)
occurrences (all)	3	1	1
Wheezing			
subjects affected / exposed	1 / 52 (1.92%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	1	0	1
Psychiatric disorders			
Anxiety			
subjects affected / exposed	1 / 52 (1.92%)	0 / 48 (0.00%)	3 / 44 (6.82%)
occurrences (all)	1	0	3
Confusional state			
subjects affected / exposed	1 / 52 (1.92%)	2 / 48 (4.17%)	1 / 44 (2.27%)
occurrences (all)	1	2	1
Delirium			
subjects affected / exposed	1 / 52 (1.92%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	1	0	1
Depression			
subjects affected / exposed	0 / 52 (0.00%)	2 / 48 (4.17%)	1 / 44 (2.27%)
occurrences (all)	0	2	1
Insomnia			
subjects affected / exposed	4 / 52 (7.69%)	2 / 48 (4.17%)	2 / 44 (4.55%)
occurrences (all)	4	2	2

Investigations			
Bilirubin conjugated increased			
subjects affected / exposed	0 / 52 (0.00%)	1 / 48 (2.08%)	0 / 44 (0.00%)
occurrences (all)	0	1	0
Aspartate aminotransferase increased			
subjects affected / exposed	4 / 52 (7.69%)	3 / 48 (6.25%)	3 / 44 (6.82%)
occurrences (all)	4	3	4
Alanine aminotransferase increased			
subjects affected / exposed	4 / 52 (7.69%)	1 / 48 (2.08%)	5 / 44 (11.36%)
occurrences (all)	4	1	5
Activated partial thromboplastin time prolonged			
subjects affected / exposed	2 / 52 (3.85%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	2	0	1
Blast cell count increased			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences (all)	0	0	0
Blood alkaline phosphatase increased			
subjects affected / exposed	2 / 52 (3.85%)	3 / 48 (6.25%)	2 / 44 (4.55%)
occurrences (all)	2	4	3
Blood bilirubin increased			
subjects affected / exposed	1 / 52 (1.92%)	3 / 48 (6.25%)	2 / 44 (4.55%)
occurrences (all)	1	3	2
Cardiac murmur			
subjects affected / exposed	2 / 52 (3.85%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences (all)	2	0	0
Blood phosphorus increased			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	0	0	1
Blood uric acid increased			
subjects affected / exposed	0 / 52 (0.00%)	1 / 48 (2.08%)	1 / 44 (2.27%)
occurrences (all)	0	1	1
Blood creatinine increased			
subjects affected / exposed	4 / 52 (7.69%)	3 / 48 (6.25%)	4 / 44 (9.09%)
occurrences (all)	5	3	5
International normalised ratio increased			

subjects affected / exposed	2 / 52 (3.85%)	1 / 48 (2.08%)	1 / 44 (2.27%)
occurrences (all)	2	1	1
Ejection fraction decreased			
subjects affected / exposed	1 / 52 (1.92%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	1	0	1
Electrocardiogram QRS complex abnormal			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	0	0	1
Electrocardiogram QT prolonged			
subjects affected / exposed	1 / 52 (1.92%)	0 / 48 (0.00%)	2 / 44 (4.55%)
occurrences (all)	1	0	2
Haematocrit increased			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences (all)	0	0	0
Lymphocyte count decreased			
subjects affected / exposed	1 / 52 (1.92%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	2	0	4
Neutrophil count decreased			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	4 / 44 (9.09%)
occurrences (all)	0	0	8
Platelet count decreased			
subjects affected / exposed	8 / 52 (15.38%)	5 / 48 (10.42%)	8 / 44 (18.18%)
occurrences (all)	12	8	9
Transaminases increased			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	0	0	1
Troponin T increased			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	0	0	1
Weight decreased			
subjects affected / exposed	14 / 52 (26.92%)	9 / 48 (18.75%)	5 / 44 (11.36%)
occurrences (all)	16	9	7
Weight increased			
subjects affected / exposed	4 / 52 (7.69%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	5	0	1

White blood cell count decreased			
subjects affected / exposed	0 / 52 (0.00%)	1 / 48 (2.08%)	2 / 44 (4.55%)
occurrences (all)	0	2	7
Injury, poisoning and procedural complications			
Contusion			
subjects affected / exposed	9 / 52 (17.31%)	4 / 48 (8.33%)	1 / 44 (2.27%)
occurrences (all)	11	4	1
Fall			
subjects affected / exposed	10 / 52 (19.23%)	3 / 48 (6.25%)	1 / 44 (2.27%)
occurrences (all)	12	3	1
Vascular access site complication			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	0	0	1
Skin laceration			
subjects affected / exposed	1 / 52 (1.92%)	2 / 48 (4.17%)	1 / 44 (2.27%)
occurrences (all)	1	2	1
Procedural site reaction			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	0	0	2
Procedural pain			
subjects affected / exposed	1 / 52 (1.92%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	1	0	1
Nail injury			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences (all)	0	0	0
Wound			
subjects affected / exposed	3 / 52 (5.77%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences (all)	3	0	0
Cardiac disorders			
Angina pectoris			
subjects affected / exposed	0 / 52 (0.00%)	1 / 48 (2.08%)	1 / 44 (2.27%)
occurrences (all)	0	1	1
Atrial fibrillation			
subjects affected / exposed	1 / 52 (1.92%)	3 / 48 (6.25%)	3 / 44 (6.82%)
occurrences (all)	1	4	3
Atrioventricular block			

subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	0	0	1
Hyperdynamic left ventricle			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	0	0	1
Palpitations			
subjects affected / exposed	3 / 52 (5.77%)	2 / 48 (4.17%)	1 / 44 (2.27%)
occurrences (all)	3	2	1
Sinus tachycardia			
subjects affected / exposed	2 / 52 (3.85%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	3	0	1
Tachycardia			
subjects affected / exposed	3 / 52 (5.77%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	3	0	1
Nervous system disorders			
Ageusia			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences (all)	0	0	0
Dysarthria			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	0	0	1
Dizziness			
subjects affected / exposed	10 / 52 (19.23%)	7 / 48 (14.58%)	3 / 44 (6.82%)
occurrences (all)	14	8	3
Cerebrovascular accident			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	0	0	1
Memory impairment			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences (all)	0	0	0
Lethargy			
subjects affected / exposed	0 / 52 (0.00%)	1 / 48 (2.08%)	1 / 44 (2.27%)
occurrences (all)	0	1	1
Hypoaesthesia			
subjects affected / exposed	2 / 52 (3.85%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences (all)	2	0	0

Headache			
subjects affected / exposed	14 / 52 (26.92%)	6 / 48 (12.50%)	8 / 44 (18.18%)
occurrences (all)	25	7	8
Dysgeusia			
subjects affected / exposed	16 / 52 (30.77%)	10 / 48 (20.83%)	9 / 44 (20.45%)
occurrences (all)	18	12	9
Presyncope			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	0	0	1
Paraesthesia			
subjects affected / exposed	1 / 52 (1.92%)	2 / 48 (4.17%)	0 / 44 (0.00%)
occurrences (all)	1	2	0
Neuropathy peripheral			
subjects affected / exposed	3 / 52 (5.77%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	3	0	1
Myoclonus			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	0	0	1
Sciatica			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences (all)	0	0	0
Migraine			
subjects affected / exposed	3 / 52 (5.77%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences (all)	3	0	0
Taste disorder			
subjects affected / exposed	2 / 52 (3.85%)	1 / 48 (2.08%)	0 / 44 (0.00%)
occurrences (all)	2	2	0
Syncope			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	0	0	1
Blood and lymphatic system disorders			
Anaemia			
subjects affected / exposed	15 / 52 (28.85%)	17 / 48 (35.42%)	12 / 44 (27.27%)
occurrences (all)	19	28	16
Leukocytosis			

subjects affected / exposed	2 / 52 (3.85%)	1 / 48 (2.08%)	3 / 44 (6.82%)
occurrences (all)	2	1	3
Lymphadenopathy			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	0	0	1
Neutropenia			
subjects affected / exposed	1 / 52 (1.92%)	1 / 48 (2.08%)	4 / 44 (9.09%)
occurrences (all)	1	1	4
Spleen disorder			
subjects affected / exposed	1 / 52 (1.92%)	3 / 48 (6.25%)	0 / 44 (0.00%)
occurrences (all)	1	3	0
Thrombocytosis			
subjects affected / exposed	4 / 52 (7.69%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences (all)	4	0	0
Thrombocytopenia			
subjects affected / exposed	12 / 52 (23.08%)	18 / 48 (37.50%)	4 / 44 (9.09%)
occurrences (all)	16	34	4
Ear and labyrinth disorders			
Ear discomfort			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	0	0	1
Ear pain			
subjects affected / exposed	1 / 52 (1.92%)	1 / 48 (2.08%)	1 / 44 (2.27%)
occurrences (all)	1	1	1
Ear haemorrhage			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	0	0	1
Hypoacusis			
subjects affected / exposed	1 / 52 (1.92%)	1 / 48 (2.08%)	1 / 44 (2.27%)
occurrences (all)	1	1	1
Tinnitus			
subjects affected / exposed	0 / 52 (0.00%)	3 / 48 (6.25%)	0 / 44 (0.00%)
occurrences (all)	0	3	0
Eye disorders			
Eye haemorrhage			

subjects affected / exposed	0 / 52 (0.00%)	1 / 48 (2.08%)	1 / 44 (2.27%)
occurrences (all)	0	1	1
Vision blurred			
subjects affected / exposed	2 / 52 (3.85%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	2	0	1
Gastrointestinal disorders			
Abdominal discomfort			
subjects affected / exposed	6 / 52 (11.54%)	2 / 48 (4.17%)	1 / 44 (2.27%)
occurrences (all)	6	3	1
Abdominal distension			
subjects affected / exposed	10 / 52 (19.23%)	6 / 48 (12.50%)	2 / 44 (4.55%)
occurrences (all)	15	6	2
Abdominal pain			
subjects affected / exposed	11 / 52 (21.15%)	11 / 48 (22.92%)	6 / 44 (13.64%)
occurrences (all)	12	14	6
Abdominal pain upper			
subjects affected / exposed	6 / 52 (11.54%)	2 / 48 (4.17%)	1 / 44 (2.27%)
occurrences (all)	7	2	1
Anal incontinence			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	0	0	1
Constipation			
subjects affected / exposed	14 / 52 (26.92%)	11 / 48 (22.92%)	4 / 44 (9.09%)
occurrences (all)	18	11	4
Diarrhoea			
subjects affected / exposed	26 / 52 (50.00%)	15 / 48 (31.25%)	17 / 44 (38.64%)
occurrences (all)	44	28	24
Dry mouth			
subjects affected / exposed	0 / 52 (0.00%)	1 / 48 (2.08%)	1 / 44 (2.27%)
occurrences (all)	0	1	1
Enterocolitis			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	0	0	1
Dysphagia			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	0	0	1

Dyspepsia			
subjects affected / exposed	1 / 52 (1.92%)	1 / 48 (2.08%)	2 / 44 (4.55%)
occurrences (all)	1	1	2
Eructation			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	0	0	1
Flatulence			
subjects affected / exposed	1 / 52 (1.92%)	3 / 48 (6.25%)	2 / 44 (4.55%)
occurrences (all)	1	3	2
Gastrooesophageal reflux disease			
subjects affected / exposed	1 / 52 (1.92%)	3 / 48 (6.25%)	1 / 44 (2.27%)
occurrences (all)	1	3	1
Gingival bleeding			
subjects affected / exposed	0 / 52 (0.00%)	1 / 48 (2.08%)	0 / 44 (0.00%)
occurrences (all)	0	2	0
Gingival pain			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	2 / 44 (4.55%)
occurrences (all)	0	0	3
Haemorrhoidal haemorrhage			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	0	0	1
Haemorrhoids			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	2 / 44 (4.55%)
occurrences (all)	0	0	2
Mouth ulceration			
subjects affected / exposed	1 / 52 (1.92%)	1 / 48 (2.08%)	0 / 44 (0.00%)
occurrences (all)	2	1	0
Mouth haemorrhage			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	0	0	1
Melaena			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	0	0	1
Nausea			
subjects affected / exposed	23 / 52 (44.23%)	18 / 48 (37.50%)	23 / 44 (52.27%)
occurrences (all)	34	24	28

Proctalgia			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	0	0	1
Oral disorder			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	0	0	1
Stomatitis			
subjects affected / exposed	2 / 52 (3.85%)	1 / 48 (2.08%)	3 / 44 (6.82%)
occurrences (all)	2	1	3
Vomiting			
subjects affected / exposed	10 / 52 (19.23%)	7 / 48 (14.58%)	13 / 44 (29.55%)
occurrences (all)	16	9	17
Hepatobiliary disorders			
Hyperbilirubinaemia			
subjects affected / exposed	1 / 52 (1.92%)	4 / 48 (8.33%)	0 / 44 (0.00%)
occurrences (all)	1	10	0
Skin and subcutaneous tissue disorders			
Dermatitis acneiform			
subjects affected / exposed	2 / 52 (3.85%)	1 / 48 (2.08%)	0 / 44 (0.00%)
occurrences (all)	4	1	0
Dry skin			
subjects affected / exposed	2 / 52 (3.85%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	2	0	1
Erythema			
subjects affected / exposed	4 / 52 (7.69%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences (all)	4	0	0
Hyperhidrosis			
subjects affected / exposed	3 / 52 (5.77%)	2 / 48 (4.17%)	0 / 44 (0.00%)
occurrences (all)	4	2	0
Night sweats			
subjects affected / exposed	7 / 52 (13.46%)	5 / 48 (10.42%)	1 / 44 (2.27%)
occurrences (all)	8	7	1
Petechiae			
subjects affected / exposed	3 / 52 (5.77%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences (all)	3	0	0
Pruritus			

subjects affected / exposed	22 / 52 (42.31%)	10 / 48 (20.83%)	2 / 44 (4.55%)
occurrences (all)	26	11	2
Skin discolouration			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	2 / 44 (4.55%)
occurrences (all)	0	0	2
Rash maculo-papular			
subjects affected / exposed	4 / 52 (7.69%)	1 / 48 (2.08%)	2 / 44 (4.55%)
occurrences (all)	7	1	2
Rash macular			
subjects affected / exposed	1 / 52 (1.92%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	1	0	1
Rash			
subjects affected / exposed	7 / 52 (13.46%)	2 / 48 (4.17%)	4 / 44 (9.09%)
occurrences (all)	11	3	4
Skin ulcer			
subjects affected / exposed	4 / 52 (7.69%)	1 / 48 (2.08%)	1 / 44 (2.27%)
occurrences (all)	4	3	1
Skin lesion			
subjects affected / exposed	2 / 52 (3.85%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences (all)	2	0	0
Urticaria			
subjects affected / exposed	0 / 52 (0.00%)	1 / 48 (2.08%)	1 / 44 (2.27%)
occurrences (all)	0	1	1
Skin induration			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	0	0	1
Renal and urinary disorders			
Acute kidney injury			
subjects affected / exposed	2 / 52 (3.85%)	2 / 48 (4.17%)	1 / 44 (2.27%)
occurrences (all)	2	2	1
Dysuria			
subjects affected / exposed	4 / 52 (7.69%)	3 / 48 (6.25%)	0 / 44 (0.00%)
occurrences (all)	5	3	0
Haematuria			
subjects affected / exposed	2 / 52 (3.85%)	1 / 48 (2.08%)	2 / 44 (4.55%)
occurrences (all)	2	1	2

Micturition urgency			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	0	0	1
Nocturia			
subjects affected / exposed	1 / 52 (1.92%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences (all)	1	0	0
Urinary incontinence			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	0	0	1
Pollakiuria			
subjects affected / exposed	3 / 52 (5.77%)	2 / 48 (4.17%)	1 / 44 (2.27%)
occurrences (all)	4	2	2
Musculoskeletal and connective tissue disorders			
Arthralgia			
subjects affected / exposed	10 / 52 (19.23%)	4 / 48 (8.33%)	1 / 44 (2.27%)
occurrences (all)	13	4	1
Bone cyst			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	0	0	1
Back pain			
subjects affected / exposed	5 / 52 (9.62%)	7 / 48 (14.58%)	3 / 44 (6.82%)
occurrences (all)	6	7	3
Bone pain			
subjects affected / exposed	7 / 52 (13.46%)	1 / 48 (2.08%)	2 / 44 (4.55%)
occurrences (all)	7	1	3
Groin pain			
subjects affected / exposed	1 / 52 (1.92%)	0 / 48 (0.00%)	2 / 44 (4.55%)
occurrences (all)	1	0	3
Gouty arthritis			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	0	0	1
Bursitis			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	0	0	1
Muscle spasms			

subjects affected / exposed	7 / 52 (13.46%)	2 / 48 (4.17%)	0 / 44 (0.00%)
occurrences (all)	8	2	0
Joint swelling			
subjects affected / exposed	1 / 52 (1.92%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences (all)	1	0	0
Muscular weakness			
subjects affected / exposed	5 / 52 (9.62%)	1 / 48 (2.08%)	1 / 44 (2.27%)
occurrences (all)	6	1	1
Musculoskeletal chest pain			
subjects affected / exposed	2 / 52 (3.85%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences (all)	2	0	0
Musculoskeletal disorder			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	0	0	1
Musculoskeletal pain			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	2 / 44 (4.55%)
occurrences (all)	0	0	2
Myalgia			
subjects affected / exposed	6 / 52 (11.54%)	3 / 48 (6.25%)	1 / 44 (2.27%)
occurrences (all)	8	3	1
Neck pain			
subjects affected / exposed	1 / 52 (1.92%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences (all)	1	0	0
Pain in extremity			
subjects affected / exposed	7 / 52 (13.46%)	2 / 48 (4.17%)	3 / 44 (6.82%)
occurrences (all)	7	2	4
Pain in jaw			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	0	0	1
Rhabdomyolysis			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	0	0	1
Infections and infestations			
Cellulitis			
subjects affected / exposed	3 / 52 (5.77%)	2 / 48 (4.17%)	1 / 44 (2.27%)
occurrences (all)	5	3	1

COVID-19			
subjects affected / exposed	6 / 52 (11.54%)	6 / 48 (12.50%)	0 / 44 (0.00%)
occurrences (all)	7	7	0
Bronchitis			
subjects affected / exposed	4 / 52 (7.69%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	5	0	1
Cystitis			
subjects affected / exposed	0 / 52 (0.00%)	1 / 48 (2.08%)	0 / 44 (0.00%)
occurrences (all)	0	1	0
Eye infection			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	0	0	1
Folliculitis			
subjects affected / exposed	1 / 52 (1.92%)	1 / 48 (2.08%)	0 / 44 (0.00%)
occurrences (all)	1	1	0
Fungal skin infection			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	0	0	1
Gastroenteritis			
subjects affected / exposed	1 / 52 (1.92%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences (all)	1	0	0
Herpes simplex reactivation			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	0	0	1
Herpes zoster			
subjects affected / exposed	4 / 52 (7.69%)	4 / 48 (8.33%)	0 / 44 (0.00%)
occurrences (all)	4	4	0
Hordeolum			
subjects affected / exposed	1 / 52 (1.92%)	1 / 48 (2.08%)	1 / 44 (2.27%)
occurrences (all)	1	1	1
Influenza			
subjects affected / exposed	0 / 52 (0.00%)	1 / 48 (2.08%)	0 / 44 (0.00%)
occurrences (all)	0	1	0
Localised infection			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences (all)	0	0	0

Lower respiratory tract infection			
subjects affected / exposed	0 / 52 (0.00%)	1 / 48 (2.08%)	0 / 44 (0.00%)
occurrences (all)	0	1	0
Orchitis			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	0	0	1
Otitis media			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	0	0	1
Pharyngitis			
subjects affected / exposed	1 / 52 (1.92%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	1	0	1
Pneumonia			
subjects affected / exposed	3 / 52 (5.77%)	2 / 48 (4.17%)	4 / 44 (9.09%)
occurrences (all)	3	2	4
Nasopharyngitis			
subjects affected / exposed	3 / 52 (5.77%)	1 / 48 (2.08%)	2 / 44 (4.55%)
occurrences (all)	4	1	3
Oral herpes			
subjects affected / exposed	0 / 52 (0.00%)	1 / 48 (2.08%)	0 / 44 (0.00%)
occurrences (all)	0	1	0
Skin infection			
subjects affected / exposed	0 / 52 (0.00%)	2 / 48 (4.17%)	0 / 44 (0.00%)
occurrences (all)	0	2	0
Sinusitis			
subjects affected / exposed	6 / 52 (11.54%)	3 / 48 (6.25%)	1 / 44 (2.27%)
occurrences (all)	6	3	1
Septic shock			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	0	0	1
Rhinitis			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences (all)	0	0	0
Respiratory tract infection			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences (all)	0	0	0

Staphylococcal infection			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	0	0	1
Urinary tract infection bacterial			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	0	0	1
Urinary tract infection			
subjects affected / exposed	7 / 52 (13.46%)	5 / 48 (10.42%)	3 / 44 (6.82%)
occurrences (all)	9	6	3
Upper respiratory tract infection			
subjects affected / exposed	6 / 52 (11.54%)	2 / 48 (4.17%)	0 / 44 (0.00%)
occurrences (all)	17	2	0
Tooth infection			
subjects affected / exposed	1 / 52 (1.92%)	1 / 48 (2.08%)	0 / 44 (0.00%)
occurrences (all)	1	1	0
Metabolism and nutrition disorders			
Decreased appetite			
subjects affected / exposed	13 / 52 (25.00%)	7 / 48 (14.58%)	18 / 44 (40.91%)
occurrences (all)	17	9	20
Dehydration			
subjects affected / exposed	1 / 52 (1.92%)	1 / 48 (2.08%)	2 / 44 (4.55%)
occurrences (all)	1	1	3
Folate deficiency			
subjects affected / exposed	0 / 52 (0.00%)	5 / 48 (10.42%)	0 / 44 (0.00%)
occurrences (all)	0	5	0
Gout			
subjects affected / exposed	3 / 52 (5.77%)	1 / 48 (2.08%)	0 / 44 (0.00%)
occurrences (all)	3	1	0
Hyperglycaemia			
subjects affected / exposed	5 / 52 (9.62%)	1 / 48 (2.08%)	6 / 44 (13.64%)
occurrences (all)	7	1	7
Hypercholesterolaemia			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences (all)	0	0	0
Hyperkalaemia			

subjects affected / exposed	4 / 52 (7.69%)	5 / 48 (10.42%)	2 / 44 (4.55%)
occurrences (all)	5	18	3
Hyperphosphataemia			
subjects affected / exposed	0 / 52 (0.00%)	1 / 48 (2.08%)	1 / 44 (2.27%)
occurrences (all)	0	1	1
Hypermagnesaemia			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	0	0	2
Hypertriglyceridaemia			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	0 / 44 (0.00%)
occurrences (all)	0	0	0
Hyperuricaemia			
subjects affected / exposed	8 / 52 (15.38%)	5 / 48 (10.42%)	6 / 44 (13.64%)
occurrences (all)	11	10	7
Hypocalcaemia			
subjects affected / exposed	2 / 52 (3.85%)	4 / 48 (8.33%)	2 / 44 (4.55%)
occurrences (all)	2	4	2
Hypokalaemia			
subjects affected / exposed	2 / 52 (3.85%)	1 / 48 (2.08%)	2 / 44 (4.55%)
occurrences (all)	2	1	2
Hypomagnesaemia			
subjects affected / exposed	2 / 52 (3.85%)	4 / 48 (8.33%)	3 / 44 (6.82%)
occurrences (all)	3	5	3
Hyponatraemia			
subjects affected / exposed	2 / 52 (3.85%)	1 / 48 (2.08%)	4 / 44 (9.09%)
occurrences (all)	2	4	6
Hypophosphataemia			
subjects affected / exposed	0 / 52 (0.00%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	0	0	1
Iron overload			
subjects affected / exposed	0 / 52 (0.00%)	4 / 48 (8.33%)	0 / 44 (0.00%)
occurrences (all)	0	4	0
Tumour lysis syndrome			
subjects affected / exposed	1 / 52 (1.92%)	0 / 48 (0.00%)	1 / 44 (2.27%)
occurrences (all)	1	0	1
Vitamin D deficiency			

subjects affected / exposed	0 / 52 (0.00%)	4 / 48 (8.33%)	0 / 44 (0.00%)
occurrences (all)	0	4	0
Type 2 diabetes mellitus			
subjects affected / exposed	0 / 52 (0.00%)	1 / 48 (2.08%)	0 / 44 (0.00%)
occurrences (all)	0	1	0

Non-serious adverse events	Phase 2 Overall	Phase 1 275 mg tablet PO daily	
Total subjects affected by non-serious adverse events			
subjects affected / exposed	289 / 292 (98.97%)	5 / 6 (83.33%)	
Neoplasms benign, malignant and unspecified (incl cysts and polyps)			
Squamous cell carcinoma			
subjects affected / exposed	8 / 292 (2.74%)	0 / 6 (0.00%)	
occurrences (all)	9	0	
Basal cell carcinoma			
subjects affected / exposed	5 / 292 (1.71%)	0 / 6 (0.00%)	
occurrences (all)	9	0	
Vascular disorders			
Flushing			
subjects affected / exposed	4 / 292 (1.37%)	0 / 6 (0.00%)	
occurrences (all)	4	0	
Haematoma			
subjects affected / exposed	12 / 292 (4.11%)	0 / 6 (0.00%)	
occurrences (all)	13	0	
Hot flush			
subjects affected / exposed	6 / 292 (2.05%)	0 / 6 (0.00%)	
occurrences (all)	6	0	
Hypotension			
subjects affected / exposed	7 / 292 (2.40%)	1 / 6 (16.67%)	
occurrences (all)	7	1	
Hypertension			
subjects affected / exposed	16 / 292 (5.48%)	0 / 6 (0.00%)	
occurrences (all)	27	0	
General disorders and administration site conditions			
Chest pain			
subjects affected / exposed	4 / 292 (1.37%)	0 / 6 (0.00%)	
occurrences (all)	4	0	

Asthenia			
subjects affected / exposed	30 / 292 (10.27%)	0 / 6 (0.00%)	
occurrences (all)	42	0	
Early satiety			
subjects affected / exposed	11 / 292 (3.77%)	0 / 6 (0.00%)	
occurrences (all)	13	0	
Chills			
subjects affected / exposed	21 / 292 (7.19%)	0 / 6 (0.00%)	
occurrences (all)	25	0	
Fatigue			
subjects affected / exposed	89 / 292 (30.48%)	1 / 6 (16.67%)	
occurrences (all)	124	2	
Gait disturbance			
subjects affected / exposed	3 / 292 (1.03%)	0 / 6 (0.00%)	
occurrences (all)	3	0	
Influenza like illness			
subjects affected / exposed	13 / 292 (4.45%)	0 / 6 (0.00%)	
occurrences (all)	22	0	
Malaise			
subjects affected / exposed	10 / 292 (3.42%)	1 / 6 (16.67%)	
occurrences (all)	10	1	
Mucosal inflammation			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences (all)	2	0	
Nodule			
subjects affected / exposed	3 / 292 (1.03%)	0 / 6 (0.00%)	
occurrences (all)	3	0	
Non-cardiac chest pain			
subjects affected / exposed	16 / 292 (5.48%)	0 / 6 (0.00%)	
occurrences (all)	22	0	
Oedema peripheral			
subjects affected / exposed	51 / 292 (17.47%)	2 / 6 (33.33%)	
occurrences (all)	58	2	
Pain			
subjects affected / exposed	5 / 292 (1.71%)	1 / 6 (16.67%)	
occurrences (all)	6	1	

Pyrexia			
subjects affected / exposed	44 / 292 (15.07%)	1 / 6 (16.67%)	
occurrences (all)	56	1	
Peripheral swelling			
subjects affected / exposed	4 / 292 (1.37%)	0 / 6 (0.00%)	
occurrences (all)	5	0	
Reproductive system and breast disorders			
Genital lesion			
subjects affected / exposed	0 / 292 (0.00%)	0 / 6 (0.00%)	
occurrences (all)	0	0	
Vaginal haemorrhage			
subjects affected / exposed	4 / 292 (1.37%)	0 / 6 (0.00%)	
occurrences (all)	4	0	
Respiratory, thoracic and mediastinal disorders			
Dyspnoea			
subjects affected / exposed	58 / 292 (19.86%)	1 / 6 (16.67%)	
occurrences (all)	66	1	
Cough			
subjects affected / exposed	73 / 292 (25.00%)	2 / 6 (33.33%)	
occurrences (all)	99	2	
Dyspnoea exertional			
subjects affected / exposed	8 / 292 (2.74%)	0 / 6 (0.00%)	
occurrences (all)	8	0	
Nasal congestion			
subjects affected / exposed	10 / 292 (3.42%)	0 / 6 (0.00%)	
occurrences (all)	11	0	
Hypoxia			
subjects affected / exposed	3 / 292 (1.03%)	1 / 6 (16.67%)	
occurrences (all)	3	1	
Epistaxis			
subjects affected / exposed	47 / 292 (16.10%)	0 / 6 (0.00%)	
occurrences (all)	109	0	
Oropharyngeal pain			
subjects affected / exposed	26 / 292 (8.90%)	0 / 6 (0.00%)	
occurrences (all)	32	0	
Pleural effusion			

subjects affected / exposed	7 / 292 (2.40%)	0 / 6 (0.00%)	
occurrences (all)	7	0	
Productive cough			
subjects affected / exposed	12 / 292 (4.11%)	0 / 6 (0.00%)	
occurrences (all)	13	0	
Pulmonary oedema			
subjects affected / exposed	3 / 292 (1.03%)	1 / 6 (16.67%)	
occurrences (all)	3	1	
Rhinorrhoea			
subjects affected / exposed	13 / 292 (4.45%)	0 / 6 (0.00%)	
occurrences (all)	13	0	
Sinus pain			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences (all)	1	0	
Upper-airway cough syndrome			
subjects affected / exposed	15 / 292 (5.14%)	0 / 6 (0.00%)	
occurrences (all)	15	0	
Wheezing			
subjects affected / exposed	3 / 292 (1.03%)	1 / 6 (16.67%)	
occurrences (all)	3	1	
Psychiatric disorders			
Anxiety			
subjects affected / exposed	9 / 292 (3.08%)	0 / 6 (0.00%)	
occurrences (all)	9	0	
Confusional state			
subjects affected / exposed	7 / 292 (2.40%)	0 / 6 (0.00%)	
occurrences (all)	7	0	
Delirium			
subjects affected / exposed	3 / 292 (1.03%)	0 / 6 (0.00%)	
occurrences (all)	3	0	
Depression			
subjects affected / exposed	8 / 292 (2.74%)	0 / 6 (0.00%)	
occurrences (all)	9	0	
Insomnia			
subjects affected / exposed	22 / 292 (7.53%)	0 / 6 (0.00%)	
occurrences (all)	23	0	

Investigations			
Bilirubin conjugated increased			
subjects affected / exposed	6 / 292 (2.05%)	0 / 6 (0.00%)	
occurrences (all)	7	0	
Aspartate aminotransferase increased			
subjects affected / exposed	24 / 292 (8.22%)	0 / 6 (0.00%)	
occurrences (all)	30	0	
Alanine aminotransferase increased			
subjects affected / exposed	24 / 292 (8.22%)	0 / 6 (0.00%)	
occurrences (all)	31	0	
Activated partial thromboplastin time prolonged			
subjects affected / exposed	2 / 292 (0.68%)	0 / 6 (0.00%)	
occurrences (all)	2	0	
Blast cell count increased			
subjects affected / exposed	5 / 292 (1.71%)	0 / 6 (0.00%)	
occurrences (all)	5	0	
Blood alkaline phosphatase increased			
subjects affected / exposed	13 / 292 (4.45%)	0 / 6 (0.00%)	
occurrences (all)	14	0	
Blood bilirubin increased			
subjects affected / exposed	20 / 292 (6.85%)	0 / 6 (0.00%)	
occurrences (all)	25	0	
Cardiac murmur			
subjects affected / exposed	5 / 292 (1.71%)	0 / 6 (0.00%)	
occurrences (all)	6	0	
Blood phosphorus increased			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences (all)	1	0	
Blood uric acid increased			
subjects affected / exposed	3 / 292 (1.03%)	0 / 6 (0.00%)	
occurrences (all)	3	0	
Blood creatinine increased			
subjects affected / exposed	25 / 292 (8.56%)	1 / 6 (16.67%)	
occurrences (all)	31	1	
International normalised ratio increased			

subjects affected / exposed	5 / 292 (1.71%)	0 / 6 (0.00%)	
occurrences (all)	5	0	
Ejection fraction decreased			
subjects affected / exposed	2 / 292 (0.68%)	0 / 6 (0.00%)	
occurrences (all)	2	0	
Electrocardiogram QRS complex abnormal			
subjects affected / exposed	0 / 292 (0.00%)	0 / 6 (0.00%)	
occurrences (all)	0	0	
Electrocardiogram QT prolonged			
subjects affected / exposed	2 / 292 (0.68%)	0 / 6 (0.00%)	
occurrences (all)	2	0	
Haematocrit increased			
subjects affected / exposed	2 / 292 (0.68%)	0 / 6 (0.00%)	
occurrences (all)	8	0	
Lymphocyte count decreased			
subjects affected / exposed	4 / 292 (1.37%)	0 / 6 (0.00%)	
occurrences (all)	9	0	
Neutrophil count decreased			
subjects affected / exposed	5 / 292 (1.71%)	1 / 6 (16.67%)	
occurrences (all)	13	1	
Platelet count decreased			
subjects affected / exposed	56 / 292 (19.18%)	1 / 6 (16.67%)	
occurrences (all)	94	1	
Transaminases increased			
subjects affected / exposed	0 / 292 (0.00%)	0 / 6 (0.00%)	
occurrences (all)	0	0	
Troponin T increased			
subjects affected / exposed	1 / 292 (0.34%)	1 / 6 (16.67%)	
occurrences (all)	1	1	
Weight decreased			
subjects affected / exposed	39 / 292 (13.36%)	3 / 6 (50.00%)	
occurrences (all)	41	5	
Weight increased			
subjects affected / exposed	8 / 292 (2.74%)	0 / 6 (0.00%)	
occurrences (all)	9	0	

White blood cell count decreased			
subjects affected / exposed	10 / 292 (3.42%)	0 / 6 (0.00%)	
occurrences (all)	17	0	
Injury, poisoning and procedural complications			
Contusion			
subjects affected / exposed	48 / 292 (16.44%)	0 / 6 (0.00%)	
occurrences (all)	62	0	
Fall			
subjects affected / exposed	33 / 292 (11.30%)	0 / 6 (0.00%)	
occurrences (all)	44	0	
Vascular access site complication			
subjects affected / exposed	0 / 292 (0.00%)	0 / 6 (0.00%)	
occurrences (all)	0	0	
Skin laceration			
subjects affected / exposed	5 / 292 (1.71%)	0 / 6 (0.00%)	
occurrences (all)	6	0	
Procedural site reaction			
subjects affected / exposed	0 / 292 (0.00%)	0 / 6 (0.00%)	
occurrences (all)	0	0	
Procedural pain			
subjects affected / exposed	6 / 292 (2.05%)	0 / 6 (0.00%)	
occurrences (all)	6	0	
Nail injury			
subjects affected / exposed	2 / 292 (0.68%)	0 / 6 (0.00%)	
occurrences (all)	3	0	
Wound			
subjects affected / exposed	7 / 292 (2.40%)	0 / 6 (0.00%)	
occurrences (all)	7	0	
Cardiac disorders			
Angina pectoris			
subjects affected / exposed	3 / 292 (1.03%)	0 / 6 (0.00%)	
occurrences (all)	3	0	
Atrial fibrillation			
subjects affected / exposed	7 / 292 (2.40%)	0 / 6 (0.00%)	
occurrences (all)	8	0	
Atrioventricular block			

subjects affected / exposed	0 / 292 (0.00%)	0 / 6 (0.00%)	
occurrences (all)	0	0	
Hyperdynamic left ventricle			
subjects affected / exposed	0 / 292 (0.00%)	0 / 6 (0.00%)	
occurrences (all)	0	0	
Palpitations			
subjects affected / exposed	14 / 292 (4.79%)	1 / 6 (16.67%)	
occurrences (all)	23	1	
Sinus tachycardia			
subjects affected / exposed	3 / 292 (1.03%)	0 / 6 (0.00%)	
occurrences (all)	4	0	
Tachycardia			
subjects affected / exposed	6 / 292 (2.05%)	1 / 6 (16.67%)	
occurrences (all)	6	1	
Nervous system disorders			
Ageusia			
subjects affected / exposed	7 / 292 (2.40%)	0 / 6 (0.00%)	
occurrences (all)	8	0	
Dysarthria			
subjects affected / exposed	2 / 292 (0.68%)	0 / 6 (0.00%)	
occurrences (all)	3	0	
Dizziness			
subjects affected / exposed	60 / 292 (20.55%)	0 / 6 (0.00%)	
occurrences (all)	78	0	
Cerebrovascular accident			
subjects affected / exposed	0 / 292 (0.00%)	0 / 6 (0.00%)	
occurrences (all)	0	0	
Memory impairment			
subjects affected / exposed	3 / 292 (1.03%)	0 / 6 (0.00%)	
occurrences (all)	3	0	
Lethargy			
subjects affected / exposed	4 / 292 (1.37%)	0 / 6 (0.00%)	
occurrences (all)	6	0	
Hypoaesthesia			
subjects affected / exposed	6 / 292 (2.05%)	0 / 6 (0.00%)	
occurrences (all)	9	0	

Headache			
subjects affected / exposed	54 / 292 (18.49%)	0 / 6 (0.00%)	
occurrences (all)	79	0	
Dysgeusia			
subjects affected / exposed	79 / 292 (27.05%)	1 / 6 (16.67%)	
occurrences (all)	95	1	
Presyncope			
subjects affected / exposed	7 / 292 (2.40%)	0 / 6 (0.00%)	
occurrences (all)	7	0	
Paraesthesia			
subjects affected / exposed	17 / 292 (5.82%)	0 / 6 (0.00%)	
occurrences (all)	22	0	
Neuropathy peripheral			
subjects affected / exposed	7 / 292 (2.40%)	0 / 6 (0.00%)	
occurrences (all)	7	0	
Myoclonus			
subjects affected / exposed	0 / 292 (0.00%)	0 / 6 (0.00%)	
occurrences (all)	0	0	
Sciatica			
subjects affected / exposed	8 / 292 (2.74%)	0 / 6 (0.00%)	
occurrences (all)	14	0	
Migraine			
subjects affected / exposed	8 / 292 (2.74%)	0 / 6 (0.00%)	
occurrences (all)	8	0	
Taste disorder			
subjects affected / exposed	7 / 292 (2.40%)	0 / 6 (0.00%)	
occurrences (all)	9	0	
Syncope			
subjects affected / exposed	5 / 292 (1.71%)	1 / 6 (16.67%)	
occurrences (all)	6	1	
Blood and lymphatic system disorders			
Anaemia			
subjects affected / exposed	99 / 292 (33.90%)	2 / 6 (33.33%)	
occurrences (all)	163	2	
Leukocytosis			

subjects affected / exposed	11 / 292 (3.77%)	0 / 6 (0.00%)	
occurrences (all)	13	0	
Lymphadenopathy			
subjects affected / exposed	4 / 292 (1.37%)	0 / 6 (0.00%)	
occurrences (all)	5	0	
Neutropenia			
subjects affected / exposed	16 / 292 (5.48%)	0 / 6 (0.00%)	
occurrences (all)	25	0	
Spleen disorder			
subjects affected / exposed	6 / 292 (2.05%)	0 / 6 (0.00%)	
occurrences (all)	6	0	
Thrombocytosis			
subjects affected / exposed	5 / 292 (1.71%)	0 / 6 (0.00%)	
occurrences (all)	12	0	
Thrombocytopenia			
subjects affected / exposed	96 / 292 (32.88%)	0 / 6 (0.00%)	
occurrences (all)	179	0	
Ear and labyrinth disorders			
Ear discomfort			
subjects affected / exposed	2 / 292 (0.68%)	0 / 6 (0.00%)	
occurrences (all)	2	0	
Ear pain			
subjects affected / exposed	6 / 292 (2.05%)	0 / 6 (0.00%)	
occurrences (all)	6	0	
Ear haemorrhage			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences (all)	1	0	
Hypoacusis			
subjects affected / exposed	3 / 292 (1.03%)	0 / 6 (0.00%)	
occurrences (all)	3	0	
Tinnitus			
subjects affected / exposed	12 / 292 (4.11%)	0 / 6 (0.00%)	
occurrences (all)	14	0	
Eye disorders			
Eye haemorrhage			

subjects affected / exposed	2 / 292 (0.68%)	0 / 6 (0.00%)	
occurrences (all)	2	0	
Vision blurred			
subjects affected / exposed	9 / 292 (3.08%)	0 / 6 (0.00%)	
occurrences (all)	10	0	
Gastrointestinal disorders			
Abdominal discomfort			
subjects affected / exposed	18 / 292 (6.16%)	0 / 6 (0.00%)	
occurrences (all)	20	0	
Abdominal distension			
subjects affected / exposed	44 / 292 (15.07%)	0 / 6 (0.00%)	
occurrences (all)	55	0	
Abdominal pain			
subjects affected / exposed	52 / 292 (17.81%)	2 / 6 (33.33%)	
occurrences (all)	65	2	
Abdominal pain upper			
subjects affected / exposed	30 / 292 (10.27%)	0 / 6 (0.00%)	
occurrences (all)	40	0	
Anal incontinence			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences (all)	1	0	
Constipation			
subjects affected / exposed	77 / 292 (26.37%)	1 / 6 (16.67%)	
occurrences (all)	89	1	
Diarrhoea			
subjects affected / exposed	143 / 292 (48.97%)	2 / 6 (33.33%)	
occurrences (all)	260	5	
Dry mouth			
subjects affected / exposed	5 / 292 (1.71%)	0 / 6 (0.00%)	
occurrences (all)	5	0	
Enterocolitis			
subjects affected / exposed	0 / 292 (0.00%)	0 / 6 (0.00%)	
occurrences (all)	0	0	
Dysphagia			
subjects affected / exposed	4 / 292 (1.37%)	0 / 6 (0.00%)	
occurrences (all)	4	0	

Dyspepsia				
subjects affected / exposed	10 / 292 (3.42%)	0 / 6 (0.00%)		
occurrences (all)	12	0		
Eructation				
subjects affected / exposed	0 / 292 (0.00%)	0 / 6 (0.00%)		
occurrences (all)	0	0		
Flatulence				
subjects affected / exposed	13 / 292 (4.45%)	0 / 6 (0.00%)		
occurrences (all)	13	0		
Gastrooesophageal reflux disease				
subjects affected / exposed	6 / 292 (2.05%)	1 / 6 (16.67%)		
occurrences (all)	6	1		
Gingival bleeding				
subjects affected / exposed	6 / 292 (2.05%)	0 / 6 (0.00%)		
occurrences (all)	8	0		
Gingival pain				
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)		
occurrences (all)	1	0		
Haemorrhoidal haemorrhage				
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)		
occurrences (all)	2	0		
Haemorrhoids				
subjects affected / exposed	5 / 292 (1.71%)	0 / 6 (0.00%)		
occurrences (all)	6	0		
Mouth ulceration				
subjects affected / exposed	9 / 292 (3.08%)	0 / 6 (0.00%)		
occurrences (all)	11	0		
Mouth haemorrhage				
subjects affected / exposed	2 / 292 (0.68%)	0 / 6 (0.00%)		
occurrences (all)	2	0		
Melaena				
subjects affected / exposed	3 / 292 (1.03%)	0 / 6 (0.00%)		
occurrences (all)	3	0		
Nausea				
subjects affected / exposed	116 / 292 (39.73%)	2 / 6 (33.33%)		
occurrences (all)	174	4		

Proctalgia			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences (all)	1	0	
Oral disorder			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences (all)	1	0	
Stomatitis			
subjects affected / exposed	14 / 292 (4.79%)	1 / 6 (16.67%)	
occurrences (all)	14	1	
Vomiting			
subjects affected / exposed	59 / 292 (20.21%)	3 / 6 (50.00%)	
occurrences (all)	81	5	
Hepatobiliary disorders			
Hyperbilirubinaemia			
subjects affected / exposed	9 / 292 (3.08%)	0 / 6 (0.00%)	
occurrences (all)	21	0	
Skin and subcutaneous tissue disorders			
Dermatitis acneiform			
subjects affected / exposed	10 / 292 (3.42%)	0 / 6 (0.00%)	
occurrences (all)	14	0	
Dry skin			
subjects affected / exposed	7 / 292 (2.40%)	0 / 6 (0.00%)	
occurrences (all)	7	0	
Erythema			
subjects affected / exposed	6 / 292 (2.05%)	0 / 6 (0.00%)	
occurrences (all)	6	0	
Hyperhidrosis			
subjects affected / exposed	10 / 292 (3.42%)	0 / 6 (0.00%)	
occurrences (all)	12	0	
Night sweats			
subjects affected / exposed	32 / 292 (10.96%)	0 / 6 (0.00%)	
occurrences (all)	44	0	
Petechiae			
subjects affected / exposed	6 / 292 (2.05%)	0 / 6 (0.00%)	
occurrences (all)	6	0	
Pruritus			

subjects affected / exposed	62 / 292 (21.23%)	2 / 6 (33.33%)	
occurrences (all)	72	2	
Skin discolouration			
subjects affected / exposed	1 / 292 (0.34%)	1 / 6 (16.67%)	
occurrences (all)	1	1	
Rash maculo-papular			
subjects affected / exposed	10 / 292 (3.42%)	0 / 6 (0.00%)	
occurrences (all)	17	0	
Rash macular			
subjects affected / exposed	2 / 292 (0.68%)	1 / 6 (16.67%)	
occurrences (all)	2	1	
Rash			
subjects affected / exposed	26 / 292 (8.90%)	0 / 6 (0.00%)	
occurrences (all)	35	0	
Skin ulcer			
subjects affected / exposed	9 / 292 (3.08%)	1 / 6 (16.67%)	
occurrences (all)	11	1	
Skin lesion			
subjects affected / exposed	15 / 292 (5.14%)	0 / 6 (0.00%)	
occurrences (all)	16	0	
Urticaria			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences (all)	1	0	
Skin induration			
subjects affected / exposed	0 / 292 (0.00%)	0 / 6 (0.00%)	
occurrences (all)	0	0	
Renal and urinary disorders			
Acute kidney injury			
subjects affected / exposed	6 / 292 (2.05%)	0 / 6 (0.00%)	
occurrences (all)	6	0	
Dysuria			
subjects affected / exposed	15 / 292 (5.14%)	0 / 6 (0.00%)	
occurrences (all)	18	0	
Haematuria			
subjects affected / exposed	9 / 292 (3.08%)	1 / 6 (16.67%)	
occurrences (all)	10	1	

Micturition urgency			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences (all)	1	0	
Nocturia			
subjects affected / exposed	5 / 292 (1.71%)	0 / 6 (0.00%)	
occurrences (all)	5	0	
Urinary incontinence			
subjects affected / exposed	0 / 292 (0.00%)	0 / 6 (0.00%)	
occurrences (all)	0	0	
Pollakiuria			
subjects affected / exposed	13 / 292 (4.45%)	0 / 6 (0.00%)	
occurrences (all)	14	0	
Musculoskeletal and connective tissue disorders			
Arthralgia			
subjects affected / exposed	56 / 292 (19.18%)	0 / 6 (0.00%)	
occurrences (all)	69	0	
Bone cyst			
subjects affected / exposed	0 / 292 (0.00%)	0 / 6 (0.00%)	
occurrences (all)	0	0	
Back pain			
subjects affected / exposed	44 / 292 (15.07%)	1 / 6 (16.67%)	
occurrences (all)	52	1	
Bone pain			
subjects affected / exposed	28 / 292 (9.59%)	0 / 6 (0.00%)	
occurrences (all)	32	0	
Groin pain			
subjects affected / exposed	5 / 292 (1.71%)	0 / 6 (0.00%)	
occurrences (all)	5	0	
Gouty arthritis			
subjects affected / exposed	0 / 292 (0.00%)	0 / 6 (0.00%)	
occurrences (all)	0	0	
Bursitis			
subjects affected / exposed	2 / 292 (0.68%)	0 / 6 (0.00%)	
occurrences (all)	2	0	
Muscle spasms			

subjects affected / exposed	50 / 292 (17.12%)	0 / 6 (0.00%)	
occurrences (all)	61	0	
Joint swelling			
subjects affected / exposed	8 / 292 (2.74%)	0 / 6 (0.00%)	
occurrences (all)	9	0	
Muscular weakness			
subjects affected / exposed	20 / 292 (6.85%)	1 / 6 (16.67%)	
occurrences (all)	25	1	
Musculoskeletal chest pain			
subjects affected / exposed	12 / 292 (4.11%)	0 / 6 (0.00%)	
occurrences (all)	13	0	
Musculoskeletal disorder			
subjects affected / exposed	0 / 292 (0.00%)	0 / 6 (0.00%)	
occurrences (all)	0	0	
Musculoskeletal pain			
subjects affected / exposed	6 / 292 (2.05%)	0 / 6 (0.00%)	
occurrences (all)	6	0	
Myalgia			
subjects affected / exposed	30 / 292 (10.27%)	0 / 6 (0.00%)	
occurrences (all)	38	0	
Neck pain			
subjects affected / exposed	8 / 292 (2.74%)	0 / 6 (0.00%)	
occurrences (all)	8	0	
Pain in extremity			
subjects affected / exposed	41 / 292 (14.04%)	0 / 6 (0.00%)	
occurrences (all)	50	0	
Pain in jaw			
subjects affected / exposed	2 / 292 (0.68%)	0 / 6 (0.00%)	
occurrences (all)	2	0	
Rhabdomyolysis			
subjects affected / exposed	0 / 292 (0.00%)	1 / 6 (16.67%)	
occurrences (all)	0	1	
Infections and infestations			
Cellulitis			
subjects affected / exposed	12 / 292 (4.11%)	0 / 6 (0.00%)	
occurrences (all)	16	0	

COVID-19			
subjects affected / exposed	52 / 292 (17.81%)	0 / 6 (0.00%)	
occurrences (all)	60	0	
Bronchitis			
subjects affected / exposed	11 / 292 (3.77%)	0 / 6 (0.00%)	
occurrences (all)	12	0	
Cystitis			
subjects affected / exposed	6 / 292 (2.05%)	0 / 6 (0.00%)	
occurrences (all)	8	0	
Eye infection			
subjects affected / exposed	2 / 292 (0.68%)	0 / 6 (0.00%)	
occurrences (all)	2	0	
Folliculitis			
subjects affected / exposed	8 / 292 (2.74%)	0 / 6 (0.00%)	
occurrences (all)	13	0	
Fungal skin infection			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences (all)	1	0	
Gastroenteritis			
subjects affected / exposed	8 / 292 (2.74%)	0 / 6 (0.00%)	
occurrences (all)	9	0	
Herpes simplex reactivation			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences (all)	1	0	
Herpes zoster			
subjects affected / exposed	31 / 292 (10.62%)	0 / 6 (0.00%)	
occurrences (all)	34	0	
Hordeolum			
subjects affected / exposed	8 / 292 (2.74%)	1 / 6 (16.67%)	
occurrences (all)	12	1	
Influenza			
subjects affected / exposed	6 / 292 (2.05%)	0 / 6 (0.00%)	
occurrences (all)	8	0	
Localised infection			
subjects affected / exposed	4 / 292 (1.37%)	0 / 6 (0.00%)	
occurrences (all)	4	0	

Lower respiratory tract infection			
subjects affected / exposed	9 / 292 (3.08%)	0 / 6 (0.00%)	
occurrences (all)	12	0	
Orchitis			
subjects affected / exposed	0 / 292 (0.00%)	0 / 6 (0.00%)	
occurrences (all)	0	0	
Otitis media			
subjects affected / exposed	3 / 292 (1.03%)	0 / 6 (0.00%)	
occurrences (all)	3	0	
Pharyngitis			
subjects affected / exposed	2 / 292 (0.68%)	0 / 6 (0.00%)	
occurrences (all)	2	0	
Pneumonia			
subjects affected / exposed	21 / 292 (7.19%)	0 / 6 (0.00%)	
occurrences (all)	24	0	
Nasopharyngitis			
subjects affected / exposed	18 / 292 (6.16%)	0 / 6 (0.00%)	
occurrences (all)	28	0	
Oral herpes			
subjects affected / exposed	13 / 292 (4.45%)	0 / 6 (0.00%)	
occurrences (all)	19	0	
Skin infection			
subjects affected / exposed	6 / 292 (2.05%)	0 / 6 (0.00%)	
occurrences (all)	6	0	
Sinusitis			
subjects affected / exposed	16 / 292 (5.48%)	0 / 6 (0.00%)	
occurrences (all)	21	0	
Septic shock			
subjects affected / exposed	0 / 292 (0.00%)	0 / 6 (0.00%)	
occurrences (all)	0	0	
Rhinitis			
subjects affected / exposed	7 / 292 (2.40%)	0 / 6 (0.00%)	
occurrences (all)	7	0	
Respiratory tract infection			
subjects affected / exposed	6 / 292 (2.05%)	0 / 6 (0.00%)	
occurrences (all)	7	0	

Staphylococcal infection			
subjects affected / exposed	0 / 292 (0.00%)	0 / 6 (0.00%)	
occurrences (all)	0	0	
Urinary tract infection bacterial			
subjects affected / exposed	0 / 292 (0.00%)	0 / 6 (0.00%)	
occurrences (all)	0	0	
Urinary tract infection			
subjects affected / exposed	39 / 292 (13.36%)	0 / 6 (0.00%)	
occurrences (all)	68	0	
Upper respiratory tract infection			
subjects affected / exposed	43 / 292 (14.73%)	0 / 6 (0.00%)	
occurrences (all)	72	0	
Tooth infection			
subjects affected / exposed	8 / 292 (2.74%)	0 / 6 (0.00%)	
occurrences (all)	9	0	
Metabolism and nutrition disorders			
Decreased appetite			
subjects affected / exposed	59 / 292 (20.21%)	2 / 6 (33.33%)	
occurrences (all)	70	3	
Dehydration			
subjects affected / exposed	5 / 292 (1.71%)	1 / 6 (16.67%)	
occurrences (all)	5	2	
Folate deficiency			
subjects affected / exposed	5 / 292 (1.71%)	0 / 6 (0.00%)	
occurrences (all)	5	0	
Gout			
subjects affected / exposed	9 / 292 (3.08%)	0 / 6 (0.00%)	
occurrences (all)	9	0	
Hyperglycaemia			
subjects affected / exposed	15 / 292 (5.14%)	1 / 6 (16.67%)	
occurrences (all)	18	2	
Hypercholesterolaemia			
subjects affected / exposed	5 / 292 (1.71%)	0 / 6 (0.00%)	
occurrences (all)	5	0	
Hyperkalaemia			

subjects affected / exposed	22 / 292 (7.53%)	0 / 6 (0.00%)	
occurrences (all)	49	0	
Hyperphosphataemia			
subjects affected / exposed	4 / 292 (1.37%)	0 / 6 (0.00%)	
occurrences (all)	9	0	
Hypermagnesaemia			
subjects affected / exposed	0 / 292 (0.00%)	0 / 6 (0.00%)	
occurrences (all)	0	0	
Hypertriglyceridaemia			
subjects affected / exposed	6 / 292 (2.05%)	0 / 6 (0.00%)	
occurrences (all)	13	0	
Hyperuricaemia			
subjects affected / exposed	27 / 292 (9.25%)	0 / 6 (0.00%)	
occurrences (all)	36	0	
Hypocalcaemia			
subjects affected / exposed	11 / 292 (3.77%)	0 / 6 (0.00%)	
occurrences (all)	14	0	
Hypokalaemia			
subjects affected / exposed	12 / 292 (4.11%)	0 / 6 (0.00%)	
occurrences (all)	12	0	
Hypomagnesaemia			
subjects affected / exposed	10 / 292 (3.42%)	0 / 6 (0.00%)	
occurrences (all)	15	0	
Hyponatraemia			
subjects affected / exposed	20 / 292 (6.85%)	0 / 6 (0.00%)	
occurrences (all)	25	0	
Hypophosphataemia			
subjects affected / exposed	3 / 292 (1.03%)	0 / 6 (0.00%)	
occurrences (all)	6	0	
Iron overload			
subjects affected / exposed	10 / 292 (3.42%)	0 / 6 (0.00%)	
occurrences (all)	10	0	
Tumour lysis syndrome			
subjects affected / exposed	1 / 292 (0.34%)	0 / 6 (0.00%)	
occurrences (all)	1	0	
Vitamin D deficiency			

subjects affected / exposed	7 / 292 (2.40%)	0 / 6 (0.00%)	
occurrences (all)	7	0	
Type 2 diabetes mellitus			
subjects affected / exposed	3 / 292 (1.03%)	0 / 6 (0.00%)	
occurrences (all)	3	0	

More information

Substantial protocol amendments (globally)

Were there any global substantial amendments to the protocol? Yes

Date	Amendment
04 January 2018	Amendment 6: A Phase 2 component was added in place of the dose expansion phase of the study to evaluate the efficacy of pelabresib with or without ruxolitinib in patients with myelofibrosis (MF). The dose escalation component of the study was subsequently referred to as Phase 1. As of Amendment 6, Phase 1 was complete, and all new patients were enrolled in Phase 2.
07 September 2018	Amendment 7: * The two arms were stratified into transfusion-dependent (TD) and non-transfusion-dependent (non-TD) cohorts. Arm 1 (Prior Janus kinase inhibitor [JAKi] Monotherapy Arm; previously the Monotherapy Arm) was stratified into Cohort 1A (TD) and Cohort 1B (non-TD), and Arm 2 (Prior JAKi Combination Arm; previously the Combination Arm) was stratified into Cohort 2A (TD) and Cohort 2B (non-TD). * A new combination arm was added for patients with myelofibrosis (MF) who had not previously been treated with a JAKi (i.e., JAKi-naïve). This arm was designated Arm 3: JAKi-Naïve Combination Arm. * Key features related to these changes were expanded, including specification of eligibility criteria and primary endpoints for TD versus non-TD cohorts. * Guidelines for dose adjustments and rules for concomitant treatments were updated.
23 September 2019	Amendment 8: * The new starting doses of pelabresib and ruxolitinib were confirmed based on the recommendations of the Safety Review Committee. * Specific instructions for dose modifications due to toxicity were added, replacing the previous approach of following institutional practice. * The number of patients in the JAKi-Naïve Combination Arm 3 was increased to more fully evaluate the efficacy and safety of pelabresib in combination with ruxolitinib. * The anemia component of the required diagnosis of myelofibrosis (MF) for study entry was removed to ensure that study patients were representative of the intended patient population. * The methodology for determining spleen volume at screening was changed to ensure more consistent assessments of baseline disease and response to study drug. * A lower-dose pelabresib and ruxolitinib combination was added for a subset of patients in the JAKi-Naïve Combination Arm 3 to study the safety and efficacy of a reduced pelabresib dose. * The Dynamic International Prognostic Scoring System (DIPSS) category in Arms 1–3 was changed to intermediate-2 or higher.
14 April 2020	Amendment 9: * Adverse event (AE) prophylaxis was added. * Guidelines for managing infection, concomitant use of systemic corticosteroids, and hyperkalemia were added. * Recommendations for the dose regimen and dose modifications in Arm 3 were updated. * The AE relatedness criteria were revised to provide more specificity in possible responses.
25 September 2020	Amendment 10: * A study arm (Arm 4) was added to enroll high-risk essential thrombocythemia (ET) patients who were resistant or intolerant to hydroxyurea (HU). * Cohort 1B was expanded to enroll up to 50 patients. * Arm 3 was reduced to a maximum of 81 patients. * Guidance for dose reductions and modifications for pelabresib and ruxolitinib was revised and clarified. * Guidelines for handling unforeseen circumstances were added to address potential risks to patients.
16 September 2022	Amendment 11: * An exploratory objective was added to evaluate overall survival in Arm 3. * Acute respiratory distress syndrome (ARDS), considered a respiratory distress syndrome (RDS) event, was added as an adverse event of special interest (AESI). * Guidance regarding COVID-19 and contraception was updated. * Assessments of acute myeloid leukemia (AML) transformation, new anti-cancer therapy, and survival follow-up were added to the schedule of events.

27 November 2023	Amendment 12: * Guidance on contraception use during the study was updated. * Confirmation was added that no further pharmacokinetic (PK) samples or allele burden blood samples would be collected.
23 February 2024	Amendment 13: * A benefit-risk assessment was added. * Dose modification guidance and withdrawal criteria for increases in peripheral blood blasts were added. * Accelerated phase (AP) and transformation to blast phase (BP) were added as adverse events of special interest (AESIs). * Adverse event (AE) reporting guidance related to AESIs of AP and transformation to BP was updated.

Notes:

Interruptions (globally)

Were there any global interruptions to the trial? No

Limitations and caveats

Limitations of the trial such as small numbers of subjects analysed or technical problems leading to unreliable data.

Due to EudraCT system limitations, which EMA is aware of, data using 999 as data points in this record are not an accurate representation of the clinical trial results. Please go to https://www.novctrd.com/#/ for complete trial results

Notes: