



Clinical trial results:

A Multicenter, Open-Label, Single-Arm, Phase IIIb, International Study Evaluating the Safety of Obinutuzumab Alone or in Combination with Chemotherapy in Patients with Previously Untreated or Relapsed/Refractory Chronic Lymphocytic Leukemia

Summary

EudraCT number	2013-000087-29
Trial protocol	IT DE ES FI LV SI IE PT EE GR SE PL SK BE LT FR
Global end of trial date	08 October 2018

Results information

Result version number	v3 (current)
This version publication date	13 November 2019
First version publication date	06 January 2018
Version creation reason	

Trial information

Trial identification

Sponsor protocol code	MO28543
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Additional study identifiers

ISRCTN number	-
ClinicalTrials.gov id (NCT number)	NCT01905943
WHO universal trial number (UTN)	-

Notes:

Sponsors

Sponsor organisation name	F. Hoffmann-La Roche AG
Sponsor organisation address	Grenzacherstrasse 124, Basel, Switzerland, CH4070
Public contact	F. Hoffmann-La Roche AG, F. Hoffmann-La Roche AG, +41 616878333, global.trial_information@roche.com
Scientific contact	F. Hoffmann-La Roche AG, F. Hoffmann-La Roche AG, +41 616878333, global.trial_information@roche.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	08 October 2018
Is this the analysis of the primary completion data?	No
Global end of trial reached?	Yes
Global end of trial date	08 October 2018
Was the trial ended prematurely?	No

Notes:

General information about the trial

Main objective of the trial:

To evaluate the safety and tolerability of obinutuzumab alone or in combination with chemotherapy.

Protection of trial subjects:

All study subjects were required to read and sign an Informed Consent Form.

Background therapy: -

Evidence for comparator: -

Actual start date of recruitment	04 November 2013
Long term follow-up planned	Yes
Long term follow-up rationale	Safety, Efficacy
Long term follow-up duration	30 Months
Independent data monitoring committee (IDMC) involvement?	Yes

Notes:

Population of trial subjects

Subjects enrolled per country

Country: Number of subjects enrolled	Argentina: 8
Country: Number of subjects enrolled	Belgium: 18
Country: Number of subjects enrolled	Brazil: 34
Country: Number of subjects enrolled	Bosnia and Herzegovina: 7
Country: Number of subjects enrolled	Canada: 69
Country: Number of subjects enrolled	Estonia: 10
Country: Number of subjects enrolled	Finland: 10
Country: Number of subjects enrolled	France: 137
Country: Number of subjects enrolled	Germany: 117
Country: Number of subjects enrolled	Greece: 27
Country: Number of subjects enrolled	Ireland: 18
Country: Number of subjects enrolled	Israel: 20
Country: Number of subjects enrolled	Italy: 81
Country: Number of subjects enrolled	Korea, Republic of: 9
Country: Number of subjects enrolled	Latvia: 10
Country: Number of subjects enrolled	Lithuania: 15
Country: Number of subjects enrolled	Mexico: 21
Country: Number of subjects enrolled	Poland: 37
Country: Number of subjects enrolled	Portugal: 18
Country: Number of subjects enrolled	Romania: 3

Country: Number of subjects enrolled	Russian Federation: 35
Country: Number of subjects enrolled	Serbia: 10
Country: Number of subjects enrolled	Slovakia: 15
Country: Number of subjects enrolled	Slovenia: 9
Country: Number of subjects enrolled	Spain: 67
Country: Number of subjects enrolled	Sweden: 16
Country: Number of subjects enrolled	Switzerland: 41
Country: Number of subjects enrolled	Thailand: 20
Country: Number of subjects enrolled	Turkey: 75
Country: Number of subjects enrolled	Macedonia, the former Yugoslav Republic of: 3
Country: Number of subjects enrolled	Egypt: 12
Worldwide total number of subjects	972
EEA total number of subjects	608

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)	434
From 65 to 84 years	523
85 years and over	15

Subject disposition

Recruitment

Recruitment details:

The study was conducted at 195 centers in 31 countries

Pre-assignment

Screening details:

A total of 1131 subjects were screened and 979 subjects were enrolled. Due to compliance issues a site in Romania was closed. Seven subjects were excluded from the analysis, because data integrity was impacted by the site's non-compliance. Hence, data analysis is reported for 972 enrolled subjects.

Period 1

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

Arms

Arm title	Obinutuzumab
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Arm description:

Participants received obinutuzumab either alone as single agent, or in combination with chemotherapy (Fludarabine/Cyclophosphamide [FC], Bendamustine or Chlorambucil).

Arm type	Experimental
Investigational medicinal product name	Obinutuzumab
Investigational medicinal product code	
Other name	
Pharmaceutical forms	Concentrate for solution for infusion
Routes of administration	Intravenous use

Dosage and administration details:

Obinutuzumab was administered by intravenous (i.v.) infusion at a dose of 1000 mg on Day 1/2, 8 and 15 of Cycle 1. From Cycle 2 to Cycle 6, it was administered at the same dose on Day 1 only. A treatment cycle was defined as lasting 28 days.

Investigational medicinal product name	Bendamustine
Investigational medicinal product code	
Other name	
Pharmaceutical forms	Powder for solution for infusion
Routes of administration	Intravenous use

Dosage and administration details:

Bendamustine: 90 milligram per millilitre square (mg/m^2) I.V. over 60 minutes once daily (QD) Day 1–2 in subjects previously untreated or $70 \text{ mg}/\text{m}^2$ I.V. over 60 minutes QD Day 1–2 in subjects with relapsed/refractory disease. In nonfit subjects only, investigators may opt at their own discretion to use lower initial doses of bendamustine, i.e., bendamustine $70 \text{ mg}/\text{m}^2$ in previously untreated subjects, and bendamustine $50 \text{ mg}/\text{m}^2$ in relapsed/refractory subjects (over 60 minutes qd Day 1–2 for each administration).

Investigational medicinal product name	Fludarabine
Investigational medicinal product code	
Other name	
Pharmaceutical forms	Concentrate for solution for infusion, Powder for oral suspension
Routes of administration	Intravenous use, Oral use

Dosage and administration details:

Fludarabine 25 mg/m² I.V. over 30 minutes QD Day 1–3 or Fludarabine 40 mg/m² per os (p.o.) QD Day 1–3 in fit subjects only.

Investigational medicinal product name	Cyclophosphamide
Investigational medicinal product code	
Other name	
Pharmaceutical forms	Concentrate for solution for infusion, Powder for solution for infusion, Powder for oral suspension
Routes of administration	Intravenous use, Oral use

Dosage and administration details:

Cyclophosphamide 250 mg/m² I.V. over 15-30 minutes qd Day 1-3 or Cyclophosphamide 250 mg/m² p.o. QD Day 1–3 in fit subjects only.

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

Dosage and administration details:

Chlorambucil 0.5 mg/kg p.o. qd on Day 1 and Day 15 in non-fit subjects only.

Number of subjects in period 1	Obinutuzumab
Started	972
Completed	658
Not completed	314
Adverse event, serious fatal	186
Adverse event, non-fatal	3
Unknown	12
Withdrawal of consent	90
Lost to follow-up	16
Investigator discretion	7

Baseline characteristics

Reporting groups

Reporting group title	Obinutuzumab
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Reporting group description:

Participants received obinutuzumab either alone as single agent, or in combination with chemotherapy (Fludarabine/Cyclophosphamide [FC], Bendamustine or Chlorambucil).

Reporting group values	Obinutuzumab	Total	
Number of subjects	972	972	
Age categorical			
Units: Subjects			
In utero	0	0	
Preterm newborn infants (gestational age < 37 wks)	0	0	
Newborns (0-27 days)	0	0	
Infants and toddlers (28 days-23 months)	0	0	
Children (2-11 years)	0	0	
Adolescents (12-17 years)	0	0	
Adults (18-64 years)	434	434	
From 65-84 years	523	523	
85 years and over	15	15	
Age Continuous			
Units: years			
arithmetic mean	65.4		
standard deviation	± 10.94	-	
Sex: Female, Male			
Units: Subjects			
Female	355	355	
Male	617	617	

Subject analysis sets

Subject analysis set title	G mono: Previously Untreated Fit
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Subject analysis set type	Intention-to-treat
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Subject analysis set description:

Participants with obinutuzumab monotherapy who were previously untreated fit. Fit participants were defined as having a total Cumulative Illness Rating Scale (CIRS) score ≤6 and creatinine clearance (CrCl) ≥70 mL/min.

Subject analysis set title	G mono: Previously Untreated Unfit
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Subject analysis set type	Intention-to-treat
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Subject analysis set description:

Participants with obinutuzumab monotherapy who were previously untreated unfit. Non fit participants were defined as having a CIRS score >6 and/or CrCl <70 mL/min.

Subject analysis set title	G mono: Relapsed/refractory
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Subject analysis set type	Intention-to-treat
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Subject analysis set description:

Participants with obinutuzumab monotherapy who had documented relapsed/refractory chronic lymphocytic leukemia (CLL).

Subject analysis set title	G-Benda: Previously Untreated Fit
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Subject analysis set type	Intention-to-treat
Subject analysis set description: Participants with obinutuzumab in combination with bendamustine who were previously untreated fit. Fit participants were defined as having a total CIRS score ≤ 6 and CrCl ≥ 70 mL/min.	
Subject analysis set title	G-Benda: Previously Untreated Unfit
Subject analysis set type	Intention-to-treat
Subject analysis set description: Participants with obinutuzumab in combination with bendamustine who were previously untreated unfit. Non fit Participants were defined as having a CIRS score > 6 and/or CrCl < 70 mL/min.	
Subject analysis set title	G-Benda: Relapsed/refractory
Subject analysis set type	Intention-to-treat
Subject analysis set description: Participants with obinutuzumab in combination with bendamustine who had documented relapsed/refractory chronic lymphocytic leukemia (CLL).	
Subject analysis set title	G-FC: Previously Untreated Fit
Subject analysis set type	Intention-to-treat
Subject analysis set description: Participants with obinutuzumab in combination with fludarabine and cyclophosphamide who were previously untreated fit. Fit subjects were defined as having a total CIRS score ≤ 6 and CrCl ≥ 70 mL/min.	
Subject analysis set title	G-FC: Previously Untreated Unfit
Subject analysis set type	Intention-to-treat
Subject analysis set description: Participants with obinutuzumab in combination with fludarabine and cyclophosphamide who were previously untreated unfit. Non fit participants were defined as having a CIRS score > 6 and/or CrCl < 70 mL/min. This is a Protocol deviation: unfit subjects were incorrectly assigned to the G-FC treatment regimen.	
Subject analysis set title	G-FC: Relapsed/refractory
Subject analysis set type	Intention-to-treat
Subject analysis set description: Participants with obinutuzumab in combination with fludarabine and cyclophosphamide who had documented relapsed/refractory chronic lymphocytic leukemia (CLL). This is a Protocol deviation: 5 unfit subjects were incorrectly assigned to this relapsed/refractory G-FC treatment regimen.	
Subject analysis set title	G-Clb: Previously Untreated Fit
Subject analysis set type	Intention-to-treat
Subject analysis set description: Participants with obinutuzumab in combination with chlorambucil who were previously untreated fit. Fit participants were defined as having a total CIRS score ≤ 6 and CrCl ≥ 70 mL/min.	
Subject analysis set title	G-Clb: Previously Untreated Unfit
Subject analysis set type	Intention-to-treat
Subject analysis set description: Participants with obinutuzumab in combination with chlorambucil who were previously untreated unfit. Non fit participants were defined as having a CIRS score > 6 and/or CrCl < 70 mL/min.	
Subject analysis set title	G-Clb: Relapsed/refractory
Subject analysis set type	Intention-to-treat
Subject analysis set description: Participants with obinutuzumab in combination with chlorambucil who had documented relapsed/refractory chronic lymphocytic leukemia (CLL).	

Reporting group values	G mono: Previously Untreated Fit	G mono: Previously Untreated Unfit	G mono: Relapsed/refractory
Number of subjects	31	32	65
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)	23	5	20
From 65-84 years	8	25	43
85 years and over	0	2	2
Age Continuous			
Units: years			
arithmetic mean	58.6	74.7	68.6
standard deviation	± 8.26	± 7.94	± 11.88
Sex: Female, Male			
Units: Subjects			
Female	16	18	28
Male	15	14	37

Reporting group values	G-Benda: Previously Untreated Fit	G-Benda: Previously Untreated Unfit	G-Benda: Relapsed/refractory
Number of subjects	168	179	190
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)	96	50	72
From 65-84 years	72	129	115
85 years and over	0	0	3
Age Continuous			
Units: years			
arithmetic mean	61.7	69.2	66.8
standard deviation	± 9.61	± 8.36	± 9.55
Sex: Female, Male			
Units: Subjects			
Female	45	73	66
Male	123	106	124

Reporting group values	G-FC: Previously Untreated Fit	G-FC: Previously Untreated Unfit	G-FC: Relapsed/refractory
Number of subjects	140	13	40
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)	119	8	31
From 65-84 years	21	5	9
85 years and over	0	0	0
Age Continuous			
Units: years			
arithmetic mean	55.8	60.2	56.7
standard deviation	± 8.29	± 8.56	± 8.69
Sex: Female, Male			
Units: Subjects			
Female	45	7	13
Male	95	6	27

Reporting group values	G-Clb: Previously Untreated Fit	G-Clb: Previously Untreated Unfit	G-Clb: Relapsed/refractory
Number of subjects	1	67	46
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	5	5
From 65-84 years	1	58	37
85 years and over	0	4	4
Age Continuous			
Units: years			
arithmetic mean	78.0	75.3	75.5
standard deviation	± 1.00	± 7.71	± 7.76
Sex: Female, Male			
Units: Subjects			
Female	0	28	16
Male	1	39	30

End points

End points reporting groups

Reporting group title	Obinutuzumab
Reporting group description: Participants received obinutuzumab either alone as single agent, or in combination with chemotherapy (Fludarabine/Cyclophosphamide [FC], Bendamustine or Chlorambucil).	
Subject analysis set title	G mono: Previously Untreated Fit
Subject analysis set type	Intention-to-treat
Subject analysis set description: Participants with obinutuzumab monotherapy who were previously untreated fit. Fit participants were defined as having a total Cumulative Illness Rating Scale (CIRS) score ≤ 6 and creatinine clearance (CrCl) ≥ 70 mL/min.	
Subject analysis set title	G mono: Previously Untreated Unfit
Subject analysis set type	Intention-to-treat
Subject analysis set description: Participants with obinutuzumab monotherapy who were previously untreated unfit. Non fit participants were defined as having a CIRS score > 6 and/or CrCl < 70 mL/min.	
Subject analysis set title	G mono: Relapsed/refractory
Subject analysis set type	Intention-to-treat
Subject analysis set description: Participants with obinutuzumab monotherapy who had documented relapsed/refractory chronic lymphocytic leukemia (CLL).	
Subject analysis set title	G-Benda: Previously Untreated Fit
Subject analysis set type	Intention-to-treat
Subject analysis set description: Participants with obinutuzumab in combination with bendamustine who were previously untreated fit. Fit participants were defined as having a total CIRS score ≤ 6 and CrCl ≥ 70 mL/min.	
Subject analysis set title	G-Benda: Previously Untreated Unfit
Subject analysis set type	Intention-to-treat
Subject analysis set description: Participants with obinutuzumab in combination with bendamustine who were previously untreated unfit. Non fit Participants were defined as having a CIRS score > 6 and/or CrCl < 70 mL/min.	
Subject analysis set title	G-Benda: Relapsed/refractory
Subject analysis set type	Intention-to-treat
Subject analysis set description: Participants with obinutuzumab in combination with bendamustine who had documented relapsed/refractory chronic lymphocytic leukemia (CLL).	
Subject analysis set title	G-FC: Previously Untreated Fit
Subject analysis set type	Intention-to-treat
Subject analysis set description: Participants with obinutuzumab in combination with fludarabine and cyclophosphamide who were previously untreated fit. Fit subjects were defined as having a total CIRS score ≤ 6 and CrCl ≥ 70 mL/min.	
Subject analysis set title	G-FC: Previously Untreated Unfit
Subject analysis set type	Intention-to-treat
Subject analysis set description: Participants with obinutuzumab in combination with fludarabine and cyclophosphamide who were previously untreated unfit. Non fit participants were defined as having a CIRS score > 6 and/or CrCl < 70 mL/min. This is a Protocol deviation: unfit subjects were incorrectly assigned to the G-FC treatment regimen.	
Subject analysis set title	G-FC: Relapsed/refractory
Subject analysis set type	Intention-to-treat

Subject analysis set description:

Participants with obinutuzumab in combination with fludarabine and cyclophosphamide who had documented relapsed/refractory chronic lymphocytic leukemia (CLL). This is a Protocol deviation: 5 unfit subjects were incorrectly assigned to this relapsed/refractory G-FC treatment regimen.

Subject analysis set title	G-Clb: Previously Untreated Fit
Subject analysis set type	Intention-to-treat

Subject analysis set description:

Participants with obinutuzumab in combination with chlorambucil who were previously untreated fit. Fit participants were defined as having a total CIRS score ≤ 6 and CrCl ≥ 70 mL/min.

Subject analysis set title	G-Clb: Previously Untreated Unfit
Subject analysis set type	Intention-to-treat

Subject analysis set description:

Participants with obinutuzumab in combination with chlorambucil who were previously untreated unfit. Non fit participants were defined as having a CIRS score > 6 and/or CrCl < 70 mL/min.

Subject analysis set title	G-Clb: Relapsed/refractory
Subject analysis set type	Intention-to-treat

Subject analysis set description:

Participants with obinutuzumab in combination with chlorambucil who had documented relapsed/refractory chronic lymphocytic leukemia (CLL).

Primary: Number of Participants with Adverse Events (AEs)

End point title	Number of Participants with Adverse Events (AEs) ^[1]
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End point description:

An adverse event is any untoward medical occurrence in a participant administered a pharmaceutical product and which does not necessarily have to have a causal relationship with the treatment. An adverse event can therefore be any unfavorable and unintended sign (including an abnormal laboratory finding, for example), symptom, or disease temporally associated with the use of a pharmaceutical product, whether or not considered related to the pharmaceutical product. Preexisting conditions which worsen during a study are also considered as adverse events. AEs, including AEs of Special Interest and AEs of Particular Interest, were reported based on the national cancer institute common terminology criteria for AEs, Version 4.0 (NCI-CTCAE, v4.0). Reported are the number of subjects with AEs, Grade 3-5 AEs, and Serious Adverse Events (SAEs). The safety population was defined as all subjects who have received at least one dose of study medication.

End point type	Primary
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End point timeframe:

Baseline up to time of primary completion (3 years)

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: No statistical analyses were performed as this study has only one arm.

End point values	Obinutuzumab			
Subject group type	Reporting group			
Number of subjects analysed	971			
Units: participants				
AEs	950			
Grade 3-5 AEs	780			
SAEs	516			

Statistical analyses

No statistical analyses for this end point

Primary: Number of Participants with Adverse Events of Special Interest (AESIs)

End point title	Number of Participants with Adverse Events of Special Interest (AESIs) ^[2]
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End point description:

The following AEs were defined as AESIs: AEs with the preferred term Tumour Lysis Syndrome (TLS), Infusion-Related Reactions (IRRs) defined as AEs that occurred during or within 24 hours of the completion of obinutuzumab infusion and were assessed as related to obinutuzumab by the Investigator, Infections defined as AEs from System Organ Class (SOC) "Infections and infestations" and AEs with the preferred term Neutropenia. Reported are number of participants with total AESIs, IRRs, Infections, Neutropenia and TLS. The safety population was defined as all subjects who have received at least one dose of study medication.

End point type	Primary
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End point timeframe:

Baseline up to time of primary completion (3 years)

Notes:

[2] - 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: No statistical analyses were performed as this study has only one arm.

End point values	Obinutuzumab			
Subject group type	Reporting group			
Number of subjects analysed	971			
Units: participants				
Total AESIs	905			
IRRs	635			
Neutropenia	599			
Infections	521			
TLS	62			

Statistical analyses

No statistical analyses for this end point

Primary: Number of Participants with Adverse Events of Particular Interest (AEPIs)

End point title	Number of Participants with Adverse Events of Particular Interest (AEPIs) ^[3]
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End point description:

The following AEs were defined as AEPIs: AEs with the preferred term Progressive multifocal leukoencephalopathy, hepatitis B reactivation defined as AEs with preferred term containing "Hepatitis B" or "hepatitis acute", thrombocytopenia defined via Roche MedDRA basket subgroup "haematopoietic thrombocytopenia", second malignancies defined as AEs from the SOC "Neoplasms benign, malignant and unspecified" starting 6 months after the first study drug intake, second malignancies based on standardised MedDRA queries (SMQ) starting 6 months after the first study drug intake based on the MedDRA SMQ "Malignant or unspecified tumours", in which benign neoplasms are not included, Cardiac events including AEs from the SOC "Cardiac disorders", and hemorrhagic events defined via Roche MedDRA basket subgroup "Haemorrhagic events". The safety population was defined as all subjects who have received at least one dose of study medication.

End point type	Primary
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End point timeframe:

Baseline up to time of primary completion (3 years)

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: No statistical analyses were performed as this study has only one arm.

End point values	Obinutuzumab			
Subject group type	Reporting group			
Number of subjects analysed	971			
Units: participants				
Total AEPs	467			
Thrombocytopenia	314			
Cardiac events	109			
Second malignancies	82			
Hemorrhagic events	69			
Hepatitis B reactivation	3			
PML	1			
Second malignancies (SMQ)	75			

Statistical analyses

No statistical analyses for this end point

Secondary: Percentage of Participants with Overall Response (OR) at Final Response Assessment (FRA)

End point title	Percentage of Participants with Overall Response (OR) at Final Response Assessment (FRA)
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End point description:

OR: percentage of participants with complete response (CR) or CR with incomplete marrow recovery (CRi), or partial response (PR), determined by investigator based on International Workshop on Chronic Lymphocytic Leukemia (IWCLL) tumor response criteria. CR: Peripheral blood lymphocytes 4,000/mcL, no significant lymphadenopathy, no hepatomegaly and splenomegaly, no disease symptoms, blood counts: neutrophils >1,500/mcL, platelets > 100,000/mcL, hemoglobin > 110 g/L and bone marrow normocellular for age. CRi: CR with persistent cytopenia. PR: $\geq$ 50% decrease in peripheral blood lymphocyte count AND $\geq$ 50% reduction in lymphadenopathy OR $\geq$ 50% reduction of liver enlargement OR $\geq$ 50% reduction of spleen PLUS one of the following: neutrophils >1,500/mcL, platelets > 100,000/mcL, Hb > 110 g/L OR $\geq$ 50% increase in neutrophils, platelets or Hb. ITT population was defined as all subjects enrolled in the study regardless of whether or not they received any study drug.

End point type	Secondary
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End point timeframe:

3 months after the last dose of study treatment (up to approximately 5 years)

End point values	G mono: Previously Untreated Fit	G mono: Previously Untreated Unfit	G mono: Relapsed/refractory	G-Benda: Previously Untreated Fit
Subject group type	Subject analysis set	Subject analysis set	Subject analysis set	Subject analysis set
Number of subjects analysed	31	32	65	168
Units: percentage of participants				
number (confidence interval 95%)	71.0 (52.0 to 85.8)	59.4 (40.6 to 76.3)	41.5 (29.4 to 54.4)	83.9 (77.5 to 89.1)

End point values	G-Benda: Previously	G-Benda: Relapsed/refractory	G-FC: Previously	G-FC: Previously
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	Untreated Unfit	Untreated Fit	Untreated Unfit	Untreated Unfit
Subject group type	Subject analysis set	Subject analysis set	Subject analysis set	Subject analysis set
Number of subjects analysed	179	190	140	13
Units: percentage of participants				
number (confidence interval 95%)	81.6 (75.1 to 87.0)	73.2 (66.3 to 79.3)	90.0 (83.8 to 94.4)	84.6 (54.6 to 98.1)

End point values	G-FC: Relapsed/refractory	G-Clb: Previously Untreated Fit	G-Clb: Previously Untreated Unfit	G-Clb: Relapsed/refractory
Subject group type	Subject analysis set	Subject analysis set	Subject analysis set	Subject analysis set
Number of subjects analysed	40	1	67	46
Units: percentage of participants				
number (confidence interval 95%)	85.0 (70.2 to 94.3)	100 (2.5 to 100.0)	82.1 (70.8 to 90.4)	56.5 (41.1 to 71.1)

Statistical analyses

No statistical analyses for this end point

Secondary: Percentage of Participants with Minimal Residual Disease (MRD)-Negativity as Assessed by Flow Cytometry

End point title	Percentage of Participants with Minimal Residual Disease (MRD)-Negativity as Assessed by Flow Cytometry
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End point description:

MRD-negativity was defined as the presence of less than 1 chronic lymphocytic leukemia (CLL) cell per 10,000 leukocytes in blood and bone marrow as assessed by flow cytometry 3 months after last dose of study treatment (i.e. at final response assessment [FRA] visit). The intent-to-ship (ITS) population included all participants from the ITT population whose MRD samples at the FRA could be shipped to the central laboratory within 48 hours. MRD ITS population was a part of the intent-to-treat population. Number of subjects analyzed indicates subjects who took part in the analysis. 9999 = Not estimable.

End point type	Secondary
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End point timeframe:

3 months after the last dose of study treatment (up to approximately 5 years)

End point values	G mono: Previously Untreated Fit	G mono: Previously Untreated Unfit	G mono: Relapsed/refractory	G-Benda: Previously Untreated Fit
Subject group type	Subject analysis set	Subject analysis set	Subject analysis set	Subject analysis set
Number of subjects analysed	24	26	49	149
Units: percentage of participants				
number (not applicable)				
Blood	8.3	23.1	4.1	63.1
Bone Marrow	4.2	3.8	2.0	31.5

End point values	G-Benda: Previously Untreated Unfit	G-Benda: Relapsed/refra ctory	G-FC: Previously Untreated Fit	G-FC: Previously Untreated Unfit
Subject group type	Subject analysis set	Subject analysis set	Subject analysis set	Subject analysis set
Number of subjects analysed	147	161	125	12
Units: percentage of participants				
number (not applicable)				
Blood	65.3	39.8	72.0	58.3
Bone Marrow	27.2	14.9	40.0	41.7

End point values	G-FC: Relapsed/refra ctory	G-Clb: Previously Untreated Fit	G-Clb: Previously Untreated Unfit	G-Clb: Relapsed/refra ctory
Subject group type	Subject analysis set	Subject analysis set	Subject analysis set	Subject analysis set
Number of subjects analysed	33	9999 ^[4]	21	8
Units: percentage of participants				
number (not applicable)				
Blood	51.5	9999	9.4	6.3
Bone Marrow	24.2	9999	5.7	3.1

Notes:

[4] - MRD- negativity could not be estimated as no patients had an event.

Statistical analyses

No statistical analyses for this end point

Secondary: Percentage of Participants with Best Overall Response (BOR)

End point title	Percentage of Participants with Best Overall Response (BOR)
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End point description:

BOR was defined as percentage of participants with best response obtained throughout the trial with CR, CRi, or PR, as determined by the investigator based on IWCLL tumor response criteria. CR: Peripheral blood lymphocytes 4,000/mcL, no significant lymphadenopathy, no hepatomegaly and splenomegaly, no disease symptoms, blood counts: neutrophils >1,500/mcL, platelets > 100,000/mcL, hemoglobin > 110 g/L and bone marrow normocellular for age. Cri: CR with persistent cytopenia. PR: >= 50% decrease in peripheral blood lymphocyte count AND >= 50% reduction in lymphadenopathy OR >= 50% reduction of liver enlargement OR >= 50% reduction of spleen PLUS one of the following: neutrophils >1,500/mcL, platelets > 100,000/mcL, Hb > 110 g/L OR >= 50% increase in neutrophils, platelets or Hb. ITT population was defined as all subjects enrolled in the study regardless of whether or not they received any study drug. Number of subjects analysed indicates subjects who took part in the analysis.

End point type	Secondary
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End point timeframe:

Baseline, Day 85, end of treatment or early termination, and follow-up, assessed up to disease progression or death, whichever occurs first (up to approximately 5 years)

End point values	G mono: Previously Untreated Fit	G mono: Previously Untreated Unfit	G mono: Relapsed/refractory	G-Benda: Previously Untreated Fit
Subject group type	Subject analysis set	Subject analysis set	Subject analysis set	Subject analysis set
Number of subjects analysed	31	32	65	168
Units: percentage of participants				
number (confidence interval 95%)	83.9 (66.3 to 94.5)	71.9 (53.3 to 86.3)	60.0 (47.1 to 72.0)	91.7 (86.4 to 95.4)

End point values	G-Benda: Previously Untreated Unfit	G-Benda: Relapsed/refractory	G-FC: Previously Untreated Fit	G-FC: Previously Untreated Unfit
Subject group type	Subject analysis set	Subject analysis set	Subject analysis set	Subject analysis set
Number of subjects analysed	179	190	140	13
Units: percentage of participants				
number (confidence interval 95%)	93.9 (89.3 to 96.9)	86.8 (81.2 to 91.3)	97.1 (92.8 to 99.2)	84.6 (54.6 to 98.1)

End point values	G-FC: Relapsed/refractory	G-Clb: Previously Untreated Fit	G-Clb: Previously Untreated Unfit	G-Clb: Relapsed/refractory
Subject group type	Subject analysis set	Subject analysis set	Subject analysis set	Subject analysis set
Number of subjects analysed	40	1	67	46
Units: percentage of participants				
number (confidence interval 95%)	97.5 (86.8 to 99.9)	100 (2.5 to 100.0)	94.0 (85.4 to 98.3)	84.8 (71.1 to 93.7)

Statistical analyses

No statistical analyses for this end point

Secondary: Median Time to Progression-Free Survival (PFS)

End point title	Median Time to Progression-Free Survival (PFS)
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End point description:

PFS was defined as time from the date of enrollment until the first documented progressive disease (PD) based on IWCLL tumor response criteria or death from any cause, whichever occurred first. PD: at least one of the following:
 >= 50% increase in the absolute number of circulating lymphocytes to at least 5,000/mcL, appearance of new palpable lymph nodes, >= 50% increase in the longest diameter of any previous site of clinically significant lymphadenopathy, >= 50% increase in the enlargement of the liver and/or spleen, transformation to more aggressive histology, progression of any cytopenia, decrease of Hb levels by > 20 g/L or to less than 100 g/L, decrease of platelet counts by > 50% or to < 100,000 /mcL, decrease of neutrophil counts by > 50% or to < 1,000/mcL. The ITT population was defined as all subjects enrolled in the study regardless of whether or not they received any study drug. 0000/9999 = Not estimable.

End point type	Secondary
End point timeframe:	
Baseline, Day 85, end of treatment or early termination, and follow-up, assessed up to disease progression or death, whichever occurs first (up to approximately 5 years)	

End point values	G mono: Previously Untreated Fit	G mono: Previously Untreated Unfit	G mono: Relapsed/refractory	G-Benda: Previously Untreated Fit
Subject group type	Subject analysis set	Subject analysis set	Subject analysis set	Subject analysis set
Number of subjects analysed	31 ^[5]	32 ^[6]	65	168
Units: months				
median (confidence interval 95%)	43.0 (17.0 to 9999)	21.2 (14.9 to 9999)	17.6 (12.5 to 20.0)	58.0 (44.5 to 58.0)

Notes:

[5] - Upper Limit 95% CI could not be estimated as too few patients had an event.

[6] - Upper Limit 95% CI could not be estimated as too few patients had an event.

End point values	G-Benda: Previously Untreated Unfit	G-Benda: Relapsed/refractory	G-FC: Previously Untreated Fit	G-FC: Previously Untreated Unfit
Subject group type	Subject analysis set	Subject analysis set	Subject analysis set	Subject analysis set
Number of subjects analysed	179 ^[7]	190 ^[8]	140	13 ^[9]
Units: months				
median (confidence interval 95%)	9999 (45.4 to 9999)	28.6 (24.2 to 31.1)	9999 (53.7 to 9999)	9999 (19.0 to 9999)

Notes:

[7] - Median and Upper Limit 95% CI could not be estimated as too few patients had an event.

[8] - Median and Upper Limit 95% CI could not be estimated as too few patients had an event.

[9] - Median and Upper Limit 95% CI could not be estimated as too few patients had an event.

End point values	G-FC: Relapsed/refractory	G-Clb: Previously Untreated Fit	G-Clb: Previously Untreated Unfit	G-Clb: Relapsed/refractory
Subject group type	Subject analysis set	Subject analysis set	Subject analysis set	Subject analysis set
Number of subjects analysed	40	1 ^[10]	67	46
Units: months				
median (confidence interval 95%)	24.8 (19.3 to 35.0)	31.3 (0000 to 9999)	31.8 (27.7 to 36.4)	14.1 (10.8 to 22.1)

Notes:

[10] - Lower and Upper Limit 95% CI could not be estimated as too few patients had an event.

Statistical analyses

No statistical analyses for this end point

Secondary: Median Time to Response (TTR)

End point title	Median Time to Response (TTR)
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End point description:

Kaplan Meier estimate of median TTR was defined as the time at which half of the participants reached CR or PR based on IWCLL tumor response criteria. CR: Peripheral blood lymphocytes 4,000/mcL, no significant lymphadenopathy, no hepatomegaly and splenomegaly, no disease symptoms, blood counts: neutrophils >1,500/mcL, platelets > 100,000/mcL, hemoglobin > 110 g/L and bone marrow normocellular for age. PR: >/= 50% decrease in peripheral blood lymphocyte count AND >/= 50%

reduction in lymphadenopathy OR $\geq$ 50% reduction of liver enlargement OR $\geq$ 50% reduction of spleen PLUS one of the following: neutrophils $>1,500/\text{mL}$, platelets $> 100,000/\text{mL}$, hemoglobin $> 110 \text{ g/L}$ OR $\geq$ 50% increase in neutrophils, platelets or hemoglobin. 0000/9999 = Not estimable. The ITT population was defined as all subjects enrolled in the study regardless of whether or not they received any study drug.

End point type	Secondary
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End point timeframe:

Baseline, Day 85, end of treatment or early termination, and follow-up, assessed up to disease progression or death, whichever occurs first (up to approximately 5 years)

End point values	G mono: Previously Untreated Fit	G mono: Previously Untreated Unfit	G mono: Relapsed/refractory	G-Benda: Previously Untreated Fit
Subject group type	Subject analysis set	Subject analysis set	Subject analysis set	Subject analysis set
Number of subjects analysed	31	32	65	168
Units: months				
median (confidence interval 95%)	3.6 (3.2 to 3.8)	3.6 (3.3 to 3.9)	3.9 (3.7 to 4.4)	3.5 (3.4 to 3.6)

End point values	G-Benda: Previously Untreated Unfit	G-Benda: Relapsed/refractory	G-FC: Previously Untreated Fit	G-FC: Previously Untreated Unfit
Subject group type	Subject analysis set	Subject analysis set	Subject analysis set	Subject analysis set
Number of subjects analysed	179	190	140	13
Units: months				
median (confidence interval 95%)	3.5 (3.5 to 3.7)	3.7 (3.5 to 3.7)	3.6 (3.5 to 3.7)	4.1 (3.3 to 7.9)

End point values	G-FC: Relapsed/refractory	G-Clb: Previously Untreated Fit	G-Clb: Previously Untreated Unfit	G-Clb: Relapsed/refractory
Subject group type	Subject analysis set	Subject analysis set	Subject analysis set	Subject analysis set
Number of subjects analysed	40	1 ^[11]	67	46
Units: months				
median (confidence interval 95%)	3.6 (3.4 to 3.7)	3.3 (0000 to 9999)	3.6 (3.4 to 3.7)	3.7 (3.5 to 3.8)

Notes:

[11] - Lower and Upper Limit 95% CI could not be estimated as too few patients had an event.

Statistical analyses

No statistical analyses for this end point

Secondary: Median Time to Event-Free Survival (EFS)

End point title	Median Time to Event-Free Survival (EFS)
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End point description:

EFS: time from enrollment to first event of PD or relapse, as assessed by the investigator based on IWCLL criteria, or start of a non-protocol-specified anti-leukemia therapy or death, whichever occurs first. PD: at least one of the following: $\geq$ 50% increase in the absolute number of circulating

lymphocytes to at least 5,000/mcL, appearance of new palpable lymph nodes, $\geq 50\%$ increase in the longest diameter of any previous site of clinically significant lymphadenopathy, $\geq 50\%$ increase in the enlargement of the liver and/or spleen, transformation to more aggressive histology, progression of any cytopenia, decrease of Hb levels by more than 20 g/L or to less than 100 g/L, decrease of platelet counts by more than 50% or to $< 100,000$ /mcL, decrease of neutrophil counts by $> 50\%$ or to $< 1,000$ /mcL. 0000/9999 = Not estimable. ITT population was defined as all subjects enrolled in the study regardless of whether or not they received any study drug.

End point type	Secondary
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End point timeframe:

Baseline, Day 85, end of treatment or early termination, and follow-up, assessed up to disease progression or death, whichever occurs first (up to approximately 5 years)

End point values	G mono: Previously Untreated Fit	G mono: Previously Untreated Unfit	G mono: Relapsed/refra ctory	G-Benda: Previously Untreated Fit
Subject group type	Subject analysis set	Subject analysis set	Subject analysis set	Subject analysis set
Number of subjects analysed	31 ^[12]	32 ^[13]	65	168
Units: months				
median (confidence interval 95%)	35.2 (15.6 to 9999)	17.9 (14.5 to 9999)	14.0 (9.1 to 17.9)	58.0 (44.5 to 58.0)

Notes:

[12] - Upper Limit 95% CI could not be estimated as too few patients had an event.

[13] - Upper Limit 95% CI could not be estimated as too few patients had an event.

End point values	G-Benda: Previously Untreated Unfit	G-Benda: Relapsed/refra ctory	G-FC: Previously Untreated Fit	G-FC: Previously Untreated Unfit
Subject group type	Subject analysis set	Subject analysis set	Subject analysis set	Subject analysis set
Number of subjects analysed	179 ^[14]	190	140 ^[15]	13 ^[16]
Units: months				
median (confidence interval 95%)	52.9 (44.7 to 9999)	25.1 (21.0 to 30.4)	9999 (53.7 to 9999)	9999 (17.8 to 9999)

Notes:

[14] - Upper Limit 95% CI could not be estimated as too few patients had an event.

[15] - Median and corresponding 95% CI could not be estimated as too few patients had an event.

[16] - Median and Upper Limit 95% CI could not be estimated as too few patients had an event.

End point values	G-FC: Relapsed/refra ctory	G-Clb: Previously Untreated Fit	G-Clb: Previously Untreated Unfit	G-Clb: Relapsed/refra ctory
Subject group type	Subject analysis set	Subject analysis set	Subject analysis set	Subject analysis set
Number of subjects analysed	40	1 ^[17]	67	46
Units: months				
median (confidence interval 95%)	24.2 (18.7 to 33.8)	31.3 (0000 to 9999)	31.8 (27.7 to 36.4)	13.7 (10.5 to 18.0)

Notes:

[17] - Lower and Upper Limit 95% CI could not be estimated as too few patients had an event.

Statistical analyses

No statistical analyses for this end point

Secondary: Median Time to Overall Survival (OS)

End point title	Median Time to Overall Survival (OS)
End point description:	
Kaplan Meier estimate of median OS was defined as the time at which half of the participants had died, regardless of the cause of death. 9999 = Not estimable. The ITT population was defined as all subjects enrolled in the study regardless of whether or not they received any study drug.	
End point type	Secondary
End point timeframe:	
Baseline until death (Approximately up to 5 years)	

End point values	G mono: Previously Untreated Fit	G mono: Previously Untreated Unfit	G mono: Relapsed/refra ctory	G-Benda: Previously Untreated Fit
Subject group type	Subject analysis set	Subject analysis set	Subject analysis set	Subject analysis set
Number of subjects analysed	31 ^[18]	32 ^[19]	65 ^[20]	168 ^[21]
Units: months				
median (confidence interval 95%)	9999 (9999 to 9999)	9999 (40.7 to 9999)	9999 (42.1 to 9999)	9999 (9999 to 9999)

Notes:

[18] - Median, Lower and Upper Limit 95% CI could not be estimated as too few patients had an event.

[19] - Median and Upper Limit 95% CI could not be estimated as too few patients had an event.

[20] - Median and Upper Limit 95% CI could not be estimated as too few patients had an event.

[21] - Median, Lower and Upper Limit 95% CI could not be estimated as too few patients had an event.

End point values	G-Benda: Previously Untreated Unfit	G-Benda: Relapsed/refra ctory	G-FC: Previously Untreated Fit	G-FC: Previously Untreated Unfit
Subject group type	Subject analysis set	Subject analysis set	Subject analysis set	Subject analysis set
Number of subjects analysed	179 ^[22]	190 ^[23]	140 ^[24]	13 ^[25]
Units: months				
median (confidence interval 95%)	9999 (9999 to 9999)	9999 (55.3 to 9999)	9999 (9999 to 9999)	9999 (9999 to 9999)

Notes:

[22] - Median, Lower and Upper Limit 95% CI could not be estimated as too few patients had an event.

[23] - Median and Upper Limit 95% CI could not be estimated as too few patients had an event.

[24] - Median, Lower and Upper Limit 95% CI could not be estimated as too few patients had an event.

[25] - Median, Lower and Upper Limit 95% CI could not be estimated as too few patients had an event.

End point values	G-FC: Relapsed/refra ctory	G-Clb: Previously Untreated Fit	G-Clb: Previously Untreated Unfit	G-Clb: Relapsed/refra ctory
Subject group type	Subject analysis set	Subject analysis set	Subject analysis set	Subject analysis set
Number of subjects analysed	40 ^[26]	1 ^[27]	67 ^[28]	46 ^[29]
Units: months				
median (confidence interval 95%)	9999 (9999 to 9999)	9999 (9999 to 9999)	9999 (9999 to 9999)	9999 (35.2 to 9999)

Notes:

[26] - Median, Lower and Upper Limit 95% CI could not be estimated as too few patients had an event.

[27] - Median, Lower and Upper Limit 95% CI could not be estimated as too few patients had an event.

[28] - Median, Lower and Upper Limit 95% CI could not be estimated as too few patients had an event.

[29] - Median and Upper Limit 95% CI could not be estimated as too few patients had an event.

Statistical analyses

No statistical analyses for this end point

Secondary: Median Time to New Anti-Leukemia Therapy (TTNT)

End point title	Median Time to New Anti-Leukemia Therapy (TTNT)
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End point description:

Kaplan Meier estimate of median TTNT was defined as the time at which half of the participants have initiated a new anti-leukemic therapy. 9999 = Not estimable. The ITT population was defined as all subjects enrolled in the study regardless of whether or not they received any study drug.

End point type	Secondary
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End point timeframe:

Baseline until end of study (up to approximately 5 years)

End point values	G mono: Previously Untreated Fit	G mono: Previously Untreated Unfit	G mono: Relapsed/refra ctory	G-Benda: Previously Untreated Fit
Subject group type	Subject analysis set	Subject analysis set	Subject analysis set	Subject analysis set
Number of subjects analysed	31 ^[30]	32 ^[31]	65	168 ^[32]
Units: months				
median (confidence interval 95%)	9999 (25.0 to 9999)	9999 (27.5 to 9999)	22.5 (14.8 to 27.0)	9999 (53.5 to 9999)

Notes:

[30] - Median and Upper Limit 95% CI could not be estimated as too few patients had an event.

[31] - Median and Upper Limit 95% CI could not be estimated as too few patients had an event.

[32] - Median and Upper Limit 95% CI could not be estimated as too few patients had an event.

End point values	G-Benda: Previously Untreated Unfit	G-Benda: Relapsed/refra ctory	G-FC: Previously Untreated Fit	G-FC: Previously Untreated Unfit
Subject group type	Subject analysis set	Subject analysis set	Subject analysis set	Subject analysis set
Number of subjects analysed	179 ^[33]	190	140 ^[34]	13 ^[35]
Units: months				
median (confidence interval 95%)	9999 (9999 to 9999)	38.3 (31.5 to 42.0)	9999 (9999 to 9999)	9999 (35.0 to 9999)

Notes:

[33] - Median, Lower and Upper Limit 95% CI could not be estimated as too few patients had an event.

[34] - Median, Lower and Upper Limit 95% CI could not be estimated as too few patients had an event.

[35] - Median and Upper Limit 95% CI could not be estimated as too few patients had an event.

End point values	G-FC: Relapsed/refra ctory	G-Clb: Previously Untreated Fit	G-Clb: Previously Untreated Unfit	G-Clb: Relapsed/refra ctory
Subject group type	Subject analysis set	Subject analysis set	Subject analysis set	Subject analysis set
Number of subjects analysed	40 ^[36]	1 ^[37]	67 ^[38]	46
Units: months				
median (confidence interval 95%)	32.6 (28.2 to 9999)	9999 (9999 to 9999)	53.7 (37.1 to 9999)	20.4 (13.7 to 43.3)

Notes:

[36] - Upper Limit 95% CI could not be estimated as too few patients had an event.

[37] - Median, Lower and Upper Limit 95% CI could not be estimated as too few patients had an event.

[38] - Upper Limit 95% CI could not be estimated as too few patients had an event.

Statistical analyses

No statistical analyses for this end point

Secondary: Median Time to Duration of Response (DoR)

End point title	Median Time to Duration of Response (DoR)
End point description:	
DoR was defined as the period from the date of initial confirmed PR or CR until the date of PD or death from any cause. PR: $\geq 50\%$ decrease in peripheral blood lymphocyte count AND $\geq 50\%$ reduction in lymphadenopathy OR $\geq 50\%$ reduction of liver enlargement OR $\geq 50\%$ reduction of spleen PLUS one of the following: neutrophils $>1,500/\text{mCL}$, platelets $>100,000/\text{mCL}$, hemoglobin $>110\text{ g/L}$ OR $\geq 50\%$ increase in neutrophils, platelets or hemoglobin. CR: Peripheral blood lymphocytes $4,000/\text{mCL}$, no significant lymphadenopathy, no hepatomegaly and splenomegaly, no disease symptoms, blood counts: neutrophils $>1,500/\text{mCL}$, platelets $>100,000/\text{mCL}$, hemoglobin $>110\text{ g/L}$ and bone marrow normocellular for age. PD: as defined in the description for Event-Free Survival outcome measure. 0000/9999 = Not estimable. The ITT population was defined as all subjects enrolled in the study regardless of whether or not they received any study drug.	
End point type	Secondary
End point timeframe:	
Baseline, Day 85, end of treatment or early termination, and follow-up, assessed up to disease progression or death, whichever occurs first (up to approximately 5 years)	

End point values	G mono: Previously Untreated Fit	G mono: Previously Untreated Unfit	G mono: Relapsed/refractory	G-Benda: Previously Untreated Fit
Subject group type	Subject analysis set	Subject analysis set	Subject analysis set	Subject analysis set
Number of subjects analysed	31 ^[39]	32 ^[40]	65	168
Units: months				
median (confidence interval 95%)	40.1 (26.3 to 9999)	20.1 (12.1 to 9999)	15.0 (12.1 to 19.9)	55.0 (44.2 to 55.0)

Notes:

[39] - Upper Limit 95% CI could not be estimated as too few patients had an event.

[40] - Upper Limit 95% CI could not be estimated as too few patients had an event.

End point values	G-Benda: Previously Untreated Unfit	G-Benda: Relapsed/refractory	G-FC: Previously Untreated Fit	G-FC: Previously Untreated Unfit
Subject group type	Subject analysis set	Subject analysis set	Subject analysis set	Subject analysis set
Number of subjects analysed	179 ^[41]	190	140 ^[42]	13 ^[43]
Units: months				
median (confidence interval 95%)	49.3 (44.2 to 9999)	25.5 (21.2 to 29.7)	9999 (50.1 to 9999)	9999 (48.3 to 9999)

Notes:

[41] - Upper Limit 95% CI could not be estimated as too few patients had an event.

[42] - Median and Upper Limit 95% CI could not be estimated as too few patients had an event.

[43] - Median and Upper Limit 95% CI could not be estimated as too few patients had an event.

End point values	G-FC: Relapsed/refractory	G-Clb: Previously Untreated Fit	G-Clb: Previously Untreated Unfit	G-Clb: Relapsed/refractory
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Subject group type	Subject analysis set	Subject analysis set	Subject analysis set	Subject analysis set
Number of subjects analysed	40	1 ^[44]	67	46
Units: months				
median (confidence interval 95%)	21.2 (15.1 to 32.1)	28.1 (0000 to 9999)	28.1 (24.3 to 34.8)	12.3 (7.3 to 20.6)

Notes:

[44] - Lower and Upper Limit 95% CI could not be estimated as too few patients had an event.

Statistical analyses

No statistical analyses for this end point

Adverse events

Adverse events information

Timeframe for reporting adverse events:

Up to 5 years

Adverse event reporting additional description:

The safety population was defined as all participants who received at least one dose of study medication. AEs that were entered into the data base at the time of the data base lock were included in the AE analysis.

Assessment type	Systematic
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Dictionary used

Dictionary name	MedDRA
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Dictionary version	21.1
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Reporting groups

Reporting group title	G-Mono
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Reporting group description:

Participants received obinutuzumab alone as single agent.

Reporting group title	G-FC
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Reporting group description:

Participants received obinutuzumab in combination with fludarabine/cyclophosphamide (FC).

Reporting group title	G-Clb
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Reporting group description:

Participants received obinutuzumab in combination with chlorambucil.

Reporting group title	G-Benda
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Reporting group description:

Participants received obinutuzumab in combination with bendamustine.

Serious adverse events	G-Mono	G-FC	G-Clb
Total subjects affected by serious adverse events			
subjects affected / exposed	71 / 127 (55.91%)	107 / 193 (55.44%)	66 / 114 (57.89%)
number of deaths (all causes)	28	20	36
number of deaths resulting from adverse events			
Neoplasms benign, malignant and unspecified (incl cysts and polyps)			
Adenocarcinoma of colon			
subjects affected / exposed	1 / 127 (0.79%)	1 / 193 (0.52%)	0 / 114 (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
Bladder transitional cell carcinoma			
subjects affected / exposed	1 / 127 (0.79%)	0 / 193 (0.00%)	0 / 114 (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

Cholangiocarcinoma			
subjects affected / exposed	1 / 127 (0.79%)	0 / 193 (0.00%)	0 / 114 (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
Clear cell renal cell carcinoma			
subjects affected / exposed	1 / 127 (0.79%)	0 / 193 (0.00%)	0 / 114 (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
Metastatic carcinoma of the bladder			
subjects affected / exposed	1 / 127 (0.79%)	0 / 193 (0.00%)	0 / 114 (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
Richter's syndrome			
subjects affected / exposed	1 / 127 (0.79%)	1 / 193 (0.52%)	2 / 114 (1.75%)
occurrences causally related to treatment / all	0 / 1	0 / 1	0 / 2
deaths causally related to treatment / all	0 / 1	0 / 1	0 / 2
Spinal meningioma benign			
subjects affected / exposed	1 / 127 (0.79%)	0 / 193 (0.00%)	0 / 114 (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
Squamous cell carcinoma			
subjects affected / exposed	1 / 127 (0.79%)	2 / 193 (1.04%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 1	1 / 2	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Squamous cell carcinoma of skin			
subjects affected / exposed	1 / 127 (0.79%)	0 / 193 (0.00%)	2 / 114 (1.75%)
occurrences causally related to treatment / all	0 / 1	0 / 0	1 / 2
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Adenocarcinoma			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Adenocarcinoma gastric			

subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	1 / 114 (0.88%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Adenosquamous cell lung cancer			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Basal cell carcinoma			
subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	2 / 114 (1.75%)
occurrences causally related to treatment / all	0 / 0	1 / 1	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Bladder cancer stage 0, with cancer in situ			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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	1 / 127 (0.79%)	1 / 193 (0.52%)	0 / 114 (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
Acute myeloid leukaemia			
subjects affected / exposed	0 / 127 (0.00%)	2 / 193 (1.04%)	1 / 114 (0.88%)
occurrences causally related to treatment / all	0 / 0	1 / 2	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 1	0 / 1
Breast cancer			
subjects affected / exposed	0 / 127 (0.00%)	3 / 193 (1.55%)	1 / 114 (0.88%)
occurrences causally related to treatment / all	0 / 0	0 / 3	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Metastatic renal cell carcinoma			
subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	0 / 114 (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
Myelodysplastic syndrome			

subjects affected / exposed	0 / 127 (0.00%)	3 / 193 (1.55%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	2 / 3	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Prostate cancer			
subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	0 / 114 (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			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	1 / 114 (0.88%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Malignant melanoma			
subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	1 / 114 (0.88%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Nasal cavity cancer			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	1 / 114 (0.88%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Primary myelofibrosis			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	1 / 114 (0.88%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Rectal adenocarcinoma			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	1 / 114 (0.88%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Bronchial carcinoma			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 myeloid leukaemia			

subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Endometrial cancer			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 carcinoma			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Intraductal proliferative breast lesion			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Laryngeal cancer			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 adenocarcinoma			
subjects affected / exposed	1 / 127 (0.79%)	0 / 193 (0.00%)	1 / 114 (0.88%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 1	0 / 0	0 / 0
Lung cancer metastatic			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 bronchial carcinoma			

subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 malignant melanoma			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	1 / 114 (0.88%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Metastatic squamous cell carcinoma			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Myelofibrosis			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Neoplasm malignant			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pancreatic carcinoma metastatic			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Prostatic adenoma			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 cancer			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 cell lung cancer metastatic			

subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Basosquamous Carcinoma of Skin			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	1 / 114 (0.88%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Bladder Neoplasm			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hodgkin's Disease			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	1 / 114 (0.88%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 1
Intraductal Papillary Mucinous Neoplasm			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Intraocular Melanoma			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Liver Carcinoma Ruptured			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 Tumour			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 T-Cell Lymphoma Unspecified			

subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 Cell Carcinoma			
subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	0 / 114 (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 Breast Carcinoma			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Transitional Cell Carcinoma			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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			
Hypertension			
subjects affected / exposed	2 / 127 (1.57%)	1 / 193 (0.52%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 2	1 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hypotension			
subjects affected / exposed	4 / 127 (3.15%)	1 / 193 (0.52%)	1 / 114 (0.88%)
occurrences causally related to treatment / all	4 / 4	1 / 1	1 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Peripheral arterial occlusive disease			
subjects affected / exposed	1 / 127 (0.79%)	0 / 193 (0.00%)	0 / 114 (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
Peripheral artery thrombosis			
subjects affected / exposed	1 / 127 (0.79%)	0 / 193 (0.00%)	0 / 114 (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
Flushing			

subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hypertensive crisis			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Orthostatic hypotension			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 vascular disorder			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Shock haemorrhagic			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Vena cava thrombosis			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Thrombophlebitis superficial			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Arterial Rupture			

subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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			
Death			
subjects affected / exposed	2 / 127 (1.57%)	1 / 193 (0.52%)	4 / 114 (3.51%)
occurrences causally related to treatment / all	0 / 2	0 / 1	0 / 4
deaths causally related to treatment / all	0 / 2	0 / 1	0 / 4
General physical health deterioration			
subjects affected / exposed	1 / 127 (0.79%)	0 / 193 (0.00%)	1 / 114 (0.88%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 1
Malaise			
subjects affected / exposed	1 / 127 (0.79%)	0 / 193 (0.00%)	0 / 114 (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
Pyrexia			
subjects affected / exposed	4 / 127 (3.15%)	11 / 193 (5.70%)	2 / 114 (1.75%)
occurrences causally related to treatment / all	1 / 7	6 / 11	1 / 2
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Asthenia			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Euthanasia			

subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hyperthermia			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 peripheral			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Catheter Site Thrombosis			
subjects affected / exposed	1 / 127 (0.79%)	0 / 193 (0.00%)	0 / 114 (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
Mucosal Inflammation			
subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	0 / 114 (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			
Anaphylactic reaction			
subjects affected / exposed	4 / 127 (3.15%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	4 / 5	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cytokine release syndrome			
subjects affected / exposed	5 / 127 (3.94%)	2 / 193 (1.04%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	5 / 5	2 / 2	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Allergy to immunoglobulin therapy			

subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 hypersensitivity			
subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	0 / 114 (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
Hypersensitivity			
subjects affected / exposed	0 / 127 (0.00%)	3 / 193 (1.55%)	1 / 114 (0.88%)
occurrences causally related to treatment / all	0 / 0	3 / 3	1 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Autoimmune disorder			
subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	0 / 114 (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
Haemophagocytic			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Reproductive system and breast disorders			
Uterine Prolapse			
subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	0 / 114 (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
Respiratory, thoracic and mediastinal disorders			
Dyspnoea			
subjects affected / exposed	2 / 127 (1.57%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	2 / 2	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pulmonary oedema			
subjects affected / exposed	1 / 127 (0.79%)	0 / 193 (0.00%)	0 / 114 (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

Bronchiectasis			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Chylothorax			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cough			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hypoxia			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Interstitial lung disease			
subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	0 / 114 (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
Lung disorder			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Obliterative bronchiolitis			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pharyngeal inflammation			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pleural effusion			

subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pleurisy			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 / 127 (0.00%)	0 / 193 (0.00%)	1 / 114 (0.88%)
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	0 / 127 (0.00%)	0 / 193 (0.00%)	1 / 114 (0.88%)
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 embolism			
subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	1 / 114 (0.88%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pulmonary necrosis			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 alkalosis			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Aspiration			

subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	0 / 114 (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
Organising pneumonia			
subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	1 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	1 / 1	0 / 0
Oropharyngeal pain			
subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	0 / 114 (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
Acute respiratory distress syndrome			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	1 / 114 (0.88%)
occurrences causally related to treatment / all	0 / 0	0 / 0	1 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	1 / 1
Acute Pulmonary Oedema			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Asthma			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 Obstructive Pulmonary Disease			
subjects affected / exposed	1 / 127 (0.79%)	0 / 193 (0.00%)	0 / 114 (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
Psychiatric disorders			
Completed suicide			
subjects affected / exposed	1 / 127 (0.79%)	0 / 193 (0.00%)	0 / 114 (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
Alcohol withdrawal syndrome			

subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 / 127 (0.00%)	0 / 193 (0.00%)	1 / 114 (0.88%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Investigations			
Hepatic enzyme increased			
subjects affected / exposed	1 / 127 (0.79%)	0 / 193 (0.00%)	0 / 114 (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
Oxygen saturation decreased			
subjects affected / exposed	1 / 127 (0.79%)	0 / 193 (0.00%)	0 / 114 (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
Weight decreased			
subjects affected / exposed	1 / 127 (0.79%)	0 / 193 (0.00%)	0 / 114 (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
Alanine aminotransferase increased			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Aspartate aminotransferase increased			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 pressure increased			

subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Gamma-glutamyltransferase increased			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Neutrophil count decreased			
subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	0 / 114 (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
Transaminases increased			
subjects affected / exposed	0 / 127 (0.00%)	2 / 193 (1.04%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	2 / 2	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Injury, poisoning and procedural complications			
Femoral neck fracture			
subjects affected / exposed	1 / 127 (0.79%)	0 / 193 (0.00%)	0 / 114 (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
Thoracic vertebral fracture			
subjects affected / exposed	1 / 127 (0.79%)	1 / 193 (0.52%)	0 / 114 (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
Ankle fracture			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cervical vertebral fracture			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Craniocerebral injury			

subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Femur fracture			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	2 / 114 (1.75%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Foot fracture			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 / 127 (0.00%)	0 / 193 (0.00%)	1 / 114 (0.88%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Radiation skin injury			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Rib fracture			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Road traffic accident			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 compression fracture			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	1 / 114 (0.88%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Toxicity to various agents			

subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Wrist fracture			
subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	0 / 114 (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
Fall			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	1 / 114 (0.88%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Head injury			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	1 / 114 (0.88%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 1
Radiation mucositis			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	1 / 114 (0.88%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Dislocation of Vertebra			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Fractured Ischium			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	1 / 114 (0.88%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hand Fracture			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 Injuries			

subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 Haematoma			
subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	0 / 114 (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
Post Procedural Haemorrhage			
subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	0 / 114 (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
Radius Fracture			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Skull Fracture			
subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	0 / 114 (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
Subdural Haematoma			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 Fracture			
subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	0 / 114 (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
Cardiac disorders			
Atrial fibrillation			
subjects affected / exposed	3 / 127 (2.36%)	0 / 193 (0.00%)	1 / 114 (0.88%)
occurrences causally related to treatment / all	1 / 3	0 / 0	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cardiac failure			

subjects affected / exposed	3 / 127 (2.36%)	0 / 193 (0.00%)	1 / 114 (0.88%)
occurrences causally related to treatment / all	1 / 3	0 / 0	0 / 5
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Sinus bradycardia			
subjects affected / exposed	1 / 127 (0.79%)	0 / 193 (0.00%)	0 / 114 (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
Acute myocardial infarction			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Aortic valve incompetence			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 flutter			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Atrioventricular block complete			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 acute			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	1 / 114 (0.88%)
occurrences causally related to treatment / all	0 / 0	0 / 0	1 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Mitral valve incompetence			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Myocardial ischaemia			

subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	0 / 114 (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
Myocardial infarction			
subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	0 / 114 (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
Ventricular arrhythmia			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Ventricular extrasystoles			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Ventricular tachycardia			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 unstable			
subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	0 / 114 (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
Cardiac arrest			
subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	1 / 114 (0.88%)
occurrences causally related to treatment / all	0 / 0	0 / 1	1 / 1
deaths causally related to treatment / all	0 / 0	0 / 1	1 / 1
Cardiopulmonary failure			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	1 / 114 (0.88%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 1
Acute Coronary Syndrome			

subjects affected / exposed	1 / 127 (0.79%)	0 / 193 (0.00%)	0 / 114 (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
Cardiac Failure Congestive			
subjects affected / exposed	1 / 127 (0.79%)	0 / 193 (0.00%)	0 / 114 (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
Myocarditis			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	1 / 114 (0.88%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 1
Nervous system disorders			
Haemorrhage intracranial			
subjects affected / exposed	2 / 127 (1.57%)	0 / 193 (0.00%)	1 / 114 (0.88%)
occurrences causally related to treatment / all	0 / 2	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 1
Paraparesis			
subjects affected / exposed	1 / 127 (0.79%)	0 / 193 (0.00%)	0 / 114 (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
Presyncope			
subjects affected / exposed	1 / 127 (0.79%)	0 / 193 (0.00%)	0 / 114 (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
Tremor			
subjects affected / exposed	1 / 127 (0.79%)	0 / 193 (0.00%)	0 / 114 (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
Balance disorder			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 haematoma			

subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 haemorrhage			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 ischaemia			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Epilepsy			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Ischaemic stroke			
subjects affected / exposed	1 / 127 (0.79%)	1 / 193 (0.52%)	0 / 114 (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
Seizure			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	1 / 114 (0.88%)
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 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Transient ischaemic attack			
subjects affected / exposed	1 / 127 (0.79%)	0 / 193 (0.00%)	0 / 114 (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
Central nervous system lesion			

subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	0 / 114 (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
Muscle spasticity			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	1 / 114 (0.88%)
occurrences causally related to treatment / all	0 / 0	0 / 0	1 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cerebrovascular Accident			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 Inflammatory Demyelinating Polyradiculoneuropathy			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Dementia Alzheimer's Type			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Dizziness			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Encephalopathy			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Intracranial Haemorrhagic Stroke			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Headache			

subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	0 / 114 (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
Peripheral Sensorimotor Neuropathy			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Polyneuropathy			
subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	0 / 114 (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
Posterior Reversible Encephalopathy Syndrome			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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	2 / 127 (1.57%)	3 / 193 (1.55%)	1 / 114 (0.88%)
occurrences causally related to treatment / all	0 / 2	1 / 3	1 / 2
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Autoimmune haemolytic anaemia			
subjects affected / exposed	3 / 127 (2.36%)	0 / 193 (0.00%)	0 / 114 (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
Febrile neutropenia			
subjects affected / exposed	7 / 127 (5.51%)	20 / 193 (10.36%)	3 / 114 (2.63%)
occurrences causally related to treatment / all	2 / 7	11 / 25	2 / 3
deaths causally related to treatment / all	0 / 1	0 / 0	0 / 0
Haemolytic anaemia			
subjects affected / exposed	1 / 127 (0.79%)	0 / 193 (0.00%)	0 / 114 (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
Neutropenia			

subjects affected / exposed	9 / 127 (7.09%)	28 / 193 (14.51%)	6 / 114 (5.26%)
occurrences causally related to treatment / all	6 / 10	38 / 50	7 / 8
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Thrombocytopenia			
subjects affected / exposed	2 / 127 (1.57%)	6 / 193 (3.11%)	5 / 114 (4.39%)
occurrences causally related to treatment / all	0 / 2	4 / 6	5 / 5
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pancytopenia			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Leukopenia			
subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	0 / 114 (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
Lymphopenia			
subjects affected / exposed	0 / 127 (0.00%)	2 / 193 (1.04%)	0 / 114 (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
Disseminated intravascular coagulation			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	1 / 114 (0.88%)
occurrences causally related to treatment / all	0 / 0	0 / 0	1 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Bicytopenia			
subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	0 / 114 (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
Coombs Negative Haemolytic Anaemia			
subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	0 / 114 (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
Cytopenia			

subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	0 / 114 (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
Eye disorders			
Retinal detachment			
subjects affected / exposed	1 / 127 (0.79%)	0 / 193 (0.00%)	0 / 114 (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
Cataract			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Retinal Degeneration			
subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	0 / 114 (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
Vitreous Haemorrhage			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	1 / 114 (0.88%)
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 disorders			
Lower gastrointestinal haemorrhage			
subjects affected / exposed	1 / 127 (0.79%)	0 / 193 (0.00%)	0 / 114 (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
Stomatitis			
subjects affected / exposed	1 / 127 (0.79%)	0 / 193 (0.00%)	0 / 114 (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
Abdominal pain			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 pain upper			

subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Ascites			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 / 127 (0.00%)	2 / 193 (1.04%)	0 / 114 (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
Diverticular perforation			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Haematochezia			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 perforation			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Mouth haemorrhage			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 / 127 (0.00%)	0 / 193 (0.00%)	1 / 114 (0.88%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Peritoneal adhesions			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Inguinal hernia			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	1 / 114 (0.88%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Neutropenic colitis			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	1 / 114 (0.88%)
occurrences causally related to treatment / all	0 / 0	0 / 0	1 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Enteritis			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Inguinal Hernia Perforation			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Large Intestinal Haemorrhage			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	1 / 114 (0.88%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Large Intestine Perforation			

subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Large Intestine Polyp			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Melaena			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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			
Cholangitis			
subjects affected / exposed	1 / 127 (0.79%)	0 / 193 (0.00%)	0 / 114 (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
Cholecystitis			
subjects affected / exposed	0 / 127 (0.00%)	2 / 193 (1.04%)	2 / 114 (1.75%)
occurrences causally related to treatment / all	0 / 0	0 / 2	1 / 2
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hepatic failure			
subjects affected / exposed	2 / 127 (1.57%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	1 / 2	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 1	0 / 0	0 / 0
Hepatocellular injury			
subjects affected / exposed	2 / 127 (1.57%)	0 / 193 (0.00%)	1 / 114 (0.88%)
occurrences causally related to treatment / all	2 / 2	0 / 0	1 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Acute hepatic failure			

subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cholecystitis acute			
subjects affected / exposed	1 / 127 (0.79%)	1 / 193 (0.52%)	0 / 114 (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
Hepatitis			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hepatitis toxic			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Liver disorder			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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-induced liver injury			
subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	1 / 114 (0.88%)
occurrences causally related to treatment / all	0 / 0	0 / 1	1 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Bile Duct Stone			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cholelithiasis			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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			
Decubitus ulcer			

subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Dermatitis allergic			
subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	0 / 114 (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
Purpura			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Diabetic foot			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	1 / 114 (0.88%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Rash maculo-papular			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 / 127 (0.00%)	0 / 193 (0.00%)	1 / 114 (0.88%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Urticaria			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	1 / 114 (0.88%)
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 and urinary disorders			
Chronic kidney disease			
subjects affected / exposed	1 / 127 (0.79%)	0 / 193 (0.00%)	0 / 114 (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 kidney injury			

subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	1 / 114 (0.88%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 1
Haematuria			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hypertensive nephropathy			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Nephrolithiasis			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 failure			
subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	0 / 114 (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
Renal impairment			
subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	0 / 114 (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
Urethral prolapse			
subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	0 / 114 (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 obstruction			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	1 / 114 (0.88%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Ureterolithiasis			

subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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			
Hypercalcaemia of malignancy			
subjects affected / exposed	1 / 127 (0.79%)	0 / 193 (0.00%)	0 / 114 (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
Hyperparathyroidism			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hyperthyroidism			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Fistula inflammation			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 chest pain			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Osteoarthritis			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0

Rheumatoid arthritis			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 column stenosis			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 pain			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Arthritis			
subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	1 / 114 (0.88%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Intervertebral Disc Protrusion			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Myositis			
subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	0 / 114 (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
Osteoporotic Fracture			
subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	0 / 114 (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
Psoriatic Arthropathy			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 Disorder			

subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	0 / 114 (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
Infections and infestations			
Appendicitis			
subjects affected / exposed	1 / 127 (0.79%)	0 / 193 (0.00%)	0 / 114 (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
Bacterial sepsis			
subjects affected / exposed	1 / 127 (0.79%)	0 / 193 (0.00%)	0 / 114 (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
Bronchitis			
subjects affected / exposed	2 / 127 (1.57%)	1 / 193 (0.52%)	0 / 114 (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
Clostridium difficile colitis			
subjects affected / exposed	1 / 127 (0.79%)	0 / 193 (0.00%)	0 / 114 (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
Enterococcal sepsis			
subjects affected / exposed	1 / 127 (0.79%)	0 / 193 (0.00%)	0 / 114 (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
Erysipelas			
subjects affected / exposed	1 / 127 (0.79%)	0 / 193 (0.00%)	0 / 114 (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
Fungal infection			
subjects affected / exposed	1 / 127 (0.79%)	0 / 193 (0.00%)	0 / 114 (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			

subjects affected / exposed	1 / 127 (0.79%)	0 / 193 (0.00%)	0 / 114 (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 zoster			
subjects affected / exposed	1 / 127 (0.79%)	2 / 193 (1.04%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	1 / 1	0 / 2	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Infection			
subjects affected / exposed	1 / 127 (0.79%)	1 / 193 (0.52%)	0 / 114 (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
Influenza			
subjects affected / exposed	1 / 127 (0.79%)	3 / 193 (1.55%)	1 / 114 (0.88%)
occurrences causally related to treatment / all	0 / 1	0 / 3	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Lower respiratory tract infection			
subjects affected / exposed	2 / 127 (1.57%)	0 / 193 (0.00%)	2 / 114 (1.75%)
occurrences causally related to treatment / all	0 / 2	0 / 0	0 / 3
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Oropharyngeal candidiasis			
subjects affected / exposed	1 / 127 (0.79%)	0 / 193 (0.00%)	0 / 114 (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
Pneumonia			
subjects affected / exposed	12 / 127 (9.45%)	15 / 193 (7.77%)	17 / 114 (14.91%)
occurrences causally related to treatment / all	1 / 13	6 / 16	8 / 18
deaths causally related to treatment / all	0 / 1	0 / 3	2 / 3
Pneumonia viral			
subjects affected / exposed	1 / 127 (0.79%)	1 / 193 (0.52%)	0 / 114 (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
Progressive multifocal leukoencephalopathy			

subjects affected / exposed	1 / 127 (0.79%)	0 / 193 (0.00%)	0 / 114 (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
Sepsis			
subjects affected / exposed	2 / 127 (1.57%)	5 / 193 (2.59%)	3 / 114 (2.63%)
occurrences causally related to treatment / all	0 / 2	3 / 5	0 / 4
deaths causally related to treatment / all	0 / 2	1 / 2	0 / 0
Septic shock			
subjects affected / exposed	2 / 127 (1.57%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 2	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 2	0 / 0	0 / 0
Sinusitis			
subjects affected / exposed	1 / 127 (0.79%)	0 / 193 (0.00%)	1 / 114 (0.88%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Staphylococcal sepsis			
subjects affected / exposed	1 / 127 (0.79%)	0 / 193 (0.00%)	0 / 114 (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
Subcutaneous abscess			
subjects affected / exposed	1 / 127 (0.79%)	0 / 193 (0.00%)	0 / 114 (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
Tracheobronchitis			
subjects affected / exposed	1 / 127 (0.79%)	0 / 193 (0.00%)	0 / 114 (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
Urinary tract infection			
subjects affected / exposed	1 / 127 (0.79%)	1 / 193 (0.52%)	2 / 114 (1.75%)
occurrences causally related to treatment / all	0 / 1	0 / 1	0 / 3
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Urosepsis			

subjects affected / exposed	1 / 127 (0.79%)	0 / 193 (0.00%)	1 / 114 (0.88%)
occurrences causally related to treatment / all	1 / 1	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Abdominal sepsis			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Abscess limb			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 hepatitis B			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Arthritis infective			
subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	0 / 114 (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
Aspergillus infection			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Atypical pneumonia			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Brain abscess			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Bronchitis bacterial			

subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	0 / 114 (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
Bronchopulmonary aspergillosis			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Campylobacter gastroenteritis			
subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	0 / 114 (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
Campylobacter infection			
subjects affected / exposed	1 / 127 (0.79%)	0 / 193 (0.00%)	0 / 114 (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 / 127 (0.00%)	1 / 193 (0.52%)	1 / 114 (0.88%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Clostridium colitis			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cryptosporidiosis infection			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cytomegalovirus infection			

subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 infection			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Eczema infected			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Encephalitis viral			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 / 127 (0.00%)	0 / 193 (0.00%)	1 / 114 (0.88%)
occurrences causally related to treatment / all	0 / 0	0 / 0	1 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Gastroenteritis			

subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	0 / 114 (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
Giardiasis			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
H1N1 influenza			
subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	0 / 114 (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
Herpes zoster oticus			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Infected skin ulcer			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Infective exacerbation chronic obstructive airways disease			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 infection			
subjects affected / exposed	0 / 127 (0.00%)	7 / 193 (3.63%)	2 / 114 (1.75%)
occurrences causally related to treatment / all	0 / 0	2 / 7	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Lymphangitis			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Meningitis			

subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Meningitis cryptococcal			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Nasopharyngitis			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Neutropenic sepsis			
subjects affected / exposed	0 / 127 (0.00%)	2 / 193 (1.04%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	2 / 2	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Oral candidiasis			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Perirectal abscess			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 cytomegaloviral			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 pseudomonal			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pseudomembranous colitis			

subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pseudomonal sepsis			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 abscess			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Upper respiratory tract infection			
subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	0 / 114 (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
Viral infection			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Visceral leishmaniasis			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
West nile viral infection			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Wound infection			

subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 / 127 (0.00%)	1 / 193 (0.52%)	0 / 114 (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
Fungal oesophagitis			
subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	0 / 114 (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
Cerebral toxoplasmosis			
subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	0 / 114 (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
Herpes virus infection			
subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	0 / 114 (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
Laryngitis viral			
subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	0 / 114 (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
Listeriosis			
subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	0 / 114 (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
Peritonitis			
subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	0 / 114 (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
Pneumococcal sepsis			

subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	0 / 114 (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
Pneumonia fungal			
subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	0 / 114 (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 pneumococcal			
subjects affected / exposed	1 / 127 (0.79%)	1 / 193 (0.52%)	0 / 114 (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
Retinitis viral			
subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	0 / 114 (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
Varicella zoster virus infection			
subjects affected / exposed	0 / 127 (0.00%)	2 / 193 (1.04%)	0 / 114 (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
Lung infection pseudomonal			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	1 / 114 (0.88%)
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 bacterial			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	1 / 114 (0.88%)
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	0 / 127 (0.00%)	0 / 193 (0.00%)	1 / 114 (0.88%)
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 enterococcal			

subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	0 / 114 (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
Hepatitis B Reactivation			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 Abscess			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Anal Abscess			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Bacterial Infection			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Bronchitis Viral			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 Sepsis			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	1 / 114 (0.88%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Febrile Infection			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Gangrene			

subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	1 / 114 (0.88%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hepatitis E			
subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	0 / 114 (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
Legionella Infection			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Meningitis Viral			
subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	0 / 114 (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
Myringitis			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Otitis Externa			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Perineal Infection			
subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	0 / 114 (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 Respiratory			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 Sepsis			

subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	0 / 114 (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
Streptococcal Bacteraemia			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Tick-Borne Viral Encephalitis			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	1 / 114 (0.88%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 1
Tuberculosis			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Varicella			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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 Device Infection			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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			
Tumour lysis syndrome			
subjects affected / exposed	2 / 127 (1.57%)	3 / 193 (1.55%)	3 / 114 (2.63%)
occurrences causally related to treatment / all	2 / 2	3 / 3	3 / 3
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Diabetes mellitus			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Diabetes mellitus inadequate control			

subjects affected / exposed	0 / 127 (0.00%)	1 / 193 (0.52%)	0 / 114 (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 / 127 (0.00%)	0 / 193 (0.00%)	1 / 114 (0.88%)
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 / 127 (0.00%)	1 / 193 (0.52%)	1 / 114 (0.88%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hyperuricaemia			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Malnutrition			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Type 2 diabetes mellitus			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	1 / 114 (0.88%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Fluid overload			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	1 / 114 (0.88%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hypoglycaemia			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	1 / 114 (0.88%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Steroid Diabetes			

subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Diabetic Metabolic Decompensation			
subjects affected / exposed	0 / 127 (0.00%)	0 / 193 (0.00%)	0 / 114 (0.00%)
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	G-Benda		
Total subjects affected by serious adverse events			
subjects affected / exposed	335 / 537 (62.38%)		
number of deaths (all causes)	103		
number of deaths resulting from adverse events			
Neoplasms benign, malignant and unspecified (incl cysts and polyps)			
Adenocarcinoma of colon			
subjects affected / exposed	2 / 537 (0.37%)		
occurrences causally related to treatment / all	0 / 2		
deaths causally related to treatment / all	0 / 1		
Bladder transitional cell carcinoma			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Cholangiocarcinoma			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Clear cell renal cell carcinoma			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Metastatic carcinoma of the bladder			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		

Richter's syndrome				
subjects affected / exposed	6 / 537 (1.12%)			
occurrences causally related to treatment / all	0 / 6			
deaths causally related to treatment / all	0 / 5			
Spinal meningioma benign				
subjects affected / exposed	0 / 537 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Squamous cell carcinoma				
subjects affected / exposed	5 / 537 (0.93%)			
occurrences causally related to treatment / all	7 / 10			
deaths causally related to treatment / all	0 / 0			
Squamous cell carcinoma of skin				
subjects affected / exposed	6 / 537 (1.12%)			
occurrences causally related to treatment / all	1 / 6			
deaths causally related to treatment / all	0 / 1			
Adenocarcinoma				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	1 / 1			
deaths causally related to treatment / all	1 / 1			
Adenocarcinoma gastric				
subjects affected / exposed	3 / 537 (0.56%)			
occurrences causally related to treatment / all	1 / 3			
deaths causally related to treatment / all	1 / 2			
Adenosquamous cell lung cancer				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Basal cell carcinoma				
subjects affected / exposed	8 / 537 (1.49%)			
occurrences causally related to treatment / all	4 / 10			
deaths causally related to treatment / all	0 / 0			
Bladder cancer stage 0, with cancer in situ				

subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Bowen's disease				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Acute myeloid leukaemia				
subjects affected / exposed	0 / 537 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Breast cancer				
subjects affected / exposed	0 / 537 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Metastatic renal cell carcinoma				
subjects affected / exposed	0 / 537 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Myelodysplastic syndrome				
subjects affected / exposed	2 / 537 (0.37%)			
occurrences causally related to treatment / all	1 / 2			
deaths causally related to treatment / all	1 / 2			
Prostate cancer				
subjects affected / exposed	5 / 537 (0.93%)			
occurrences causally related to treatment / all	0 / 5			
deaths causally related to treatment / all	0 / 0			
Colon cancer				
subjects affected / exposed	0 / 537 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Malignant melanoma				

subjects affected / exposed	4 / 537 (0.74%)		
occurrences causally related to treatment / all	0 / 5		
deaths causally related to treatment / all	0 / 0		
Nasal cavity cancer			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Primary myelofibrosis			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Rectal adenocarcinoma			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Bronchial carcinoma			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 1		
Chronic myeloid leukaemia			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Endometrial cancer			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Hepatocellular carcinoma			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Intraductal proliferative breast lesion			

subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	1 / 1		
deaths causally related to treatment / all	0 / 0		
Laryngeal cancer			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Lung adenocarcinoma			
subjects affected / exposed	3 / 537 (0.56%)		
occurrences causally related to treatment / all	0 / 3		
deaths causally related to treatment / all	0 / 1		
Lung cancer metastatic			
subjects affected / exposed	2 / 537 (0.37%)		
occurrences causally related to treatment / all	0 / 2		
deaths causally related to treatment / all	0 / 1		
Lung neoplasm malignant			
subjects affected / exposed	2 / 537 (0.37%)		
occurrences causally related to treatment / all	0 / 2		
deaths causally related to treatment / all	0 / 1		
Metastatic bronchial carcinoma			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Metastatic malignant melanoma			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Metastatic squamous cell carcinoma			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	2 / 2		
deaths causally related to treatment / all	0 / 0		
Myelofibrosis			

subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Neoplasm malignant				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	1 / 1			
deaths causally related to treatment / all	1 / 1			
Pancreatic carcinoma metastatic				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 1			
Prostatic adenoma				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Skin cancer				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 2			
deaths causally related to treatment / all	0 / 1			
Small cell lung cancer metastatic				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Basosquamous Carcinoma of Skin				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Bladder Neoplasm				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Hodgkin's Disease				

subjects affected / exposed	0 / 537 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Intraductal Papillary Mucinous Neoplasm				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Intraocular Melanoma				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Liver Carcinoma Ruptured				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Neuroendocrine Tumour				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Peripheral T-Cell Lymphoma Unspecified				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 1			
Renal Cell Carcinoma				
subjects affected / exposed	0 / 537 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Squamous Cell Breast Carcinoma				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Transitional Cell Carcinoma				

subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Vascular disorders			
Hypertension			
subjects affected / exposed	2 / 537 (0.37%)		
occurrences causally related to treatment / all	1 / 2		
deaths causally related to treatment / all	0 / 0		
Hypotension			
subjects affected / exposed	4 / 537 (0.74%)		
occurrences causally related to treatment / all	4 / 4		
deaths causally related to treatment / all	0 / 0		
Peripheral arterial occlusive disease			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Peripheral artery thrombosis			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Flushing			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	1 / 1		
deaths causally related to treatment / all	0 / 0		
Haematoma			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Hypertensive crisis			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Orthostatic hypotension			

subjects affected / exposed	2 / 537 (0.37%)		
occurrences causally related to treatment / all	2 / 2		
deaths causally related to treatment / all	0 / 0		
Peripheral vascular disorder			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Shock haemorrhagic			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Vena cava thrombosis			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Thrombophlebitis superficial			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Arterial Rupture			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
General disorders and administration site conditions			
Death			
subjects affected / exposed	7 / 537 (1.30%)		
occurrences causally related to treatment / all	1 / 7		
deaths causally related to treatment / all	1 / 7		
General physical health deterioration			
subjects affected / exposed	4 / 537 (0.74%)		
occurrences causally related to treatment / all	0 / 4		
deaths causally related to treatment / all	0 / 3		
Malaise			

subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Pyrexia				
subjects affected / exposed	23 / 537 (4.28%)			
occurrences causally related to treatment / all	10 / 27			
deaths causally related to treatment / all	0 / 0			
Asthenia				
subjects affected / exposed	2 / 537 (0.37%)			
occurrences causally related to treatment / all	0 / 2			
deaths causally related to treatment / all	0 / 0			
Chest pain				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Chills				
subjects affected / exposed	4 / 537 (0.74%)			
occurrences causally related to treatment / all	5 / 5			
deaths causally related to treatment / all	0 / 0			
Euthanasia				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 1			
Hyperthermia				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	1 / 1			
deaths causally related to treatment / all	0 / 0			
Multiple organ dysfunction syndrome				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	1 / 1			
deaths causally related to treatment / all	0 / 0			
Oedema peripheral				

subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Catheter Site Thrombosis			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Mucosal Inflammation			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Immune system disorders			
Anaphylactic reaction			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	1 / 1		
deaths causally related to treatment / all	0 / 0		
Cytokine release syndrome			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	1 / 1		
deaths causally related to treatment / all	0 / 0		
Allergy to immunoglobulin therapy			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Drug hypersensitivity			
subjects affected / exposed	4 / 537 (0.74%)		
occurrences causally related to treatment / all	1 / 4		
deaths causally related to treatment / all	0 / 0		
Hypersensitivity			
subjects affected / exposed	2 / 537 (0.37%)		
occurrences causally related to treatment / all	2 / 2		
deaths causally related to treatment / all	0 / 0		
Autoimmune disorder			

subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Haemophagocytic			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 1		
Reproductive system and breast disorders			
Uterine Prolapse			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Respiratory, thoracic and mediastinal disorders			
Dyspnoea			
subjects affected / exposed	2 / 537 (0.37%)		
occurrences causally related to treatment / all	1 / 2		
deaths causally related to treatment / all	0 / 0		
Pulmonary oedema			
subjects affected / exposed	2 / 537 (0.37%)		
occurrences causally related to treatment / all	0 / 2		
deaths causally related to treatment / all	0 / 0		
Bronchiectasis			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Chylothorax			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Cough			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		

Hypoxia				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	1 / 1			
deaths causally related to treatment / all	0 / 0			
Interstitial lung disease				
subjects affected / exposed	2 / 537 (0.37%)			
occurrences causally related to treatment / all	1 / 2			
deaths causally related to treatment / all	0 / 1			
Lung disorder				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	1 / 1			
deaths causally related to treatment / all	0 / 0			
Obliterative bronchiolitis				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	1 / 1			
deaths causally related to treatment / all	0 / 0			
Pharyngeal inflammation				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Pleural effusion				
subjects affected / exposed	3 / 537 (0.56%)			
occurrences causally related to treatment / all	1 / 3			
deaths causally related to treatment / all	0 / 0			
Pleurisy				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Pneumonia aspiration				
subjects affected / exposed	3 / 537 (0.56%)			
occurrences causally related to treatment / all	0 / 3			
deaths causally related to treatment / all	0 / 2			
Pneumonitis				

subjects affected / exposed	2 / 537 (0.37%)		
occurrences causally related to treatment / all	1 / 2		
deaths causally related to treatment / all	0 / 0		
Pneumothorax			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 1		
Pulmonary embolism			
subjects affected / exposed	5 / 537 (0.93%)		
occurrences causally related to treatment / all	1 / 5		
deaths causally related to treatment / all	0 / 1		
Pulmonary necrosis			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	1 / 1		
deaths causally related to treatment / all	0 / 0		
Respiratory alkalosis			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	1 / 1		
deaths causally related to treatment / all	0 / 0		
Aspiration			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Organising pneumonia			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Oropharyngeal pain			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Acute respiratory distress syndrome			

subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Acute Pulmonary Oedema			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Asthma			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Chronic Obstructive Pulmonary Disease			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Psychiatric disorders			
Completed suicide			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Alcohol withdrawal syndrome			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Depression			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Investigations			
Hepatic enzyme increased			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		

Oxygen saturation decreased				
subjects affected / exposed	0 / 537 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Weight decreased				
subjects affected / exposed	0 / 537 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Alanine aminotransferase increased				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	1 / 1			
deaths causally related to treatment / all	0 / 0			
Aspartate aminotransferase increased				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	1 / 1			
deaths causally related to treatment / all	0 / 0			
Blood creatinine increased				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	1 / 1			
deaths causally related to treatment / all	0 / 0			
Blood pressure increased				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	1 / 1			
deaths causally related to treatment / all	0 / 0			
Gamma-glutamyltransferase increased				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	1 / 1			
deaths causally related to treatment / all	0 / 0			
Neutrophil count decreased				
subjects affected / exposed	0 / 537 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Transaminases increased				

subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Injury, poisoning and procedural complications			
Femoral neck fracture			
subjects affected / exposed	2 / 537 (0.37%)		
occurrences causally related to treatment / all	0 / 2		
deaths causally related to treatment / all	0 / 0		
Thoracic vertebral fracture			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Ankle fracture			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Cervical vertebral fracture			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Craniocerebral injury			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Femur fracture			
subjects affected / exposed	2 / 537 (0.37%)		
occurrences causally related to treatment / all	0 / 2		
deaths causally related to treatment / all	0 / 0		
Foot fracture			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Hip fracture			

subjects affected / exposed	2 / 537 (0.37%)			
occurrences causally related to treatment / all	0 / 2			
deaths causally related to treatment / all	0 / 0			
Radiation skin injury				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Rib fracture				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Road traffic accident				
subjects affected / exposed	2 / 537 (0.37%)			
occurrences causally related to treatment / all	0 / 2			
deaths causally related to treatment / all	0 / 1			
Spinal compression fracture				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Toxicity to various agents				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Wrist fracture				
subjects affected / exposed	0 / 537 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Fall				
subjects affected / exposed	0 / 537 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Head injury				

subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Radiation mucositis			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Dislocation of Vertebra			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Fractured Ischium			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Hand Fracture			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Multiple Injuries			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Post Procedural Haematoma			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Post Procedural Haemorrhage			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Radius Fracture			

subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Skull Fracture			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Subdural Haematoma			
subjects affected / exposed	2 / 537 (0.37%)		
occurrences causally related to treatment / all	0 / 2		
deaths causally related to treatment / all	0 / 0		
Spinal Fracture			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Cardiac disorders			
Atrial fibrillation			
subjects affected / exposed	4 / 537 (0.74%)		
occurrences causally related to treatment / all	0 / 9		
deaths causally related to treatment / all	0 / 0		
Cardiac failure			
subjects affected / exposed	5 / 537 (0.93%)		
occurrences causally related to treatment / all	0 / 5		
deaths causally related to treatment / all	0 / 0		
Sinus bradycardia			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Acute myocardial infarction			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Aortic valve incompetence			

subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Atrial flutter			
subjects affected / exposed	2 / 537 (0.37%)		
occurrences causally related to treatment / all	0 / 2		
deaths causally related to treatment / all	0 / 0		
Atrioventricular block complete			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Cardiac failure acute			
subjects affected / exposed	2 / 537 (0.37%)		
occurrences causally related to treatment / all	0 / 2		
deaths causally related to treatment / all	0 / 1		
Mitral valve incompetence			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Myocardial ischaemia			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Myocardial infarction			
subjects affected / exposed	3 / 537 (0.56%)		
occurrences causally related to treatment / all	0 / 3		
deaths causally related to treatment / all	0 / 0		
Ventricular arrhythmia			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	1 / 1		
deaths causally related to treatment / all	1 / 1		
Ventricular extrasystoles			

subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Ventricular tachycardia			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	1 / 1		
deaths causally related to treatment / all	0 / 0		
Angina unstable			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Cardiac arrest			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Cardiopulmonary failure			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Acute Coronary Syndrome			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Cardiac Failure Congestive			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Myocarditis			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Nervous system disorders			
Haemorrhage intracranial			

subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Paraparesis			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Presyncope			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Tremor			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Balance disorder			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Cerebral haematoma			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Cerebral haemorrhage			
subjects affected / exposed	3 / 537 (0.56%)		
occurrences causally related to treatment / all	0 / 3		
deaths causally related to treatment / all	0 / 1		
Cerebral ischaemia			
subjects affected / exposed	2 / 537 (0.37%)		
occurrences causally related to treatment / all	0 / 2		
deaths causally related to treatment / all	0 / 0		
Epilepsy			

subjects affected / exposed	2 / 537 (0.37%)			
occurrences causally related to treatment / all	0 / 2			
deaths causally related to treatment / all	0 / 0			
Ischaemic stroke				
subjects affected / exposed	3 / 537 (0.56%)			
occurrences causally related to treatment / all	0 / 3			
deaths causally related to treatment / all	0 / 0			
Seizure				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Syncope				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	1 / 1			
deaths causally related to treatment / all	0 / 0			
Transient ischaemic attack				
subjects affected / exposed	2 / 537 (0.37%)			
occurrences causally related to treatment / all	0 / 2			
deaths causally related to treatment / all	0 / 0			
Central nervous system lesion				
subjects affected / exposed	0 / 537 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Muscle spasticity				
subjects affected / exposed	0 / 537 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Cerebrovascular Accident				
subjects affected / exposed	2 / 537 (0.37%)			
occurrences causally related to treatment / all	0 / 2			
deaths causally related to treatment / all	0 / 0			
Chronic Inflammatory Demyelinating Polyradiculoneuropathy				

subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Dementia Alzheimer's Type				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Dizziness				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Encephalopathy				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 1			
Intracranial Haemorrhagic Stroke				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Headache				
subjects affected / exposed	0 / 537 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Peripheral Sensorimotor Neuropathy				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Polyneuropathy				
subjects affected / exposed	0 / 537 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Posterior Reversible Encephalopathy Syndrome				

subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Blood and lymphatic system disorders			
Anaemia			
subjects affected / exposed	5 / 537 (0.93%)		
occurrences causally related to treatment / all	3 / 5		
deaths causally related to treatment / all	0 / 0		
Autoimmune haemolytic anaemia			
subjects affected / exposed	2 / 537 (0.37%)		
occurrences causally related to treatment / all	1 / 2		
deaths causally related to treatment / all	0 / 0		
Febrile neutropenia			
subjects affected / exposed	43 / 537 (8.01%)		
occurrences causally related to treatment / all	34 / 49		
deaths causally related to treatment / all	1 / 1		
Haemolytic anaemia			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Neutropenia			
subjects affected / exposed	62 / 537 (11.55%)		
occurrences causally related to treatment / all	61 / 100		
deaths causally related to treatment / all	0 / 0		
Thrombocytopenia			
subjects affected / exposed	17 / 537 (3.17%)		
occurrences causally related to treatment / all	16 / 17		
deaths causally related to treatment / all	0 / 0		
Pancytopenia			
subjects affected / exposed	6 / 537 (1.12%)		
occurrences causally related to treatment / all	3 / 6		
deaths causally related to treatment / all	0 / 0		
Leukopenia			

subjects affected / exposed	3 / 537 (0.56%)		
occurrences causally related to treatment / all	3 / 4		
deaths causally related to treatment / all	0 / 0		
Lymphopenia			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Disseminated intravascular coagulation			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Bicytopenia			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Coombs Negative Haemolytic Anaemia			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Cytopenia			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Eye disorders			
Retinal detachment			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Cataract			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Retinal Degeneration			

subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Vitreous Haemorrhage			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Gastrointestinal disorders			
Lower gastrointestinal haemorrhage			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Stomatitis			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Abdominal pain			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Abdominal pain upper			
subjects affected / exposed	2 / 537 (0.37%)		
occurrences causally related to treatment / all	0 / 2		
deaths causally related to treatment / all	0 / 0		
Ascites			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Colitis			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Diarrhoea			

subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 2		
deaths causally related to treatment / all	0 / 0		
Diverticular perforation			
subjects affected / exposed	2 / 537 (0.37%)		
occurrences causally related to treatment / all	0 / 2		
deaths causally related to treatment / all	0 / 1		
Haematochezia			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Intestinal perforation			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Mouth haemorrhage			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Nausea			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Peritoneal adhesions			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Vomiting			
subjects affected / exposed	3 / 537 (0.56%)		
occurrences causally related to treatment / all	1 / 3		
deaths causally related to treatment / all	0 / 0		
Inguinal hernia			

subjects affected / exposed	3 / 537 (0.56%)			
occurrences causally related to treatment / all	0 / 3			
deaths causally related to treatment / all	0 / 0			
Neutropenic colitis				
subjects affected / exposed	0 / 537 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Enteritis				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Inguinal Hernia Perforation				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Large Intestinal Haemorrhage				
subjects affected / exposed	0 / 537 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Large Intestine Perforation				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Large Intestine Polyp				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Melaena				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Rectal Haemorrhage				

subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Hepatobiliary disorders			
Cholangitis			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Cholecystitis			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Hepatic failure			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	1 / 1		
deaths causally related to treatment / all	0 / 0		
Hepatocellular injury			
subjects affected / exposed	3 / 537 (0.56%)		
occurrences causally related to treatment / all	3 / 3		
deaths causally related to treatment / all	0 / 0		
Acute hepatic failure			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	1 / 1		
deaths causally related to treatment / all	1 / 1		
Cholecystitis acute			
subjects affected / exposed	2 / 537 (0.37%)		
occurrences causally related to treatment / all	0 / 2		
deaths causally related to treatment / all	0 / 0		
Hepatitis			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	1 / 1		
deaths causally related to treatment / all	0 / 0		
Hepatitis toxic			

subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	1 / 1		
deaths causally related to treatment / all	0 / 0		
Liver disorder			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	1 / 1		
deaths causally related to treatment / all	0 / 0		
Drug-induced liver injury			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Bile Duct Stone			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Cholelithiasis			
subjects affected / exposed	2 / 537 (0.37%)		
occurrences causally related to treatment / all	0 / 2		
deaths causally related to treatment / all	0 / 0		
Skin and subcutaneous tissue disorders			
Decubitus ulcer			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Dermatitis allergic			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Purpura			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	1 / 1		
deaths causally related to treatment / all	0 / 0		
Diabetic foot			

subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Rash maculo-papular			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Drug eruption			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Urticaria			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Renal and urinary disorders			
Chronic kidney disease			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Acute kidney injury			
subjects affected / exposed	6 / 537 (1.12%)		
occurrences causally related to treatment / all	2 / 7		
deaths causally related to treatment / all	0 / 0		
Haematuria			
subjects affected / exposed	2 / 537 (0.37%)		
occurrences causally related to treatment / all	0 / 2		
deaths causally related to treatment / all	0 / 0		
Hypertensive nephropathy			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Nephrolithiasis			

subjects affected / exposed	2 / 537 (0.37%)		
occurrences causally related to treatment / all	0 / 2		
deaths causally related to treatment / all	0 / 0		
Renal failure			
subjects affected / exposed	2 / 537 (0.37%)		
occurrences causally related to treatment / all	0 / 2		
deaths causally related to treatment / all	0 / 0		
Renal impairment			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Urethral prolapse			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Urinary tract obstruction			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Ureterolithiasis			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Endocrine disorders			
Hypercalcaemia of malignancy			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Hyperparathyroidism			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Hyperthyroidism			

subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Musculoskeletal and connective tissue disorders			
Back pain			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Fistula inflammation			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Musculoskeletal chest pain			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Osteoarthritis			
subjects affected / exposed	4 / 537 (0.74%)		
occurrences causally related to treatment / all	0 / 4		
deaths causally related to treatment / all	0 / 0		
Rheumatoid arthritis			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Spinal column stenosis			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Spinal pain			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	1 / 1		
deaths causally related to treatment / all	0 / 0		
Arthritis			

subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Intervertebral Disc Protrusion			
subjects affected / exposed	2 / 537 (0.37%)		
occurrences causally related to treatment / all	0 / 2		
deaths causally related to treatment / all	0 / 0		
Myositis			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Osteoporotic Fracture			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Psoriatic Arthropathy			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Spinal Disorder			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Infections and infestations			
Appendicitis			
subjects affected / exposed	2 / 537 (0.37%)		
occurrences causally related to treatment / all	0 / 2		
deaths causally related to treatment / all	0 / 0		
Bacterial sepsis			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Bronchitis			

subjects affected / exposed	3 / 537 (0.56%)			
occurrences causally related to treatment / all	0 / 5			
deaths causally related to treatment / all	0 / 0			
Clostridium difficile colitis				
subjects affected / exposed	0 / 537 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Enterococcal sepsis				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Erysipelas				
subjects affected / exposed	2 / 537 (0.37%)			
occurrences causally related to treatment / all	0 / 2			
deaths causally related to treatment / all	0 / 0			
Fungal infection				
subjects affected / exposed	2 / 537 (0.37%)			
occurrences causally related to treatment / all	0 / 2			
deaths causally related to treatment / all	0 / 0			
Herpes simplex				
subjects affected / exposed	0 / 537 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Herpes zoster				
subjects affected / exposed	4 / 537 (0.74%)			
occurrences causally related to treatment / all	1 / 4			
deaths causally related to treatment / all	0 / 0			
Infection				
subjects affected / exposed	5 / 537 (0.93%)			
occurrences causally related to treatment / all	0 / 5			
deaths causally related to treatment / all	0 / 1			
Influenza				

subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Lower respiratory tract infection				
subjects affected / exposed	2 / 537 (0.37%)			
occurrences causally related to treatment / all	0 / 5			
deaths causally related to treatment / all	0 / 0			
Oropharyngeal candidiasis				
subjects affected / exposed	0 / 537 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Pneumonia				
subjects affected / exposed	65 / 537 (12.10%)			
occurrences causally related to treatment / all	21 / 90			
deaths causally related to treatment / all	0 / 9			
Pneumonia viral				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Progressive multifocal leukoencephalopathy				
subjects affected / exposed	0 / 537 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Sepsis				
subjects affected / exposed	10 / 537 (1.86%)			
occurrences causally related to treatment / all	3 / 10			
deaths causally related to treatment / all	0 / 3			
Septic shock				
subjects affected / exposed	4 / 537 (0.74%)			
occurrences causally related to treatment / all	0 / 4			
deaths causally related to treatment / all	0 / 2			
Sinusitis				

subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Staphylococcal sepsis			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	1 / 1		
deaths causally related to treatment / all	0 / 0		
Subcutaneous abscess			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Tracheobronchitis			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Urinary tract infection			
subjects affected / exposed	7 / 537 (1.30%)		
occurrences causally related to treatment / all	1 / 8		
deaths causally related to treatment / all	0 / 0		
Urosepsis			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Abdominal sepsis			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 1		
Abscess limb			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Acute hepatitis B			

subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Arthritis infective			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Aspergillus infection			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 1		
Atypical pneumonia			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	1 / 1		
deaths causally related to treatment / all	0 / 0		
Brain abscess			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 1		
Bronchitis bacterial			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	1 / 1		
deaths causally related to treatment / all	0 / 0		
Bronchopulmonary aspergillosis			
subjects affected / exposed	4 / 537 (0.74%)		
occurrences causally related to treatment / all	3 / 5		
deaths causally related to treatment / all	1 / 1		
Campylobacter gastroenteritis			
subjects affected / exposed	2 / 537 (0.37%)		
occurrences causally related to treatment / all	1 / 2		
deaths causally related to treatment / all	0 / 0		
Campylobacter infection			

subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Cellulitis			
subjects affected / exposed	4 / 537 (0.74%)		
occurrences causally related to treatment / all	1 / 5		
deaths causally related to treatment / all	0 / 0		
Clostridium colitis			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	1 / 1		
deaths causally related to treatment / all	0 / 0		
Clostridium difficile infection			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Cryptosporidiosis infection			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 1		
Cytomegalovirus infection			
subjects affected / exposed	5 / 537 (0.93%)		
occurrences causally related to treatment / all	1 / 6		
deaths causally related to treatment / all	0 / 0		
Device related infection			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	1 / 1		
deaths causally related to treatment / all	0 / 0		
Diverticulitis			
subjects affected / exposed	2 / 537 (0.37%)		
occurrences causally related to treatment / all	0 / 3		
deaths causally related to treatment / all	0 / 0		
Ear infection			

subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Eczema infected				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Encephalitis viral				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Endocarditis				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Escherichia bacteraemia				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Gastroenteritis				
subjects affected / exposed	2 / 537 (0.37%)			
occurrences causally related to treatment / all	0 / 2			
deaths causally related to treatment / all	0 / 0			
Giardiasis				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
H1N1 influenza				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Herpes zoster oticus				

subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Infected skin ulcer				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Infective exacerbation chronic obstructive airways disease				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	1 / 4			
deaths causally related to treatment / all	0 / 0			
Lung infection				
subjects affected / exposed	5 / 537 (0.93%)			
occurrences causally related to treatment / all	0 / 5			
deaths causally related to treatment / all	0 / 1			
Lymphangitis				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Meningitis				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Meningitis cryptococcal				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 2			
deaths causally related to treatment / all	0 / 0			
Nasopharyngitis				
subjects affected / exposed	2 / 537 (0.37%)			
occurrences causally related to treatment / all	0 / 2			
deaths causally related to treatment / all	0 / 0			
Neutropenic sepsis				

subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	1 / 1		
deaths causally related to treatment / all	0 / 0		
Oral candidiasis			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	1 / 1		
deaths causally related to treatment / all	0 / 0		
Perirectal abscess			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Pneumonia cytomegaloviral			
subjects affected / exposed	2 / 537 (0.37%)		
occurrences causally related to treatment / all	0 / 2		
deaths causally related to treatment / all	0 / 0		
Pneumonia pseudomonal			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	2 / 2		
deaths causally related to treatment / all	0 / 0		
Pseudomembranous colitis			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Pseudomonal sepsis			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 1		
Pyelonephritis			
subjects affected / exposed	2 / 537 (0.37%)		
occurrences causally related to treatment / all	0 / 4		
deaths causally related to treatment / all	0 / 0		
Splenic abscess			

subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Upper respiratory tract infection			
subjects affected / exposed	2 / 537 (0.37%)		
occurrences causally related to treatment / all	0 / 2		
deaths causally related to treatment / all	0 / 0		
Viral infection			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Visceral leishmaniasis			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	1 / 1		
deaths causally related to treatment / all	1 / 1		
West Nile viral infection			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	1 / 1		
deaths causally related to treatment / all	1 / 1		
Wound infection			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Escherichia infection			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Fungal oesophagitis			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Cerebral toxoplasmosis			

subjects affected / exposed	0 / 537 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Herpes virus infection				
subjects affected / exposed	0 / 537 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Laryngitis viral				
subjects affected / exposed	0 / 537 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Listeriosis				
subjects affected / exposed	0 / 537 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Peritonitis				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Pneumococcal sepsis				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 1			
Pneumonia fungal				
subjects affected / exposed	0 / 537 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Pneumonia pneumococcal				
subjects affected / exposed	0 / 537 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Retinitis viral				

subjects affected / exposed	0 / 537 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Varicella zoster virus infection				
subjects affected / exposed	0 / 537 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Lung infection pseudomonal				
subjects affected / exposed	0 / 537 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Pneumonia bacterial				
subjects affected / exposed	0 / 537 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Respiratory tract infection				
subjects affected / exposed	0 / 537 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Urinary tract infection enterococcal				
subjects affected / exposed	0 / 537 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Hepatitis B Reactivation				
subjects affected / exposed	3 / 537 (0.56%)			
occurrences causally related to treatment / all	1 / 3			
deaths causally related to treatment / all	0 / 0			
Abdominal Abscess				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Anal Abscess				

subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Bacterial Infection				
subjects affected / exposed	2 / 537 (0.37%)			
occurrences causally related to treatment / all	0 / 2			
deaths causally related to treatment / all	0 / 0			
Bronchitis Viral				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Escherichia Sepsis				
subjects affected / exposed	0 / 537 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Febrile Infection				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Gangrene				
subjects affected / exposed	0 / 537 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Hepatitis E				
subjects affected / exposed	0 / 537 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Legionella Infection				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 1			
Meningitis Viral				

subjects affected / exposed	0 / 537 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Myringitis				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	1 / 1			
deaths causally related to treatment / all	0 / 0			
Otitis Externa				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Perineal Infection				
subjects affected / exposed	0 / 537 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Pneumonia Respiratory				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Pulmonary Sepsis				
subjects affected / exposed	0 / 537 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Streptococcal Bacteraemia				
subjects affected / exposed	1 / 537 (0.19%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Tick-Borne Viral Encephalitis				
subjects affected / exposed	0 / 537 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Tuberculosis				

subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Varicella			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Vascular Device Infection			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Metabolism and nutrition disorders			
Tumour lysis syndrome			
subjects affected / exposed	30 / 537 (5.59%)		
occurrences causally related to treatment / all	28 / 30		
deaths causally related to treatment / all	2 / 2		
Diabetes mellitus			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Diabetes mellitus inadequate control			
subjects affected / exposed	2 / 537 (0.37%)		
occurrences causally related to treatment / all	1 / 2		
deaths causally related to treatment / all	0 / 0		
Hypercalcaemia			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Hyperglycaemia			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	1 / 1		
deaths causally related to treatment / all	0 / 0		
Hyperuricaemia			

subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	1 / 1		
deaths causally related to treatment / all	0 / 0		
Malnutrition			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Type 2 diabetes mellitus			
subjects affected / exposed	3 / 537 (0.56%)		
occurrences causally related to treatment / all	0 / 3		
deaths causally related to treatment / all	0 / 0		
Fluid overload			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Hypoglycaemia			
subjects affected / exposed	0 / 537 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Steroid Diabetes			
subjects affected / exposed	1 / 537 (0.19%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Diabetic Metabolic Decompensation			
subjects affected / exposed	2 / 537 (0.37%)		
occurrences causally related to treatment / all	0 / 2		
deaths causally related to treatment / all	0 / 0		

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

Non-serious adverse events	G-Mono	G-FC	G-Clb
Total subjects affected by non-serious adverse events			
subjects affected / exposed	108 / 127 (85.04%)	184 / 193 (95.34%)	107 / 114 (93.86%)
Investigations			
Alanine aminotransferase increased			
subjects affected / exposed	4 / 127 (3.15%)	11 / 193 (5.70%)	3 / 114 (2.63%)
occurrences (all)	4	12	3
Aspartate aminotransferase increased			
subjects affected / exposed	4 / 127 (3.15%)	10 / 193 (5.18%)	5 / 114 (4.39%)
occurrences (all)	4	10	6
Vascular disorders			
Hypertension			
subjects affected / exposed	7 / 127 (5.51%)	16 / 193 (8.29%)	5 / 114 (4.39%)
occurrences (all)	7	17	5
Hypotension			
subjects affected / exposed	10 / 127 (7.87%)	17 / 193 (8.81%)	16 / 114 (14.04%)
occurrences (all)	12	20	18
Nervous system disorders			
Headache			
subjects affected / exposed	10 / 127 (7.87%)	23 / 193 (11.92%)	10 / 114 (8.77%)
occurrences (all)	18	25	10
Tremor			
subjects affected / exposed	6 / 127 (4.72%)	2 / 193 (1.04%)	6 / 114 (5.26%)
occurrences (all)	6	2	6
Dizziness			
subjects affected / exposed	3 / 127 (2.36%)	4 / 193 (2.07%)	9 / 114 (7.89%)
occurrences (all)	5	4	9
Blood and lymphatic system disorders			
Anaemia			
subjects affected / exposed	19 / 127 (14.96%)	50 / 193 (25.91%)	27 / 114 (23.68%)
occurrences (all)	28	76	35
Leukopenia			
subjects affected / exposed	7 / 127 (5.51%)	27 / 193 (13.99%)	8 / 114 (7.02%)
occurrences (all)	10	45	12
Neutropenia			
subjects affected / exposed	45 / 127 (35.43%)	138 / 193 (71.50%)	58 / 114 (50.88%)
occurrences (all)	130	416	127

Thrombocytopenia subjects affected / exposed occurrences (all)	27 / 127 (21.26%) 34	67 / 193 (34.72%) 116	34 / 114 (29.82%) 54
Lymphopenia subjects affected / exposed occurrences (all)	2 / 127 (1.57%) 2	11 / 193 (5.70%) 19	2 / 114 (1.75%) 3
General disorders and administration site conditions			
Asthenia subjects affected / exposed occurrences (all)	8 / 127 (6.30%) 9	30 / 193 (15.54%) 42	14 / 114 (12.28%) 17
Chest discomfort subjects affected / exposed occurrences (all)	7 / 127 (5.51%) 7	7 / 193 (3.63%) 7	3 / 114 (2.63%) 3
Chills subjects affected / exposed occurrences (all)	18 / 127 (14.17%) 19	30 / 193 (15.54%) 32	19 / 114 (16.67%) 22
Fatigue subjects affected / exposed occurrences (all)	7 / 127 (5.51%) 7	19 / 193 (9.84%) 25	23 / 114 (20.18%) 27
Pyrexia subjects affected / exposed occurrences (all)	28 / 127 (22.05%) 33	61 / 193 (31.61%) 93	28 / 114 (24.56%) 39
Oedema peripheral subjects affected / exposed occurrences (all)	4 / 127 (3.15%) 5	3 / 193 (1.55%) 4	9 / 114 (7.89%) 15
Gastrointestinal disorders			
Constipation subjects affected / exposed occurrences (all)	8 / 127 (6.30%) 10	24 / 193 (12.44%) 30	15 / 114 (13.16%) 18
Diarrhoea subjects affected / exposed occurrences (all)	6 / 127 (4.72%) 8	38 / 193 (19.69%) 44	11 / 114 (9.65%) 12
Nausea subjects affected / exposed occurrences (all)	23 / 127 (18.11%) 27	75 / 193 (38.86%) 147	26 / 114 (22.81%) 33
Vomiting			

subjects affected / exposed occurrences (all)	9 / 127 (7.09%) 9	44 / 193 (22.80%) 67	15 / 114 (13.16%) 16
Abdominal pain subjects affected / exposed occurrences (all)	3 / 127 (2.36%) 5	10 / 193 (5.18%) 10	4 / 114 (3.51%) 5
Dyspepsia subjects affected / exposed occurrences (all)	3 / 127 (2.36%) 3	5 / 193 (2.59%) 5	6 / 114 (5.26%) 6
Respiratory, thoracic and mediastinal disorders Cough subjects affected / exposed occurrences (all)	17 / 127 (13.39%) 18	28 / 193 (14.51%) 33	17 / 114 (14.91%) 18
Dyspnoea subjects affected / exposed occurrences (all)	9 / 127 (7.09%) 9	24 / 193 (12.44%) 24	11 / 114 (9.65%) 13
Skin and subcutaneous tissue disorders Hyperhidrosis subjects affected / exposed occurrences (all)	5 / 127 (3.94%) 5	8 / 193 (4.15%) 8	7 / 114 (6.14%) 7
Pruritus subjects affected / exposed occurrences (all)	4 / 127 (3.15%) 4	11 / 193 (5.70%) 11	2 / 114 (1.75%) 2
Rash subjects affected / exposed occurrences (all)	4 / 127 (3.15%) 4	24 / 193 (12.44%) 31	11 / 114 (9.65%) 15
Musculoskeletal and connective tissue disorders Arthralgia subjects affected / exposed occurrences (all)	9 / 127 (7.09%) 11	7 / 193 (3.63%) 9	8 / 114 (7.02%) 10
Back pain subjects affected / exposed occurrences (all)	7 / 127 (5.51%) 8	13 / 193 (6.74%) 15	10 / 114 (8.77%) 11
Infections and infestations Bronchitis subjects affected / exposed occurrences (all)	11 / 127 (8.66%) 12	18 / 193 (9.33%) 23	6 / 114 (5.26%) 8

Influenza			
subjects affected / exposed	5 / 127 (3.94%)	12 / 193 (6.22%)	5 / 114 (4.39%)
occurrences (all)	5	12	5
Nasopharyngitis			
subjects affected / exposed	9 / 127 (7.09%)	19 / 193 (9.84%)	4 / 114 (3.51%)
occurrences (all)	11	24	4
Upper respiratory tract infection			
subjects affected / exposed	6 / 127 (4.72%)	14 / 193 (7.25%)	5 / 114 (4.39%)
occurrences (all)	7	17	7
Urinary tract infection			
subjects affected / exposed	6 / 127 (4.72%)	7 / 193 (3.63%)	11 / 114 (9.65%)
occurrences (all)	9	9	16
Sinusitis			
subjects affected / exposed	4 / 127 (3.15%)	14 / 193 (7.25%)	5 / 114 (4.39%)
occurrences (all)	4	19	8
Respiratory tract infection			
subjects affected / exposed	5 / 127 (3.94%)	6 / 193 (3.11%)	7 / 114 (6.14%)
occurrences (all)	6	7	8
Pneumonia			
subjects affected / exposed	4 / 127 (3.15%)	9 / 193 (4.66%)	4 / 114 (3.51%)
occurrences (all)	6	10	4
Metabolism and nutrition disorders			
Decreased appetite			
subjects affected / exposed	4 / 127 (3.15%)	11 / 193 (5.70%)	11 / 114 (9.65%)
occurrences (all)	4	15	12
Hyperuricaemia			
subjects affected / exposed	0 / 127 (0.00%)	7 / 193 (3.63%)	10 / 114 (8.77%)
occurrences (all)	0	8	11

Non-serious adverse events	G-Benda		
Total subjects affected by non-serious adverse events			
subjects affected / exposed	503 / 537 (93.67%)		
Investigations			
Alanine aminotransferase increased			
subjects affected / exposed	13 / 537 (2.42%)		
occurrences (all)	14		
Aspartate aminotransferase increased			

subjects affected / exposed occurrences (all)	15 / 537 (2.79%) 15		
Vascular disorders			
Hypertension			
subjects affected / exposed	44 / 537 (8.19%)		
occurrences (all)	51		
Hypotension			
subjects affected / exposed	53 / 537 (9.87%)		
occurrences (all)	61		
Nervous system disorders			
Headache			
subjects affected / exposed	46 / 537 (8.57%)		
occurrences (all)	57		
Tremor			
subjects affected / exposed	11 / 537 (2.05%)		
occurrences (all)	11		
Dizziness			
subjects affected / exposed	22 / 537 (4.10%)		
occurrences (all)	23		
Blood and lymphatic system disorders			
Anaemia			
subjects affected / exposed	134 / 537 (24.95%)		
occurrences (all)	174		
Leukopenia			
subjects affected / exposed	70 / 537 (13.04%)		
occurrences (all)	119		
Neutropenia			
subjects affected / exposed	288 / 537 (53.63%)		
occurrences (all)	645		
Thrombocytopenia			
subjects affected / exposed	162 / 537 (30.17%)		
occurrences (all)	253		
Lymphopenia			
subjects affected / exposed	53 / 537 (9.87%)		
occurrences (all)	88		
General disorders and administration site conditions			

Asthenia			
subjects affected / exposed	48 / 537 (8.94%)		
occurrences (all)	61		
Chest discomfort			
subjects affected / exposed	15 / 537 (2.79%)		
occurrences (all)	15		
Chills			
subjects affected / exposed	92 / 537 (17.13%)		
occurrences (all)	110		
Fatigue			
subjects affected / exposed	71 / 537 (13.22%)		
occurrences (all)	85		
Pyrexia			
subjects affected / exposed	173 / 537 (32.22%)		
occurrences (all)	247		
Oedema peripheral			
subjects affected / exposed	31 / 537 (5.77%)		
occurrences (all)	37		
Gastrointestinal disorders			
Constipation			
subjects affected / exposed	72 / 537 (13.41%)		
occurrences (all)	84		
Diarrhoea			
subjects affected / exposed	83 / 537 (15.46%)		
occurrences (all)	114		
Nausea			
subjects affected / exposed	144 / 537 (26.82%)		
occurrences (all)	206		
Vomiting			
subjects affected / exposed	68 / 537 (12.66%)		
occurrences (all)	87		
Abdominal pain			
subjects affected / exposed	24 / 537 (4.47%)		
occurrences (all)	30		
Dyspepsia			

subjects affected / exposed occurrences (all)	24 / 537 (4.47%) 26		
Respiratory, thoracic and mediastinal disorders Cough subjects affected / exposed occurrences (all) Dyspnoea subjects affected / exposed occurrences (all)	 57 / 537 (10.61%) 70 53 / 537 (9.87%) 66		
Skin and subcutaneous tissue disorders Hyperhidrosis subjects affected / exposed occurrences (all) Pruritus subjects affected / exposed occurrences (all) Rash subjects affected / exposed occurrences (all)	 21 / 537 (3.91%) 22 25 / 537 (4.66%) 30 46 / 537 (8.57%) 63		
Musculoskeletal and connective tissue disorders Arthralgia subjects affected / exposed occurrences (all) Back pain subjects affected / exposed occurrences (all)	 19 / 537 (3.54%) 21 35 / 537 (6.52%) 37		
Infections and infestations Bronchitis subjects affected / exposed occurrences (all) Influenza subjects affected / exposed occurrences (all) Nasopharyngitis subjects affected / exposed occurrences (all)	 39 / 537 (7.26%) 43 13 / 537 (2.42%) 16 25 / 537 (4.66%) 28		

Upper respiratory tract infection subjects affected / exposed occurrences (all)	48 / 537 (8.94%) 62		
Urinary tract infection subjects affected / exposed occurrences (all)	29 / 537 (5.40%) 35		
Sinusitis subjects affected / exposed occurrences (all)	16 / 537 (2.98%) 18		
Respiratory tract infection subjects affected / exposed occurrences (all)	12 / 537 (2.23%) 19		
Pneumonia subjects affected / exposed occurrences (all)	28 / 537 (5.21%) 33		
Metabolism and nutrition disorders			
Decreased appetite subjects affected / exposed occurrences (all)	44 / 537 (8.19%) 45		
Hyperuricaemia subjects affected / exposed occurrences (all)	33 / 537 (6.15%) 36		

More information

Substantial protocol amendments (globally)

Were there any global substantial amendments to the protocol? Yes

Date	Amendment
17 March 2013	1. The original protocol was amended to reflect changes in the safety profile of obinutuzumab with regard to a higher incidence of thrombocytopenia and hemorrhagic events during the first cycle in subjects with CLL treated with obinutuzumab plus chlorambucil, as compared to subjects treated with rituximab plus chlorambucil or chlorambucil alone, in the Phase 3 pivotal study NCT01010061 2. Investigators were requested to closely monitor all subjects treated with obinutuzumab for thrombocytopenia and signs of hemorrhage, especially during the first cycle.
09 October 2014	1. To ensure that subject safety was not put at risk by clarifying inconsistencies and thereby reducing protocol violations 2. The amendment also included criteria for defining tumor lysis syndrome (TLS) and additional guidance regarding the risk of TLS, reinforcing adherence to the protocol when treating and monitoring subjects in the study.
06 May 2015	1. Addition of an interim efficacy analysis in a subgroup of subjects in Cohort 1 treated with obinutuzumab plus bendamustine, amendment of the secondary endpoint regarding response assessment from "ORR at end of treatment" to "CR and ORR at the time of FRA visit" for clarity, addition of Cohort 3 design per recommendations of the study IDMC, and removal of short-duration infusion-related assessments 2. Short-duration infusion was defined as the full 1000 mg obinutuzumab dose administered over 90 minutes, i.e., at a rate of approximately 667 mg/hour, rather than at the prior maximum rate of 400 mg/hour. The exploratory objective to examine the safety and PK profile of subjects who were to receive the short-duration infusion was removed.
27 October 2015	Addition of gastro-intestinal (GI) perforation as an important risk based on data from a trial in subjects with non Hodgkin's lymphoma (NHL) treated with obinutuzumab in combination with bendamustine.
06 March 2016	1. To implement additional risk minimization measures in subjects with CLL at risk of TLS treated with a combination of obinutuzumab and bendamustine. This followed the occurrence of a second fatal case of TLS in a subject at risk of TLS treated for CLL with obinutuzumab and bendamustine. More detailed criteria, including renal impairment in addition to criteria relating to tumor load, were added to identify subjects in this treatment group considered to be at high risk of TLS. The risk minimization measures included additional monitoring and laboratory assessments during Cycle 1.

Notes:

Interruptions (globally)

Were there any global interruptions to the trial? No

Limitations and caveats

None reported