Clinical trial results:
A Phase 3, Randomized, Open Label Trial of Lenalidomide/dexamethasone With or Without Elotuzumab in Relapsed or Refractory Multiple Myeloma

Summary

EudraCT number	2010-020347-12
Trial protocol	HU DE BE FR CZ DK IE AT GB GR ES PL IT
Global end of trial date	21 April 2021

Results information

Result version number	v1 (current)
This version publication date	06 April 2022
First version publication date	06 April 2022

Trial information

Trial identification

Sponsor protocol code	CA204-004

Additional study identifiers

ISRCTN number	-
ClinicalTrials.gov id (NCT number)	NCT01239797
WHO universal trial number (UTN)	-

Notes:

Sponsors

Sponsor organisation name	Bristol-Myers Squibb
Sponsor organisation address	Chaussée de la Hulpe 185, Brussels, Belgium, 1170
Public contact	EU Study Start-Up Unit, Bristol-Myers Squibb International Corporation, Clinical.Trials@bms.com
Scientific contact	Bristol-Myers Squibb Study Director, Bristol-Myers Squibb, Clinical.Trials@bms.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	Interim
Date of interim/final analysis	09 August 2021
Is this the analysis of the primary completion data?	No

Global end of trial reached?	Yes
Global end of trial date	21 April 2021
Was the trial ended prematurely?	No

Notes:

General information about the trial

Main objective of the trial:

To compare PFS of lenalidomide/low-dose dexamethasone + elotuzumab (LdE) versus lenalidomide/low-dose dexamethasone (Ld) in subjects with relapsed or refractory multiple myeloma (MM) and to compare the ORR of LdE versus Ld.

Protection of trial subjects:

The study was in compliance with the ethical principles derived from the Declaration of Helsinki and in compliance with all International Conference on Harmonization Good Clinical Practice Guidelines. All the local regulatory requirements pertinent to safety of trial participants were followed.

Background therapy: -

Evidence for comparator: -

Actual start date of recruitment	20 June 2011
Long term follow-up planned	No
Independent data monitoring committee (IDMC) involvement?	Yes

Notes:

Population of trial subjects

Subjects enrolled per country

Country: Number of subjects enrolled	Australia: 28
Country: Number of subjects enrolled	Austria: 3
Country: Number of subjects enrolled	Belgium: 17
Country: Number of subjects enrolled	Canada: 67
Country: Number of subjects enrolled	Czechia: 24
Country: Number of subjects enrolled	Denmark: 13
Country: Number of subjects enrolled	France: 50
Country: Number of subjects enrolled	Germany: 46
Country: Number of subjects enrolled	Greece: 43
Country: Number of subjects enrolled	Hungary: 7
Country: Number of subjects enrolled	Ireland: 2
Country: Number of subjects enrolled	Israel: 22
Country: Number of subjects enrolled	Italy: 41
Country: Number of subjects enrolled	Japan: 60
Country: Number of subjects enrolled	Poland: 48
Country: Number of subjects enrolled	Romania: 24
Country: Number of subjects enrolled	Spain: 36
Country: Number of subjects enrolled	Switzerland: 1
Country: Number of subjects enrolled	Turkey: 12

Country: Number of subjects enrolled	United Kingdom: 35
Country: Number of subjects enrolled	United States: 67
Worldwide total number of subjects	646
EEA total number of subjects	354

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)	276
From 65 to 84 years	363
85 years and over	7

Subject disposition

Recruitment

Recruitment details: -

Pre-assignment

Screening details:

646 participants were randomized and 635 were treated. 1 participant was randomized to treatment E-Ld but received treatment Ld.

Period 1

Period 1 title	Randomized Participants
Is this the baseline period?	Yes
Allocation method	Randomised - controlled
Blinding used	Not blinded

Arms

Are arms mutually exclusive?	Yes
Arm title	Lenalidomide + Dexamethasone + Elotuzumab

Arm description:

Lenalidomide: Capsules, Oral, 25 mg, once daily, on Days 1-21 Repeat every 28 days until participants met criteria for discontinuation of study drug Dexamethasone (Oral): On weeks without Elotuzumab dosing: Tablets, Oral, 40mg Repeat every 28 days until participants met criteria for discontinuation of study drug On weeks with Elotuzumab dosing: Tablets, Oral, 28 mg Repeat every 28 days until participants met criteria for discontinuation of study drug Dexamethasone (IV): On weeks without Elotuzumab dosing: Not Applicable (N/A) On weeks with Elotuzumab dosing: Solution, Intravenous (IV), 8 mg, weekly Repeat every 28 days until participants met criteria for discontinuation of study drug Elotuzumab (BMS-901608; HuLuc63): Solution, IV, 10 mg/kg, weekly, on Days 1, 8, 15, 22 (cycles 1&2); Days 1 and 15 (cycles 3 and beyond) Repeat every 28 days until participants met criteria for discontinuation of study drug

Arm type	Experimental
Investigational medicinal product name	Elotuzumab
Investigational medicinal product code	
Other name	BMS-901608
Pharmaceutical forms	Powder and solvent for concentrate for solution for infusion
Routes of administration	Intravenous use

Dosage and administration details:

Solution, IV, 10 mg/kg, weekly, on Days 1, 8, 15, 22 (cycles 1&2); Days 1 and 15 (cycles 3 and beyond) Repeat every 28 days.

Investigational medicinal product name	Lenalidomide
Investigational medicinal product code	
Other name	
Pharmaceutical forms	Capsule
Routes of administration	Oral use

Dosage and administration details:

25 mg, once daily, on Days 1-21 Repeat every 28 days

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

Dosage and administration details:

On weeks without Elotuzumab dosing: Tablets, Oral, 40mg Repeat every 28 days. On weeks with Elotuzumab dosing: Tablets, Oral, 28 mg Repeat every 28 days.

Investigational medicinal product name	Dexamethasone
Investigational medicinal product code	
Other name	
Pharmaceutical forms	Solution for infusion
Routes of administration	Intravenous use

Dosage and administration details:

On weeks without Elotuzumab dosing: Not Applicable (N/A) On weeks with Elotuzumab dosing: Solution, Intravenous (IV), 8 mg, weekly Repeat every 28 days

Arm title	Lenalidomide + Dexamethasone

Arm description:

Lenalidomide: Capsules, Oral, 25 mg, once daily, on Days 1-21 Regimen repeated every 28 days until participants met criteria for discontinuation of study drug Dexamethasone: Tablets, Oral, 40 mg, weekly, on Days 1, 8, 15, 22 Regimen repeated every 28 days until participants met criteria for discontinuation of study drug

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

Dosage and administration details:

Oral, 40 mg, weekly, on Days 1, 8, 15, 22 Regimen repeated every 28 days

Investigational medicinal product name	Lenalidomide
Investigational medicinal product code	
Other name	
Pharmaceutical forms	Capsule
Routes of administration	Oral use

Dosage and administration details:

25 mg, once daily, on Days 1-21 Repeat every 28 days

Number of subjects in period 1	Lenalidomide + Dexamethasone + Elotuzumab	Lenalidomide + Dexamethasone
Started	321	325
Completed	319	316
Not completed	2	9
Participant withdrew consent	1	7
Participant no longer meets study criteria	1	1
Adverse event unrelated to study drug	-	1

Period 2

Period 2 title	Treated Participants
Is this the baseline period?	No
Allocation method	Randomised - controlled
Blinding used	Not blinded

Arms

Are arms mutually exclusive?	Yes
Arm title	Lenalidomide + Dexamethasone + Elotuzumab

Arm description:

Lenalidomide: Capsules, Oral, 25 mg, once daily, on Days 1-21 Repeat every 28 days until participants met criteria for discontinuation of study drug Dexamethasone (Oral): On weeks without Elotuzumab dosing: Tablets, Oral, 40mg Repeat every 28 days until participants met criteria for discontinuation of study drug On weeks with Elotuzumab dosing: Tablets, Oral, 28 mg Repeat every 28 days until participants met criteria for discontinuation of study drug Dexamethasone (IV): On weeks without Elotuzumab dosing: Not Applicable (N/A) On weeks with Elotuzumab dosing: Solution, Intravenous (IV), 8 mg, weekly Repeat every 28 days until participants met criteria for discontinuation of study drug Elotuzumab (BMS-901608; HuLuc63): Solution, IV, 10 mg/kg, weekly, on Days 1, 8, 15, 22 (cycles 1&2); Days 1 and 15 (cycles 3 and beyond) Repeat every 28 days until participants met criteria for discontinuation of study drug

Arm type	Experimental
Investigational medicinal product name	Elotuzumab
Investigational medicinal product code	
Other name	BMS-901608
Pharmaceutical forms	Powder and solvent for concentrate for solution for infusion
Routes of administration	Intravenous use

Dosage and administration details:

Solution, IV, 10 mg/kg, weekly, on Days 1, 8, 15, 22 (cycles 1&2); Days 1 and 15 (cycles 3 and beyond) Repeat every 28 days.

Investigational medicinal product name	Lenalidomide
Investigational medicinal product code	
Other name	
Pharmaceutical forms	Capsule
Routes of administration	Oral use

Dosage and administration details:

25 mg, once daily, on Days 1-21 Repeat every 28 days

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

Dosage and administration details:

On weeks without Elotuzumab dosing: Tablets, Oral, 40mg Repeat every 28 days. On weeks with Elotuzumab dosing: Tablets, Oral, 28 mg Repeat every 28 days.

Investigational medicinal product name	Dexamethasone
Investigational medicinal product code	
Other name	
Pharmaceutical forms	Solution for infusion
Routes of administration	Intravenous use

Dosage and administration details:

On weeks without Elotuzumab dosing: Not Applicable (N/A) On weeks with Elotuzumab dosing: Solution, Intravenous (IV), 8 mg, weekly Repeat every 28 days.

Arm title	Lenalidomide + Dexamethasone

Arm description:

Lenalidomide: Capsules, Oral, 25 mg, once daily, on Days 1-21 Regimen repeated every 28 days until

participants met criteria for discontinuation of study drug Dexamethasone: Tablets, Oral, 40 mg, weekly, on Days 1, 8, 15, 22 Regimen repeated every 28 days until participants met criteria for discontinuation of study drug

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

Dosage and administration details:

Oral, 40 mg, weekly, on Days 1, 8, 15, 22 Regimen repeated every 28 days

Investigational medicinal product name	Lenalidomide
Investigational medicinal product code	
Other name	
Pharmaceutical forms	Capsule
Routes of administration	Oral use

Dosage and administration details:

25 mg, once daily, on Days 1-21 Repeat every 28 days

Number of subjects in period 2	Lenalidomide + Dexamethasone + Elotuzumab	Lenalidomide + Dexamethasone
Started	319	316
Completed	0	0
Not completed	319	316
Adverse event, serious fatal	1	1
Disease progression	185	185
Participant withdrew consent	6	11
Study drug toxicity	39	45
Participant no longer meets study criteria	2	-
Adverse event unrelated to study drug	31	37
Other reasons	14	14
Participants request to discontinue treatment	28	18
Poor/non-compliance	-	1
Administrative reason by sponsor	13	4

Baseline characteristics

Reporting groups

Reporting group title	Lenalidomide + Dexamethasone + Elotuzumab

Reporting group description:

Lenalidomide: Capsules, Oral, 25 mg, once daily, on Days 1-21 Repeat every 28 days until participants met criteria for discontinuation of study drug Dexamethasone (Oral): On weeks without Elotuzumab dosing: Tablets, Oral, 40mg Repeat every 28 days until participants met criteria for discontinuation of study drug On weeks with Elotuzumab dosing: Tablets, Oral, 28 mg Repeat every 28 days until participants met criteria for discontinuation of study drug Dexamethasone (IV): On weeks without Elotuzumab dosing: Not Applicable (N/A) On weeks with Elotuzumab dosing: Solution, Intravenous (IV), 8 mg, weekly Repeat every 28 days until participants met criteria for discontinuation of study drug Elotuzumab (BMS-901608; HuLuc63): Solution, IV, 10 mg/kg, weekly, on Days 1, 8, 15, 22 (cycles 1&2); Days 1 and 15 (cycles 3 and beyond) Repeat every 28 days until participants met criteria for discontinuation of study drug

Reporting group title	Lenalidomide + Dexamethasone

Reporting group description:

Lenalidomide: Capsules, Oral, 25 mg, once daily, on Days 1-21 Regimen repeated every 28 days until participants met criteria for discontinuation of study drug Dexamethasone: Tablets, Oral, 40 mg, weekly, on Days 1, 8, 15, 22 Regimen repeated every 28 days until participants met criteria for discontinuation of study drug

Reporting group values	Lenalidomide + Dexamethasone + Elotuzumab	Lenalidomide + Dexamethasone	Total
Number of subjects	321	325	646
Age Categorical Units:			
< 65 years old	134	142	276
>= 65 and < 75 years old	119	122	241
>= 75 years old	68	61	129
Age Continuous Units: years			
arithmetic mean	66.2	65.3	
standard deviation	± 9.34	± 10.26	-
Sex: Female, Male Units:			
Female	129	132	261
Male	192	193	385
Ethnicity (NIH/OMB) Units: Subjects			
Hispanic or Latino	5	1	6
Not Hispanic or Latino	28	33	61
Unknown or Not Reported	288	291	579
Race/Ethnicity, Customized Units: Subjects			
White	264	280	544
Black or African American	13	10	23
American Indian or Alaska Native	0	0	0
Asian	33	31	64
Native Hawaiian or Other Pacific Islander	1	0	1
Other	9	4	13

Not Reported	1	0	1

End points

End points reporting groups

Reporting group title	Lenalidomide + Dexamethasone + Elotuzumab

Reporting group description:

Lenalidomide: Capsules, Oral, 25 mg, once daily, on Days 1-21 Repeat every 28 days until participants met criteria for discontinuation of study drug Dexamethasone (Oral): On weeks without Elotuzumab dosing: Tablets, Oral, 40mg Repeat every 28 days until participants met criteria for discontinuation of study drug On weeks with Elotuzumab dosing: Tablets, Oral, 28 mg Repeat every 28 days until participants met criteria for discontinuation of study drug Dexamethasone (IV): On weeks without Elotuzumab dosing: Not Applicable (N/A) On weeks with Elotuzumab dosing: Solution, Intravenous (IV), 8 mg, weekly Repeat every 28 days until participants met criteria for discontinuation of study drug Elotuzumab (BMS-901608; HuLuc63): Solution, IV, 10 mg/kg, weekly, on Days 1, 8, 15, 22 (cycles 1&2); Days 1 and 15 (cycles 3 and beyond) Repeat every 28 days until participants met criteria for discontinuation of study drug

Reporting group title	Lenalidomide + Dexamethasone

Reporting group description:

Lenalidomide: Capsules, Oral, 25 mg, once daily, on Days 1-21 Regimen repeated every 28 days until participants met criteria for discontinuation of study drug Dexamethasone: Tablets, Oral, 40 mg, weekly, on Days 1, 8, 15, 22 Regimen repeated every 28 days until participants met criteria for discontinuation of study drug

Reporting group title	Lenalidomide + Dexamethasone + Elotuzumab

Reporting group description:

Lenalidomide: Capsules, Oral, 25 mg, once daily, on Days 1-21 Repeat every 28 days until participants met criteria for discontinuation of study drug Dexamethasone (Oral): On weeks without Elotuzumab dosing: Tablets, Oral, 40mg Repeat every 28 days until participants met criteria for discontinuation of study drug On weeks with Elotuzumab dosing: Tablets, Oral, 28 mg Repeat every 28 days until participants met criteria for discontinuation of study drug Dexamethasone (IV): On weeks without Elotuzumab dosing: Not Applicable (N/A) On weeks with Elotuzumab dosing: Solution, Intravenous (IV), 8 mg, weekly Repeat every 28 days until participants met criteria for discontinuation of study drug Elotuzumab (BMS-901608; HuLuc63): Solution, IV, 10 mg/kg, weekly, on Days 1, 8, 15, 22 (cycles 1&2); Days 1 and 15 (cycles 3 and beyond) Repeat every 28 days until participants met criteria for discontinuation of study drug

Reporting group title	Lenalidomide + Dexamethasone

Reporting group description:

Lenalidomide: Capsules, Oral, 25 mg, once daily, on Days 1-21 Regimen repeated every 28 days until participants met criteria for discontinuation of study drug Dexamethasone: Tablets, Oral, 40 mg, weekly, on Days 1, 8, 15, 22 Regimen repeated every 28 days until participants met criteria for discontinuation of study drug

Primary: Median Progression Free Survival (PFS)

End point title	Median Progression Free Survival (PFS)

End point description:

Primary definition of Progression-free survival (PFS) defined as the time from randomization to the date of first documented tumor progression or death due to any cause. Participants were censored at the last adequate assessment prior to the start of any subsequent systemic-therapy or at the last adequate assessment prior to 2 missing assessments (> 10 weeks). Participants who died more than 10 weeks after the randomization date and had no on-treatment assessment were censored at the randomization date. Clinical deterioration was not considered progression. The primary analysis of PFS was based on the primary definition using the Independent Review Committee (IRC) tumor assessment using the European Group for Blood and Bone Marrow Transplant (EBMT) criteria. Tumor assessments were made every 4 weeks (±1 week) relative to the first dose of study medication.

End point type	Primary

End point timeframe:

From randomization up to to the date of first documented tumor progression or death (approximately 85 months)

End point values	Lenalidomide + Dexamethasone + Elotuzumab	Lenalidomide + Dexamethasone		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	321	325		
Units: Months				
median (confidence interval 95%)	19.42 (16.62 to 22.31)	14.92 (12.25 to 17.31)		

Statistical analyses

Statistical analysis title	LdE versus Ld
Statistical analysis description:	
Hazard Ratio of Lenalidomide + Dexamethasone + Elotuzumab to Lenalidomide + Dexamethasone	
Comparison groups	Lenalidomide + Dexamethasone + Elotuzumab v Lenalidomide + Dexamethasone
Number of subjects included in analysis	646
Analysis specification	Pre-specified
Analysis type	
P-value	= 0.0005
Method	Logrank
Parameter estimate	Hazard ratio (HR)
Point estimate	0.72
Confidence interval	
level	95 %
sides	2-sided
lower limit	0.6
upper limit	0.87

Primary: Objective Response Rate (ORR)

End point title	Objective Response Rate (ORR)
End point description:	
Objective response rate (ORR) defined as the number of participants with a best response on-study of partial response (PR) or better (stringent CR [sCR], complete response [CR], very good partial response [VGPR], and partial response [PR]) based on the Independent Review Committee (IRC) assessment of best response using the European Group for Blood and Bone Marrow Transplant (EBMT) assessment criteria. Participants were censored at the last adequate assessment prior to the start of any subsequent systemic-therapy or at the last adequate assessment prior to 2 missing assessments (> 10 weeks). Participants who died more than 10 weeks after the randomization date and had no on-treatment assessment were censored at the randomization date. Clinical deterioration was not considered progression. Assessments were made every 4 weeks.	
End point type	Primary
End point timeframe:	
Randomization up to approximately 52 months	

End point values	Lenalidomide + Dexamethasone + Elotuzumab	Lenalidomide + Dexamethasone		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	321	325		
Units: Participants	272	239		

Statistical analyses

Statistical analysis title	Ratio of LdE Ld
Comparison groups	Lenalidomide + Dexamethasone + Elotuzumab v Lenalidomide + Dexamethasone
Number of subjects included in analysis	646
Analysis specification	Pre-specified
Analysis type	
P-value	= 0.0004
Method	Cochran-Mantel-Haenszel
Parameter estimate	Odds ratio (OR)
Point estimate	2.03
Confidence interval	
level	95 %
sides	2-sided
lower limit	1.37
upper limit	3

Statistical analysis title	LdE versus Ld
Statistical analysis description:	
Difference of Lenalidomide + Dexamethasone + Elotuzumab minus Lenalidomide + Dexamethasone computed using the method of DerSimonian and Laird (weighted average over the strata)	
Comparison groups	Lenalidomide + Dexamethasone + Elotuzumab v Lenalidomide + Dexamethasone
Number of subjects included in analysis	646
Analysis specification	Pre-specified
Analysis type	
Parameter estimate	Difference in ORR
Point estimate	10.4
Confidence interval	
level	95 %
sides	2-sided
lower limit	4.4
upper limit	16.4

Secondary: Median Overall Survival (OS)

End point title	Median Overall Survival (OS)

End point description:

Overall survival is defined as the time from randomization to the date of death from any cause. If a subject has not died, their survival time will be censored at the date of last contact ("last known alive date"). A subject will be censored at the date of randomization if they were randomized but had no follow-up. (Based on Kaplan Meier estimates)

End point type	Secondary

End point timeframe:

Randomization to the date of death from any cause (up to approximately 9 years)

End point values	Lenalidomide + Dexamethasone + Elotuzumab	Lenalidomide + Dexamethasone		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	321	325		
Units: Months				
median (confidence interval 95%)	48.30 (40.34 to 51.94)	39.62 (33.25 to 45.27)		

Statistical analyses

No statistical analyses for this end point

Secondary: Change from baseline of mean score pain severity (BPI-SF)

End point title	Change from baseline of mean score pain severity (BPI-SF)

End point description:

The change from baseline of the mean score of pain severity at the end of treatment using the Brief Pain Inventory-Short Form (BPI-SF). The BPI-SF is a self administered questionnaire developed to assess the severity of pain (the sensory dimension) as well as the degree to which pain interferes with function (the reactive dimension). The BPI-SF uses 0 ("No pain", "No interference") to 10 ("Pain as bad as you can imagine", "Highest imaginable interference") numeric rating scale.

End point type	Secondary

End point timeframe:

From baseline up to approximately 38 months

End point values	Lenalidomide + Dexamethasone + Elotuzumab	Lenalidomide + Dexamethasone		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	114	152		
Units: Score on a scale				
arithmetic mean (standard deviation)	0.52 (± 2.237)	-0.04 (± 2.408)		

Statistical analyses

No statistical analyses for this end point

Secondary: Change from baseline of mean score pain Interference (BPI-SF)

End point title	Change from baseline of mean score pain Interference (BPI-SF)

End point description:

The change from baseline of the mean score of pain interference at the end of treatment using the Brief Pain Inventory-Short Form (BPI-SF). The BPI-SF is a self administered questionnaire developed to assess the severity of pain (the sensory dimension) as well as the degree to which pain interferes with function (the reactive dimension). The BPI-SF uses 0 ("No pain", "No interference") to 10 ("Pain as bad as you can imagine", "Highest imaginable interference") numeric rating scale.

End point type	Secondary

End point timeframe:

From baseline up to approximately 38 months

End point values	Lenalidomide + Dexamethasone + Elotuzumab	Lenalidomide + Dexamethasone		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	113	150		
Units: Score on a scale				
arithmetic mean (standard deviation)	0.95 (± 2.466)	0.48 (± 2.868)		

Statistical analyses

No statistical analyses for this end point

Adverse events

Adverse events information	
Timeframe for reporting adverse events:	
From first dose up to 60 days post last dose of study therapy (Up to approximately 9 years)	
Assessment type	Systematic

Dictionary used	
Dictionary name	MedDRA
Dictionary version	24.0

Reporting groups	
Reporting group title	Lenalidomide + Dexamethasone
Reporting group description:	
Lenalidomide: Capsules, Oral, 25 mg, once daily, on Days 1-21 Regimen repeated every 28 days until participants met criteria for discontinuation of study drug Dexamethasone: Tablets, Oral, 40 mg, weekly, on Days 1, 8, 15, 22 Regimen repeated every 28 days until participants met criteria for discontinuation of study drug	
Reporting group title	Lenalidomide + Dexamethasone + Elotuzumab
Reporting group description:	
Lenalidomide: Capsules, Oral, 25 mg, once daily, on Days 1-21 Repeat every 28 days until participants met criteria for discontinuation of study drug Dexamethasone (Oral): On weeks without Elotuzumab dosing: Tablets, Oral, 40mg Repeat every 28 days until participants met criteria for discontinuation of study drug On weeks with Elotuzumab dosing: Tablets, Oral, 28 mg Repeat every 28 days until participants met criteria for discontinuation of study drug Dexamethasone (IV): On weeks without Elotuzumab dosing: Not Applicable (N/A) On weeks with Elotuzumab dosing: Solution, Intravenous (IV), 8 mg, weekly Repeat every 28 days until participants met criteria for discontinuation of study drug Elotuzumab (BMS-901608; HuLuc63): Solution, IV, 10 mg/kg, weekly, on Days 1, 8, 15, 22 (cycles 1&2); Days 1 and 15 (cycles 3 and beyond) Repeat every 28 days until participants met criteria for discontinuation of study drug	

Serious adverse events	Lenalidomide + Dexamethasone	Lenalidomide + Dexamethasone + Elotuzumab	
Total subjects affected by serious adverse events			
subjects affected / exposed	194 / 317 (61.20%)	238 / 318 (74.84%)	
number of deaths (all causes)	249	226	
number of deaths resulting from adverse events			
Neoplasms benign, malignant and unspecified (incl cysts and polyps)			
Adenocarcinoma of colon			
subjects affected / exposed	1 / 317 (0.32%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 1	0 / 0	
Bladder transitional cell carcinoma			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	

Haemangioma of bone			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Malignant melanoma in situ			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	1 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Malignant neoplasm of unknown primary site			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 1	0 / 0	
Squamous cell carcinoma			
subjects affected / exposed	3 / 317 (0.95%)	2 / 318 (0.63%)	
occurrences causally related to treatment / all	1 / 6	1 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Lung neoplasm malignant			
subjects affected / exposed	0 / 317 (0.00%)	4 / 318 (1.26%)	
occurrences causally related to treatment / all	0 / 0	0 / 5	
deaths causally related to treatment / all	0 / 0	0 / 3	
Malignant neoplasm progression			
subjects affected / exposed	4 / 317 (1.26%)	5 / 318 (1.57%)	
occurrences causally related to treatment / all	0 / 4	0 / 5	
deaths causally related to treatment / all	0 / 4	0 / 3	
Myelodysplastic syndrome			
subjects affected / exposed	3 / 317 (0.95%)	2 / 318 (0.63%)	
occurrences causally related to treatment / all	2 / 3	2 / 2	
deaths causally related to treatment / all	0 / 1	1 / 1	
Prostatic adenoma			
subjects affected / exposed	1 / 317 (0.32%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Plasmacytoma			

subjects affected / exposed	3 / 317 (0.95%)	2 / 318 (0.63%)	
occurrences causally related to treatment / all	0 / 3	0 / 2	
deaths causally related to treatment / all	0 / 1	0 / 0	
Tumour associated fever			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Basal cell carcinoma			
subjects affected / exposed	4 / 317 (1.26%)	10 / 318 (3.14%)	
occurrences causally related to treatment / all	1 / 4	5 / 12	
deaths causally related to treatment / all	0 / 0	0 / 0	
Endometrial cancer			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Gastrointestinal neoplasm			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	1 / 1	
deaths causally related to treatment / all	0 / 0	1 / 1	
Meningioma			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Prostate cancer			
subjects affected / exposed	2 / 317 (0.63%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 2	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Plasma cell myeloma			
subjects affected / exposed	5 / 317 (1.58%)	6 / 318 (1.89%)	
occurrences causally related to treatment / all	0 / 5	0 / 6	
deaths causally related to treatment / all	0 / 3	0 / 2	
Haemangioma			

subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Squamous cell carcinoma of skin			
subjects affected / exposed	3 / 317 (0.95%)	12 / 318 (3.77%)	
occurrences causally related to treatment / all	0 / 8	4 / 15	
deaths causally related to treatment / all	0 / 0	0 / 0	
Tonsil cancer			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	1 / 1	0 / 0	
deaths causally related to treatment / all	1 / 1	0 / 0	
Chronic lymphocytic leukaemia			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Malignant pleural effusion			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Mesothelioma			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Lung cancer metastatic			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Acute erythroid leukaemia			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Acute myeloid leukaemia			

subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Atypical fibroxanthoma			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Breast cancer stage I			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cervix carcinoma			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Colorectal adenocarcinoma			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
External ear neoplasm malignant			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Lip squamous cell carcinoma			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	1 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Neoplasm malignant			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 1	
Non-small cell lung cancer			

subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 1	
Plasma cell myeloma recurrent			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 1	0 / 0	
Rectal adenocarcinoma			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	1 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Sarcoma			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Vulval cancer			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Vascular disorders			
Aortic aneurysm rupture			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 1	
Orthostatic hypotension			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Thrombosis			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	1 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Deep vein thrombosis			

subjects affected / exposed	4 / 317 (1.26%)	10 / 318 (3.14%)	
occurrences causally related to treatment / all	3 / 4	10 / 10	
deaths causally related to treatment / all	0 / 0	0 / 0	
Subclavian vein thrombosis			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Thrombophlebitis			
subjects affected / exposed	2 / 317 (0.63%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	1 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Peripheral ischaemia			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Thrombophlebitis superficial			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	1 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Embolism			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	1 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hypotension			
subjects affected / exposed	1 / 317 (0.32%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 1	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Peripheral artery thrombosis			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Venous thrombosis			

subjects affected / exposed	2 / 317 (0.63%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	2 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pelvic venous thrombosis			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Venous thrombosis limb			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hypovolaemic shock			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 1	
Surgical and medical procedures			
Hip arthroplasty			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
General disorders and administration site conditions			
Chest pain			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pain			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pyrexia			
subjects affected / exposed	17 / 317 (5.36%)	25 / 318 (7.86%)	
occurrences causally related to treatment / all	5 / 24	9 / 26	
deaths causally related to treatment / all	0 / 0	0 / 0	

Asthenia			
subjects affected / exposed	0 / 317 (0.00%)	4 / 318 (1.26%)	
occurrences causally related to treatment / all	0 / 0	0 / 4	
deaths causally related to treatment / all	0 / 0	0 / 0	
Chills			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Fatigue			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Death			
subjects affected / exposed	1 / 317 (0.32%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 1	0 / 1	
Disease progression			
subjects affected / exposed	10 / 317 (3.15%)	15 / 318 (4.72%)	
occurrences causally related to treatment / all	0 / 10	0 / 15	
deaths causally related to treatment / all	0 / 8	0 / 11	
Malaise			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
General physical health deterioration			
subjects affected / exposed	4 / 317 (1.26%)	5 / 318 (1.57%)	
occurrences causally related to treatment / all	1 / 4	1 / 5	
deaths causally related to treatment / all	1 / 4	0 / 2	
Hernia			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Multiple organ dysfunction syndrome			

subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Immune system disorders			
Haemophagocytic lymphohistiocytosis			
subjects affected / exposed	1 / 317 (0.32%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	1 / 1	1 / 1	
deaths causally related to treatment / all	0 / 0	1 / 1	
Reproductive system and breast disorders			
Benign prostatic hyperplasia			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	1 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Ovarian cyst			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Respiratory, thoracic and mediastinal disorders			
Epistaxis			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pulmonary alveolar haemorrhage			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Obliterative bronchiolitis			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cough			

subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Dyspnoea			
subjects affected / exposed	4 / 317 (1.26%)	4 / 318 (1.26%)	
occurrences causally related to treatment / all	0 / 4	1 / 6	
deaths causally related to treatment / all	0 / 0	0 / 0	
Haemoptysis			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Organising pneumonia			
subjects affected / exposed	0 / 317 (0.00%)	2 / 318 (0.63%)	
occurrences causally related to treatment / all	0 / 0	1 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pleural effusion			
subjects affected / exposed	1 / 317 (0.32%)	2 / 318 (0.63%)	
occurrences causally related to treatment / all	0 / 1	1 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Respiratory failure			
subjects affected / exposed	1 / 317 (0.32%)	2 / 318 (0.63%)	
occurrences causally related to treatment / all	0 / 1	1 / 4	
deaths causally related to treatment / all	0 / 0	0 / 0	
Acute respiratory failure			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Bronchial disorder			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Chronic obstructive pulmonary disease			

subjects affected / exposed	3 / 317 (0.95%)	5 / 318 (1.57%)	
occurrences causally related to treatment / all	1 / 4	0 / 7	
deaths causally related to treatment / all	0 / 1	0 / 0	
Lung disorder			
subjects affected / exposed	1 / 317 (0.32%)	4 / 318 (1.26%)	
occurrences causally related to treatment / all	1 / 1	2 / 4	
deaths causally related to treatment / all	0 / 0	0 / 1	
Bronchial hyperreactivity			
subjects affected / exposed	0 / 317 (0.00%)	2 / 318 (0.63%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pneumonitis			
subjects affected / exposed	2 / 317 (0.63%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	2 / 2	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pulmonary embolism			
subjects affected / exposed	8 / 317 (2.52%)	11 / 318 (3.46%)	
occurrences causally related to treatment / all	6 / 9	10 / 11	
deaths causally related to treatment / all	1 / 1	1 / 1	
Hypoxia			
subjects affected / exposed	1 / 317 (0.32%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Interstitial lung disease			
subjects affected / exposed	3 / 317 (0.95%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	1 / 3	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Acute pulmonary oedema			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Asthma			

subjects affected / exposed	1 / 317 (0.32%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Acute respiratory distress syndrome			
subjects affected / exposed	1 / 317 (0.32%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 1	0 / 0	
Pulmonary artery thrombosis			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pulmonary fibrosis			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Psychiatric disorders			
Completed suicide			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 1	
Depression			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Mental status changes			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Confusional state			
subjects affected / exposed	1 / 317 (0.32%)	4 / 318 (1.26%)	
occurrences causally related to treatment / all	1 / 1	1 / 5	
deaths causally related to treatment / all	0 / 0	0 / 0	
Delirium			

subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Product issues			
Device dislocation			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Investigations			
International normalised ratio increased			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Blood creatinine increased			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 1	0 / 0	
Viral test positive			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Liver function test abnormal			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Escherichia test positive			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Haemoglobin decreased			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	

Clostridium test positive			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hepatic enzyme increased			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Influenza A virus test positive			
subjects affected / exposed	1 / 317 (0.32%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	1 / 1	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
General physical condition abnormal			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Influenza B virus test positive			
subjects affected / exposed	1 / 317 (0.32%)	2 / 318 (0.63%)	
occurrences causally related to treatment / all	1 / 1	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Polymerase chain reaction positive			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Injury, poisoning and procedural complications			
Femoral neck fracture			
subjects affected / exposed	1 / 317 (0.32%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hip fracture			
subjects affected / exposed	2 / 317 (0.63%)	2 / 318 (0.63%)	
occurrences causally related to treatment / all	0 / 2	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	

Humerus fracture			
subjects affected / exposed	0 / 317 (0.00%)	2 / 318 (0.63%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Rib fracture			
subjects affected / exposed	0 / 317 (0.00%)	2 / 318 (0.63%)	
occurrences causally related to treatment / all	0 / 0	1 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Subdural haematoma			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 1	0 / 0	
Upper limb fracture			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Compression fracture			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Fall			
subjects affected / exposed	1 / 317 (0.32%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Ilium fracture			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Lumbar vertebral fracture			
subjects affected / exposed	1 / 317 (0.32%)	2 / 318 (0.63%)	
occurrences causally related to treatment / all	0 / 1	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Stenosis of vesicourethral anastomosis			

subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Ankle fracture			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pelvic fracture			
subjects affected / exposed	2 / 317 (0.63%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 2	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Seroma			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Spinal compression fracture			
subjects affected / exposed	1 / 317 (0.32%)	3 / 318 (0.94%)	
occurrences causally related to treatment / all	0 / 1	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Femur fracture			
subjects affected / exposed	1 / 317 (0.32%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Craniocerebral injury			
subjects affected / exposed	2 / 317 (0.63%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 1	0 / 0	
Soft tissue injury			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Thoracic vertebral fracture			

subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Lip injury			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Post procedural haemorrhage			
subjects affected / exposed	2 / 317 (0.63%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 3	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Tendon rupture			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Traumatic fracture			
subjects affected / exposed	2 / 317 (0.63%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Road traffic accident			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 1	0 / 0	
Accidental overdose			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Chemical burn			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Joint dislocation			

subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Limb injury			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Overdose			
subjects affected / exposed	1 / 317 (0.32%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Procedural pain			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Skin laceration			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Spinal fracture			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Toxicity to various agents			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cardiac disorders			
Acute coronary syndrome			
subjects affected / exposed	2 / 317 (0.63%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 2	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Arrhythmia			

subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cardio-respiratory arrest			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 1	
Myocardial infarction			
subjects affected / exposed	3 / 317 (0.95%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 3	0 / 0	
deaths causally related to treatment / all	0 / 1	0 / 0	
Left ventricular failure			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	1 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cardiac failure congestive			
subjects affected / exposed	1 / 317 (0.32%)	2 / 318 (0.63%)	
occurrences causally related to treatment / all	0 / 1	2 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Acute myocardial infarction			
subjects affected / exposed	2 / 317 (0.63%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 2	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Angina pectoris			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Atrial fibrillation			
subjects affected / exposed	9 / 317 (2.84%)	6 / 318 (1.89%)	
occurrences causally related to treatment / all	1 / 9	0 / 7	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cardiac failure			

subjects affected / exposed	3 / 317 (0.95%)	3 / 318 (0.94%)	
occurrences causally related to treatment / all	2 / 5	0 / 4	
deaths causally related to treatment / all	0 / 0	0 / 2	
Atrioventricular block complete			
subjects affected / exposed	0 / 317 (0.00%)	2 / 318 (0.63%)	
occurrences causally related to treatment / all	0 / 0	1 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cardiac arrest			
subjects affected / exposed	1 / 317 (0.32%)	3 / 318 (0.94%)	
occurrences causally related to treatment / all	0 / 1	1 / 3	
deaths causally related to treatment / all	0 / 1	0 / 2	
Myocardial ischaemia			
subjects affected / exposed	2 / 317 (0.63%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	1 / 2	0 / 0	
deaths causally related to treatment / all	0 / 1	0 / 0	
Supraventricular tachycardia			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Acute left ventricular failure			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Angina unstable			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Brugada syndrome			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cardiac aneurysm			

subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pericarditis			
subjects affected / exposed	1 / 317 (0.32%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	1 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cardiogenic shock			
subjects affected / exposed	0 / 317 (0.00%)	2 / 318 (0.63%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Sinus node dysfunction			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Nervous system disorders			
Cerebrovascular accident			
subjects affected / exposed	3 / 317 (0.95%)	2 / 318 (0.63%)	
occurrences causally related to treatment / all	1 / 4	1 / 2	
deaths causally related to treatment / all	0 / 0	0 / 1	
Hemiparesis			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	2 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hypoaesthesia			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Syncope			
subjects affected / exposed	2 / 317 (0.63%)	3 / 318 (0.94%)	
occurrences causally related to treatment / all	0 / 2	0 / 4	
deaths causally related to treatment / all	0 / 0	0 / 0	
Paraesthesia			

subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Somnolence			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Transient ischaemic attack			
subjects affected / exposed	1 / 317 (0.32%)	2 / 318 (0.63%)	
occurrences causally related to treatment / all	1 / 1	1 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cerebral ischaemia			
subjects affected / exposed	1 / 317 (0.32%)	2 / 318 (0.63%)	
occurrences causally related to treatment / all	0 / 1	1 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Dementia			
subjects affected / exposed	0 / 317 (0.00%)	2 / 318 (0.63%)	
occurrences causally related to treatment / all	0 / 0	1 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Dizziness			
subjects affected / exposed	0 / 317 (0.00%)	2 / 318 (0.63%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Presyncope			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Status epilepticus			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	1 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Optic neuritis			

subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Encephalopathy			
subjects affected / exposed	2 / 317 (0.63%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 1	0 / 0	
Paraparesis			
subjects affected / exposed	1 / 317 (0.32%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cerebral infarction			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	3 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Neuropathy peripheral			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	1 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Spinal cord disorder			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cerebral haemorrhage			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 1	0 / 0	
Spinal cord compression			
subjects affected / exposed	2 / 317 (0.63%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 2	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cerebrospinal fluid leakage			

subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hepatic encephalopathy			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Ischaemic stroke			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Nervous system disorder			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Sciatica			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Seizure			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Blood and lymphatic system disorders			
Febrile neutropenia			
subjects affected / exposed	4 / 317 (1.26%)	5 / 318 (1.57%)	
occurrences causally related to treatment / all	2 / 5	4 / 6	
deaths causally related to treatment / all	0 / 0	0 / 1	
Neutropenia			
subjects affected / exposed	4 / 317 (1.26%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	4 / 5	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pancytopenia			

subjects affected / exposed	0 / 317 (0.00%)	2 / 318 (0.63%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Splenic infarction			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	1 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Eosinophilia			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Anaemia			
subjects affected / exposed	8 / 317 (2.52%)	11 / 318 (3.46%)	
occurrences causally related to treatment / all	2 / 9	2 / 17	
deaths causally related to treatment / all	0 / 0	0 / 0	
Bone marrow failure			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Thrombocytopenia			
subjects affected / exposed	2 / 317 (0.63%)	5 / 318 (1.57%)	
occurrences causally related to treatment / all	0 / 2	1 / 5	
deaths causally related to treatment / all	0 / 0	0 / 0	
Microangiopathic haemolytic anaemia			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Eye disorders			
Visual impairment			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cataract nuclear			

subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cataract			
subjects affected / exposed	6 / 317 (1.89%)	6 / 318 (1.89%)	
occurrences causally related to treatment / all	4 / 7	4 / 7	
deaths causally related to treatment / all	0 / 0	0 / 0	
Gastrointestinal disorders			
Ileus			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Ileus paralytic			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Large intestinal ulcer			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	1 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Abdominal pain			
subjects affected / exposed	0 / 317 (0.00%)	3 / 318 (0.94%)	
occurrences causally related to treatment / all	0 / 0	0 / 4	
deaths causally related to treatment / all	0 / 0	0 / 0	
Constipation			
subjects affected / exposed	2 / 317 (0.63%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	1 / 2	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Gastrointestinal haemorrhage			
subjects affected / exposed	1 / 317 (0.32%)	2 / 318 (0.63%)	
occurrences causally related to treatment / all	0 / 1	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Intestinal obstruction			

subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Large intestine perforation			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pancreatitis acute			
subjects affected / exposed	0 / 317 (0.00%)	2 / 318 (0.63%)	
occurrences causally related to treatment / all	0 / 0	1 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Colitis			
subjects affected / exposed	2 / 317 (0.63%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	1 / 2	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Enteritis			
subjects affected / exposed	0 / 317 (0.00%)	2 / 318 (0.63%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Gastric ulcer			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Haemorrhoids			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	1 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Intestinal haemorrhage			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 1	0 / 0	
Mouth haemorrhage			

subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pancreatitis			
subjects affected / exposed	1 / 317 (0.32%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	1 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Upper gastrointestinal haemorrhage			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	1 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Vomiting			
subjects affected / exposed	4 / 317 (1.26%)	2 / 318 (0.63%)	
occurrences causally related to treatment / all	3 / 5	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Nausea			
subjects affected / exposed	1 / 317 (0.32%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	2 / 2	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Inguinal hernia, obstructive			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Lower gastrointestinal haemorrhage			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	1 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Subileus			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	1 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Abdominal pain upper			

subjects affected / exposed	0 / 317 (0.00%)	2 / 318 (0.63%)	
occurrences causally related to treatment / all	0 / 0	1 / 2	
deaths causally related to treatment / all	0 / 0	0 / 1	
Diarrhoea			
subjects affected / exposed	12 / 317 (3.79%)	7 / 318 (2.20%)	
occurrences causally related to treatment / all	2 / 13	5 / 9	
deaths causally related to treatment / all	0 / 0	0 / 0	
Diverticulum			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Enterocolitis			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Enterocutaneous fistula			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hiatus hernia			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Gastric haemorrhage			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Intestinal ischaemia			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 1	
Inguinal hernia			

subjects affected / exposed	1 / 317 (0.32%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Irritable bowel syndrome			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Rectal haemorrhage			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hepatobiliary disorders			
Cholelithiasis			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hepatitis			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cholangitis			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hepatic failure			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hepatic function abnormal			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hyperbilirubinaemia			

subjects affected / exposed	0 / 317 (0.00%)	2 / 318 (0.63%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cholecystitis			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cholecystitis acute			
subjects affected / exposed	0 / 317 (0.00%)	4 / 318 (1.26%)	
occurrences causally related to treatment / all	0 / 0	0 / 4	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cholecystitis chronic			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Skin and subcutaneous tissue disorders			
Pustular psoriasis			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Skin haemorrhage			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Rash maculo-papular			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Skin ulcer			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Dermal cyst			

subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Renal and urinary disorders			
Haemorrhage urinary tract			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Nephrolithiasis			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Urinary retention			
subjects affected / exposed	3 / 317 (0.95%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 3	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Neurogenic bladder			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Renal tubular acidosis			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Prerenal failure			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 1	
Renal failure			
subjects affected / exposed	5 / 317 (1.58%)	4 / 318 (1.26%)	
occurrences causally related to treatment / all	1 / 7	0 / 5	
deaths causally related to treatment / all	0 / 0	0 / 0	
Renal impairment			

subjects affected / exposed	0 / 317 (0.00%)	3 / 318 (0.94%)	
occurrences causally related to treatment / all	0 / 0	1 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Urinary tract obstruction			
subjects affected / exposed	1 / 317 (0.32%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Acute kidney injury			
subjects affected / exposed	8 / 317 (2.52%)	11 / 318 (3.46%)	
occurrences causally related to treatment / all	2 / 8	2 / 12	
deaths causally related to treatment / all	0 / 2	0 / 0	
Haematuria			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Myeloma cast nephropathy			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Endocrine disorders			
Hypothyroidism			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	1 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Musculoskeletal and connective tissue disorders			
Arthralgia			
subjects affected / exposed	2 / 317 (0.63%)	2 / 318 (0.63%)	
occurrences causally related to treatment / all	0 / 2	1 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Bone pain			
subjects affected / exposed	0 / 317 (0.00%)	3 / 318 (0.94%)	
occurrences causally related to treatment / all	0 / 0	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	

Musculoskeletal pain			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pathological fracture			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cervical spinal stenosis			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Muscle mass			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Muscular weakness			
subjects affected / exposed	0 / 317 (0.00%)	2 / 318 (0.63%)	
occurrences causally related to treatment / all	0 / 0	1 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Musculoskeletal chest pain			
subjects affected / exposed	1 / 317 (0.32%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pain in extremity			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Rhabdomyolysis			
subjects affected / exposed	3 / 317 (0.95%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 3	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Back pain			

subjects affected / exposed	5 / 317 (1.58%)	7 / 318 (2.20%)	
occurrences causally related to treatment / all	0 / 5	0 / 8	
deaths causally related to treatment / all	0 / 0	0 / 0	
Bone lesion			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Osteonecrosis of jaw			
subjects affected / exposed	6 / 317 (1.89%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 6	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Exostosis			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Osteonecrosis			
subjects affected / exposed	1 / 317 (0.32%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 1	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Arthritis			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Bone cyst			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Osteoarthritis			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Spinal stenosis			

subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Tendonitis			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Infections and infestations			
Diverticulitis			
subjects affected / exposed	0 / 317 (0.00%)	2 / 318 (0.63%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pyelonephritis			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Respiratory syncytial virus infection			
subjects affected / exposed	1 / 317 (0.32%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 1	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Skin infection			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	1 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Streptococcal bacteraemia			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	1 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Bronchitis			
subjects affected / exposed	8 / 317 (2.52%)	8 / 318 (2.52%)	
occurrences causally related to treatment / all	3 / 8	2 / 11	
deaths causally related to treatment / all	0 / 0	0 / 0	
Clostridium difficile infection			

subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cytomegalovirus infection			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Endocarditis			
subjects affected / exposed	2 / 317 (0.63%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 2	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Enteritis infectious			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Meningitis staphylococcal			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 1	
Neutropenic sepsis			
subjects affected / exposed	1 / 317 (0.32%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	1 / 1	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pyelonephritis acute			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Staphylococcal sepsis			
subjects affected / exposed	1 / 317 (0.32%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	1 / 2	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Urinary tract infection			

subjects affected / exposed	5 / 317 (1.58%)	3 / 318 (0.94%)	
occurrences causally related to treatment / all	1 / 5	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Urosepsis			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	2 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Abscess oral			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cerebral aspergillosis			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Osteomyelitis			
subjects affected / exposed	1 / 317 (0.32%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 1	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pneumonia pneumococcal			
subjects affected / exposed	1 / 317 (0.32%)	3 / 318 (0.94%)	
occurrences causally related to treatment / all	0 / 1	2 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Staphylococcal bacteraemia			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Tooth abscess			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Bacterial sepsis			

subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cellulitis			
subjects affected / exposed	1 / 317 (0.32%)	8 / 318 (2.52%)	
occurrences causally related to treatment / all	0 / 2	1 / 8	
deaths causally related to treatment / all	0 / 0	0 / 0	
Enterocolitis viral			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Gastroenteritis			
subjects affected / exposed	2 / 317 (0.63%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	1 / 2	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pneumonia fungal			
subjects affected / exposed	0 / 317 (0.00%)	2 / 318 (0.63%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 1	
Viral diarrhoea			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Viral infection			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Wound infection			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Atypical pneumonia			

subjects affected / exposed	0 / 317 (0.00%)	3 / 318 (0.94%)	
occurrences causally related to treatment / all	0 / 0	1 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Bacteraemia			
subjects affected / exposed	1 / 317 (0.32%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Device related sepsis			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	1 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Febrile infection			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Herpes zoster			
subjects affected / exposed	1 / 317 (0.32%)	4 / 318 (1.26%)	
occurrences causally related to treatment / all	1 / 1	2 / 4	
deaths causally related to treatment / all	0 / 0	0 / 0	
Infection			
subjects affected / exposed	1 / 317 (0.32%)	3 / 318 (0.94%)	
occurrences causally related to treatment / all	0 / 1	1 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Listeriosis			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	1 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Lower respiratory tract infection			
subjects affected / exposed	4 / 317 (1.26%)	8 / 318 (2.52%)	
occurrences causally related to treatment / all	1 / 5	4 / 9	
deaths causally related to treatment / all	0 / 0	1 / 1	
Periodontitis			

subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Peritonitis			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	1 / 1	0 / 0	
deaths causally related to treatment / all	1 / 1	0 / 0	
Pharyngitis			
subjects affected / exposed	2 / 317 (0.63%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 2	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pneumonia			
subjects affected / exposed	40 / 317 (12.62%)	56 / 318 (17.61%)	
occurrences causally related to treatment / all	14 / 49	30 / 74	
deaths causally related to treatment / all	0 / 3	0 / 3	
Septic shock			
subjects affected / exposed	5 / 317 (1.58%)	2 / 318 (0.63%)	
occurrences causally related to treatment / all	3 / 6	0 / 2	
deaths causally related to treatment / all	0 / 1	0 / 1	
Sinusitis			
subjects affected / exposed	1 / 317 (0.32%)	2 / 318 (0.63%)	
occurrences causally related to treatment / all	1 / 1	1 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Influenza			
subjects affected / exposed	4 / 317 (1.26%)	3 / 318 (0.94%)	
occurrences causally related to treatment / all	0 / 4	2 / 4	
deaths causally related to treatment / all	0 / 0	1 / 1	
Pneumocystis jirovecii pneumonia			
subjects affected / exposed	1 / 317 (0.32%)	2 / 318 (0.63%)	
occurrences causally related to treatment / all	1 / 1	1 / 2	
deaths causally related to treatment / all	1 / 1	0 / 0	
Pneumonia bacterial			

subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pneumonia influenzal			
subjects affected / exposed	2 / 317 (0.63%)	2 / 318 (0.63%)	
occurrences causally related to treatment / all	0 / 2	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Respiratory tract infection			
subjects affected / exposed	4 / 317 (1.26%)	11 / 318 (3.46%)	
occurrences causally related to treatment / all	2 / 4	6 / 11	
deaths causally related to treatment / all	0 / 0	0 / 0	
Sepsis			
subjects affected / exposed	8 / 317 (2.52%)	5 / 318 (1.57%)	
occurrences causally related to treatment / all	3 / 8	1 / 5	
deaths causally related to treatment / all	2 / 5	0 / 2	
Upper respiratory tract infection			
subjects affected / exposed	1 / 317 (0.32%)	2 / 318 (0.63%)	
occurrences causally related to treatment / all	0 / 1	1 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Bronchopulmonary aspergillosis			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	1 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Device related infection			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Lung abscess			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pneumococcal sepsis			

subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Acute sinusitis			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Arthritis bacterial			
subjects affected / exposed	2 / 317 (0.63%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Campylobacter gastroenteritis			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Gastroenteritis viral			
subjects affected / exposed	4 / 317 (1.26%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	2 / 4	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
H1N1 influenza			
subjects affected / exposed	1 / 317 (0.32%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	1 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Ophthalmic herpes zoster			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Urinary tract infection enterococcal			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Abdominal abscess			

subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Appendicitis			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Biliary sepsis			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Brain abscess			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	1 / 1	
deaths causally related to treatment / all	0 / 0	1 / 1	
Bursitis infective			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Catheter site abscess			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Clostridium colitis			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Clostridium difficile colitis			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cytomegalovirus chorioretinitis			

subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Epididymitis			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Escherichia urinary tract infection			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Gastroenteritis clostridial			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Gastroenteritis norovirus			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hepatitis B			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Genital abscess			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	1 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Herpes zoster reactivation			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Infective exacerbation of chronic obstructive airways disease			

subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Intervertebral discitis			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	1 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Large intestine infection			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Metapneumovirus infection			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Localised infection			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Otitis media chronic			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Parvovirus B19 infection			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	1 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Post procedural infection			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pseudomonal sepsis			

subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pseudomonas infection			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pulmonary sepsis			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Retinitis			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Rotavirus infection			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Systemic infection			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Varicella zoster virus infection			
subjects affected / exposed	0 / 317 (0.00%)	2 / 318 (0.63%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Metabolism and nutrition disorders			
Dehydration			
subjects affected / exposed	1 / 317 (0.32%)	2 / 318 (0.63%)	
occurrences causally related to treatment / all	0 / 1	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hyponatraemia			

subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	1 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Failure to thrive			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hypoglycaemia			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hypocalcaemia			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hypokalaemia			
subjects affected / exposed	2 / 317 (0.63%)	3 / 318 (0.94%)	
occurrences causally related to treatment / all	0 / 2	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hypercalcaemia			
subjects affected / exposed	2 / 317 (0.63%)	3 / 318 (0.94%)	
occurrences causally related to treatment / all	1 / 2	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Tumour lysis syndrome			
subjects affected / exposed	1 / 317 (0.32%)	0 / 318 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Decreased appetite			
subjects affected / exposed	1 / 317 (0.32%)	2 / 318 (0.63%)	
occurrences causally related to treatment / all	1 / 1	1 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Diabetes mellitus			

subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hyperglycaemia			
subjects affected / exposed	1 / 317 (0.32%)	3 / 318 (0.94%)	
occurrences causally related to treatment / all	1 / 1	2 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Fluid retention			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Ketoacidosis			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Type 2 diabetes mellitus			
subjects affected / exposed	0 / 317 (0.00%)	1 / 318 (0.31%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	

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

Non-serious adverse events	Lenalidomide + Dexamethasone	Lenalidomide + Dexamethasone + Elotuzumab	
Total subjects affected by non-serious adverse events			
subjects affected / exposed	309 / 317 (97.48%)	314 / 318 (98.74%)	
Vascular disorders			
Hypertension			
subjects affected / exposed	23 / 317 (7.26%)	40 / 318 (12.58%)	
occurrences (all)	33	62	
Deep vein thrombosis			
subjects affected / exposed	10 / 317 (3.15%)	19 / 318 (5.97%)	
occurrences (all)	10	21	
Hypotension			

subjects affected / exposed	12 / 317 (3.79%)	33 / 318 (10.38%)	
occurrences (all)	13	44	
Flushing			
subjects affected / exposed	6 / 317 (1.89%)	16 / 318 (5.03%)	
occurrences (all)	6	21	
Haematoma			
subjects affected / exposed	8 / 317 (2.52%)	17 / 318 (5.35%)	
occurrences (all)	9	18	
General disorders and administration site conditions			
Oedema peripheral			
subjects affected / exposed	78 / 317 (24.61%)	97 / 318 (30.50%)	
occurrences (all)	100	172	
Chest pain			
subjects affected / exposed	14 / 317 (4.42%)	28 / 318 (8.81%)	
occurrences (all)	14	32	
Influenza like illness			
subjects affected / exposed	16 / 317 (5.05%)	27 / 318 (8.49%)	
occurrences (all)	24	38	
Pyrexia			
subjects affected / exposed	74 / 317 (23.34%)	121 / 318 (38.05%)	
occurrences (all)	106	235	
Asthenia			
subjects affected / exposed	54 / 317 (17.03%)	80 / 318 (25.16%)	
occurrences (all)	85	128	
Chills			
subjects affected / exposed	12 / 317 (3.79%)	31 / 318 (9.75%)	
occurrences (all)	12	38	
Fatigue			
subjects affected / exposed	131 / 317 (41.32%)	154 / 318 (48.43%)	
occurrences (all)	159	229	
Malaise			
subjects affected / exposed	12 / 317 (3.79%)	19 / 318 (5.97%)	
occurrences (all)	12	24	
Pain			

subjects affected / exposed	7 / 317 (2.21%)	17 / 318 (5.35%)	
occurrences (all)	8	19	
Respiratory, thoracic and mediastinal disorders			
Dyspnoea exertional			
subjects affected / exposed	15 / 317 (4.73%)	20 / 318 (6.29%)	
occurrences (all)	15	21	
Epistaxis			
subjects affected / exposed	19 / 317 (5.99%)	22 / 318 (6.92%)	
occurrences (all)	22	24	
Oropharyngeal pain			
subjects affected / exposed	15 / 317 (4.73%)	32 / 318 (10.06%)	
occurrences (all)	20	48	
Cough			
subjects affected / exposed	62 / 317 (19.56%)	109 / 318 (34.28%)	
occurrences (all)	95	188	
Dyspnoea			
subjects affected / exposed	60 / 317 (18.93%)	72 / 318 (22.64%)	
occurrences (all)	74	108	
Productive cough			
subjects affected / exposed	5 / 317 (1.58%)	25 / 318 (7.86%)	
occurrences (all)	5	32	
Dysphonia			
subjects affected / exposed	32 / 317 (10.09%)	25 / 318 (7.86%)	
occurrences (all)	33	28	
Psychiatric disorders			
Mood altered			
subjects affected / exposed	8 / 317 (2.52%)	23 / 318 (7.23%)	
occurrences (all)	8	25	
Depression			
subjects affected / exposed	14 / 317 (4.42%)	18 / 318 (5.66%)	
occurrences (all)	15	19	
Insomnia			
subjects affected / exposed	83 / 317 (26.18%)	79 / 318 (24.84%)	
occurrences (all)	112	90	
Anxiety			

subjects affected / exposed	24 / 317 (7.57%)	26 / 318 (8.18%)	
occurrences (all)	26	27	
Confusional state			
subjects affected / exposed	10 / 317 (3.15%)	18 / 318 (5.66%)	
occurrences (all)	10	28	
Investigations			
Blood creatinine increased			
subjects affected / exposed	27 / 317 (8.52%)	36 / 318 (11.32%)	
occurrences (all)	38	76	
Aspartate aminotransferase increased			
subjects affected / exposed	31 / 317 (9.78%)	21 / 318 (6.60%)	
occurrences (all)	49	26	
Weight decreased			
subjects affected / exposed	20 / 317 (6.31%)	52 / 318 (16.35%)	
occurrences (all)	21	54	
Alanine aminotransferase increased			
subjects affected / exposed	33 / 317 (10.41%)	25 / 318 (7.86%)	
occurrences (all)	70	37	
Platelet count decreased			
subjects affected / exposed	5 / 317 (1.58%)	16 / 318 (5.03%)	
occurrences (all)	8	28	
C-reactive protein increased			
subjects affected / exposed	16 / 317 (5.05%)	13 / 318 (4.09%)	
occurrences (all)	25	25	
Injury, poisoning and procedural complications			
Contusion			
subjects affected / exposed	28 / 317 (8.83%)	44 / 318 (13.84%)	
occurrences (all)	33	54	
Fall			
subjects affected / exposed	14 / 317 (4.42%)	21 / 318 (6.60%)	
occurrences (all)	21	27	
Nervous system disorders			
Hypoaesthesia			
subjects affected / exposed	12 / 317 (3.79%)	28 / 318 (8.81%)	
occurrences (all)	12	31	
Headache			

subjects affected / exposed	29 / 317 (9.15%)	54 / 318 (16.98%)	
occurrences (all)	53	104	
Paraesthesia			
subjects affected / exposed	29 / 317 (9.15%)	35 / 318 (11.01%)	
occurrences (all)	35	40	
Peripheral sensory neuropathy			
subjects affected / exposed	39 / 317 (12.30%)	33 / 318 (10.38%)	
occurrences (all)	47	37	
Dizziness			
subjects affected / exposed	38 / 317 (11.99%)	48 / 318 (15.09%)	
occurrences (all)	45	76	
Tremor			
subjects affected / exposed	29 / 317 (9.15%)	30 / 318 (9.43%)	
occurrences (all)	35	34	
Neuropathy peripheral			
subjects affected / exposed	31 / 317 (9.78%)	51 / 318 (16.04%)	
occurrences (all)	34	56	
Taste disorder			
subjects affected / exposed	14 / 317 (4.42%)	18 / 318 (5.66%)	
occurrences (all)	14	19	
Blood and lymphatic system disorders			
Leukopenia			
subjects affected / exposed	26 / 317 (8.20%)	26 / 318 (8.18%)	
occurrences (all)	73	72	
Neutropenia			
subjects affected / exposed	137 / 317 (43.22%)	114 / 318 (35.85%)	
occurrences (all)	476	365	
Anaemia			
subjects affected / exposed	118 / 317 (37.22%)	135 / 318 (42.45%)	
occurrences (all)	229	289	
Thrombocytopenia			
subjects affected / exposed	72 / 317 (22.71%)	91 / 318 (28.62%)	
occurrences (all)	173	215	
Lymphopenia			
subjects affected / exposed	23 / 317 (7.26%)	41 / 318 (12.89%)	
occurrences (all)	60	92	

Eye disorders			
Vision blurred			
subjects affected / exposed	18 / 317 (5.68%)	31 / 318 (9.75%)	
occurrences (all)	21	33	
Cataract			
subjects affected / exposed	36 / 317 (11.36%)	58 / 318 (18.24%)	
occurrences (all)	37	65	
Gastrointestinal disorders			
Abdominal pain			
subjects affected / exposed	31 / 317 (9.78%)	44 / 318 (13.84%)	
occurrences (all)	39	49	
Constipation			
subjects affected / exposed	89 / 317 (28.08%)	115 / 318 (36.16%)	
occurrences (all)	117	184	
Dyspepsia			
subjects affected / exposed	19 / 317 (5.99%)	36 / 318 (11.32%)	
occurrences (all)	26	50	
Vomiting			
subjects affected / exposed	32 / 317 (10.09%)	58 / 318 (18.24%)	
occurrences (all)	42	88	
Stomatitis			
subjects affected / exposed	14 / 317 (4.42%)	30 / 318 (9.43%)	
occurrences (all)	16	36	
Nausea			
subjects affected / exposed	73 / 317 (23.03%)	82 / 318 (25.79%)	
occurrences (all)	99	129	
Abdominal pain upper			
subjects affected / exposed	20 / 317 (6.31%)	30 / 318 (9.43%)	
occurrences (all)	25	33	
Diarrhoea			
subjects affected / exposed	122 / 317 (38.49%)	159 / 318 (50.00%)	
occurrences (all)	247	451	
Dry mouth			
subjects affected / exposed	10 / 317 (3.15%)	16 / 318 (5.03%)	
occurrences (all)	10	16	
Toothache			

subjects affected / exposed	11 / 317 (3.47%)	18 / 318 (5.66%)	
occurrences (all)	12	23	
Skin and subcutaneous tissue disorders			
Pruritus			
subjects affected / exposed	29 / 317 (9.15%)	36 / 318 (11.32%)	
occurrences (all)	31	47	
Rash			
subjects affected / exposed	58 / 317 (18.30%)	63 / 318 (19.81%)	
occurrences (all)	93	105	
Hyperhidrosis			
subjects affected / exposed	25 / 317 (7.89%)	38 / 318 (11.95%)	
occurrences (all)	33	46	
Erythema			
subjects affected / exposed	16 / 317 (5.05%)	26 / 318 (8.18%)	
occurrences (all)	22	28	
Night sweats			
subjects affected / exposed	10 / 317 (3.15%)	24 / 318 (7.55%)	
occurrences (all)	13	34	
Renal and urinary disorders			
Dysuria			
subjects affected / exposed	9 / 317 (2.84%)	16 / 318 (5.03%)	
occurrences (all)	9	19	
Musculoskeletal and connective tissue disorders			
Arthralgia			
subjects affected / exposed	63 / 317 (19.87%)	98 / 318 (30.82%)	
occurrences (all)	78	135	
Bone pain			
subjects affected / exposed	43 / 317 (13.56%)	33 / 318 (10.38%)	
occurrences (all)	54	42	
Myalgia			
subjects affected / exposed	22 / 317 (6.94%)	26 / 318 (8.18%)	
occurrences (all)	25	29	
Muscle spasms			
subjects affected / exposed	85 / 317 (26.81%)	99 / 318 (31.13%)	
occurrences (all)	112	145	
Neck pain			

subjects affected / exposed	13 / 317 (4.10%)	23 / 318 (7.23%)	
occurrences (all)	14	27	
Muscular weakness			
subjects affected / exposed	28 / 317 (8.83%)	42 / 318 (13.21%)	
occurrences (all)	31	45	
Musculoskeletal chest pain			
subjects affected / exposed	30 / 317 (9.46%)	39 / 318 (12.26%)	
occurrences (all)	31	47	
Pain in extremity			
subjects affected / exposed	35 / 317 (11.04%)	64 / 318 (20.13%)	
occurrences (all)	43	88	
Back pain			
subjects affected / exposed	97 / 317 (30.60%)	105 / 318 (33.02%)	
occurrences (all)	122	144	
Infections and infestations			
Bronchitis			
subjects affected / exposed	52 / 317 (16.40%)	66 / 318 (20.75%)	
occurrences (all)	81	124	
Nasopharyngitis			
subjects affected / exposed	60 / 317 (18.93%)	83 / 318 (26.10%)	
occurrences (all)	128	236	
Urinary tract infection			
subjects affected / exposed	31 / 317 (9.78%)	36 / 318 (11.32%)	
occurrences (all)	64	65	
Rhinitis			
subjects affected / exposed	13 / 317 (4.10%)	30 / 318 (9.43%)	
occurrences (all)	17	34	
Herpes zoster			
subjects affected / exposed	5 / 317 (1.58%)	18 / 318 (5.66%)	
occurrences (all)	5	19	
Lower respiratory tract infection			
subjects affected / exposed	16 / 317 (5.05%)	32 / 318 (10.06%)	
occurrences (all)	21	70	
Pharyngitis			
subjects affected / exposed	16 / 317 (5.05%)	17 / 318 (5.35%)	
occurrences (all)	29	18	

Pneumonia			
subjects affected / exposed	23 / 317 (7.26%)	30 / 318 (9.43%)	
occurrences (all)	29	36	
Sinusitis			
subjects affected / exposed	15 / 317 (4.73%)	23 / 318 (7.23%)	
occurrences (all)	37	52	
Influenza			
subjects affected / exposed	17 / 317 (5.36%)	25 / 318 (7.86%)	
occurrences (all)	26	32	
Respiratory tract infection			
subjects affected / exposed	32 / 317 (10.09%)	36 / 318 (11.32%)	
occurrences (all)	63	67	
Upper respiratory tract infection			
subjects affected / exposed	62 / 317 (19.56%)	86 / 318 (27.04%)	
occurrences (all)	146	209	
Oral herpes			
subjects affected / exposed	13 / 317 (4.10%)	20 / 318 (6.29%)	
occurrences (all)	16	30	
Gastroenteritis			
subjects affected / exposed	9 / 317 (2.84%)	18 / 318 (5.66%)	
occurrences (all)	13	20	
Viral infection			
subjects affected / exposed	10 / 317 (3.15%)	18 / 318 (5.66%)	
occurrences (all)	16	36	
Metabolism and nutrition disorders			
Hyponatraemia			
subjects affected / exposed	10 / 317 (3.15%)	18 / 318 (5.66%)	
occurrences (all)	17	41	
Hypomagnesaemia			
subjects affected / exposed	5 / 317 (1.58%)	22 / 318 (6.92%)	
occurrences (all)	5	46	
Hypocalcaemia			
subjects affected / exposed	28 / 317 (8.83%)	48 / 318 (15.09%)	
occurrences (all)	71	135	
Hypokalaemia			

subjects affected / exposed	47 / 317 (14.83%)	67 / 318 (21.07%)	
occurrences (all)	103	124	
Decreased appetite			
subjects affected / exposed	42 / 317 (13.25%)	71 / 318 (22.33%)	
occurrences (all)	48	93	
Hyperglycaemia			
subjects affected / exposed	45 / 317 (14.20%)	61 / 318 (19.18%)	
occurrences (all)	87	129	
Hypoalbuminaemia			
subjects affected / exposed	11 / 317 (3.47%)	17 / 318 (5.35%)	
occurrences (all)	21	22	
Hypophosphataemia			
subjects affected / exposed	10 / 317 (3.15%)	19 / 318 (5.97%)	
occurrences (all)	33	45	

More information

Substantial protocol amendments (globally)

Were there any global substantial amendments to the protocol? Yes

Date	Amendment
15 March 2012	Clarification of subject eligibility or study procedures
14 April 2014	Addition of formal interim analyses.
14 April 2014	The addition of a secondary and exploratory objectives and the removal Brief Pain Inventory assessment as exploratory endpoint.

Notes:

Interruptions (globally)

Were there any global interruptions to the trial? No

Limitations and caveats

None reported