



Clinical trial results:

An international, multicenter, randomized, double-blind, placebo-controlled phase 3 trial investigating the efficacy and safety of rivaroxaban to reduce the risk of major thrombotic vascular events in patients with symptomatic peripheral artery disease undergoing lower extremity revascularization procedures

Summary

EudraCT number	2014-005569-58
Trial protocol	CZ AT ES HU GB FR PT SE BE BG DK DE NL PL SK LT LV RO EE
Global end of trial date	551 09 January 2020

Results information

Result version number	v1 (current)
This version publication date	26 November 2020
First version publication date	26 November 2020

Trial information

Trial identification

Sponsor protocol code	17454
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Additional study identifiers

ISRCTN number	-
ClinicalTrials.gov id (NCT number)	NCT02504216
WHO universal trial number (UTN)	-

Notes:

Sponsors

Sponsor organisation name	Bayer AG
Sponsor organisation address	Kaiser-Wilhelm-Allee, Leverkusen, Germany, D-51368
Public contact	Therapeutic Area Head, Bayer AG, clinical-trials-contact@bayer.com
Scientific contact	Therapeutic Area Head, Bayer AG, clinical-trials-contact@bayer.com

Notes:

Paediatric regulatory details

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

Notes:

Results analysis stage

Analysis stage	Final
Date of interim/final analysis	31 March 2020
Is this the analysis of the primary completion data?	No
Global end of trial reached?	Yes
Global end of trial date	09 January 2020
Was the trial ended prematurely?	No

Notes:

General information about the trial

Main objective of the trial:

The primary efficacy objective of the study was: - to evaluate whether rivaroxaban added to ASA is superior to ASA alone in reducing the risk of major thrombotic vascular events (defined as MI, ischemic stroke, CV (Cardiovascular) death, ALI (Acute limb ischemia), and major amputation of a vascular etiology) in symptomatic PAD (Peripheral artery disease) patients with a recent lower extremity revascularization procedure. The primary safety objective of the study was: - to evaluate the overall safety and tolerability of rivaroxaban added to ASA compared to ASA alone.

Protection of trial subjects:

The conduct of this clinical study met all local legal and regulatory requirements. The study was conducted in accordance with ethical principles that have their origin in the Declaration of Helsinki and the International Council for Harmonization guideline E6: Good Clinical Practice. Before entering the study, the informed consent was read by and explained to all the subjects. Participating subjects signed informed consent form and could withdraw from the study at any time without any disadvantage and without having to provide a reason for this decision. Only investigators qualified by training and experience were selected as appropriate experts to investigate the study drug.

Background therapy: -

Evidence for comparator: -

Actual start date of recruitment	18 August 2015
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	China: 287
Country: Number of subjects enrolled	Japan: 459
Country: Number of subjects enrolled	Korea, Republic of: 148
Country: Number of subjects enrolled	Bulgaria: 737
Country: Number of subjects enrolled	Czechia: 243
Country: Number of subjects enrolled	Estonia: 9
Country: Number of subjects enrolled	Hungary: 261
Country: Number of subjects enrolled	Latvia: 202
Country: Number of subjects enrolled	Lithuania: 84
Country: Number of subjects enrolled	Poland: 168
Country: Number of subjects enrolled	Romania: 214
Country: Number of subjects enrolled	Russian Federation: 188
Country: Number of subjects enrolled	Serbia: 68
Country: Number of subjects enrolled	Slovakia: 126

Country: Number of subjects enrolled	Ukraine: 299
Country: Number of subjects enrolled	Canada: 170
Country: Number of subjects enrolled	United States: 524
Country: Number of subjects enrolled	Argentina: 299
Country: Number of subjects enrolled	Brazil: 185
Country: Number of subjects enrolled	Austria: 212
Country: Number of subjects enrolled	Belgium: 126
Country: Number of subjects enrolled	Denmark: 78
Country: Number of subjects enrolled	Thailand: 67
Country: Number of subjects enrolled	Finland: 17
Country: Number of subjects enrolled	France: 107
Country: Number of subjects enrolled	Germany: 593
Country: Number of subjects enrolled	Italy: 184
Country: Number of subjects enrolled	Netherlands: 77
Country: Number of subjects enrolled	Portugal: 74
Country: Number of subjects enrolled	Spain: 96
Country: Number of subjects enrolled	Sweden: 42
Country: Number of subjects enrolled	Switzerland: 91
Country: Number of subjects enrolled	United Kingdom: 129
Worldwide total number of subjects	6564
EEA total number of subjects	3779

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)	2622
From 65 to 84 years	3801
85 years and over	141

Subject disposition

Recruitment

Recruitment details:

Study was conducted at 548 centers which screened subjects (534 sites randomized subjects) world-wide between 31 Jul 2015 (first patient's first visit) and 27 Nov 2019 (last patient's last visit).

Pre-assignment

Screening details:

A total of 6772 subjects were screened, of which 6564 were randomly assigned to either of the two treatment arms.

Period 1

Period 1 title	Overall (overall period)
Is this the baseline period?	Yes
Allocation method	Randomised - controlled
Blinding used	Double blind
Roles blinded	Subject, Investigator, Monitor, Data analyst, Carer, Assessor

Arms

Are arms mutually exclusive?	Yes
Arm title	Rivaroxaban 2.5 mg bid + Aspirin 100 mg od

Arm description:

Subjects were treated with Rivaroxaban 2.5 mg twice-daily (bid) and aspirin (ASA: Acetylsalicylic Acid) 100 mg once-daily (od)

Arm type	Experimental
Investigational medicinal product name	Acetylsalicylic Acid (ASA)
Investigational medicinal product code	
Other name	
Pharmaceutical forms	Tablet
Routes of administration	Oral use

Dosage and administration details:

100 mg once-daily (od)

Investigational medicinal product name	Rivaroxaban (Xarelto, BAY59-7939)
Investigational medicinal product code	
Other name	
Pharmaceutical forms	Tablet
Routes of administration	Oral use

Dosage and administration details:

2.5 mg twice-daily (bid) (5 mg/day)

Arm title	Rivaroxaban placebo bid + Aspirin 100 mg od
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Arm description:

Subjects were treated with Rivaroxaban-placebo twice-daily and aspirin (ASA: Acetylsalicylic Acid) 100 mg once-daily

Arm type	Placebo
Investigational medicinal product name	Acetylsalicylic Acid (ASA)
Investigational medicinal product code	
Other name	
Pharmaceutical forms	Tablet
Routes of administration	Oral use

Dosage and administration details:

100 mg once-daily (od)

Investigational medicinal product name	Rivaroxaban-Placebo
Investigational medicinal product code	
Other name	
Pharmaceutical forms	Tablet
Routes of administration	Oral use

Dosage and administration details:

Matching placebo, twice daily,

Number of subjects in period 1	Rivaroxaban 2.5 mg bid + Aspirin 100 mg od	Rivaroxaban placebo bid + Aspirin 100 mg od
Started	3286	3278
Received treatment	3256 ^[1]	3248 ^[2]
Completed	3275	3263
Not completed	11	15
Consent withdrawn by subject	8	12
Lost to follow-up	3	3

Notes:

[1] - The number of subjects at this milestone seems inconsistent with the number of subjects in the arm. It is expected that the number of subjects will be greater than, or equal to the number that completed, minus those who left.

Justification: 30 subjects were never administered study drug.

[2] - The number of subjects at this milestone seems inconsistent with the number of subjects in the arm. It is expected that the number of subjects will be greater than, or equal to the number that completed, minus those who left.

Justification: 30 subjects were never administered study drug.

Baseline characteristics

Reporting groups

Reporting group title	Rivaroxaban 2.5 mg bid + Aspirin 100 mg od
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Reporting group description:

Subjects were treated with Rivaroxaban 2.5 mg twice-daily (bid) and aspirin (ASA: Acetylsalicylic Acid) 100 mg once-daily (od)

Reporting group title	Rivaroxaban placebo bid + Aspirin 100 mg od
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Reporting group description:

Subjects were treated with Rivaroxaban-placebo twice-daily and aspirin (ASA: Acetylsalicylic Acid) 100 mg once-daily

Reporting group values	Rivaroxaban 2.5 mg bid + Aspirin 100 mg od	Rivaroxaban placebo bid + Aspirin 100 mg od	Total
Number of subjects	3286	3278	6564
Age Categorical Units: Subjects			

Age Continuous Units: years median full range (min-max)	67.0 49 to 93	67.0 50 to 95	-
Gender Categorical Units: Subjects			
Female	847	857	1704
Male	2439	2421	4860
Race Units: Subjects			
White	2647	2656	5303
Black or African American	84	71	155
Asian	484	482	966
American Indian or Alaska native	1	4	5
Native Hawaiian or other Pacific islander	0	1	1
Not reported	69	62	131
Multiple	1	2	3

End points

End points reporting groups

Reporting group title	Rivaroxaban 2.5 mg bid + Aspirin 100 mg od
Reporting group description: Subjects were treated with Rivaroxaban 2.5 mg twice-daily (bid) and aspirin (ASA: Acetylsalicylic Acid) 100 mg once-daily (od)	
Reporting group title	Rivaroxaban placebo bid + Aspirin 100 mg od
Reporting group description: Subjects were treated with Rivaroxaban-placebo twice-daily and aspirin (ASA: Acetylsalicylic Acid) 100 mg once-daily	
Subject analysis set title	Intention-to-treat analysis set (ITT Set)
Subject analysis set type	Intention-to-treat
Subject analysis set description: The ITT population comprised all randomized subjects.	
Subject analysis set title	Safety analysis set (SAF Set)
Subject analysis set type	Safety analysis
Subject analysis set description: The SAF comprised all treated subjects, i.e. randomized subjects who received at least one dose of study drug (rivaroxaban or rivaroxaban placebo).	

Primary: Primary efficacy outcome: Composite of Myocardial infarction (MI), ischemic stroke, cardiovascular death, acute limb ischemia (ALI) and major amputation due to a vascular etiology

End point title	Primary efficacy outcome: Composite of Myocardial infarction (MI), ischemic stroke, cardiovascular death, acute limb ischemia (ALI) and major amputation due to a vascular etiology
End point description: Only the first occurrence of the outcome event under analysis within the data scope from a subject is considered.	
End point type	Primary
End point timeframe: For each participant, the first occurrence of the composite primary efficacy outcome after randomization up until the efficacy cut-off date (08-Sep-2019) was considered. The mean time in follow-up survival time until ECOD that date was 1109.76 days.	

End point values	Rivaroxaban 2.5 mg bid + Aspirin 100 mg od	Rivaroxaban placebo bid + Aspirin 100 mg od		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	3286	3278		
Units: Subject	508	584		

Statistical analyses

Statistical analysis title	Primary efficacy outcome
Comparison groups	Rivaroxaban placebo bid + Aspirin 100 mg od v Rivaroxaban

	2.5 mg bid + Aspirin 100 mg od
Number of subjects included in analysis	6564
Analysis specification	Pre-specified
Analysis type	superiority
P-value	= 0.0043 ^[1]
Method	Logrank
Parameter estimate	Hazard ratio (HR)
Point estimate	0.85
Confidence interval	
level	95 %
sides	2-sided
lower limit	0.76
upper limit	0.96

Notes:

[1] - P-value (1-sided) is based on the log rank test stratified by type of procedure and clopidogrel use per IxRS (Interactive web/voice response system) assignment with treatment as a factor.

Primary: Primary safety outcome: Number of participants with TIMI Major bleeding

End point title	Primary safety outcome: Number of participants with TIMI Major bleeding
End point description:	TIMI: Thrombolysis in Myocardial Infarction Only the first occurrence of the outcome event under analysis within the data scope from a subject is considered.
End point type	Primary
End point timeframe:	For each participant, the first occurrence of the primary safety outcome after randomization up until 2 days after permanent stop of study drug (rivaroxaban or rivaroxaban placebo).

End point values	Rivaroxaban 2.5 mg bid + Aspirin 100 mg od	Rivaroxaban placebo bid + Aspirin 100 mg od		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	3256	3248		
Units: Subject	62	44		

Statistical analyses

Statistical analysis title	TIMI major bleeding
Comparison groups	Rivaroxaban 2.5 mg bid + Aspirin 100 mg od v Rivaroxaban placebo bid + Aspirin 100 mg od
Number of subjects included in analysis	6504
Analysis specification	Pre-specified
Analysis type	other
P-value	= 0.0695 ^[2]
Method	Logrank
Parameter estimate	Hazard ratio (HR)
Point estimate	1.43

Confidence interval	
level	95 %
sides	2-sided
lower limit	0.97
upper limit	2.1

Notes:

[2] - P-value (2-sided) is based on the log rank test stratified by type of procedure and clopidogrel use per IxRS assignment with treatment as a factor.

Secondary: Composite of MI, ischemic stroke, Coronary heart disease (CHD) death, ALI, and major amputation of a vascular etiology

End point title	Composite of MI, ischemic stroke, Coronary heart disease (CHD) death, ALI, and major amputation of a vascular etiology
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End point description:

MI, stroke, mortality, ALI, and major amputation of a vascular etiology, and events leading to hospitalization are adjudicated.

End point type	Secondary
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End point timeframe:

For each participant, the first occurrence of the composite primary efficacy outcome after randomization up until the efficacy cut-off date (08-Sep-2019) was considered. The mean survival time until ECOD was 1108.79 days.

End point values	Rivaroxaban 2.5 mg bid + Aspirin 100 mg od	Rivaroxaban placebo bid + Aspirin 100 mg od		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	3286	3278		
Units: Subject	433	528		

Statistical analyses

Statistical analysis title	Secondary efficacy outcome
Comparison groups	Rivaroxaban 2.5 mg bid + Aspirin 100 mg od v Rivaroxaban placebo bid + Aspirin 100 mg od
Number of subjects included in analysis	6564
Analysis specification	Pre-specified
Analysis type	superiority
P-value	= 0.0004 [3]
Method	Logrank
Parameter estimate	Hazard ratio (HR)
Point estimate	0.8
Confidence interval	
level	95 %
sides	2-sided
lower limit	0.71
upper limit	0.91

Notes:

[3] - P-value (1-sided) is based on the log rank test stratified by type of procedure and clopidogrel use per IxRS assignment with treatment as a factor.

Secondary: Number of participants with an unplanned index limb revascularization for recurrent limb ischemia (subsequent index leg revascularization that was not planned or considered as part of the initial treatment plan at the time of randomization)

End point title	Number of participants with an unplanned index limb revascularization for recurrent limb ischemia (subsequent index leg revascularization that was not planned or considered as part of the initial treatment plan at the time of randomization)
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End point description:

Index limb revascularization is defined as subsequent index leg revascularization that was not planned or considered as part of the initial treatment plan at the time of randomization. index limb revascularization were reported by investigator only.

End point type	Secondary
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End point timeframe:

For each participant, the first occurrence of the composite primary efficacy outcome after randomization up until the efficacy cut-off date (08-Sep-2019) was considered. The mean survival time until ECOD was 1062.48 days.

End point values	Rivaroxaban 2.5 mg bid + Aspirin 100 mg od	Rivaroxaban placebo bid + Aspirin 100 mg od		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	3286	3278		
Units: Subject	584	655		

Statistical analyses

Statistical analysis title	Secondary efficacy outcome
Comparison groups	Rivaroxaban 2.5 mg bid + Aspirin 100 mg od v Rivaroxaban placebo bid + Aspirin 100 mg od
Number of subjects included in analysis	6564
Analysis specification	Pre-specified
Analysis type	superiority
P-value	= 0.014 [4]
Method	Logrank
Parameter estimate	Hazard ratio (HR)
Point estimate	0.88
Confidence interval	
level	95 %
sides	2-sided
lower limit	0.79
upper limit	0.99

Notes:

[4] - P-value (1-sided) is based on the log rank test stratified by type of procedure and clopidogrel use per IxRS assignment with treatment as a factor.

Secondary: Number of participants with hospitalization for a coronary or peripheral cause (either lower limb) of a thrombotic nature

End point title	Number of participants with hospitalization for a coronary or peripheral cause (either lower limb) of a thrombotic nature
End point description:	Events leading to hospitalization are adjudicated.
End point type	Secondary
End point timeframe:	For each participant, the first occurrence of the composite primary efficacy outcome after randomization up until the efficacy cut-off date (08-Sep-2019) was considered. The mean survival time until ECOD was 1154.04 days

End point values	Rivaroxaban 2.5 mg bid + Aspirin 100 mg od	Rivaroxaban placebo bid + Aspirin 100 mg od		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	3286	3278		
Units: Subject	262	356		

Statistical analyses

Statistical analysis title	Secondary efficacy outcome
Comparison groups	Rivaroxaban 2.5 mg bid + Aspirin 100 mg od v Rivaroxaban placebo bid + Aspirin 100 mg od
Number of subjects included in analysis	6564
Analysis specification	Pre-specified
Analysis type	superiority
P-value	< 0.0001 ^[5]
Method	Logrank
Parameter estimate	Hazard ratio (HR)
Point estimate	0.72
Confidence interval	
level	95 %
sides	2-sided
lower limit	0.62
upper limit	0.85

Notes:

[5] - P-value (1-sided) is based on the log rank test stratified by type of procedure and clopidogrel use per IxRS assignment with treatment as a factor.

Secondary: Composite of MI, ischemic stroke, All-cause mortality (ACM), ALI, and major amputation of a vascular etiology

End point title	Composite of MI, ischemic stroke, All-cause mortality (ACM), ALI, and major amputation of a vascular etiology
End point description:	MI, stroke, mortality, ALI, and major amputation of a vascular etiology are adjudicated.
End point type	Secondary

End point timeframe:

For each participant, the first occurrence of the composite primary efficacy outcome after randomization up until the efficacy cut-off date (08-Sep-2019) was considered. The mean survival time until ECOD was 1085.13 days

End point values	Rivaroxaban 2.5 mg bid + Aspirin 100 mg od	Rivaroxaban placebo bid + Aspirin 100 mg od		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	3286	3278		
Units: Subject	614	679		

Statistical analyses

Statistical analysis title	Secondary efficacy outcome
Comparison groups	Rivaroxaban 2.5 mg bid + Aspirin 100 mg od v Rivaroxaban placebo bid + Aspirin 100 mg od
Number of subjects included in analysis	6564
Analysis specification	Pre-specified
Analysis type	superiority
P-value	= 0.0145 [6]
Method	Logrank
Parameter estimate	Hazard ratio (HR)
Point estimate	0.89
Confidence interval	
level	95 %
sides	2-sided
lower limit	0.79
upper limit	0.99

Notes:

[6] - P-value (1-sided) is based on the log rank test stratified by type of procedure and clopidogrel use per IxRS assignment with treatment as a factor.

Secondary: Composite of MI, all-cause stroke, Cardiovascular (CV) death, Acute limb ischemia (ALI), and major amputation of a vascular etiology

End point title	Composite of MI, all-cause stroke, Cardiovascular (CV) death, Acute limb ischemia (ALI), and major amputation of a vascular etiology
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End point description:

MI, stroke, mortality, ALI, and major amputation of a vascular etiology are adjudicated.

End point type	Secondary
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End point timeframe:

For each participant, the first occurrence of the composite primary efficacy outcome after randomization up until the efficacy cut-off date (08-Sep-2019) was considered. The mean survival time until ECOD was 1108.29 days

End point values	Rivaroxaban 2.5 mg bid + Aspirin 100 mg od	Rivaroxaban placebo bid + Aspirin 100 mg od		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	3286	3278		
Units: Subject	514	588		

Statistical analyses

Statistical analysis title	Secondary efficacy outcome
Comparison groups	Rivaroxaban 2.5 mg bid + Aspirin 100 mg od v Rivaroxaban placebo bid + Aspirin 100 mg od
Number of subjects included in analysis	6564
Analysis specification	Pre-specified
Analysis type	superiority
P-value	= 0.0051 ^[7]
Method	Logrank
Parameter estimate	Hazard ratio (HR)
Point estimate	0.86
Confidence interval	
level	95 %
sides	2-sided
lower limit	0.76
upper limit	0.96

Notes:

[7] - P-value (1-sided) is based on the log rank test stratified by type of procedure and clopidogrel use per IxRS assignment with treatment as a factor.

Secondary: Mortality (all-cause)

End point title	Mortality (all-cause)
End point description:	Mortality are adjudicated.
End point type	Secondary
End point timeframe:	For each participant, the first occurrence of the composite primary efficacy outcome after randomization up until the efficacy cut-off date (08-Sep-2019) was considered. The mean survival time until ECOD was 1188.48 days

End point values	Rivaroxaban 2.5 mg bid + Aspirin 100 mg od	Rivaroxaban placebo bid + Aspirin 100 mg od		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	3286	3278		
Units: Subject	321	297		

Statistical analyses

Statistical analysis title	Secondary efficacy outcome
Comparison groups	Rivaroxaban 2.5 mg bid + Aspirin 100 mg od v Rivaroxaban placebo bid + Aspirin 100 mg od
Number of subjects included in analysis	6564
Analysis specification	Pre-specified
Analysis type	superiority
P-value	= 0.8322 [8]
Method	Logrank
Parameter estimate	Hazard ratio (HR)
Point estimate	1.08
Confidence interval	
level	95 %
sides	2-sided
lower limit	0.92
upper limit	1.27

Notes:

[8] - P-value (1-sided) is based on the log rank test stratified by type of procedure and clopidogrel use per IxRS assignment with treatment as a factor.

Secondary: Number of participants with venous thromboembolic (VTE) events

End point title	Number of participants with venous thromboembolic (VTE) events
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End point description:

Venous thromboembolic events were reported by investigator only.

End point type	Secondary
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End point timeframe:

For each participant, the first occurrence of the composite primary efficacy outcome after randomization up until the efficacy cut-off date (08-Sep-2019) was considered. The mean survival time until ECOD was 1187.65 days

End point values	Rivaroxaban 2.5 mg bid + Aspirin 100 mg od	Rivaroxaban placebo bid + Aspirin 100 mg od		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	3286	3278		
Units: Subject	25	41		

Statistical analyses

Statistical analysis title	Secondary efficacy outcome
Comparison groups	Rivaroxaban 2.5 mg bid + Aspirin 100 mg od v Rivaroxaban placebo bid + Aspirin 100 mg od

Number of subjects included in analysis	6564
Analysis specification	Pre-specified
Analysis type	superiority
P-value	= 0.0235 [9]
Method	Logrank
Parameter estimate	Hazard ratio (HR)
Point estimate	0.61
Confidence interval	
level	95 %
sides	2-sided
lower limit	0.37
upper limit	1

Notes:

[9] - P-value (1-sided) is based on the log rank test stratified by type of procedure and clopidogrel use per IxRS assignment with treatment as a factor.

Secondary: Secondary safety outcome: Number of participants with ISTH major bleeding

End point title	Secondary safety outcome: Number of participants with ISTH major bleeding
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End point description:

ISTH: International Society on Thrombosis and Haemostasis Only the first occurrence of the outcome event under analysis within the data scope from a subject is considered.

End point type	Secondary
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End point timeframe:

For each participant, the first occurrence of the major bleeding events according to the ISTH classification after randomization up until 2 days after permanent stop of study drug (rivaroxaban or rivaroxaban placebo).

End point values	Rivaroxaban 2.5 mg bid + Aspirin 100 mg od	Rivaroxaban placebo bid + Aspirin 100 mg od		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	3256	3248		
Units: Subject	140	100		

Statistical analyses

Statistical analysis title	ISTH major bleeding
Comparison groups	Rivaroxaban 2.5 mg bid + Aspirin 100 mg od v Rivaroxaban placebo bid + Aspirin 100 mg od
Number of subjects included in analysis	6504
Analysis specification	Pre-specified
Analysis type	other
P-value	= 0.0068
Method	Logrank
Parameter estimate	Hazard ratio (HR)
Point estimate	1.42

Confidence interval	
level	95 %
sides	2-sided
lower limit	1.1
upper limit	1.84

Secondary: Secondary safety outcome: Number of participants with BARC Type 3b and above bleeding events

End point title	Secondary safety outcome: Number of participants with BARC Type 3b and above bleeding events
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End point description:

BARC: Bleeding Academic Research Consortium Only the first occurrence of the outcome event under analysis within the data scope from a subject is considered

End point type	Secondary
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End point timeframe:

For each participant, the first occurrence of the type 3b and above bleeding events according to the BARC classification after randomization up until 2 days after permanent stop of study drug (rivaroxaban or rivaroxaban placebo)

End point values	Rivaroxaban 2.5 mg bid + Aspirin 100 mg od	Rivaroxaban placebo bid + Aspirin 100 mg od		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	3256	3248		
Units: Subject	93	73		

Statistical analyses

Statistical analysis title	BARC Type 3b and above bleeding events
Comparison groups	Rivaroxaban 2.5 mg bid + Aspirin 100 mg od v Rivaroxaban placebo bid + Aspirin 100 mg od
Number of subjects included in analysis	6504
Analysis specification	Pre-specified
Analysis type	other
P-value	= 0.0979
Method	Logrank
Parameter estimate	Hazard ratio (HR)
Point estimate	1.29
Confidence interval	
level	95 %
sides	2-sided
lower limit	0.95
upper limit	1.76

Adverse events

Adverse events information

Timeframe for reporting adverse events:

From randomization until 2 days following permanent discontinuation of the study drug

Assessment type	Non-systematic
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Dictionary used

Dictionary name	MedDRA
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Dictionary version	22.1
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Reporting groups

Reporting group title	Rivaroxaban 2.5 mg bid + ASA 100 mg od
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Reporting group description:

Subjects were treated with Rivaroxaban 2.5 mg twice-daily (bid) and ASA (Acetylsalicylic Acid) 100 mg once-daily (od)

Reporting group title	Rivaroxaban placebo bid + ASA 100 mg od
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Reporting group description:

Subjects were treated with Rivaroxaban-placebo twice-daily and ASA (Acetylsalicylic Acid) 100 mg once-daily

Serious adverse events	Rivaroxaban 2.5 mg bid + ASA 100 mg od	Rivaroxaban placebo bid + ASA 100 mg od	
Total subjects affected by serious adverse events			
subjects affected / exposed	948 / 3256 (29.12%)	927 / 3248 (28.54%)	
number of deaths (all causes)	327	304	
number of deaths resulting from adverse events	76	79	
Neoplasms benign, malignant and unspecified (incl cysts and polyps)			
Acute myeloid leukaemia			
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Adenocarcinoma			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Adenocarcinoma gastric			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 1	
Adenocarcinoma of colon			

subjects affected / exposed	3 / 3256 (0.09%)	2 / 3248 (0.06%)
occurrences causally related to treatment / all	0 / 3	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0
B-cell lymphoma		
subjects affected / exposed	1 / 3256 (0.03%)	2 / 3248 (0.06%)
occurrences causally related to treatment / all	0 / 1	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 1
Basal cell carcinoma		
subjects affected / exposed	3 / 3256 (0.09%)	7 / 3248 (0.22%)
occurrences causally related to treatment / all	0 / 3	0 / 7
deaths causally related to treatment / all	0 / 0	0 / 0
Benign laryngeal neoplasm		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Benign neoplasm of skin		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Benign ovarian tumour		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Bile duct cancer		
subjects affected / exposed	0 / 3256 (0.00%)	2 / 3248 (0.06%)
occurrences causally related to treatment / all	0 / 0	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0
Bladder cancer		
subjects affected / exposed	4 / 3256 (0.12%)	4 / 3248 (0.12%)
occurrences causally related to treatment / all	0 / 4	0 / 4
deaths causally related to treatment / all	0 / 0	0 / 0
Bladder cancer stage IV		

subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 1
Bladder neoplasm		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Bladder transitional cell carcinoma		
subjects affected / exposed	2 / 3256 (0.06%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 2	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Bone cancer		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 1
Breast cancer		
subjects affected / exposed	2 / 3256 (0.06%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 2	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Bronchial carcinoma		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Central nervous system lymphoma		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 1	0 / 0
Cervix carcinoma		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Cholangiocarcinoma		

subjects affected / exposed	2 / 3256 (0.06%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 2	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Cholesteatoma		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Chronic myeloid leukaemia		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Chronic myelomonocytic leukaemia		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 1
Colon cancer		
subjects affected / exposed	5 / 3256 (0.15%)	4 / 3248 (0.12%)
occurrences causally related to treatment / all	0 / 5	0 / 4
deaths causally related to treatment / all	0 / 1	0 / 0
Endometrial cancer		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Follicular thyroid cancer		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Gallbladder cancer		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Gastric cancer		

subjects affected / exposed	5 / 3256 (0.15%)	4 / 3248 (0.12%)
occurrences causally related to treatment / all	0 / 5	0 / 4
deaths causally related to treatment / all	0 / 0	0 / 1
Gastric cancer stage I		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Gastric cancer stage II		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Hodgkin's disease		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Laryngeal cancer		
subjects affected / exposed	1 / 3256 (0.03%)	3 / 3248 (0.09%)
occurrences causally related to treatment / all	0 / 2	0 / 3
deaths causally related to treatment / all	0 / 0	0 / 0
Laryngeal neoplasm		
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 2	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Lung adenocarcinoma		
subjects affected / exposed	4 / 3256 (0.12%)	3 / 3248 (0.09%)
occurrences causally related to treatment / all	0 / 4	0 / 3
deaths causally related to treatment / all	0 / 1	0 / 0
Lung adenocarcinoma recurrent		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 1	0 / 0
Lung adenocarcinoma stage IV		

subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 1
Lung carcinoma cell type unspecified recurrent		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 1	0 / 0
Lung carcinoma cell type unspecified stage IV		
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 1
Lung squamous cell carcinoma stage IV		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Malignant melanoma		
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Metastases to liver		
subjects affected / exposed	3 / 3256 (0.09%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 3	0 / 1
deaths causally related to treatment / all	0 / 1	0 / 1
Metastases to lung		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Metastases to lymph nodes		
subjects affected / exposed	2 / 3256 (0.06%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 2	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Metastases to neck		

subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 1
Myelodysplastic syndrome		
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	1 / 1	0 / 1
deaths causally related to treatment / all	1 / 1	0 / 0
Neoplasm malignant		
subjects affected / exposed	1 / 3256 (0.03%)	2 / 3248 (0.06%)
occurrences causally related to treatment / all	0 / 1	0 / 2
deaths causally related to treatment / all	0 / 1	0 / 1
Neoplasm skin		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Oesophageal adenocarcinoma		
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 2	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Oesophageal carcinoma		
subjects affected / exposed	3 / 3256 (0.09%)	2 / 3248 (0.06%)
occurrences causally related to treatment / all	0 / 3	0 / 2
deaths causally related to treatment / all	0 / 1	0 / 0
Oesophageal carcinoma recurrent		
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Oropharyngeal cancer		
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Pancreatic carcinoma		

subjects affected / exposed	2 / 3256 (0.06%)	3 / 3248 (0.09%)
occurrences causally related to treatment / all	0 / 2	0 / 3
deaths causally related to treatment / all	0 / 2	0 / 2
Papillary serous endometrial carcinoma		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Plasma cell myeloma		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Pleural mesothelioma		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 1	0 / 0
Prostate cancer stage III		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0
Rectal adenocarcinoma		
subjects affected / exposed	2 / 3256 (0.06%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 2	0 / 0
deaths causally related to treatment / all	0 / 1	0 / 0
Rectal cancer		
subjects affected / exposed	2 / 3256 (0.06%)	2 / 3248 (0.06%)
occurrences causally related to treatment / all	0 / 2	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 1
Rectal cancer stage I		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Rectal cancer stage IV		

subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 1	0 / 0
Rectosigmoid cancer		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 1
Renal cancer		
subjects affected / exposed	3 / 3256 (0.09%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 3	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Small cell carcinoma		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Small cell lung cancer		
subjects affected / exposed	3 / 3256 (0.09%)	2 / 3248 (0.06%)
occurrences causally related to treatment / all	0 / 3	0 / 2
deaths causally related to treatment / all	0 / 3	0 / 1
Squamous cell carcinoma		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Squamous cell carcinoma of lung		
subjects affected / exposed	1 / 3256 (0.03%)	2 / 3248 (0.06%)
occurrences causally related to treatment / all	0 / 1	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 1
Squamous cell carcinoma of skin		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Squamous cell carcinoma of the oral cavity		

subjects affected / exposed	1 / 3256 (0.03%)	2 / 3248 (0.06%)
occurrences causally related to treatment / all	0 / 1	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0
Squamous cell carcinoma of the tongue		
subjects affected / exposed	0 / 3256 (0.00%)	2 / 3248 (0.06%)
occurrences causally related to treatment / all	0 / 0	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0
T-cell lymphoma		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 1	0 / 0
Transitional cell carcinoma		
subjects affected / exposed	2 / 3256 (0.06%)	3 / 3248 (0.09%)
occurrences causally related to treatment / all	0 / 2	0 / 3
deaths causally related to treatment / all	0 / 0	0 / 0
Rectal adenoma		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Colon adenoma		
subjects affected / exposed	2 / 3256 (0.06%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 2	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Lung cancer metastatic		
subjects affected / exposed	3 / 3256 (0.09%)	3 / 3248 (0.09%)
occurrences causally related to treatment / all	0 / 3	0 / 3
deaths causally related to treatment / all	0 / 3	0 / 2
Salivary gland adenoma		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Metastases to peritoneum		

subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
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 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 1
Adenocarcinoma pancreas		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 1
Oesophageal cancer metastatic		
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 2	0 / 1
deaths causally related to treatment / all	0 / 1	0 / 1
Neuroendocrine carcinoma		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 1	0 / 0
Lung neoplasm malignant		
subjects affected / exposed	8 / 3256 (0.25%)	14 / 3248 (0.43%)
occurrences causally related to treatment / all	0 / 8	0 / 16
deaths causally related to treatment / all	0 / 4	0 / 7
Metastases to central nervous system		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Pancreatic carcinoma stage IV		
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 1	0 / 1
Non-small cell lung cancer metastatic		

subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 1
Prostate cancer		
subjects affected / exposed	9 / 3256 (0.28%)	2 / 3248 (0.06%)
occurrences causally related to treatment / all	0 / 9	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0
Brain neoplasm		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Thymoma malignant		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Metastatic neoplasm		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 1
Colorectal cancer		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Renal neoplasm		
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Respiratory tract neoplasm		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 1
Lip and/or oral cavity cancer		

subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Adrenal neoplasm		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Hodgkin's disease stage IV		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Non-small cell lung cancer		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Oesophageal neoplasm		
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 2	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 1
Pancreatic neoplasm		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 1
Lung neoplasm		
subjects affected / exposed	4 / 3256 (0.12%)	4 / 3248 (0.12%)
occurrences causally related to treatment / all	0 / 4	0 / 4
deaths causally related to treatment / all	0 / 1	0 / 3
Rectal neoplasm		
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Metastatic squamous cell carcinoma		

subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 1
Gingival cancer		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 3	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Head and neck cancer		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Renal cell carcinoma		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Inflammatory pseudotumour		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Adenolymphoma		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Lung squamous cell carcinoma metastatic		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 1
Hepatic cancer		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 1
Hepatocellular carcinoma		

subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Ductal adenocarcinoma of pancreas			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 1	
Malignant genitourinary tract neoplasm			
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Malignant neoplasm papilla of Vater			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Gastrointestinal adenocarcinoma			
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Skin squamous cell carcinoma recurrent			
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Vascular disorders			
Aortic aneurysm			
subjects affected / exposed	2 / 3256 (0.06%)	4 / 3248 (0.12%)	
occurrences causally related to treatment / all	0 / 3	0 / 4	
deaths causally related to treatment / all	0 / 0	0 / 0	
Aortic dissection			
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Aortic stenosis			

subjects affected / exposed	6 / 3256 (0.18%)	3 / 3248 (0.09%)
occurrences causally related to treatment / all	0 / 6	0 / 3
deaths causally related to treatment / all	0 / 0	0 / 0
Arteriovenous fistula		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Circulatory collapse		
subjects affected / exposed	1 / 3256 (0.03%)	2 / 3248 (0.06%)
occurrences causally related to treatment / all	0 / 1	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0
Haematoma		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	1 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Hypertension		
subjects affected / exposed	1 / 3256 (0.03%)	5 / 3248 (0.15%)
occurrences causally related to treatment / all	0 / 1	0 / 5
deaths causally related to treatment / all	0 / 0	0 / 0
Hypertensive crisis		
subjects affected / exposed	5 / 3256 (0.15%)	6 / 3248 (0.18%)
occurrences causally related to treatment / all	0 / 5	0 / 8
deaths causally related to treatment / all	0 / 0	0 / 0
Hypotension		
subjects affected / exposed	4 / 3256 (0.12%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 4	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Hypovolaemic shock		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Intermittent claudication		

subjects affected / exposed	43 / 3256 (1.32%)	52 / 3248 (1.60%)
occurrences causally related to treatment / all	0 / 51	0 / 58
deaths causally related to treatment / all	0 / 0	0 / 0
Lymphoedema		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Orthostatic hypotension		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Peripheral coldness		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Peripheral ischaemia		
subjects affected / exposed	34 / 3256 (1.04%)	29 / 3248 (0.89%)
occurrences causally related to treatment / all	0 / 44	1 / 46
deaths causally related to treatment / all	0 / 0	0 / 0
Peripheral vascular disorder		
subjects affected / exposed	3 / 3256 (0.09%)	5 / 3248 (0.15%)
occurrences causally related to treatment / all	0 / 4	0 / 5
deaths causally related to treatment / all	0 / 0	0 / 0
Poor peripheral circulation		
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 1	0 / 0
Renovascular hypertension		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Temporal arteritis		

subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Thrombophlebitis superficial		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Varicose vein		
subjects affected / exposed	1 / 3256 (0.03%)	2 / 3248 (0.06%)
occurrences causally related to treatment / all	0 / 1	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0
Lymphocele		
subjects affected / exposed	1 / 3256 (0.03%)	2 / 3248 (0.06%)
occurrences causally related to treatment / all	0 / 1	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0
Dry gangrene		
subjects affected / exposed	2 / 3256 (0.06%)	3 / 3248 (0.09%)
occurrences causally related to treatment / all	0 / 3	0 / 3
deaths causally related to treatment / all	0 / 0	0 / 0
Subclavian artery stenosis		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Lymphorrhoea		
subjects affected / exposed	2 / 3256 (0.06%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 2	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Iliac artery stenosis		
subjects affected / exposed	4 / 3256 (0.12%)	3 / 3248 (0.09%)
occurrences causally related to treatment / all	0 / 4	0 / 3
deaths causally related to treatment / all	0 / 0	0 / 0
Vascular stenosis		

subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Peripheral artery aneurysm		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Peripheral artery occlusion		
subjects affected / exposed	7 / 3256 (0.21%)	9 / 3248 (0.28%)
occurrences causally related to treatment / all	0 / 9	0 / 10
deaths causally related to treatment / all	0 / 0	0 / 0
Hypertensive emergency		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Hypertensive urgency		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Venous occlusion		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	1 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Angiopathy		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Extremity necrosis		
subjects affected / exposed	12 / 3256 (0.37%)	8 / 3248 (0.25%)
occurrences causally related to treatment / all	0 / 14	0 / 9
deaths causally related to treatment / all	0 / 1	0 / 0
Peripheral embolism		

subjects affected / exposed	1 / 3256 (0.03%)	2 / 3248 (0.06%)	
occurrences causally related to treatment / all	0 / 1	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Peripheral arterial occlusive disease			
subjects affected / exposed	125 / 3256 (3.84%)	130 / 3248 (4.00%)	
occurrences causally related to treatment / all	1 / 156	0 / 182	
deaths causally related to treatment / all	0 / 0	0 / 0	
Arterial occlusive disease			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Ischaemic limb pain			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Lymphatic fistula			
subjects affected / exposed	0 / 3256 (0.00%)	3 / 3248 (0.09%)	
occurrences causally related to treatment / all	0 / 0	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Peripheral artery stenosis			
subjects affected / exposed	25 / 3256 (0.77%)	14 / 3248 (0.43%)	
occurrences causally related to treatment / all	0 / 29	0 / 17	
deaths causally related to treatment / all	0 / 0	0 / 0	
Peripheral artery thrombosis			
subjects affected / exposed	2 / 3256 (0.06%)	2 / 3248 (0.06%)	
occurrences causally related to treatment / all	0 / 2	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Peripheral venous disease			
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Surgical and medical procedures			
Bladder calculus removal			

subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Blepharoplasty		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Carotid endarterectomy		
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Foot amputation		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Knee arthroplasty		
subjects affected / exposed	0 / 3256 (0.00%)	2 / 3248 (0.06%)
occurrences causally related to treatment / all	0 / 0	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0
Umbilical hernia repair		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Vitrectomy		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Radical prostatectomy		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Diabetes mellitus management		

subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Transurethral bladder resection		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Peripheral revascularisation		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Brain tumour operation		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Cystoprostatectomy		
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Peripheral artery angioplasty		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Amputation		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Bladder polypectomy		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Bladder neoplasm surgery		

subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Limb reconstructive surgery			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Peripheral artery bypass			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Peripheral artery stent insertion			
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Eventration repair			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
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			
Asthenia			
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Chest discomfort			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Chest pain			
subjects affected / exposed	4 / 3256 (0.12%)	3 / 3248 (0.09%)	
occurrences causally related to treatment / all	0 / 4	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Crepitations			

subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Death		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 1	0 / 0
Gait disturbance		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Granuloma		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Hernia		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Impaired healing		
subjects affected / exposed	3 / 3256 (0.09%)	7 / 3248 (0.22%)
occurrences causally related to treatment / all	0 / 3	0 / 7
deaths causally related to treatment / all	0 / 0	0 / 0
Oedema		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Oedema peripheral		
subjects affected / exposed	1 / 3256 (0.03%)	2 / 3248 (0.06%)
occurrences causally related to treatment / all	0 / 1	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0
Pain		

subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Pyrexia		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Peripheral swelling		
subjects affected / exposed	0 / 3256 (0.00%)	2 / 3248 (0.06%)
occurrences causally related to treatment / all	0 / 0	0 / 3
deaths causally related to treatment / all	0 / 0	0 / 0
General physical health deterioration		
subjects affected / exposed	2 / 3256 (0.06%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 2	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Puncture site haemorrhage		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	1 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Systemic inflammatory response syndrome		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 1	0 / 0
Organ failure		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 1	0 / 0
Inflammation		
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Drug intolerance		

subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Non-cardiac chest pain			
subjects affected / exposed	3 / 3256 (0.09%)	3 / 3248 (0.09%)	
occurrences causally related to treatment / all	0 / 3	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Vascular stent thrombosis			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Stent-graft endoleak			
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Vascular stent occlusion			
subjects affected / exposed	1 / 3256 (0.03%)	2 / 3248 (0.06%)	
occurrences causally related to treatment / all	0 / 1	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Vascular stent stenosis			
subjects affected / exposed	11 / 3256 (0.34%)	8 / 3248 (0.25%)	
occurrences causally related to treatment / all	2 / 11	0 / 10	
deaths causally related to treatment / all	0 / 0	0 / 0	
Multiple organ dysfunction syndrome			
subjects affected / exposed	0 / 3256 (0.00%)	2 / 3248 (0.06%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 2	
Vascular device occlusion			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Immune system disorders			
Anaphylactic reaction			

subjects affected / exposed	0 / 3256 (0.00%)	2 / 3248 (0.06%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hypersensitivity			
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 1	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Social circumstances			
Disease risk factor			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Reproductive system and breast disorders			
Benign prostatic hyperplasia			
subjects affected / exposed	3 / 3256 (0.09%)	2 / 3248 (0.06%)	
occurrences causally related to treatment / all	0 / 3	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cervical polyp			
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pelvic prolapse			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Spermatocele			
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Uterine prolapse			
subjects affected / exposed	2 / 3256 (0.06%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	

Vulvar dysplasia			
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Prostatic obstruction			
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Vulval eczema			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Respiratory, thoracic and mediastinal disorders			
Acute pulmonary oedema			
subjects affected / exposed	2 / 3256 (0.06%)	2 / 3248 (0.06%)	
occurrences causally related to treatment / all	0 / 2	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Acute respiratory distress syndrome			
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Acute respiratory failure			
subjects affected / exposed	2 / 3256 (0.06%)	5 / 3248 (0.15%)	
occurrences causally related to treatment / all	0 / 2	0 / 5	
deaths causally related to treatment / all	0 / 0	0 / 1	
Aspiration			
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (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 / 3256 (0.03%)	2 / 3248 (0.06%)	
occurrences causally related to treatment / all	0 / 1	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	

Chronic obstructive pulmonary disease			
subjects affected / exposed	16 / 3256 (0.49%)	9 / 3248 (0.28%)	
occurrences causally related to treatment / all	0 / 23	0 / 12	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cough			
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Dyspnoea			
subjects affected / exposed	4 / 3256 (0.12%)	6 / 3248 (0.18%)	
occurrences causally related to treatment / all	0 / 5	0 / 6	
deaths causally related to treatment / all	0 / 0	0 / 0	
Emphysema			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hypoxia			
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Interstitial lung disease			
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 2	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Nasal polyps			
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Nasal septum deviation			
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pleural effusion			

subjects affected / exposed	3 / 3256 (0.09%)	3 / 3248 (0.09%)
occurrences causally related to treatment / all	0 / 3	0 / 3
deaths causally related to treatment / all	0 / 0	0 / 0
Pneumonia aspiration		
subjects affected / exposed	5 / 3256 (0.15%)	6 / 3248 (0.18%)
occurrences causally related to treatment / all	0 / 6	0 / 6
deaths causally related to treatment / all	0 / 1	0 / 3
Pneumothorax		
subjects affected / exposed	3 / 3256 (0.09%)	2 / 3248 (0.06%)
occurrences causally related to treatment / all	0 / 4	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0
Pneumothorax spontaneous		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Pulmonary congestion		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Pulmonary embolism		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Pulmonary fibrosis		
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	1 / 1	0 / 1
deaths causally related to treatment / all	1 / 1	0 / 0
Pulmonary hypertension		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Pulmonary oedema		

subjects affected / exposed	1 / 3256 (0.03%)	2 / 3248 (0.06%)
occurrences causally related to treatment / all	0 / 1	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0
Respiratory distress		
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 1
Respiratory failure		
subjects affected / exposed	5 / 3256 (0.15%)	4 / 3248 (0.12%)
occurrences causally related to treatment / all	0 / 5	0 / 4
deaths causally related to treatment / all	0 / 2	0 / 3
Sinus polyp		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Vocal cord polyp		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Pharyngeal lesion		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Pulmonary mass		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Tonsillar cyst		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Chronic respiratory disease		

subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 1	0 / 0	
Acute interstitial pneumonitis			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Psychiatric disorders			
Acute psychosis			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Alcohol abuse			
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Alcoholism			
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Completed suicide			
subjects affected / exposed	4 / 3256 (0.12%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 4	0 / 0	
deaths causally related to treatment / all	0 / 4	0 / 0	
Depression			
subjects affected / exposed	3 / 3256 (0.09%)	5 / 3248 (0.15%)	
occurrences causally related to treatment / all	0 / 3	0 / 5	
deaths causally related to treatment / all	0 / 0	0 / 0	
Drug abuse			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Intentional self-injury			

subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Organic brain syndrome			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Mental status changes			
subjects affected / exposed	1 / 3256 (0.03%)	4 / 3248 (0.12%)	
occurrences causally related to treatment / all	0 / 1	0 / 4	
deaths causally related to treatment / all	0 / 0	0 / 0	
Alcohol withdrawal syndrome			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Transient psychosis			
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Product issues			
Device breakage			
subjects affected / exposed	2 / 3256 (0.06%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Device dislocation			
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Device occlusion			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Patient-device incompatibility			

subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Device battery issue			
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hepatobiliary disorders			
Bile duct stone			
subjects affected / exposed	3 / 3256 (0.09%)	2 / 3248 (0.06%)	
occurrences causally related to treatment / all	0 / 4	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cholangitis			
subjects affected / exposed	1 / 3256 (0.03%)	3 / 3248 (0.09%)	
occurrences causally related to treatment / all	0 / 2	0 / 6	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cholangitis acute			
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cholecystitis			
subjects affected / exposed	7 / 3256 (0.21%)	3 / 3248 (0.09%)	
occurrences causally related to treatment / all	0 / 7	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cholecystitis acute			
subjects affected / exposed	2 / 3256 (0.06%)	12 / 3248 (0.37%)	
occurrences causally related to treatment / all	0 / 2	0 / 12	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cholelithiasis			
subjects affected / exposed	2 / 3256 (0.06%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 2	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cholestasis			

subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Cirrhosis alcoholic		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Hepatic cirrhosis		
subjects affected / exposed	3 / 3256 (0.09%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 3	0 / 0
deaths causally related to treatment / all	0 / 1	0 / 0
Hepatic failure		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
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	2 / 3256 (0.06%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	1 / 2	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Hepatic steatosis		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Hepatorenal syndrome		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Bile duct obstruction		
subjects affected / exposed	0 / 3256 (0.00%)	3 / 3248 (0.09%)
occurrences causally related to treatment / all	0 / 0	0 / 3
deaths causally related to treatment / all	0 / 0	0 / 0
Biliary dyskinesia		

subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Steatohepatitis			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Primary biliary cholangitis			
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Investigations			
Blood creatinine increased			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Blood glucose increased			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Blood potassium increased			
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Blood pressure increased			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cardiovascular function test abnormal			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cystoscopy			

subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Immunoglobulins decreased			
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Carbohydrate antigen 19-9 increased			
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cardiac stress test abnormal			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
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 / 3256 (0.03%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Vascular test			
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (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			
Accidental overdose			
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Alcohol poisoning			
subjects affected / exposed	2 / 3256 (0.06%)	0 / 3248 (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	2 / 3256 (0.06%)	3 / 3248 (0.09%)
occurrences causally related to treatment / all	0 / 2	0 / 4
deaths causally related to treatment / all	0 / 0	0 / 0
Brain herniation		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Carbon monoxide poisoning		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Compression fracture		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Concussion		
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Facial bones fracture		
subjects affected / exposed	0 / 3256 (0.00%)	2 / 3248 (0.06%)
occurrences causally related to treatment / all	0 / 0	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0
Failure to anastomose		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Fall		
subjects affected / exposed	5 / 3256 (0.15%)	2 / 3248 (0.06%)
occurrences causally related to treatment / all	0 / 5	0 / 2
deaths causally related to treatment / all	0 / 1	0 / 0
Femoral neck fracture		

subjects affected / exposed	4 / 3256 (0.12%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 4	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Femur fracture		
subjects affected / exposed	10 / 3256 (0.31%)	12 / 3248 (0.37%)
occurrences causally related to treatment / all	0 / 10	0 / 12
deaths causally related to treatment / all	0 / 0	0 / 0
Fibula fracture		
subjects affected / exposed	3 / 3256 (0.09%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 3	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Foot fracture		
subjects affected / exposed	2 / 3256 (0.06%)	2 / 3248 (0.06%)
occurrences causally related to treatment / all	0 / 2	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0
Forearm fracture		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Fractured sacrum		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Head injury		
subjects affected / exposed	2 / 3256 (0.06%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 2	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Hip fracture		
subjects affected / exposed	5 / 3256 (0.15%)	5 / 3248 (0.15%)
occurrences causally related to treatment / all	0 / 5	0 / 5
deaths causally related to treatment / all	0 / 0	0 / 0
Humerus fracture		

subjects affected / exposed	1 / 3256 (0.03%)	3 / 3248 (0.09%)
occurrences causally related to treatment / all	0 / 1	0 / 3
deaths causally related to treatment / all	0 / 0	0 / 0
Incisional hernia		
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Injury		
subjects affected / exposed	2 / 3256 (0.06%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 2	0 / 0
deaths causally related to treatment / all	0 / 1	0 / 0
Intentional overdose		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Joint dislocation		
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Multiple injuries		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Open fracture		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Patella fracture		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Penetrating abdominal trauma		

subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Radius fracture		
subjects affected / exposed	3 / 3256 (0.09%)	2 / 3248 (0.06%)
occurrences causally related to treatment / all	0 / 3	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0
Rib fracture		
subjects affected / exposed	2 / 3256 (0.06%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 2	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Road traffic accident		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 1	0 / 0
Seroma		
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 1	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0
Skull fractured base		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (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 / 3256 (0.03%)	4 / 3248 (0.12%)
occurrences causally related to treatment / all	0 / 1	0 / 5
deaths causally related to treatment / all	0 / 0	0 / 0
Spinal fracture		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Tendon rupture		

subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Tibia fracture		
subjects affected / exposed	3 / 3256 (0.09%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 3	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Traumatic ulcer		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Wound dehiscence		
subjects affected / exposed	4 / 3256 (0.12%)	3 / 3248 (0.09%)
occurrences causally related to treatment / all	0 / 4	0 / 3
deaths causally related to treatment / all	0 / 0	0 / 0
Vascular pseudoaneurysm		
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	1 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Vascular graft occlusion		
subjects affected / exposed	1 / 3256 (0.03%)	3 / 3248 (0.09%)
occurrences causally related to treatment / all	0 / 1	0 / 3
deaths causally related to treatment / all	0 / 0	0 / 0
Traumatic fracture		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
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	2 / 3256 (0.06%)	4 / 3248 (0.12%)
occurrences causally related to treatment / all	0 / 2	0 / 4
deaths causally related to treatment / all	0 / 0	0 / 0
Thoracic vertebral fracture		

subjects affected / exposed	2 / 3256 (0.06%)	2 / 3248 (0.06%)
occurrences causally related to treatment / all	0 / 2	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0
Contusion		
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Anastomotic stenosis		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Graft thrombosis		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Thermal burn		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Wound complication		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Wound necrosis		
subjects affected / exposed	1 / 3256 (0.03%)	2 / 3248 (0.06%)
occurrences causally related to treatment / all	0 / 1	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0
Coronary artery restenosis		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Postoperative thoracic procedure complication		

subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Post laminectomy syndrome		
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Underdose		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	1 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Vascular procedure complication		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 1	0 / 0
Post procedural bile leak		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Post procedural discharge		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Procedural complication		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 1
Chemical burn of skin		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Post procedural complication		

subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (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	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Pelvic fracture		
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Limb injury		
subjects affected / exposed	2 / 3256 (0.06%)	4 / 3248 (0.12%)
occurrences causally related to treatment / all	0 / 3	0 / 4
deaths causally related to treatment / all	0 / 0	0 / 0
Chest injury		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 1
Traumatic intracranial haemorrhage		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	1 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Upper limb fracture		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Lower limb fracture		
subjects affected / exposed	1 / 3256 (0.03%)	2 / 3248 (0.06%)
occurrences causally related to treatment / all	0 / 1	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0
Post procedural oedema		

subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Vascular graft complication		
subjects affected / exposed	5 / 3256 (0.15%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 5	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Cerebral hyperperfusion syndrome		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Gastrointestinal anastomotic leak		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Bone contusion		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Periprosthetic fracture		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Peripheral arterial reocclusion		
subjects affected / exposed	4 / 3256 (0.12%)	2 / 3248 (0.06%)
occurrences causally related to treatment / all	0 / 6	0 / 3
deaths causally related to treatment / all	0 / 0	0 / 0
Spinal column injury		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Vascular graft thrombosis		

subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Pubis fracture		
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)
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 / 3256 (0.06%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 2	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Peripheral artery restenosis		
subjects affected / exposed	16 / 3256 (0.49%)	15 / 3248 (0.46%)
occurrences causally related to treatment / all	0 / 22	0 / 18
deaths causally related to treatment / all	0 / 0	0 / 0
Meniscus injury		
subjects affected / exposed	2 / 3256 (0.06%)	4 / 3248 (0.12%)
occurrences causally related to treatment / all	0 / 2	0 / 4
deaths causally related to treatment / all	0 / 0	0 / 0
Skin scar contracture		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Procedural intestinal perforation		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Vascular graft stenosis		
subjects affected / exposed	2 / 3256 (0.06%)	2 / 3248 (0.06%)
occurrences causally related to treatment / all	0 / 2	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0
Vascular graft restenosis		

subjects affected / exposed	0 / 3256 (0.00%)	2 / 3248 (0.06%)	
occurrences causally related to treatment / all	0 / 0	0 / 4	
deaths causally related to treatment / all	0 / 0	0 / 0	
Vascular access site haematoma			
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Arterial bypass thrombosis			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Arterial bypass occlusion			
subjects affected / exposed	2 / 3256 (0.06%)	2 / 3248 (0.06%)	
occurrences causally related to treatment / all	0 / 2	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Arterial bypass stenosis			
subjects affected / exposed	1 / 3256 (0.03%)	2 / 3248 (0.06%)	
occurrences causally related to treatment / all	0 / 1	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Foreign body in gastrointestinal tract			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Congenital, familial and genetic disorders			
Congenital bladder neck obstruction			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hydrocele			
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Phimosis			

subjects affected / exposed	0 / 3256 (0.00%)	2 / 3248 (0.06%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Adenomatous polyposis coli			
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (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 myocardial infarction			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Angina pectoris			
subjects affected / exposed	10 / 3256 (0.31%)	10 / 3248 (0.31%)	
occurrences causally related to treatment / all	0 / 11	0 / 10	
deaths causally related to treatment / all	0 / 0	0 / 0	
Angina unstable			
subjects affected / exposed	2 / 3256 (0.06%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 2	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Aortic valve stenosis			
subjects affected / exposed	2 / 3256 (0.06%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Arrhythmia			
subjects affected / exposed	3 / 3256 (0.09%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 3	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Arrhythmia supraventricular			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Arteriosclerosis coronary artery			

subjects affected / exposed	2 / 3256 (0.06%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 2	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Atrial fibrillation		
subjects affected / exposed	19 / 3256 (0.58%)	20 / 3248 (0.62%)
occurrences causally related to treatment / all	0 / 19	0 / 21
deaths causally related to treatment / all	0 / 0	0 / 0
Atrial flutter		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Atrioventricular block complete		
subjects affected / exposed	3 / 3256 (0.09%)	2 / 3248 (0.06%)
occurrences causally related to treatment / all	0 / 3	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0
Atrioventricular block first degree		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Atrioventricular block second degree		
subjects affected / exposed	0 / 3256 (0.00%)	2 / 3248 (0.06%)
occurrences causally related to treatment / all	0 / 0	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0
Bradycardia		
subjects affected / exposed	2 / 3256 (0.06%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 2	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Cardiac arrest		
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Cardiac failure		

subjects affected / exposed	14 / 3256 (0.43%)	14 / 3248 (0.43%)
occurrences causally related to treatment / all	0 / 16	0 / 15
deaths causally related to treatment / all	0 / 1	0 / 2
Cardiac failure acute		
subjects affected / exposed	3 / 3256 (0.09%)	6 / 3248 (0.18%)
occurrences causally related to treatment / all	0 / 3	0 / 6
deaths causally related to treatment / all	0 / 0	0 / 0
Cardiac failure chronic		
subjects affected / exposed	6 / 3256 (0.18%)	3 / 3248 (0.09%)
occurrences causally related to treatment / all	0 / 8	0 / 4
deaths causally related to treatment / all	0 / 0	0 / 0
Cardiac failure congestive		
subjects affected / exposed	10 / 3256 (0.31%)	8 / 3248 (0.25%)
occurrences causally related to treatment / all	0 / 11	0 / 8
deaths causally related to treatment / all	0 / 1	0 / 0
Cardiac tamponade		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (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	2 / 3256 (0.06%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 2	0 / 0
deaths causally related to treatment / all	0 / 2	0 / 0
Cardiogenic shock		
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 1	0 / 0
Coronary artery disease		
subjects affected / exposed	15 / 3256 (0.46%)	21 / 3248 (0.65%)
occurrences causally related to treatment / all	0 / 18	0 / 21
deaths causally related to treatment / all	0 / 1	0 / 0
Coronary artery occlusion		

subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Coronary artery stenosis		
subjects affected / exposed	4 / 3256 (0.12%)	2 / 3248 (0.06%)
occurrences causally related to treatment / all	0 / 4	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0
Cyanosis		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Hypertensive heart disease		
subjects affected / exposed	0 / 3256 (0.00%)	2 / 3248 (0.06%)
occurrences causally related to treatment / all	0 / 0	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0
Myocardial infarction		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Myocardial ischaemia		
subjects affected / exposed	2 / 3256 (0.06%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 2	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Pericardial effusion		
subjects affected / exposed	0 / 3256 (0.00%)	2 / 3248 (0.06%)
occurrences causally related to treatment / all	0 / 0	0 / 3
deaths causally related to treatment / all	0 / 0	0 / 0
Pericardial haemorrhage		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	1 / 1
deaths causally related to treatment / all	0 / 0	1 / 1
Pericarditis		

subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Right ventricular failure		
subjects affected / exposed	0 / 3256 (0.00%)	2 / 3248 (0.06%)
occurrences causally related to treatment / all	0 / 0	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0
Sinus arrest		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Sinus bradycardia		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Supraventricular tachycardia		
subjects affected / exposed	3 / 3256 (0.09%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 3	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Ventricular arrhythmia		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Ventricular fibrillation		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Ventricular tachycardia		
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Ischaemic cardiomyopathy		

subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 3	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cardiopulmonary failure			
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 1	0 / 0	
Congestive cardiomyopathy			
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Acute left ventricular failure			
subjects affected / exposed	2 / 3256 (0.06%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 2	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Supraventricular tachyarrhythmia			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Stress cardiomyopathy			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Sinus node dysfunction			
subjects affected / exposed	0 / 3256 (0.00%)	2 / 3248 (0.06%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Nervous system disorders			
Carotid artery stenosis			
subjects affected / exposed	22 / 3256 (0.68%)	24 / 3248 (0.74%)	
occurrences causally related to treatment / all	0 / 23	0 / 26	
deaths causally related to treatment / all	0 / 0	0 / 0	
Carpal tunnel syndrome			

subjects affected / exposed	1 / 3256 (0.03%)	2 / 3248 (0.06%)
occurrences causally related to treatment / all	0 / 1	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0
Cerebral atrophy		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Cerebral haemorrhage		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	1 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Cerebral infarction		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Dementia		
subjects affected / exposed	3 / 3256 (0.09%)	2 / 3248 (0.06%)
occurrences causally related to treatment / all	0 / 3	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0
Diabetic neuropathy		
subjects affected / exposed	2 / 3256 (0.06%)	2 / 3248 (0.06%)
occurrences causally related to treatment / all	0 / 2	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0
Dizziness		
subjects affected / exposed	5 / 3256 (0.15%)	3 / 3248 (0.09%)
occurrences causally related to treatment / all	0 / 5	0 / 3
deaths causally related to treatment / all	0 / 0	0 / 0
Dizziness postural		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Drop attacks		

subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Epilepsy		
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Guillain-Barre syndrome		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Hemiparesis		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Hydrocephalus		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Intracranial aneurysm		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Mental impairment		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Myelopathy		
subjects affected / exposed	1 / 3256 (0.03%)	2 / 3248 (0.06%)
occurrences causally related to treatment / all	0 / 1	0 / 3
deaths causally related to treatment / all	0 / 0	0 / 0
Nerve compression		

subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Neuralgia		
subjects affected / exposed	0 / 3256 (0.00%)	2 / 3248 (0.06%)
occurrences causally related to treatment / all	0 / 0	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0
Parkinsonism		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Polyneuropathy		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Presyncope		
subjects affected / exposed	2 / 3256 (0.06%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 2	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Sciatica		
subjects affected / exposed	2 / 3256 (0.06%)	4 / 3248 (0.12%)
occurrences causally related to treatment / all	0 / 2	0 / 4
deaths causally related to treatment / all	0 / 0	0 / 0
Somnolence		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Spinal claudication		
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Status epilepticus		

subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Subarachnoid haemorrhage		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	1 / 1
deaths causally related to treatment / all	0 / 0	1 / 1
Syncope		
subjects affected / exposed	9 / 3256 (0.28%)	11 / 3248 (0.34%)
occurrences causally related to treatment / all	1 / 9	0 / 12
deaths causally related to treatment / all	0 / 1	0 / 0
Tension headache		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Tonic convulsion		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Vertebral artery stenosis		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Carotid artery occlusion		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Cervical radiculopathy		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Lumbar radiculopathy		

subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 2	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Facial paresis		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Ischaemic neuropathy		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Restless legs syndrome		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Putamen haemorrhage		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	1 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Cranial nerve disorder		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
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 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 1
Parkinson's disease		
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Carotid artery disease		

subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Vascular encephalopathy			
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pseudoradicular syndrome			
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Brain injury			
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 1	0 / 0	
Colloid brain cyst			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Sacral radiculopathy			
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Blood and lymphatic system disorders			
Anaemia			
subjects affected / exposed	14 / 3256 (0.43%)	10 / 3248 (0.31%)	
occurrences causally related to treatment / all	10 / 16	4 / 13	
deaths causally related to treatment / all	0 / 0	0 / 0	
Iron deficiency anaemia			
subjects affected / exposed	4 / 3256 (0.12%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	2 / 4	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Leukocytosis			

subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Microcytic anaemia			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Normochromic normocytic anaemia			
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Normocytic anaemia			
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Thrombocytopenia			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Thrombocytosis			
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Nephrogenic anaemia			
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Blood loss anaemia			
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	1 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Ear and labyrinth disorders			
Deafness			

subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Vertigo			
subjects affected / exposed	0 / 3256 (0.00%)	2 / 3248 (0.06%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Sudden hearing loss			
subjects affected / exposed	2 / 3256 (0.06%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Eye disorders			
Cataract			
subjects affected / exposed	8 / 3256 (0.25%)	8 / 3248 (0.25%)	
occurrences causally related to treatment / all	0 / 9	0 / 10	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cataract subcapsular			
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Diabetic retinopathy			
subjects affected / exposed	3 / 3256 (0.09%)	2 / 3248 (0.06%)	
occurrences causally related to treatment / all	0 / 3	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Eyelid ptosis			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Glaucoma			
subjects affected / exposed	2 / 3256 (0.06%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Iridocyclitis			

subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Macular degeneration		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Macular oedema		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Myopia		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Retinal detachment		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Retinal haemorrhage		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Vitreous haemorrhage		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	1 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Macular hole		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Detachment of macular retinal pigment epithelium		

subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Vitreoretinal traction syndrome			
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Macular detachment			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Gastrointestinal disorders			
Abdominal adhesions			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Abdominal mass			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Abdominal pain			
subjects affected / exposed	3 / 3256 (0.09%)	2 / 3248 (0.06%)	
occurrences causally related to treatment / all	0 / 4	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Abdominal pain lower			
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Abdominal pain upper			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Anal fissure			

subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Anal stenosis		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Ascites		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Chronic gastritis		
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Colitis		
subjects affected / exposed	1 / 3256 (0.03%)	2 / 3248 (0.06%)
occurrences causally related to treatment / all	0 / 1	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0
Colitis ischaemic		
subjects affected / exposed	2 / 3256 (0.06%)	2 / 3248 (0.06%)
occurrences causally related to treatment / all	0 / 2	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0
Constipation		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Crohn's disease		
subjects affected / exposed	2 / 3256 (0.06%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 2	0 / 0
deaths causally related to treatment / all	0 / 1	0 / 0
Dental alveolar anomaly		

subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Diaphragmatic hernia		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Diarrhoea		
subjects affected / exposed	3 / 3256 (0.09%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 3	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Diverticulum intestinal haemorrhagic		
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	1 / 1	1 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Dry mouth		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	1 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Duodenal ulcer		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Dysphagia		
subjects affected / exposed	3 / 3256 (0.09%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 3	0 / 1
deaths causally related to treatment / all	0 / 1	0 / 0
Gallstone ileus		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Gastric haemorrhage		

subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Gastric polyps		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Gastric ulcer		
subjects affected / exposed	3 / 3256 (0.09%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 3	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Gastric ulcer haemorrhage		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	1 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Gastritis		
subjects affected / exposed	3 / 3256 (0.09%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 3	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Gastritis erosive		
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Gastroesophageal reflux disease		
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Gastrointestinal necrosis		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (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	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Hiatus hernia		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Ileus		
subjects affected / exposed	2 / 3256 (0.06%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 2	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Impaired gastric emptying		
subjects affected / exposed	2 / 3256 (0.06%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 2	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Inguinal hernia		
subjects affected / exposed	4 / 3256 (0.12%)	4 / 3248 (0.12%)
occurrences causally related to treatment / all	0 / 4	0 / 4
deaths causally related to treatment / all	0 / 0	0 / 0
Intestinal angina		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0
Intestinal ischaemia		
subjects affected / exposed	4 / 3256 (0.12%)	3 / 3248 (0.09%)
occurrences causally related to treatment / all	0 / 4	0 / 3
deaths causally related to treatment / all	0 / 1	0 / 2
Intestinal obstruction		
subjects affected / exposed	3 / 3256 (0.09%)	3 / 3248 (0.09%)
occurrences causally related to treatment / all	0 / 3	0 / 3
deaths causally related to treatment / all	0 / 0	0 / 1
Intestinal perforation		

subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Intestinal ulcer		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 3
deaths causally related to treatment / all	0 / 0	0 / 0
Large intestine perforation		
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Nausea		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Oesophagitis		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Pancreatitis		
subjects affected / exposed	4 / 3256 (0.12%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 4	1 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Pancreatitis acute		
subjects affected / exposed	1 / 3256 (0.03%)	4 / 3248 (0.12%)
occurrences causally related to treatment / all	0 / 1	0 / 5
deaths causally related to treatment / all	0 / 0	0 / 0
Pancreatitis chronic		
subjects affected / exposed	0 / 3256 (0.00%)	4 / 3248 (0.12%)
occurrences causally related to treatment / all	0 / 0	1 / 4
deaths causally related to treatment / all	0 / 0	0 / 0
Pancreatitis necrotising		

subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Peptic ulcer perforation		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Periodontal disease		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Small intestinal obstruction		
subjects affected / exposed	0 / 3256 (0.00%)	3 / 3248 (0.09%)
occurrences causally related to treatment / all	0 / 0	0 / 3
deaths causally related to treatment / all	0 / 0	0 / 0
Small intestinal perforation		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Tooth loss		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Umbilical hernia		
subjects affected / exposed	2 / 3256 (0.06%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 2	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Umbilical hernia, obstructive		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Upper gastrointestinal haemorrhage		

subjects affected / exposed	2 / 3256 (0.06%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	2 / 2	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Volvulus		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Vomiting		
subjects affected / exposed	0 / 3256 (0.00%)	2 / 3248 (0.06%)
occurrences causally related to treatment / all	0 / 0	1 / 2
deaths causally related to treatment / all	0 / 0	0 / 0
Pancreatic mass		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Subileus		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (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 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	2 / 2	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Mechanical ileus		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Gastric antral vascular ectasia		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	1 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Large intestine polyp		

subjects affected / exposed	10 / 3256 (0.31%)	4 / 3248 (0.12%)	
occurrences causally related to treatment / all	0 / 10	0 / 4	
deaths causally related to treatment / all	0 / 0	0 / 0	
Oedematous pancreatitis			
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Alcoholic pancreatitis			
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Large intestinal obstruction			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Enterovesical fistula			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pancreatic duct stenosis			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Abdominal incarcerated hernia			
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Obstructive pancreatitis			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
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			
Angioedema			

subjects affected / exposed	0 / 3256 (0.00%)	2 / 3248 (0.06%)
occurrences causally related to treatment / all	0 / 0	0 / 3
deaths causally related to treatment / all	0 / 0	0 / 0
Decubitus ulcer		
subjects affected / exposed	0 / 3256 (0.00%)	2 / 3248 (0.06%)
occurrences causally related to treatment / all	0 / 0	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0
Dermal cyst		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Dermatitis		
subjects affected / exposed	2 / 3256 (0.06%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 2	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Dermatitis allergic		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Dermatitis contact		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Haemorrhage subcutaneous		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	1 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Hidradenitis		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Pemphigoid		

subjects affected / exposed	2 / 3256 (0.06%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 2	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Pemphigus		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Skin exfoliation		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Skin lesion		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Skin necrosis		
subjects affected / exposed	1 / 3256 (0.03%)	2 / 3248 (0.06%)
occurrences causally related to treatment / all	0 / 1	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0
Skin ulcer		
subjects affected / exposed	10 / 3256 (0.31%)	12 / 3248 (0.37%)
occurrences causally related to treatment / all	0 / 11	0 / 14
deaths causally related to treatment / all	0 / 0	0 / 0
Urticaria		
subjects affected / exposed	2 / 3256 (0.06%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	1 / 2	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Diabetic ulcer		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Toxic skin eruption		

subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Diabetic foot			
subjects affected / exposed	9 / 3256 (0.28%)	9 / 3248 (0.28%)	
occurrences causally related to treatment / all	0 / 9	0 / 11	
deaths causally related to treatment / all	0 / 0	0 / 0	
Drug reaction with eosinophilia and systemic symptoms			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Ischaemic skin ulcer			
subjects affected / exposed	1 / 3256 (0.03%)	3 / 3248 (0.09%)	
occurrences causally related to treatment / all	0 / 1	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Myxoid cyst			
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Renal and urinary disorders			
Calculus bladder			
subjects affected / exposed	3 / 3256 (0.09%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 3	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Calculus urinary			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
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 / 3256 (0.03%)	2 / 3248 (0.06%)	
occurrences causally related to treatment / all	0 / 1	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pelvi-ureteric obstruction			

subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Renal artery stenosis		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Renal colic		
subjects affected / exposed	4 / 3256 (0.12%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 4	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Renal failure		
subjects affected / exposed	6 / 3256 (0.18%)	3 / 3248 (0.09%)
occurrences causally related to treatment / all	1 / 8	0 / 3
deaths causally related to treatment / all	0 / 0	0 / 1
Ureteric fistula		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Urinary incontinence		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (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	1 / 3256 (0.03%)	2 / 3248 (0.06%)
occurrences causally related to treatment / all	0 / 1	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0
Urinary tract obstruction		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Renal impairment		

subjects affected / exposed	3 / 3256 (0.09%)	3 / 3248 (0.09%)	
occurrences causally related to treatment / all	1 / 3	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Chronic kidney disease			
subjects affected / exposed	5 / 3256 (0.15%)	2 / 3248 (0.06%)	
occurrences causally related to treatment / all	0 / 5	0 / 2	
deaths causally related to treatment / all	0 / 1	0 / 0	
Urethral stenosis			
subjects affected / exposed	1 / 3256 (0.03%)	2 / 3248 (0.06%)	
occurrences causally related to treatment / all	0 / 1	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Acute kidney injury			
subjects affected / exposed	9 / 3256 (0.28%)	17 / 3248 (0.52%)	
occurrences causally related to treatment / all	0 / 9	1 / 17	
deaths causally related to treatment / all	0 / 0	0 / 1	
End stage renal disease			
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 1	0 / 0	
Ureterolithiasis			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Endocrine disorders			
Goitre			
subjects affected / exposed	2 / 3256 (0.06%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 2	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Thyroid mass			
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Musculoskeletal and connective tissue disorders			

Arthritis			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Back pain			
subjects affected / exposed	3 / 3256 (0.09%)	2 / 3248 (0.06%)	
occurrences causally related to treatment / all	0 / 3	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Bone disorder			
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Bone formation increased			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Bursitis			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cervical spinal stenosis			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Compartment syndrome			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Fracture nonunion			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Gouty arthritis			

subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Jaw cyst		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Lumbar spinal stenosis		
subjects affected / exposed	3 / 3256 (0.09%)	3 / 3248 (0.09%)
occurrences causally related to treatment / all	0 / 3	0 / 4
deaths causally related to treatment / all	0 / 0	0 / 0
Metatarsalgia		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Muscle disorder		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Muscle spasms		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Muscular weakness		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Musculoskeletal pain		
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Osteitis		

subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Osteoarthritis		
subjects affected / exposed	14 / 3256 (0.43%)	13 / 3248 (0.40%)
occurrences causally related to treatment / all	2 / 18	0 / 14
deaths causally related to treatment / all	0 / 0	0 / 0
Osteonecrosis		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Osteoporotic fracture		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Pain in extremity		
subjects affected / exposed	6 / 3256 (0.18%)	14 / 3248 (0.43%)
occurrences causally related to treatment / all	0 / 6	1 / 15
deaths causally related to treatment / all	0 / 0	0 / 0
Pathological fracture		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Polymyalgia rheumatica		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Rhabdomyolysis		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Rotator cuff syndrome		

subjects affected / exposed	0 / 3256 (0.00%)	4 / 3248 (0.12%)
occurrences causally related to treatment / all	0 / 0	0 / 4
deaths causally related to treatment / all	0 / 0	0 / 0
Spinal osteoarthritis		
subjects affected / exposed	2 / 3256 (0.06%)	4 / 3248 (0.12%)
occurrences causally related to treatment / all	0 / 3	0 / 4
deaths causally related to treatment / all	0 / 0	0 / 0
Synovial cyst		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Tendon disorder		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Pseudarthrosis		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Intervertebral disc protrusion		
subjects affected / exposed	6 / 3256 (0.18%)	6 / 3248 (0.18%)
occurrences causally related to treatment / all	0 / 7	0 / 7
deaths causally related to treatment / all	0 / 0	0 / 0
Musculoskeletal chest pain		
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Spondyloarthropathy		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Vertebral column mass		

subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Acquired claw toe		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Foot deformity		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Ligament disorder		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Soft tissue disorder		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Intervertebral disc disorder		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Spondylolisthesis		
subjects affected / exposed	1 / 3256 (0.03%)	2 / 3248 (0.06%)
occurrences causally related to treatment / all	0 / 1	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0
Bone loss		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Soft tissue necrosis		

subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Spinal pain			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Systemic scleroderma			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
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	6 / 3256 (0.18%)	2 / 3248 (0.06%)	
occurrences causally related to treatment / all	0 / 6	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Infections and infestations			
Abscess			
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Acute sinusitis			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Appendicitis			
subjects affected / exposed	4 / 3256 (0.12%)	5 / 3248 (0.15%)	
occurrences causally related to treatment / all	0 / 4	0 / 5	
deaths causally related to treatment / all	0 / 0	0 / 0	
Arteriosclerotic gangrene			
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Bacteraemia			

subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Blister infected		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Bronchiolitis		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Bronchitis		
subjects affected / exposed	2 / 3256 (0.06%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 2	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Carbuncle		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0
Cellulitis		
subjects affected / exposed	19 / 3256 (0.58%)	18 / 3248 (0.55%)
occurrences causally related to treatment / all	0 / 19	1 / 20
deaths causally related to treatment / all	0 / 0	0 / 0
Cellulitis gangrenous		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
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	2 / 3256 (0.06%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 2	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Cystitis		

subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Diabetic gangrene		
subjects affected / exposed	2 / 3256 (0.06%)	2 / 3248 (0.06%)
occurrences causally related to treatment / all	0 / 2	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0
Diarrhoea infectious		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Diverticulitis		
subjects affected / exposed	4 / 3256 (0.12%)	5 / 3248 (0.15%)
occurrences causally related to treatment / all	0 / 4	0 / 5
deaths causally related to treatment / all	0 / 0	0 / 0
Ear infection		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0
Eczema infected		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Endocarditis		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Epididymitis		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Erysipelas		

subjects affected / exposed	1 / 3256 (0.03%)	10 / 3248 (0.31%)
occurrences causally related to treatment / all	0 / 1	0 / 11
deaths causally related to treatment / all	0 / 0	0 / 0
Escherichia sepsis		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Fournier's gangrene		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Fungal cystitis		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Fungal skin infection		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Gangrene		
subjects affected / exposed	16 / 3256 (0.49%)	20 / 3248 (0.62%)
occurrences causally related to treatment / all	0 / 18	0 / 22
deaths causally related to treatment / all	0 / 0	0 / 0
Gastroenteritis		
subjects affected / exposed	3 / 3256 (0.09%)	2 / 3248 (0.06%)
occurrences causally related to treatment / all	0 / 3	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0
Gastroenteritis salmonella		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Gastroenteritis viral		

subjects affected / exposed	2 / 3256 (0.06%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 2	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Infected skin ulcer		
subjects affected / exposed	6 / 3256 (0.18%)	4 / 3248 (0.12%)
occurrences causally related to treatment / all	0 / 6	0 / 4
deaths causally related to treatment / all	0 / 0	0 / 0
Influenza		
subjects affected / exposed	2 / 3256 (0.06%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 2	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Localised infection		
subjects affected / exposed	19 / 3256 (0.58%)	11 / 3248 (0.34%)
occurrences causally related to treatment / all	0 / 19	0 / 11
deaths causally related to treatment / all	0 / 0	0 / 0
Lower respiratory tract infection		
subjects affected / exposed	0 / 3256 (0.00%)	2 / 3248 (0.06%)
occurrences causally related to treatment / all	0 / 0	0 / 3
deaths causally related to treatment / all	0 / 0	0 / 0
Lymphangitis		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Mastoiditis		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Oesophageal candidiasis		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Ophthalmic herpes zoster		

subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Orchitis		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Osteomyelitis		
subjects affected / exposed	11 / 3256 (0.34%)	12 / 3248 (0.37%)
occurrences causally related to treatment / all	0 / 15	0 / 14
deaths causally related to treatment / all	0 / 0	0 / 0
Osteomyelitis chronic		
subjects affected / exposed	3 / 3256 (0.09%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 3	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Otitis media		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Perinephric abscess		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 2	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Periodontitis		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Peritonitis		
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Pharyngitis		

subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Pneumonia		
subjects affected / exposed	40 / 3256 (1.23%)	45 / 3248 (1.39%)
occurrences causally related to treatment / all	0 / 49	0 / 50
deaths causally related to treatment / all	0 / 5	0 / 4
Postoperative wound infection		
subjects affected / exposed	27 / 3256 (0.83%)	25 / 3248 (0.77%)
occurrences causally related to treatment / all	0 / 28	0 / 26
deaths causally related to treatment / all	0 / 0	0 / 0
Pseudomembranous colitis		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Pulmonary tuberculosis		
subjects affected / exposed	2 / 3256 (0.06%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 2	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Pyelonephritis		
subjects affected / exposed	1 / 3256 (0.03%)	3 / 3248 (0.09%)
occurrences causally related to treatment / all	0 / 1	0 / 3
deaths causally related to treatment / all	0 / 0	0 / 0
Pyelonephritis acute		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Sepsis		
subjects affected / exposed	12 / 3256 (0.37%)	13 / 3248 (0.40%)
occurrences causally related to treatment / all	0 / 12	0 / 13
deaths causally related to treatment / all	0 / 3	0 / 4
Septic shock		

subjects affected / exposed	1 / 3256 (0.03%)	5 / 3248 (0.15%)
occurrences causally related to treatment / all	0 / 1	0 / 5
deaths causally related to treatment / all	0 / 1	0 / 4
Sinusitis		
subjects affected / exposed	0 / 3256 (0.00%)	2 / 3248 (0.06%)
occurrences causally related to treatment / all	0 / 0	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0
Skin infection		
subjects affected / exposed	2 / 3256 (0.06%)	2 / 3248 (0.06%)
occurrences causally related to treatment / all	0 / 2	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0
Subcutaneous abscess		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Tonsillitis		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Tuberculosis		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Urinary tract infection		
subjects affected / exposed	13 / 3256 (0.40%)	11 / 3248 (0.34%)
occurrences causally related to treatment / all	0 / 17	0 / 13
deaths causally related to treatment / all	0 / 0	0 / 0
Wound infection		
subjects affected / exposed	17 / 3256 (0.52%)	15 / 3248 (0.46%)
occurrences causally related to treatment / all	0 / 19	0 / 16
deaths causally related to treatment / all	0 / 0	0 / 0
Urosepsis		

subjects affected / exposed	0 / 3256 (0.00%)	5 / 3248 (0.15%)
occurrences causally related to treatment / all	0 / 0	0 / 5
deaths causally related to treatment / all	0 / 0	0 / 0
Streptococcal sepsis		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 1
Scrotal abscess		
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Groin abscess		
subjects affected / exposed	3 / 3256 (0.09%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 3	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Abscess limb		
subjects affected / exposed	6 / 3256 (0.18%)	5 / 3248 (0.15%)
occurrences causally related to treatment / all	0 / 8	0 / 5
deaths causally related to treatment / all	0 / 0	0 / 0
Groin infection		
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 1	0 / 0
Lymph gland infection		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Periumbilical abscess		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Burn infection		

subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 2	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Haematoma infection		
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Pulmonary sepsis		
subjects affected / exposed	1 / 3256 (0.03%)	2 / 3248 (0.06%)
occurrences causally related to treatment / all	0 / 1	0 / 2
deaths causally related to treatment / all	0 / 1	0 / 1
Perineal abscess		
subjects affected / exposed	2 / 3256 (0.06%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 2	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Arthritis bacterial		
subjects affected / exposed	2 / 3256 (0.06%)	2 / 3248 (0.06%)
occurrences causally related to treatment / all	0 / 2	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0
Abdominal infection		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Graft infection		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Gastric infection		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Infective exacerbation of chronic obstructive airways disease		

subjects affected / exposed	3 / 3256 (0.09%)	2 / 3248 (0.06%)
occurrences causally related to treatment / all	0 / 4	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0
Infected lymphocele		
subjects affected / exposed	2 / 3256 (0.06%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 2	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Intervertebral discitis		
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Diabetic foot infection		
subjects affected / exposed	1 / 3256 (0.03%)	6 / 3248 (0.18%)
occurrences causally related to treatment / all	0 / 3	0 / 6
deaths causally related to treatment / all	0 / 0	0 / 0
Pneumonia bacterial		
subjects affected / exposed	2 / 3256 (0.06%)	3 / 3248 (0.09%)
occurrences causally related to treatment / all	0 / 2	0 / 3
deaths causally related to treatment / all	0 / 0	0 / 0
Arthritis infective		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Extradural abscess		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 1	0 / 0
Soft tissue infection		
subjects affected / exposed	2 / 3256 (0.06%)	3 / 3248 (0.09%)
occurrences causally related to treatment / all	0 / 2	0 / 3
deaths causally related to treatment / all	0 / 1	0 / 0
Respiratory tract infection		

subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Cholecystitis infective		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Device related infection		
subjects affected / exposed	2 / 3256 (0.06%)	2 / 3248 (0.06%)
occurrences causally related to treatment / all	0 / 2	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0
Cystitis bacterial		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Osteomyelitis bacterial		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Dermo-hypodermatitis		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 1	0 / 0
Post procedural sepsis		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Superinfection bacterial		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Post procedural infection		

subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Bone abscess		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Infectious pleural effusion		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Pneumocystis jirovecii pneumonia		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Infected seroma		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Medical device site infection		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 2	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Medical device site joint infection		
subjects affected / exposed	0 / 3256 (0.00%)	3 / 3248 (0.09%)
occurrences causally related to treatment / all	0 / 0	0 / 4
deaths causally related to treatment / all	0 / 0	0 / 0
Perineal cellulitis		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Vascular device infection		

subjects affected / exposed	2 / 3256 (0.06%)	2 / 3248 (0.06%)	
occurrences causally related to treatment / all	0 / 3	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Complicated appendicitis			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Large intestine infection			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Metabolism and nutrition disorders			
Dehydration			
subjects affected / exposed	6 / 3256 (0.18%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 6	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Diabetes mellitus			
subjects affected / exposed	6 / 3256 (0.18%)	8 / 3248 (0.25%)	
occurrences causally related to treatment / all	0 / 6	0 / 9	
deaths causally related to treatment / all	0 / 0	0 / 0	
Diabetes mellitus inadequate control			
subjects affected / exposed	2 / 3256 (0.06%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Diabetic ketoacidosis			
subjects affected / exposed	2 / 3256 (0.06%)	0 / 3248 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hyperglycaemia			
subjects affected / exposed	1 / 3256 (0.03%)	3 / 3248 (0.09%)	
occurrences causally related to treatment / all	0 / 1	0 / 4	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hyperkalaemia			

subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Hypoglycaemia		
subjects affected / exposed	3 / 3256 (0.09%)	6 / 3248 (0.18%)
occurrences causally related to treatment / all	0 / 3	0 / 7
deaths causally related to treatment / all	0 / 0	0 / 0
Hypokalaemia		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Hyponatraemia		
subjects affected / exposed	3 / 3256 (0.09%)	5 / 3248 (0.15%)
occurrences causally related to treatment / all	0 / 3	0 / 8
deaths causally related to treatment / all	0 / 0	0 / 0
Ketoacidosis		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0
Lactic acidosis		
subjects affected / exposed	1 / 3256 (0.03%)	0 / 3248 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 1	0 / 0
Metabolic disorder		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Decreased appetite		
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)
occurrences causally related to treatment / all	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0
Hyperglycaemic hyperosmolar nonketotic syndrome		

subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hypophagia			
subjects affected / exposed	0 / 3256 (0.00%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 1	
Type 1 diabetes mellitus			
subjects affected / exposed	0 / 3256 (0.00%)	2 / 3248 (0.06%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Type 2 diabetes mellitus			
subjects affected / exposed	3 / 3256 (0.09%)	5 / 3248 (0.15%)	
occurrences causally related to treatment / all	0 / 3	0 / 6	
deaths causally related to treatment / all	0 / 0	0 / 0	
Diabetic metabolic decompensation			
subjects affected / exposed	1 / 3256 (0.03%)	1 / 3248 (0.03%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	

Frequency threshold for reporting non-serious adverse events: 0.1 %

Non-serious adverse events	Rivaroxaban 2.5 mg bid + ASA 100 mg od	Rivaroxaban placebo bid + ASA 100 mg od	
Total subjects affected by non-serious adverse events			
subjects affected / exposed	725 / 3256 (22.27%)	729 / 3248 (22.44%)	
Neoplasms benign, malignant and unspecified (incl cysts and polyps)			
Prostate cancer			
subjects affected / exposed	1 / 3256 (0.03%)	4 / 3248 (0.12%)	
occurrences (all)	1	4	
Vascular disorders			
Hypertension			
subjects affected / exposed	49 / 3256 (1.50%)	50 / 3248 (1.54%)	
occurrences (all)	54	56	

Hypotension			
subjects affected / exposed	7 / 3256 (0.21%)	3 / 3248 (0.09%)	
occurrences (all)	7	3	
Hypertensive crisis			
subjects affected / exposed	11 / 3256 (0.34%)	13 / 3248 (0.40%)	
occurrences (all)	13	15	
Peripheral ischaemia			
subjects affected / exposed	3 / 3256 (0.09%)	11 / 3248 (0.34%)	
occurrences (all)	3	14	
Intermittent claudication			
subjects affected / exposed	39 / 3256 (1.20%)	43 / 3248 (1.32%)	
occurrences (all)	48	48	
Peripheral vascular disorder			
subjects affected / exposed	6 / 3256 (0.18%)	5 / 3248 (0.15%)	
occurrences (all)	8	5	
Dry gangrene			
subjects affected / exposed	1 / 3256 (0.03%)	4 / 3248 (0.12%)	
occurrences (all)	1	4	
Peripheral artery occlusion			
subjects affected / exposed	6 / 3256 (0.18%)	12 / 3248 (0.37%)	
occurrences (all)	6	13	
Extremity necrosis			
subjects affected / exposed	6 / 3256 (0.18%)	5 / 3248 (0.15%)	
occurrences (all)	7	5	
Peripheral arterial occlusive disease			
subjects affected / exposed	34 / 3256 (1.04%)	31 / 3248 (0.95%)	
occurrences (all)	45	37	
Peripheral artery stenosis			
subjects affected / exposed	13 / 3256 (0.40%)	10 / 3248 (0.31%)	
occurrences (all)	16	13	
Surgical and medical procedures			
Tooth extraction			
subjects affected / exposed	6 / 3256 (0.18%)	5 / 3248 (0.15%)	
occurrences (all)	7	5	
Cataract operation			

subjects affected / exposed occurrences (all)	5 / 3256 (0.15%) 6	8 / 3248 (0.25%) 10	
General disorders and administration site conditions			
Asthenia			
subjects affected / exposed	6 / 3256 (0.18%)	4 / 3248 (0.12%)	
occurrences (all)	6	4	
Chest discomfort			
subjects affected / exposed	6 / 3256 (0.18%)	1 / 3248 (0.03%)	
occurrences (all)	6	1	
Chest pain			
subjects affected / exposed	11 / 3256 (0.34%)	16 / 3248 (0.49%)	
occurrences (all)	12	16	
Fatigue			
subjects affected / exposed	11 / 3256 (0.34%)	12 / 3248 (0.37%)	
occurrences (all)	11	14	
Gait disturbance			
subjects affected / exposed	5 / 3256 (0.15%)	2 / 3248 (0.06%)	
occurrences (all)	5	2	
Impaired healing			
subjects affected / exposed	4 / 3256 (0.12%)	2 / 3248 (0.06%)	
occurrences (all)	4	2	
Oedema peripheral			
subjects affected / exposed	16 / 3256 (0.49%)	20 / 3248 (0.62%)	
occurrences (all)	17	21	
Peripheral swelling			
subjects affected / exposed	9 / 3256 (0.28%)	10 / 3248 (0.31%)	
occurrences (all)	10	10	
Non-cardiac chest pain			
subjects affected / exposed	5 / 3256 (0.15%)	3 / 3248 (0.09%)	
occurrences (all)	5	3	
Vascular stent stenosis			
subjects affected / exposed	9 / 3256 (0.28%)	10 / 3248 (0.31%)	
occurrences (all)	10	10	
Immune system disorders			

Hypersensitivity subjects affected / exposed occurrences (all)	1 / 3256 (0.03%) 1	5 / 3248 (0.15%) 5	
Reproductive system and breast disorders Benign prostatic hyperplasia subjects affected / exposed occurrences (all)	5 / 3256 (0.15%) 5	9 / 3248 (0.28%) 9	
Respiratory, thoracic and mediastinal disorders Chronic obstructive pulmonary disease subjects affected / exposed occurrences (all) Cough subjects affected / exposed occurrences (all) Dyspnoea subjects affected / exposed occurrences (all) Dyspnoea exertional subjects affected / exposed occurrences (all) Epistaxis subjects affected / exposed occurrences (all)	6 / 3256 (0.18%) 6 8 / 3256 (0.25%) 8 11 / 3256 (0.34%) 11 3 / 3256 (0.09%) 3 9 / 3256 (0.28%) 14	4 / 3248 (0.12%) 5 14 / 3248 (0.43%) 14 15 / 3248 (0.46%) 15 5 / 3248 (0.15%) 5 3 / 3248 (0.09%) 3	
Psychiatric disorders Delirium subjects affected / exposed occurrences (all) Depression subjects affected / exposed occurrences (all)	0 / 3256 (0.00%) 0 11 / 3256 (0.34%) 12	4 / 3248 (0.12%) 4 14 / 3248 (0.43%) 14	
Investigations Blood pressure increased subjects affected / exposed occurrences (all) Haemoglobin decreased	4 / 3256 (0.12%) 5	6 / 3248 (0.18%) 6	

subjects affected / exposed occurrences (all)	5 / 3256 (0.15%) 5	2 / 3248 (0.06%) 2	
Injury, poisoning and procedural complications			
Fall			
subjects affected / exposed occurrences (all)	6 / 3256 (0.18%) 6	6 / 3248 (0.18%) 6	
Foot fracture			
subjects affected / exposed occurrences (all)	5 / 3256 (0.15%) 5	2 / 3248 (0.06%) 2	
Ligament sprain			
subjects affected / exposed occurrences (all)	0 / 3256 (0.00%) 0	5 / 3248 (0.15%) 5	
Rib fracture			
subjects affected / exposed occurrences (all)	2 / 3256 (0.06%) 2	6 / 3248 (0.18%) 6	
Muscle strain			
subjects affected / exposed occurrences (all)	4 / 3256 (0.12%) 4	3 / 3248 (0.09%) 3	
Contusion			
subjects affected / exposed occurrences (all)	9 / 3256 (0.28%) 11	10 / 3248 (0.31%) 11	
Limb injury			
subjects affected / exposed occurrences (all)	11 / 3256 (0.34%) 11	11 / 3248 (0.34%) 11	
Peripheral artery restenosis			
subjects affected / exposed occurrences (all)	5 / 3256 (0.15%) 8	6 / 3248 (0.18%) 8	
Meniscus injury			
subjects affected / exposed occurrences (all)	2 / 3256 (0.06%) 2	4 / 3248 (0.12%) 4	
Arterial bypass occlusion			
subjects affected / exposed occurrences (all)	2 / 3256 (0.06%) 2	5 / 3248 (0.15%) 5	
Cardiac disorders			

Angina pectoris			
subjects affected / exposed	9 / 3256 (0.28%)	7 / 3248 (0.22%)	
occurrences (all)	10	7	
Atrial fibrillation			
subjects affected / exposed	31 / 3256 (0.95%)	48 / 3248 (1.48%)	
occurrences (all)	33	49	
Atrial flutter			
subjects affected / exposed	7 / 3256 (0.21%)	2 / 3248 (0.06%)	
occurrences (all)	7	2	
Cardiac failure			
subjects affected / exposed	3 / 3256 (0.09%)	7 / 3248 (0.22%)	
occurrences (all)	4	7	
Coronary artery disease			
subjects affected / exposed	12 / 3256 (0.37%)	9 / 3248 (0.28%)	
occurrences (all)	13	9	
Myocardial ischaemia			
subjects affected / exposed	6 / 3256 (0.18%)	1 / 3248 (0.03%)	
occurrences (all)	6	1	
Palpitations			
subjects affected / exposed	3 / 3256 (0.09%)	4 / 3248 (0.12%)	
occurrences (all)	3	4	
Sinus tachycardia			
subjects affected / exposed	4 / 3256 (0.12%)	0 / 3248 (0.00%)	
occurrences (all)	4	0	
Nervous system disorders			
Carotid artery stenosis			
subjects affected / exposed	3 / 3256 (0.09%)	8 / 3248 (0.25%)	
occurrences (all)	3	10	
Carpal tunnel syndrome			
subjects affected / exposed	2 / 3256 (0.06%)	6 / 3248 (0.18%)	
occurrences (all)	2	6	
Diabetic neuropathy			
subjects affected / exposed	3 / 3256 (0.09%)	4 / 3248 (0.12%)	
occurrences (all)	3	4	
Dizziness			

subjects affected / exposed occurrences (all)	23 / 3256 (0.71%) 27	26 / 3248 (0.80%) 28	
Headache subjects affected / exposed occurrences (all)	18 / 3256 (0.55%) 20	19 / 3248 (0.58%) 21	
Hypoaesthesia subjects affected / exposed occurrences (all)	6 / 3256 (0.18%) 7	8 / 3248 (0.25%) 9	
Neuropathy peripheral subjects affected / exposed occurrences (all)	5 / 3256 (0.15%) 5	1 / 3248 (0.03%) 1	
Paraesthesia subjects affected / exposed occurrences (all)	4 / 3256 (0.12%) 4	9 / 3248 (0.28%) 9	
Sciatica subjects affected / exposed occurrences (all)	6 / 3256 (0.18%) 6	13 / 3248 (0.40%) 14	
Syncope subjects affected / exposed occurrences (all)	3 / 3256 (0.09%) 3	9 / 3248 (0.28%) 10	
Blood and lymphatic system disorders			
Anaemia subjects affected / exposed occurrences (all)	29 / 3256 (0.89%) 29	23 / 3248 (0.71%) 23	
Iron deficiency anaemia subjects affected / exposed occurrences (all)	8 / 3256 (0.25%) 9	3 / 3248 (0.09%) 3	
Ear and labyrinth disorders			
Tinnitus subjects affected / exposed occurrences (all)	4 / 3256 (0.12%) 5	3 / 3248 (0.09%) 3	
Vertigo subjects affected / exposed occurrences (all)	7 / 3256 (0.21%) 7	11 / 3248 (0.34%) 13	
Eye disorders			

Cataract			
subjects affected / exposed	14 / 3256 (0.43%)	9 / 3248 (0.28%)	
occurrences (all)	17	9	
Diabetic retinopathy			
subjects affected / exposed	4 / 3256 (0.12%)	0 / 3248 (0.00%)	
occurrences (all)	4	0	
Gastrointestinal disorders			
Abdominal discomfort			
subjects affected / exposed	6 / 3256 (0.18%)	3 / 3248 (0.09%)	
occurrences (all)	7	3	
Abdominal pain			
subjects affected / exposed	6 / 3256 (0.18%)	6 / 3248 (0.18%)	
occurrences (all)	6	6	
Abdominal pain upper			
subjects affected / exposed	19 / 3256 (0.58%)	15 / 3248 (0.46%)	
occurrences (all)	19	16	
Constipation			
subjects affected / exposed	11 / 3256 (0.34%)	13 / 3248 (0.40%)	
occurrences (all)	11	13	
Diarrhoea			
subjects affected / exposed	23 / 3256 (0.71%)	22 / 3248 (0.68%)	
occurrences (all)	24	22	
Diverticulum intestinal			
subjects affected / exposed	1 / 3256 (0.03%)	4 / 3248 (0.12%)	
occurrences (all)	1	4	
Duodenal ulcer			
subjects affected / exposed	4 / 3256 (0.12%)	0 / 3248 (0.00%)	
occurrences (all)	4	0	
Dyspepsia			
subjects affected / exposed	15 / 3256 (0.46%)	8 / 3248 (0.25%)	
occurrences (all)	16	8	
Gastric ulcer			
subjects affected / exposed	4 / 3256 (0.12%)	5 / 3248 (0.15%)	
occurrences (all)	4	5	
Gastritis			

subjects affected / exposed occurrences (all)	15 / 3256 (0.46%) 15	8 / 3248 (0.25%) 8	
Gastritis erosive subjects affected / exposed occurrences (all)	2 / 3256 (0.06%) 2	5 / 3248 (0.15%) 5	
Gastroesophageal reflux disease subjects affected / exposed occurrences (all)	8 / 3256 (0.25%) 8	8 / 3248 (0.25%) 9	
Hiatus hernia subjects affected / exposed occurrences (all)	4 / 3256 (0.12%) 4	2 / 3248 (0.06%) 2	
Inguinal hernia subjects affected / exposed occurrences (all)	1 / 3256 (0.03%) 1	5 / 3248 (0.15%) 5	
Nausea subjects affected / exposed occurrences (all)	14 / 3256 (0.43%) 14	10 / 3248 (0.31%) 10	
Vomiting subjects affected / exposed occurrences (all)	7 / 3256 (0.21%) 7	13 / 3248 (0.40%) 13	
Large intestine polyp subjects affected / exposed occurrences (all)	6 / 3256 (0.18%) 6	4 / 3248 (0.12%) 4	
Hepatobiliary disorders Hepatic steatosis subjects affected / exposed occurrences (all)	4 / 3256 (0.12%) 4	2 / 3248 (0.06%) 2	
Skin and subcutaneous tissue disorders Dermatitis allergic subjects affected / exposed occurrences (all)	1 / 3256 (0.03%) 1	4 / 3248 (0.12%) 4	
Ecchymosis subjects affected / exposed occurrences (all)	5 / 3256 (0.15%) 5	2 / 3248 (0.06%) 2	
Eczema			

subjects affected / exposed occurrences (all)	2 / 3256 (0.06%) 2	4 / 3248 (0.12%) 4	
Erythema subjects affected / exposed occurrences (all)	2 / 3256 (0.06%) 2	5 / 3248 (0.15%) 5	
Pruritus subjects affected / exposed occurrences (all)	10 / 3256 (0.31%) 11	7 / 3248 (0.22%) 7	
Rash subjects affected / exposed occurrences (all)	16 / 3256 (0.49%) 16	18 / 3248 (0.55%) 20	
Skin ulcer subjects affected / exposed occurrences (all)	16 / 3256 (0.49%) 17	18 / 3248 (0.55%) 18	
Urticaria subjects affected / exposed occurrences (all)	7 / 3256 (0.21%) 7	5 / 3248 (0.15%) 5	
Diabetic foot subjects affected / exposed occurrences (all)	6 / 3256 (0.18%) 6	1 / 3248 (0.03%) 1	
Renal and urinary disorders			
Dysuria subjects affected / exposed occurrences (all)	3 / 3256 (0.09%) 3	4 / 3248 (0.12%) 4	
Haematuria subjects affected / exposed occurrences (all)	4 / 3256 (0.12%) 4	2 / 3248 (0.06%) 2	
Renal failure subjects affected / exposed occurrences (all)	2 / 3256 (0.06%) 2	5 / 3248 (0.15%) 6	
Renal impairment subjects affected / exposed occurrences (all)	4 / 3256 (0.12%) 5	6 / 3248 (0.18%) 6	
Chronic kidney disease subjects affected / exposed occurrences (all)	8 / 3256 (0.25%) 8	1 / 3248 (0.03%) 1	

Acute kidney injury subjects affected / exposed occurrences (all)	7 / 3256 (0.21%) 8	4 / 3248 (0.12%) 4	
Musculoskeletal and connective tissue disorders			
Arthralgia subjects affected / exposed occurrences (all)	18 / 3256 (0.55%) 19	32 / 3248 (0.99%) 37	
Bursitis subjects affected / exposed occurrences (all)	5 / 3256 (0.15%) 6	7 / 3248 (0.22%) 7	
Back pain subjects affected / exposed occurrences (all)	23 / 3256 (0.71%) 25	30 / 3248 (0.92%) 33	
Joint swelling subjects affected / exposed occurrences (all)	6 / 3256 (0.18%) 6	2 / 3248 (0.06%) 2	
Lumbar spinal stenosis subjects affected / exposed occurrences (all)	5 / 3256 (0.15%) 6	2 / 3248 (0.06%) 4	
Muscle spasms subjects affected / exposed occurrences (all)	13 / 3256 (0.40%) 14	13 / 3248 (0.40%) 16	
Muscular weakness subjects affected / exposed occurrences (all)	4 / 3256 (0.12%) 4	5 / 3248 (0.15%) 6	
Musculoskeletal pain subjects affected / exposed occurrences (all)	10 / 3256 (0.31%) 10	11 / 3248 (0.34%) 11	
Myalgia subjects affected / exposed occurrences (all)	6 / 3256 (0.18%) 6	4 / 3248 (0.12%) 4	
Osteoarthritis subjects affected / exposed occurrences (all)	12 / 3256 (0.37%) 12	14 / 3248 (0.43%) 14	
Pain in extremity			

subjects affected / exposed	47 / 3256 (1.44%)	52 / 3248 (1.60%)
occurrences (all)	58	53
Rotator cuff syndrome		
subjects affected / exposed	4 / 3256 (0.12%)	2 / 3248 (0.06%)
occurrences (all)	4	3
Spinal osteoarthritis		
subjects affected / exposed	2 / 3256 (0.06%)	5 / 3248 (0.15%)
occurrences (all)	2	6
Intervertebral disc protrusion		
subjects affected / exposed	7 / 3256 (0.21%)	5 / 3248 (0.15%)
occurrences (all)	7	5
Spinal pain		
subjects affected / exposed	1 / 3256 (0.03%)	4 / 3248 (0.12%)
occurrences (all)	1	4
Spinal stenosis		
subjects affected / exposed	4 / 3256 (0.12%)	5 / 3248 (0.15%)
occurrences (all)	4	5
Infections and infestations		
Bronchitis		
subjects affected / exposed	15 / 3256 (0.46%)	17 / 3248 (0.52%)
occurrences (all)	17	21
Cellulitis		
subjects affected / exposed	10 / 3256 (0.31%)	6 / 3248 (0.18%)
occurrences (all)	11	7
Gangrene		
subjects affected / exposed	5 / 3256 (0.15%)	2 / 3248 (0.06%)
occurrences (all)	5	2
Herpes zoster		
subjects affected / exposed	5 / 3256 (0.15%)	7 / 3248 (0.22%)
occurrences (all)	5	7
Influenza		
subjects affected / exposed	10 / 3256 (0.31%)	7 / 3248 (0.22%)
occurrences (all)	11	8
Localised infection		
subjects affected / exposed	11 / 3256 (0.34%)	7 / 3248 (0.22%)
occurrences (all)	11	7

Nasopharyngitis			
subjects affected / exposed	21 / 3256 (0.64%)	30 / 3248 (0.92%)	
occurrences (all)	23	32	
Osteomyelitis			
subjects affected / exposed	4 / 3256 (0.12%)	1 / 3248 (0.03%)	
occurrences (all)	4	1	
Pneumonia			
subjects affected / exposed	9 / 3256 (0.28%)	14 / 3248 (0.43%)	
occurrences (all)	9	14	
Postoperative wound infection			
subjects affected / exposed	9 / 3256 (0.28%)	9 / 3248 (0.28%)	
occurrences (all)	9	9	
Sinusitis			
subjects affected / exposed	6 / 3256 (0.18%)	3 / 3248 (0.09%)	
occurrences (all)	6	4	
Upper respiratory tract infection			
subjects affected / exposed	4 / 3256 (0.12%)	2 / 3248 (0.06%)	
occurrences (all)	4	2	
Urinary tract infection			
subjects affected / exposed	22 / 3256 (0.68%)	21 / 3248 (0.65%)	
occurrences (all)	23	27	
Wound infection			
subjects affected / exposed	9 / 3256 (0.28%)	4 / 3248 (0.12%)	
occurrences (all)	9	4	
Respiratory tract infection viral			
subjects affected / exposed	2 / 3256 (0.06%)	4 / 3248 (0.12%)	
occurrences (all)	2	4	
Respiratory tract infection			
subjects affected / exposed	5 / 3256 (0.15%)	2 / 3248 (0.06%)	
occurrences (all)	5	3	
Metabolism and nutrition disorders			
Diabetes mellitus			
subjects affected / exposed	6 / 3256 (0.18%)	12 / 3248 (0.37%)	
occurrences (all)	6	12	
Gout			

subjects affected / exposed occurrences (all)	5 / 3256 (0.15%) 5	11 / 3248 (0.34%) 11
Hyperkalaemia subjects affected / exposed occurrences (all)	0 / 3256 (0.00%) 0	4 / 3248 (0.12%) 4
Vitamin D deficiency subjects affected / exposed occurrences (all)	5 / 3256 (0.15%) 5	1 / 3248 (0.03%) 1
Dyslipidaemia subjects affected / exposed occurrences (all)	1 / 3256 (0.03%) 1	5 / 3248 (0.15%) 5
Decreased appetite subjects affected / exposed occurrences (all)	6 / 3256 (0.18%) 6	4 / 3248 (0.12%) 4
Type 2 diabetes mellitus subjects affected / exposed occurrences (all)	13 / 3256 (0.40%) 13	7 / 3248 (0.22%) 7

More information

Substantial protocol amendments (globally)

Were there any global substantial amendments to the protocol? Yes

Date	Amendment
09 February 2016	First global amendment including 25 changes.
20 March 2017	Second global amendment including 7 changes.

Notes:

Interruptions (globally)

Were there any global interruptions to the trial? No

Limitations and caveats

None reported