Clinical trial results:
A Phase III Randomized, Double-blind, Placebo-controlled Multicenter Study in Adults to Determine the Safety, Efficacy, and Immunogenicity of AZD1222, a Non-replicating ChAdOx1 Vector Vaccine, for the Prevention of COVID-19

Summary

EudraCT number	2020-005226-28
Trial protocol	FR DE IT
Global end of trial date	21 March 2023

Results information

Result version number	v1 (current)
This version publication date	23 November 2023
First version publication date	23 November 2023

Trial information

Trial identification

Sponsor protocol code	D8110C00001

Additional study identifiers

ISRCTN number	-
ClinicalTrials.gov id (NCT number)	NCT04516746
WHO universal trial number (UTN)	-

Notes:

Sponsors

Sponsor organisation name	AstraZeneca AB
Sponsor organisation address	Karlebyhusentren, B674 Astraallen, Södertälje, Sweden, SE-151 85
Public contact	Global Clinical Lead, AstraZeneca, +1 877-240-9479, information.center@astrazeneca.com
Scientific contact	Global Clinical Lead, AstraZeneca, +1 877-240-9479, information.center@astrazeneca.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	21 March 2023
Is this the analysis of the primary completion data?	No

Global end of trial reached?	Yes
Global end of trial date	21 March 2023
Was the trial ended prematurely?	No

Notes:

General information about the trial

Main objective of the trial:

To estimate the efficacy and assess the safety, tolerability and reactogenicity (for substudy only) of 2 intramuscular (IM) doses of AZD1222 compared to saline placebo for the prevention of coronavirus disease-2019 (COVID-19).

Protection of trial subjects:

This study was performed in accordance with the ethical principles that have their origin in the Declaration of Helsinki and that are consistent with International Council for Harmonisation/Good Clinical Practice, applicable regulatory requirements, and the AstraZeneca policy on Bioethics.

Background therapy: -

Evidence for comparator: -

Actual start date of recruitment	28 August 2020
Long term follow-up planned	Yes
Long term follow-up rationale	Safety
Long term follow-up duration	2 Years
Independent data monitoring committee (IDMC) involvement?	Yes

Notes:

Population of trial subjects

Subjects enrolled per country

Country: Number of subjects enrolled	Chile: 2206
Country: Number of subjects enrolled	Peru: 1470
Country: Number of subjects enrolled	United States: 28774
Worldwide total number of subjects	32450
EEA total number of subjects	0

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)	25195
From 65 to 84 years	7136
85 years and over	119

Subject disposition

Recruitment

Recruitment details:

This Phase III randomized study was conducted in adult participants who were healthy or had medically stable chronic diseases and were at increased risk for severe acute respiratory syndrome-coronavirus-2 (SARS-CoV-2) acquisition and coronavirus disease-2019 (COVID-19). One participant was enrolled at 2 sites and excluded from all analysis sets.

Pre-assignment

Screening details:

Study had a screening period (14 days), followed by a treatment and follow-up period (up to 760 days). A total of 32450 participants were randomized in a 2:1 ratio to receive AZD1222 or placebo. The first participants randomized in each age group in the United States of America participated in a substudy to assess immunogenicity and reactogenicity.

Period 1

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

Arms

Are arms mutually exclusive?	Yes
Arm title	AZD1222

Arm description:

Participants were randomized to receive 2 IM doses of 5*10^10 viral particles (vp) (nominal, ± 1.5*10^10 vp) AZD1222 on Days 1 and 29.

Arm type	Experimental
Investigational medicinal product name	AZD1222
Investigational medicinal product code	ChAdOx1 nCoV-19
Other name	Vaxzevria
Pharmaceutical forms	Solution for injection
Routes of administration	Intramuscular use

Dosage and administration details:

AZD1222 5*10^10 vp was administered on Days 1 and 29.

Arm title	Placebo

Arm description:

Participants were randomized to receive 2 IM doses of placebo matching with AZD1222 on Days 1 and 29.

Arm type	Placebo
Investigational medicinal product name	Placebo
Investigational medicinal product code	
Other name	
Pharmaceutical forms	Solution for injection
Routes of administration	Intramuscular use

Dosage and administration details:

Placebo matching with AZD1222 was administered on Days 1 and 29.

Number of subjects in period 1	AZD1222	Placebo
Started	21634	10816
Participants who received first dose	21583	10797
Participants who received second dose	20770	9954
Completed	16099	7213
Not completed	5535	3603
Consent withdrawn by subject	1946	1696
Physician decision	10	8
Adverse event, non-fatal	-	1
Death	62	32
Randomized but not Treated	51	19
Unspecified	35	16
Lost to follow-up	3431	1831

Baseline characteristics

Reporting groups	
Reporting group title	AZD1222
Reporting group description:	
Participants were randomized to receive 2 IM doses of 5*10^10 viral particles (vp) (nominal, ± 1.5*10^10 vp) AZD1222 on Days 1 and 29.	
Reporting group title	Placebo
Reporting group description:	
Participants were randomized to receive 2 IM doses of placebo matching with AZD1222 on Days 1 and 29.	

Reporting group values	AZD1222	Placebo	Total
Number of subjects	21634	10816	32450
Age Categorical Units: Subjects			
Age Continuous Units: years			
arithmetic mean	50.2	50.2	
standard deviation	± 15.92	± 15.86	-
Gender Categorical Units: Subjects			
Female	9603	4797	14400
Male	12031	6019	18050
Ethnicity Units: Subjects			
Hispanic or Latino	4787	2456	7243
Not Hispanic or Latino	16506	8217	24723
Not reported	293	127	420
Unknown	48	16	64
Race Units: Subjects			
Multiple	513	258	771
White	17100	8534	25634
Black or African American	1798	899	2697
Asian	947	483	1430
American Indian or Alaska Native	855	430	1285
Native Hawaiian or Other Pacific Islander	60	21	81
Not reported	260	138	398
Unknown	101	53	154

End points

End points reporting groups

Reporting group title	AZD1222
Reporting group description:	
Participants were randomized to receive 2 IM doses of 5*10^10 viral particles (vp) (nominal, ± 1.5*10^10 vp) AZD1222 on Days 1 and 29.	
Reporting group title	Placebo
Reporting group description:	
Participants were randomized to receive 2 IM doses of placebo matching with AZD1222 on Days 1 and 29.	

Primary: Number of Participants With Binary Response

End point title	Number of Participants With Binary Response
End point description:	
A binary response, whereby a participant with negative serostatus at baseline is defined as a COVID-19 case if their first case of SARS-CoV-2 RT-PCR-positive symptomatic illness occurs ≥ 15 days post second dose of study intervention. Otherwise, a participant is not defined as a COVID-19 case. The primary efficacy analysis was performed once approximately 150 events meeting the primary efficacy outcome measure definition had occurred across the AZD1222 and placebo groups. The fully vaccinated analysis set (FVS) included all participants in the full analysis set (FAS) who were seronegative at baseline, received 2 doses of study intervention, and who remained on-study 15 days after their second dose without having had a prior SARS-CoV-2 RT-PCR-positive confirmed COVID-19 infection.	
End point type	Primary
End point timeframe:	
From 15 days post second dose up to data cut-off date (DCO) of 05 March 2021 or study discontinuation or unblinding or receipt of non-study COVID-19 vaccination, up to a maximum of 17 weeks	

End point values	AZD1222	Placebo		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	17662	8550		
Units: participants	73	130		

Statistical analyses

Statistical analysis title	Treatment ratio in binary response
Statistical analysis description:	
The 95% confidence interval (CI) and p-value were estimated based on Poisson regression with robust variance (including study arm and stratification factor [age group at informed consent] as covariates, and log of the follow up time as an offset).	
Comparison groups	AZD1222 v Placebo

Number of subjects included in analysis	26212
Analysis specification	Pre-specified
Analysis type	superiority
P-value	< 0.001
Method	Poisson regression with robust variance
Parameter estimate	Vaccine efficacy
Point estimate	73.98
Confidence interval	
level	95 %
sides	2-sided
lower limit	65.34
upper limit	80.47

Primary: Number of Participants With Adverse Events (AEs) Post Each Dose of Study Intervention

End point title	Number of Participants With Adverse Events (AEs) Post Each Dose of Study Intervention[1]

End point description:

An AE is the development of any untoward medical occurrence in a clinical study participant administered medicinal product and which does not necessarily have a causal relationship with this medicinal product. An AE can therefore be any unfavorable and unintended sign, symptom, or disease temporally associated with the use of a medicinal product, whether or not considered related to the medicinal product. The safety analysis set included all participants who received at least 1 dose of study intervention. Here, 'n' is number of participants analyzed at specific time point.

End point type	Primary

End point timeframe:

From Day 1 up to 28 days post second dose of study intervention, approximately 57 days

Notes:

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

End point values	AZD1222	Placebo		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	21587	10792		
Units: participants				
After first dose (n= 21587, 10792)	5736	1926		
After second dose (n= 20773, 9947)	5074	1797		
After any dose (n= 21587, 10792)	8771	3201		

Statistical analyses

No statistical analyses for this end point

Primary: Number of Participants With Serious Adverse Events (SAE), Medically Attended Adverse Events (MAAE), and Adverse Event of Special Interest (AESI) Throughout the Study

End point title	Number of Participants With Serious Adverse Events (SAE), Medically Attended Adverse Events (MAAE), and Adverse Event

	of Special Interest (AESI) Throughout the Study[2]

End point description:

An SAE is an AE occurring during any study phase that fulfils 1 or more of the following criteria: death; immediately life-threatening; in-participant hospitalization or prolongation of existing hospitalization; persistent or significant disability or incapacity; congenital abnormality or birth defect; an important medical event. AESIs were events of scientific and medical interest specific to the further understanding of the study intervention safety profile and required close monitoring and rapid communication by the investigators to the sponsor. MAAEs are defined as AEs leading to medically-attended visits that were not routine visits for physical examination or vaccination, such as an emergency room visit, or an otherwise unscheduled visit to or from medical personnel (medical doctor) for any reason. Different follow-up time between AZD1222 and Placebo groups (20223 versus 3893 participant years). The safety analysis set.

End point type	Primary

End point timeframe:

From Day 1 up to 2 years of follow-up post first dose or study discontinuation or receipt of non-study COVID-19 vaccination, a maximum of approximately 760 days

Notes:

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

End point values	AZD1222	Placebo		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	21587	10793		
Units: participants				
SAEs	621	136		
MAAEs	4750	1256		
AESIs	2516	591		

Statistical analyses

No statistical analyses for this end point

Primary: Number of Participants With Local and Systemic Solicited AEs in the Substudy Only

End point title	Number of Participants With Local and Systemic Solicited AEs in the Substudy Only[3]

End point description:

Solicited AEs are local or systemic predefined events for assessment of reactogenicity. Solicited AEs were collected in a e-Diary only for participants in the substudy. The safety analysis set included all participants who received at least 1 dose of study intervention. Only participants included in the substudy were analyzed. Here, 'n' is number of participants analyzed at specific time point.

End point type	Primary

End point timeframe:

From Day 1 up to 7 days post each dose of study intervention, approximately 14 days

Notes:

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

End point values	AZD1222	Placebo		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	2037	1013		
Units: participants				
Solicited local AEs: After 1st dose(n=1830,920)	1250	173		
Solicited local AEs: After 2nd dose(n=1829,908)	977	120		
Solicited local AEs: After any dose(n=1944,979)	1440	239		
Solicited systemic AEs: After 1st dose(n=1842,924)	1191	415		
Solicited systemic AEs: After 2nd dose(n=1821,905)	862	314		
Solicited systemic AEs: After any dose(n=1947,980)	1395	519		

Statistical analyses

No statistical analyses for this end point

Secondary: Number of Participants With First COVID-19 Symptomatic Illness Using Centers for Disease Control and Prevention (CDC) Criteria Post Second Dose of Study Intervention

End point title	Number of Participants With First COVID-19 Symptomatic Illness Using Centers for Disease Control and Prevention (CDC) Criteria Post Second Dose of Study Intervention

End point description:

The incidence of the first case of SARS-CoV-2 RT-PCR-positive symptomatic illness occurring ≥ 15 days post second dose of study intervention using CDC criteria. Participant must present with at least 1 of the following symptoms per CDC criteria: fever, shortness of breath, difficulty breathing, chills, cough, fatigue, muscle aches, body aches, headache, new loss of taste, new loss of smell, sore throat, congestion, runny nose, nausea, vomiting, or diarrhea. The FVS included all participants in the full analysis set who were seronegative at baseline, received 2 doses of study intervention, and who remained on-study 15 days after their second dose without having had a prior SARS-CoV-2 RT-PCR-positive confirmed COVID-19 infection.

End point type	Secondary

End point timeframe:

From 15 days post second dose up to DCO of 05 March 2021 or study discontinuation or unblinding or receipt of non-study COVID-19 vaccination, up to a maximum of 17 weeks

End point values	AZD1222	Placebo		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	17662	8550		
Units: participants	95	145		

Statistical analyses

Statistical analysis title	Treatment difference: COVID-19 symptomatic illness

Statistical analysis description:

The 95% CI and p-value were estimated based on Poisson regression with robust variance (including study arm and stratification factor [age group at informed consent] as covariates, and log of the follow up time as an offset).

Comparison groups	AZD1222 v Placebo
Number of subjects included in analysis	26212
Analysis specification	Pre-specified
Analysis type	superiority
P-value	< 0.001
Method	Poisson regression with robust variance
Parameter estimate	Vaccine efficacy
Point estimate	69.65
Confidence interval	
level	95 %
sides	2-sided
lower limit	60.68
upper limit	76.57

Secondary: Number of Participants With First Post-intervention Response for SARS-CoV-2 Nucleocapsid Antibodies Post Second Dose of Study

End point title	Number of Participants With First Post-intervention Response for SARS-CoV-2 Nucleocapsid Antibodies Post Second Dose of Study

End point description:

The incidence of the first post-intervention response (negative at baseline to positive post intervention with study intervention) for SARS-CoV-2 nucleocapsid antibodies occurring ≥ 15 days post second dose of study intervention (key secondary endpoint). The FVS included all participants in the FAS who were seronegative at baseline, received 2 doses of study intervention, and who remained on-study 15 days after their second dose without having had a prior SARS-CoV-2 RT-PCR-positive confirmed COVID-19 infection.

End point type	Secondary

End point timeframe:

From 15 days post second dose up to DCO of 05 March 2021 or study discontinuation or unblinding or receipt of non-study COVID-19 vaccination, up to a maximum of 17 weeks

End point values	AZD1222	Placebo		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	17662	8550		
Units: participants	156	202		

Statistical analyses

Statistical analysis title	Treatment difference: Nucleocapsid antibodies

Statistical analysis description:

The 95% CI and p-value were estimated based on Poisson regression with robust variance (including study arm and stratification factor [age group at informed consent] as covariates, and log of the follow

up time as an offset).

Comparison groups	AZD1222 v Placebo
Number of subjects included in analysis	26212
Analysis specification	Pre-specified
Analysis type	superiority
P-value	< 0.001
Method	Poisson regression with robust variance
Parameter estimate	Vaccine efficacy
Point estimate	64.32
Confidence interval	
level	95 %
sides	2-sided
lower limit	56.05
upper limit	71.03

Secondary: Number of Participants With First COVID-19 Symptomatic Illness Using University of Oxford-Defined Symptom Criteria Post Second Dose of Study Intervention

End point title	Number of Participants With First COVID-19 Symptomatic Illness Using University of Oxford-Defined Symptom Criteria Post Second Dose of Study Intervention

End point description:

The incidence of the first case of SARS-CoV-2 RT-PCR-positive symptomatic illness occurring ≥ 15 days post second dose of study intervention using University of Oxford-defined symptom criteria: new onset of fever (> 100 °Fahrenheit [> 37.8 °Celsius]), cough, shortness of breath, or anosmia/ageusia. The FVS included all participants in the FAS who were seronegative at baseline, received 2 doses of study intervention, and who remained on-study 15 days after their second dose without having had a prior SARS-CoV-2 RT-PCR-positive confirmed COVID-19 infection.

End point type	Secondary

End point timeframe:

From 15 days post second dose up to DCO of 05 March 2021 or study discontinuation or unblinding or receipt of non-study COVID-19 vaccination, up to a maximum of 17 weeks

End point values	AZD1222	Placebo		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	17662	8550		
Units: participants	86	136		

Statistical analyses

Statistical analysis title	Treatment difference: COVID-19 symptomatic illness

Statistical analysis description:

The 95% CI and p-value were estimated based on Poisson regression with robust variance (including study arm and stratification factor [age group at informed consent] as covariates, and log of the follow up time as an offset).

Comparison groups	AZD1222 v Placebo

Number of subjects included in analysis	26212
Analysis specification	Pre-specified
Analysis type	superiority
P-value	< 0.001
Method	Poisson regression with robust variance
Parameter estimate	Vaccine efficacy
Point estimate	70.7
Confidence interval	
level	95 %
sides	2-sided
lower limit	61.62
upper limit	77.64

Secondary: Number of Participants With First Symptomatic COVID-19 Regardless of Evidence of Prior SARS-CoV-2 Infection Post Second Dose of Study Intervention

End point title	Number of Participants With First Symptomatic COVID-19 Regardless of Evidence of Prior SARS-CoV-2 Infection Post Second Dose of Study Intervention

End point description:

The incidence of the first case of SARS-CoV-2 RT-PCR-positive symptomatic illness occurring ≥ 15 days post second dose of study intervention regardless of evidence of prior SARS-CoV-2 infection (key secondary endpoint). The FVS regardless of prior SARS-COV-2 infection included all participants in the FAS who were seronegative at baseline, received 2 doses of study intervention, and who remained on-study 15 days after their second dose.

End point type	Secondary

End point timeframe:

From 15 days post second dose up to DCO of 05 March 2021 or study discontinuation or unblinding or receipt of non-study COVID-19 vaccination, up to a maximum of 17 weeks

End point values	AZD1222	Placebo		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	18563	9031		
Units: participants	76	135		

Statistical analyses

Statistical analysis title	Treatment difference: First symptomatic COVID-19

Statistical analysis description:

The 95% CI and p-value were estimated based on Poisson regression with robust variance (including study arm and stratification factor [age group at informed consent] as covariates, and log of the follow up time as an offset).

Comparison groups	AZD1222 v Placebo

Number of subjects included in analysis	27594
Analysis specification	Pre-specified
Analysis type	superiority
P-value	< 0.001
Method	Poisson regression with robust variance
Parameter estimate	Vaccine efficacy
Point estimate	73.68
Confidence interval	
level	95 %
sides	2-sided
lower limit	65.13
upper limit	80.13

Secondary: Number of Participants With COVID-19 Severe or Critical Symptomatic Illness Post Second Dose of Study Intervention

End point title	Number of Participants With COVID-19 Severe or Critical Symptomatic Illness Post Second Dose of Study Intervention

End point description:

The incidence of SARS-CoV-2 RT-PCR-positive severe or critical symptomatic illness occurring ≥ 15 days post second dose of study intervention. The severity of COVID-19 was evaluated in participants with symptoms of COVID-19. Following are the findings regarding severe of critical symptomatic COVID-19: clinical signs at rest indicative of severe systemic illness; respiratory failure; evidence of shock; significant acute renal, hepatic, or neurologic dysfunction; admission to an intensive care unit; and death (key secondary endpoint). The FVS included all participants in the FAS who were seronegative at baseline, received 2 doses of study intervention, and who remained on-study 15 days after their second dose without having had a prior SARS-CoV-2 RT-PCR-positive confirmed COVID-19 infection.

End point type	Secondary

End point timeframe:

From 15 days post second dose up to DCO of 05 March 2021 or study discontinuation or unblinding or receipt of non-study COVID-19 vaccination, up to a maximum of 17 weeks

End point values	AZD1222	Placebo		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	17662	8550		
Units: participants	0	8		

Statistical analyses

Statistical analysis title	Treatment difference: COVID-19 symptomatic illness

Statistical analysis description:

The exact 1-sided 97.5% CI and p-value were estimated based on stratified Poisson regression with exact conditional method (including study arm and stratification factor [age group at informed consent] as strata factor and log of total number of participants for each combination of study arm and strata as an offset).

Comparison groups	AZD1222 v Placebo

Number of subjects included in analysis	26212
Analysis specification	Pre-specified
Analysis type	superiority
P-value	< 0.001
Method	Poisson regression exact conditional
Parameter estimate	Vaccine efficacy
Point estimate	100
Confidence interval	
level	Other: 97.5 %
sides	1-sided
lower limit	71.62

Secondary: Number of Participants With COVID-19-Related Emergency Department Visits Post Second Dose of Study Intervention

End point title	Number of Participants With COVID-19-Related Emergency Department Visits Post Second Dose of Study Intervention

End point description:

The incidence of COVID-19-related emergency department visits occurring ≥ 15 days post second dose of study intervention (key secondary endpoint). The FVS included all participants in the FAS who were seronegative at baseline, received 2 doses of study intervention, and who remained on-study 15 days after their second dose without having had a prior SARS-CoV-2 RT-PCR-positive confirmed COVID-19 infection.

End point type	Secondary

End point timeframe:

From 15 days post second dose up to DCO of 05 March 2021 or study discontinuation or unblinding or receipt of non-study COVID-19 vaccination, up to a maximum of 17 weeks

End point values	AZD1222	Placebo		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	17662	8550		
Units: participants	1	9		

Statistical analyses

Statistical analysis title	Treatment difference: Emergency department visits

Statistical analysis description:

The 95% CI and p-value were estimated based on Poisson regression with robust variance (including study arm and stratification factor [age group at informed consent] as covariates, and log of the follow up time as an offset).

Comparison groups	AZD1222 v Placebo

Number of subjects included in analysis	26212
Analysis specification	Pre-specified
Analysis type	superiority
P-value	= 0.005
Method	Poisson regression with robust variance
Parameter estimate	Vaccine efficacy
Point estimate	94.8
Confidence interval	
level	95 %
sides	2-sided
lower limit	58.98
upper limit	99.34

Secondary: Number of Participants With COVID-19 Severe or Critical Symptomatic Illness Post First Dose of Study Intervention

End point title	Number of Participants With COVID-19 Severe or Critical Symptomatic Illness Post First Dose of Study Intervention

End point description:

The incidence of SARS-CoV-2 RT-PCR-positive severe or critical symptomatic illness occurring post first dose of study intervention. The severity of COVID-19 was evaluated in participants with symptoms of COVID-19. Following are the findings regarding severe of critical symptomatic COVID-19: clinical signs at rest indicative of severe systemic illness; respiratory failure; evidence of shock; significant acute renal, hepatic, or neurologic dysfunction; admission to an intensive care unit; and death. The FAS included all randomized participants who received at least 1 dose of study intervention, irrespective of their protocol adherence and continued participation in the study. Only participants who are seronegative at baseline were analyzed.

End point type	Secondary

End point timeframe:

From 15 days post second dose up to DCO of 05 March 2021 or study discontinuation or unblinding or receipt of non-study COVID-19 vaccination, up to a maximum of 17 weeks

End point values	AZD1222	Placebo		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	20589	10300		
Units: participants	5	16		

Statistical analyses

Statistical analysis title	Treatment difference: COVID-19 symptomatic illness

Statistical analysis description:

The 95% CI were estimated based on Poisson regression with robust variance (including study arm and age group at screening (18-65 years, ≥ 65 years) as covariates and log of the follow-up time as an offset).

Comparison groups	AZD1222 v Placebo

Number of subjects included in analysis	30889
Analysis specification	Pre-specified
Analysis type	superiority
P-value	< 0.001
Method	Poisson regression with robust variance
Parameter estimate	Vaccine efficacy
Point estimate	84.97
Confidence interval	
level	95 %
sides	2-sided
lower limit	58.97
upper limit	94.5

Secondary: Geometric Mean Titers (GMTs) for SARS-CoV-2 Spike (S) and Receptor Binding Domain (RBD) Antibodies as Measured by Meso Scale Discovery (MSD) Serology Assay

End point title	Geometric Mean Titers (GMTs) for SARS-CoV-2 Spike (S) and Receptor Binding Domain (RBD) Antibodies as Measured by Meso Scale Discovery (MSD) Serology Assay

End point description:

The GMT was calculated as the antilogarithm of Σ(log base 2 transformed titer/n), i.e. as the anti-logarithm transformation of the mean of the log-transformed titer, where 'n' is the number of participants with titer information. The immunogenicity analysis population included all participants in the safety analysis set who had no protocol deviations judged to have the potential to interfere with the generation or interpretation of an immune response. Only participants included in the substudy were analyzed. Here, 'n' is number of participants analyzed at specific time point.

End point type	Secondary

End point timeframe:

Baseline (Day 1) and Days 15, 29, 43, 57, 90, 180, 360, and 730

End point values	AZD1222	Placebo		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	2008	1002		
Units: arbitrary units per milliliter (AU/mL)				
geometric mean (confidence interval 95%)				
S Antibody Titer: Day 1 (n=2008, 1002)	53.18 (50.37 to 56.14)	54.68 (50.10 to 59.69)		
S Antibody Titer: Day 15 (n=1897, 949)	1820.10 (1696.48 to 1952.72)	53.47 (48.88 to 58.50)		
S Antibody Titer: Day 29 (n=1428, 708)	5782.09 (5389.88 to 6202.84)	53.64 (47.98 to 59.97)		
S Antibody Titer: Day 43 (n=1809, 899)	24105.87 (22945.04 to 25325.44)	58.15 (52.68 to 64.18)		
S Antibody Titer: Day 57 (n=1760, 829)	19488.64 (18521.81 to 20505.92)	58.30 (52.35 to 64.94)		

S Antibody Titer: Day 90 (n=1675, 643)	14583.30 (13839.22 to 15367.38)	87.51 (74.78 to 102.42)		
S Antibody Titer: Day 180 (n=1217, 147)	7483.04 (6911.35 to 8102.02)	266.47 (161.19 to 440.49)		
S Antibody Titer: Day 360 (n=874, 47)	6686.81 (5779.74 to 7736.24)	1268.45 (489.80 to 3284.98)		
S Antibody Titer: Day 730 (n=276, 22)	186727.78 (154395.93 to 225830.20)	19093.94 (5685.04 to 64129.44)		
RBD Antibody Titer: Day 1 (n=2008, 1002)	133.6 (129.1 to 138.4)	138.9 (130.9 to 147.3)		
RBD Antibody Titer: Day 15 (n=1897, 949)	965.0 (897.0 to 1038.1)	138.6 (130.5 to 147.1)		
RBD Antibody Titer: Day 29 (n=1427, 708)	5176.6 (4794.8 to 5588.9)	143.0 (132.4 to 154.3)		
RBD Antibody Titer: Day 43 (n=1809, 899)	29351.9 (27875.6 to 30906.4)	145.7 (136.0 to 156.1)		
RBD Antibody Titer: Day 57 (n=1760, 829)	23840.6 (22614.3 to 25133.3)	151.6 (140.3 to 163.8)		
RBD Antibody Titer: Day 90 (n=1675, 643)	17499.9 (16559.1 to 18494.1)	192.6 (171.0 to 217.0)		
RBD Antibody Titer: Day 180 (n=1217, 147)	8333.1 (7647.9 to 9079.7)	481.6 (314.8 to 736.7)		
RBD Antibody Titer: Day 360 (n=874, 47)	7499.2 (6417.7 to 8763.1)	1912.3 (839.0 to 4358.5)		
RBD Antibody Titer: Day 730 (n=276, 22)	276381.8 (227665.9 to 335521.9)	19319.4 (5290.3 to 70551.9)		

Statistical analyses

No statistical analyses for this end point

Secondary: Geometric Mean Fold Rise (GMFR) for SARS-CoV-2 S and RBD Antibodies as Measured by MSD Serology Assay

End point title	Geometric Mean Fold Rise (GMFR) for SARS-CoV-2 S and RBD Antibodies as Measured by MSD Serology Assay

End point description:

The fold rise was calculated as the ratio of the post-vaccination titer level to the pre-vaccination titer level. GMFR was calculated as anti-logarithm of Σ (log base 2 transformed (post-vaccination titer/ pre-vaccination titer)/n). Where 'n' is the number of participants with titer information. The immunogenicity analysis population included all participants in the safety analysis set who had no protocol deviations judged to have the potential to interfere with the generation or interpretation of an immune response. Only participants included in the substudy were analyzed. Here, 'n' is number of participants analyzed at specific time point.

End point type	Secondary

End point timeframe:

Days 15, 29, 43, 57, 90, 180, 360, and 730

End point values	AZD1222	Placebo		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	1885	943		
Units: ratio				
geometric mean (confidence interval 95%)				
S Antibody Titer: Day 15 (n=1885, 943)	34.28 (31.87 to 36.86)	0.97 (0.94 to 1.00)		
S Antibody Titer: Day 29 (n=1420, 706)	108.35 (99.95 to 117.45)	0.92 (0.87 to 0.96)		
S Antibody Titer: Day 43 (n=1797, 894)	455.38 (425.71 to 487.12)	1.08 (1.02 to 1.14)		
S Antibody Titer: Day 57 (n=1746, 823)	373.44 (348.12 to 400.60)	1.10 (1.02 to 1.19)		
S Antibody Titer: Day 90 (n=1664, 638)	274.05 (255.01 to 294.50)	1.61 (1.41 to 1.85)		
S Antibody Titer: Day 180 (n=1207, 146)	138.44 (125.37 to 152.88)	3.33 (2.07 to 5.37)		
S Antibody Titer: Day 360 (n=866, 46)	121.12 (102.89 to 142.58)	17.44 (6.75 to 45.07)		
S Antibody Titer: Day 730 (n=273, 22)	3506.74 (2808.52 to 4378.55)	295.66 (89.07 to 981.35)		
RBD Antibody Titer: Day 15 (n=1885, 943)	7.24 (6.76 to 7.76)	1.00 (0.98 to 1.02)		
RBD Antibody Titer: Day 29 (n=1419, 706)	39.01 (36.10 to 42.17)	1.01 (0.98 to 1.03)		
RBD Antibody Titer: Day 43 (n=1797, 894)	219.94 (207.28 to 233.37)	1.05 (1.01 to 1.09)		
RBD Antibody Titer: Day 57 (n=1746, 823)	178.61 (167.97 to 189.92)	1.10 (1.04 to 1.16)		
RBD Antibody Titer: Day 90 (n=1664, 638)	130.46 (122.53 to 138.89)	1.41 (1.27 to 1.57)		
RBD Antibody Titer: Day 180 (n=1207, 146)	61.69 (56.27 to 67.62)	3.13 (2.11 to 4.65)		
RBD Antibody Titer: Day 360 (n=866, 46)	53.91 (45.86 to 63.37)	11.18 (4.91 to 25.45)		
RBD Antibody Titer: Day 730 (n=273, 22)	2103.65 (1706.88 to 2592.64)	106.35 (29.93 to 377.87)		

Statistical analyses

No statistical analyses for this end point

Secondary: Percentage of Participants With Seroresponse to the S and RBD Antigens of AZD1222 as Measured by MSD Serology Assay

End point title	Percentage of Participants With Seroresponse to the S and RBD Antigens of AZD1222 as Measured by MSD Serology Assay

End point description:

The fold rise was calculated as the ratio of the post-vaccination titer level to the pre-vaccination titer level. The percentage of participants with a post-intervention seroresponse (≥ 4-fold rise in titers from baseline value to 28 days post each dose) to the S and RBD antigens of AZD1222 as measured by MSD serology assay is reported. The immunogenicity analysis population included all participants in the safety analysis set who had no protocol deviations judged to have the potential to interfere with the generation or interpretation of an immune response. Only participants included in the substudy were analyzed. Here, 'n' is number of participants analyzed at specific time point.

End point type	Secondary

End point timeframe:

Days 15, 29, 43, 57, 90, 180, 360, and 730

End point values	AZD1222	Placebo		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	1885	943		
Units: percentage of participants				
number (not applicable)				
S Antibody Titer: Day 15 (n=1885, 943)	89.8	1.1		
S Antibody Titer: Day 29 (n=1420, 706)	97.1	1.6		
S Antibody Titer: Day 43 (n=1797, 894)	99.4	2.5		
S Antibody Titer: Day 57 (n=1746, 823)	99.3	3.2		
S Antibody Titer: Day 90 (n=1664, 638)	99.3	8.9		
S Antibody Titer: Day 180 (n=1207, 146)	98.3	23.3		
S Antibody Titer: Day 360 (n=866, 46)	96.1	54.3		
S Antibody Titer: Day 730 (n=273, 22)	100.0	90.9		
RBD Antibody Titer: Day 15 (n=1885, 943)	61.3	0.4		
RBD Antibody Titer: Day 29 (n=1419, 706)	92.2	1.0		
RBD Antibody Titer: Day 43 (n=1797, 894)	98.8	2.0		
RBD Antibody Titer: Day 57 (n=1746, 823)	98.7	2.6		
RBD Antibody Titer: Day 90 (n=1664, 638)	98.6	7.7		
RBD Antibody Titer: Day 180 (n=1207, 146)	96.3	22.6		
RBD Antibody Titer: Day 360 (n=866, 46)	89.8	50.0		
RBD Antibody Titer: Day 730 (n=273, 22)	99.3	86.4		

Statistical analyses

No statistical analyses for this end point

Secondary: GMTs for SARS-CoV-2 Neutralizing Antibodies as Measured by Pseudo-neutralization Assay

End point title	GMTs for SARS-CoV-2 Neutralizing Antibodies as Measured by Pseudo-neutralization Assay

End point description:

The GMT was calculated as the antilogarithm of Σ(log base 2 transformed titer/n), i.e. as the anti-logarithm transformation of the mean of the log-transformed titer, where 'n' is the number of participants with titer information. The immunogenicity analysis population included all participants in the safety analysis set who had no protocol deviations judged to have the potential to interfere with the generation or interpretation of an immune response. Only participants included in the substudy were analyzed. Here, 'n' is number of participants analyzed at specific time point.

End point type	Secondary

End point timeframe:

Baseline (Day 1) and Days 15, 29, 43, 57, 90, 180, and 360

End point values	AZD1222	Placebo		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	1943	973		
Units: AU/mL				
geometric mean (confidence interval 95%)				
Day 1 (n=1943, 973)	20.6 (20.3 to 20.9)	21.4 (20.7 to 22.0)		
Day 15 (n=1747, 920)	41.7 (39.3 to 44.2)	21.7 (20.9 to 22.4)		
Day 29 (n=1227, 682)	65.9 (60.8 to 71.4)	23.0 (21.8 to 24.3)		
Day 43 (n=1642, 876)	221.3 (208.4 to 235.0)	22.6 (21.6 to 23.5)		
Day 57 (n=1469, 777)	251.8 (235.7 to 269.0)	23.3 (22.1 to 24.6)		
Day 90 (n=60, 21)	206.1 (150.3 to 282.6)	37.5 (15.0 to 94.1)		
Day 180 (n=283, 127)	112.9 (90.4 to 141.0)	51.3 (37.4 to 70.4)		
Day 360 (n=27, 14)	265.7 (94.0 to 751.1)	46.3 (20.1 to 106.4)		

Statistical analyses

No statistical analyses for this end point

Secondary: GMFR for SARS-CoV-2 Neutralizing Antibodies as Measured by Pseudo-neutralization Assay

End point title	GMFR for SARS-CoV-2 Neutralizing Antibodies as Measured by Pseudo-neutralization Assay

End point description:

The fold rise was calculated as the ratio of the post-vaccination titer level to the pre-vaccination titer level. GMFR was calculated as anti-logarithm of Σ (log base 2 transformed (post-vaccination titer/ pre-vaccination titer)/n). Where 'n' is the number of participants with titer information. The immunogenicity analysis population included all participants in the safety analysis set who had no protocol deviations judged to have the potential to interfere with the generation or interpretation of an immune response.

Only participants included in the substudy were analyzed. Here, 'n' is number of participants analyzed at specific time point.

End point type	Secondary

End point timeframe:

Days 15, 29, 43, 57, 90, 180, and 360

End point values	AZD1222	Placebo		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	1705	896		
Units: ratio				
geometric mean (confidence interval 95%)				
Day 15 (n=1705, 896)	2.03 (1.92 to 2.15)	1.01 (0.99 to 1.03)		
Day 29 (n=1200, 670)	3.24 (2.99 to 3.50)	1.07 (1.02 to 1.11)		
Day 43 (n=1597, 854)	10.91 (10.27 to 11.58)	1.05 (1.01 to 1.08)		
Day 57 (n=1432, 759)	12.32 (11.54 to 13.16)	1.08 (1.03 to 1.14)		
Day 90 (n=60, 20)	9.95 (7.33 to 13.51)	1.94 (0.74 to 5.09)		
Day 180 (n=274, 123)	5.37 (4.30 to 6.71)	2.22 (1.65 to 3.00)		
Day 360 (n=26, 14)	12.64 (4.31 to 37.11)	2.31 (1.01 to 5.32)		

Statistical analyses

No statistical analyses for this end point

Secondary: Percentage of Participants With Seroresponse to SARS-CoV-2 Neutralizing Antibodies of AZD1222 as Measured by Pseudo-neutralization Assay

End point title	Percentage of Participants With Seroresponse to SARS-CoV-2 Neutralizing Antibodies of AZD1222 as Measured by Pseudo-neutralization Assay

End point description:

The fold rise was calculated as the ratio of the post-vaccination titer level to the pre-vaccination titer level. The percentage of participants with a post-intervention seroresponse (≥ 4-fold rise in titers from baseline value to 28 days post each dose) to SARS-CoV-2 neutralizing antibodies of AZD1222 as measured by pseudo-neutralization assay is reported. The immunogenicity analysis population included all participants in the safety analysis set who had no protocol deviations judged to have the potential to interfere with the generation or interpretation of an immune response. Only participants included in the substudy were analyzed. Here, 'n' is number of participants analyzed at specific time point.

End point type	Secondary

End point timeframe:

Days 15, 29, 43, 57, 90, 180, and 360

End point values	AZD1222	Placebo		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	1705	896		
Units: percentage of participants				
number (not applicable)				
Day 15 (n=1705, 896)	24.8	0.3		
Day 29 (n=1200, 670)	41.1	1.8		
Day 43 (n=1597, 854)	84.4	1.9		
Day 57 (n=1432, 759)	84.4	2.2		
Day 90 (n=60, 20)	83.3	10.0		
Day 180 (n=274, 123)	51.5	22.0		
Day 360 (n=26, 14)	53.8	28.6		

Statistical analyses

No statistical analyses for this end point

Secondary: Number of Participants With COVID-19 Symptomatic Illness Post First Dose of Study Intervention

End point title	Number of Participants With COVID-19 Symptomatic Illness Post First Dose of Study Intervention

End point description:

The incidence of SARS-CoV-2 RT-PCR-positive symptomatic illness occurring post first dose of study intervention. The FAS included all randomized participants who received at least 1 dose of study intervention, irrespective of their protocol adherence and continued participation in the study. Only participants who are seronegative at baseline were analyzed.

End point type	Secondary

End point timeframe:

From Day 1 up to DCO of 05 March 2021 or study discontinuation or unblinding or receipt of non-study COVID-19 vaccination, up to a maximum of approximately 27 weeks

End point values	AZD1222	Placebo		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	20589	10300		
Units: participants	287	303		

Statistical analyses

Statistical analysis title	Treatment difference: COVID-19 symptomatic illness

Statistical analysis description:

The 95% CI were estimated based on Poisson regression with robust variance (including study arm and age group at screening (18-65 years, ≥ 65 years) as covariates and log of the follow-up time as an offset).

Comparison groups	AZD1222 v Placebo

Number of subjects included in analysis	30889
Analysis specification	Pre-specified
Analysis type	superiority
Parameter estimate	Vaccine efficacy
Point estimate	54.47
Confidence interval	
level	95 %
sides	2-sided
lower limit	46.48
upper limit	61.26

Adverse events

Adverse events information

Timeframe for reporting adverse events:

For SAEs: From first dose of study intervention up to 2 years of follow-up post first dose, a maximum of approximately 760 days;
For non-serious AEs: From first dose of study intervention up to 28 days post each dose, a maximum of approximately 57 days.

Adverse event reporting additional description:

The safety analysis set included all participants who received at least 1 dose of study intervention. Adverse events data reported for double-blind period only. All-cause mortality data reported for overall period (double-blind period and unblinded period).

Assessment type	Systematic

Dictionary used

Dictionary name	MedDRA
Dictionary version	23.1

Reporting groups

Reporting group title	Placebo

Reporting group description:

Participants received 2 IM doses of placebo matching with AZD1222 on Days 1 and 29.

Reporting group title	AZD1222

Reporting group description:

Participants received 2 IM doses of 5*10^10 vp (nominal, ± 1.5*10^10 vp) AZD1222 on Days 1 and 29.

Serious adverse events	Placebo	AZD1222	
Total subjects affected by serious adverse events			
subjects affected / exposed	467 / 10793 (4.33%)	1039 / 21587 (4.81%)	
number of deaths (all causes)	33	62	
number of deaths resulting from adverse events	33	62	
Neoplasms benign, malignant and unspecified (incl cysts and polyps)			
Bladder cancer			
subjects affected / exposed	2 / 10793 (0.02%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 2	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Acral lentiginous melanoma			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Acute monocytic leukaemia			

subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Acute myeloid leukaemia			
subjects affected / exposed	1 / 10793 (0.01%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 1	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Adenocarcinoma of colon			
subjects affected / exposed	3 / 10793 (0.03%)	7 / 21587 (0.03%)	
occurrences causally related to treatment / all	0 / 3	0 / 7	
deaths causally related to treatment / all	0 / 0	0 / 0	
Adenocarcinoma pancreas			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Astrocytoma			
subjects affected / exposed	2 / 10793 (0.02%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
B-cell lymphoma			
subjects affected / exposed	1 / 10793 (0.01%)	3 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 2	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Basal cell carcinoma			
subjects affected / exposed	1 / 10793 (0.01%)	3 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 1	0 / 4	
deaths causally related to treatment / all	0 / 0	0 / 0	
Benign neoplasm of bladder			
subjects affected / exposed	1 / 10793 (0.01%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Benign salivary gland neoplasm			

subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Chronic lymphocytic leukaemia			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Bladder neoplasm			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Bladder transitional cell carcinoma			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 1	
Borderline mucinous tumour of ovary			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Brain neoplasm			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Breast cancer			
subjects affected / exposed	7 / 10793 (0.06%)	11 / 21587 (0.05%)	
occurrences causally related to treatment / all	0 / 7	0 / 11	
deaths causally related to treatment / all	0 / 0	0 / 0	
Breast cancer female			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Breast cancer male			

subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Breast cancer metastatic			
subjects affected / exposed	1 / 10793 (0.01%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 1	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Breast cancer recurrent			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Breast cancer stage I			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Breast cancer stage II			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cerebral haemangioma			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cervix carcinoma			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cervix carcinoma stage IV			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
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 / 10793 (0.02%)	3 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 2	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 1	
Choroid melanoma			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Colorectal cancer metastatic			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Glioblastoma			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Colorectal adenocarcinoma			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 1	
Colon cancer stage IV			
subjects affected / exposed	2 / 10793 (0.02%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Colon cancer stage III			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Colon cancer metastatic			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Colon cancer			

subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Colon adenoma			
subjects affected / exposed	1 / 10793 (0.01%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 1	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Clear cell renal cell carcinoma			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Diffuse large B-cell lymphoma			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (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 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Endometrial cancer			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Ependymoma			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Follicular thyroid cancer			
subjects affected / exposed	1 / 10793 (0.01%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Gastrointestinal tract adenoma			

subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Colorectal cancer			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Invasive ductal breast carcinoma			
subjects affected / exposed	7 / 10793 (0.06%)	13 / 21587 (0.06%)	
occurrences causally related to treatment / all	0 / 7	0 / 13	
deaths causally related to treatment / all	0 / 0	0 / 0	
Invasive breast carcinoma			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Intraductal proliferative breast lesion			
subjects affected / exposed	2 / 10793 (0.02%)	7 / 21587 (0.03%)	
occurrences causally related to treatment / all	0 / 2	0 / 7	
deaths causally related to treatment / all	0 / 0	0 / 0	
Intraductal papillary breast neoplasm			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Huerthle cell carcinoma			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hormone receptor positive breast cancer			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Haemangioma of bone			

subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Glioma			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Glioblastoma multiforme			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Invasive lobular breast carcinoma			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Lung adenocarcinoma stage III			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Lung adenocarcinoma			
subjects affected / exposed	1 / 10793 (0.01%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 1	0 / 2	
deaths causally related to treatment / all	0 / 1	0 / 0	
Lipoma			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Lip squamous cell carcinoma			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Lip and/or oral cavity cancer			

subjects affected / exposed	1 / 10793 (0.01%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 1	
Lentigo maligna			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Laryngeal squamous cell carcinoma			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Lung carcinoma cell type unspecified stage IV			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Mucoepidermoid carcinoma			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Lymphoma			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Malignant anorectal neoplasm			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Malignant melanoma			
subjects affected / exposed	2 / 10793 (0.02%)	9 / 21587 (0.04%)	
occurrences causally related to treatment / all	0 / 3	0 / 9	
deaths causally related to treatment / all	0 / 0	0 / 0	
Malignant neoplasm of ampulla of Vater			

subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Malignant neoplasm of pleura			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Malignant neoplasm of thymus			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Malignant splenic neoplasm			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 1	
Mantle cell lymphoma			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Metastatic squamous cell carcinoma			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Metastatic carcinoma of the bladder			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Metastases to peritoneum			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Metastases to liver			

subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Metastases to bone			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Meningioma benign			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Meningioma			
subjects affected / exposed	1 / 10793 (0.01%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Lung neoplasm malignant			
subjects affected / exposed	2 / 10793 (0.02%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Myelodysplastic syndrome			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pancreatic carcinoma metastatic			
subjects affected / exposed	1 / 10793 (0.01%)	3 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 1	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 2	
Neuroendocrine carcinoma of the skin			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Neuroendocrine tumour			

subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Non-Hodgkin's lymphoma			
subjects affected / exposed	2 / 10793 (0.02%)	4 / 21587 (0.02%)	
occurrences causally related to treatment / all	0 / 2	0 / 5	
deaths causally related to treatment / all	0 / 0	0 / 0	
Non-Hodgkin's lymphoma stage III			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Non-small cell lung cancer			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
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 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Oesophageal adenocarcinoma stage III			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Oesophageal cancer metastatic			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Oesophageal carcinoma			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Oesophageal squamous cell carcinoma stage II			

subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Oligodendroglioma			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Ovarian cancer			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Ovarian cancer metastatic			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Ovarian cancer stage III			
subjects affected / exposed	2 / 10793 (0.02%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Neuroendocrine carcinoma			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 1	
Pancreatic carcinoma stage IV			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Renal cell carcinoma			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Parathyroid tumour benign			

subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pituitary tumour benign			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
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 / 10793 (0.01%)	3 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 1	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Plasmacytoma			
subjects affected / exposed	1 / 10793 (0.01%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Polycythaemia vera			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Prostate cancer			
subjects affected / exposed	22 / 10793 (0.20%)	32 / 21587 (0.15%)	
occurrences causally related to treatment / all	0 / 22	0 / 34	
deaths causally related to treatment / all	0 / 0	0 / 0	
Prostate cancer metastatic			
subjects affected / exposed	1 / 10793 (0.01%)	4 / 21587 (0.02%)	
occurrences causally related to treatment / all	0 / 1	0 / 4	
deaths causally related to treatment / all	0 / 0	0 / 0	
Prostate cancer recurrent			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Prostate cancer stage I			

subjects affected / exposed	1 / 10793 (0.01%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Prostate cancer stage II			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Prostate cancer stage III			
subjects affected / exposed	2 / 10793 (0.02%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Prostate cancer stage IV			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Rectal adenocarcinoma			
subjects affected / exposed	1 / 10793 (0.01%)	3 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 1	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Renal cancer			
subjects affected / exposed	1 / 10793 (0.01%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Papillary thyroid cancer			
subjects affected / exposed	0 / 10793 (0.00%)	7 / 21587 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 7	
deaths causally related to treatment / all	0 / 0	0 / 0	
Renal cell carcinoma stage IV			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Triple negative breast cancer			

subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Sarcoma uterus			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Seminoma			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Signet-ring cell carcinoma			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Sinonasal papilloma			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
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	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 1	
Spinal cord neoplasm			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Squamous cell carcinoma			
subjects affected / exposed	1 / 10793 (0.01%)	2 / 21587 (0.01%)	
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 / 10793 (0.00%)	6 / 21587 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 6	
deaths causally related to treatment / all	0 / 0	0 / 0	
Squamous cell carcinoma of the oral cavity			
subjects affected / exposed	1 / 10793 (0.01%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 1	0 / 0	
T-cell lymphoma			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Testicular seminoma (pure)			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Testis cancer			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Throat cancer			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Tonsil cancer			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Transitional cell carcinoma			
subjects affected / exposed	1 / 10793 (0.01%)	3 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 2	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Retinal melanocytoma			

subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Tumour associated fever			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Uterine leiomyoma			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Vascular disorders			
Haematoma			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Granulomatosis with polyangiitis			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hypertensive crisis			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hypertensive emergency			
subjects affected / exposed	2 / 10793 (0.02%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hypotension			
subjects affected / exposed	4 / 10793 (0.04%)	3 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 4	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Aortic aneurysm			

subjects affected / exposed	1 / 10793 (0.01%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Aortic aneurysm rupture			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Aortic dilatation			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Aortic dissection			
subjects affected / exposed	1 / 10793 (0.01%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Aortic stenosis			
subjects affected / exposed	1 / 10793 (0.01%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Arterial haemorrhage			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Arteriosclerosis			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Deep vein thrombosis			
subjects affected / exposed	3 / 10793 (0.03%)	13 / 21587 (0.06%)	
occurrences causally related to treatment / all	0 / 3	0 / 14	
deaths causally related to treatment / all	0 / 0	0 / 1	
Embolism			

subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hypertension			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hypovolaemic shock			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pelvic venous thrombosis			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Peripheral arterial occlusive disease			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Peripheral artery haematoma			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
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	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Shock haemorrhagic			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Venous thrombosis limb			

subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pregnancy, puerperium and perinatal conditions			
Ruptured ectopic pregnancy			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Abortion spontaneous			
subjects affected / exposed	3 / 10793 (0.03%)	15 / 21587 (0.07%)	
occurrences causally related to treatment / all	0 / 3	1 / 17	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cervical incompetence			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Ectopic pregnancy			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Gestational hypertension			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Postpartum haemorrhage			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pre-eclampsia			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Premature delivery			

subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Premature labour			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Stillbirth			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
General disorders and administration site conditions			
Asthenia			
subjects affected / exposed	1 / 10793 (0.01%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Catheter site haematoma			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Catheter site haemorrhage			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Chest discomfort			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Chest pain			
subjects affected / exposed	2 / 10793 (0.02%)	7 / 21587 (0.03%)	
occurrences causally related to treatment / all	0 / 2	0 / 7	
deaths causally related to treatment / all	0 / 0	0 / 0	
Death			

subjects affected / exposed	4 / 10793 (0.04%)	7 / 21587 (0.03%)	
occurrences causally related to treatment / all	0 / 4	0 / 7	
deaths causally related to treatment / all	0 / 4	0 / 7	
Drowning			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 1	0 / 0	
Hernia			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hypothermia			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Incarcerated hernia			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Non-cardiac chest pain			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Oedema peripheral			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Procedural failure			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (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	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Sudden cardiac death			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 1	
Swelling face			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Multiple organ dysfunction syndrome			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 1	
Immune system disorders			
Drug hypersensitivity			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hypersensitivity			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Anaphylactic reaction			
subjects affected / exposed	2 / 10793 (0.02%)	4 / 21587 (0.02%)	
occurrences causally related to treatment / all	0 / 2	0 / 4	
deaths causally related to treatment / all	0 / 0	0 / 0	
Reproductive system and breast disorders			
Adenomyosis			
subjects affected / exposed	2 / 10793 (0.02%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	

Rectocele			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Prostatitis			
subjects affected / exposed	1 / 10793 (0.01%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 1	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Benign prostatic hyperplasia			
subjects affected / exposed	2 / 10793 (0.02%)	4 / 21587 (0.02%)	
occurrences causally related to treatment / all	0 / 2	0 / 6	
deaths causally related to treatment / all	0 / 0	0 / 0	
Breast haematoma			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cervical dysplasia			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Endometriosis			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Female genital tract fistula			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Menorrhagia			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Ovarian cyst			

subjects affected / exposed	1 / 10793 (0.01%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Ovarian cyst ruptured			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Vaginal haemorrhage			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Respiratory, thoracic and mediastinal disorders			
Asthmatic crisis			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Chronic obstructive pulmonary disease			
subjects affected / exposed	1 / 10793 (0.01%)	6 / 21587 (0.03%)	
occurrences causally related to treatment / all	0 / 1	0 / 7	
deaths causally related to treatment / all	0 / 0	0 / 2	
Chronic respiratory failure			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Diaphragmatic paralysis			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Dyspnoea			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Dyspnoea exertional			

subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Epistaxis			
subjects affected / exposed	1 / 10793 (0.01%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Acute pulmonary oedema			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Acute respiratory distress syndrome			
subjects affected / exposed	0 / 10793 (0.00%)	3 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 1	
Acute respiratory failure			
subjects affected / exposed	3 / 10793 (0.03%)	11 / 21587 (0.05%)	
occurrences causally related to treatment / all	0 / 3	0 / 11	
deaths causally related to treatment / all	0 / 1	0 / 2	
Asphyxia			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 1	0 / 0	
Aspiration			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 1	
Asthma			
subjects affected / exposed	2 / 10793 (0.02%)	3 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 2	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Haemoptysis			

subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
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	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hypoxia			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Idiopathic pulmonary fibrosis			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 1	
Laryngeal leukoplakia			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Laryngeal obstruction			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pleural effusion			
subjects affected / exposed	0 / 10793 (0.00%)	4 / 21587 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 4	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pleuritic pain			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pneumonia aspiration			

subjects affected / exposed	1 / 10793 (0.01%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 1	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pneumothorax			
subjects affected / exposed	1 / 10793 (0.01%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 1	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pulmonary embolism			
subjects affected / exposed	7 / 10793 (0.06%)	29 / 21587 (0.13%)	
occurrences causally related to treatment / all	0 / 7	0 / 32	
deaths causally related to treatment / all	0 / 1	0 / 3	
Pulmonary fibrosis			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pulmonary granuloma			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pulmonary haemorrhage			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pulmonary hypertension			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pulmonary infarction			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Haemothorax			

subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
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	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Respiratory failure			
subjects affected / exposed	0 / 10793 (0.00%)	4 / 21587 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 4	
deaths causally related to treatment / all	0 / 0	0 / 2	
Sleep apnoea syndrome			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Respiratory arrest			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Psychiatric disorders			
Acute psychosis			
subjects affected / exposed	1 / 10793 (0.01%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Adjustment disorder with depressed mood			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Affective disorder			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Alcohol withdrawal syndrome			

subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Alcoholism			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Anxiety			
subjects affected / exposed	2 / 10793 (0.02%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Bipolar I disorder			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Bipolar II disorder			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Suicidal ideation			
subjects affected / exposed	5 / 10793 (0.05%)	5 / 21587 (0.02%)	
occurrences causally related to treatment / all	0 / 6	0 / 5	
deaths causally related to treatment / all	0 / 0	0 / 0	
Completed suicide			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 2	
Delirium			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Delirium tremens			

subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Depression			
subjects affected / exposed	1 / 10793 (0.01%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Depression suicidal			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Major depression			
subjects affected / exposed	1 / 10793 (0.01%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Mania			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Mental status changes			
subjects affected / exposed	1 / 10793 (0.01%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 1	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Post-traumatic stress disorder			
subjects affected / exposed	1 / 10793 (0.01%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Schizoaffective disorder			
subjects affected / exposed	2 / 10793 (0.02%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 4	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Schizophrenia			

subjects affected / exposed	2 / 10793 (0.02%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Substance abuse			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Substance-induced psychotic disorder			
subjects affected / exposed	0 / 10793 (0.00%)	3 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Bipolar disorder			
subjects affected / exposed	3 / 10793 (0.03%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 3	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Suicide attempt			
subjects affected / exposed	3 / 10793 (0.03%)	7 / 21587 (0.03%)	
occurrences causally related to treatment / all	0 / 3	0 / 7	
deaths causally related to treatment / all	0 / 0	0 / 0	
Product issues			
Device breakage			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (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 stenosis			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Bile duct stone			
subjects affected / exposed	1 / 10793 (0.01%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 1	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	

Cholangitis			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cholecystitis			
subjects affected / exposed	2 / 10793 (0.02%)	8 / 21587 (0.04%)	
occurrences causally related to treatment / all	0 / 2	0 / 8	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cholecystitis acute			
subjects affected / exposed	3 / 10793 (0.03%)	9 / 21587 (0.04%)	
occurrences causally related to treatment / all	0 / 3	0 / 9	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cholelithiasis			
subjects affected / exposed	3 / 10793 (0.03%)	8 / 21587 (0.04%)	
occurrences causally related to treatment / all	0 / 3	0 / 8	
deaths causally related to treatment / all	0 / 0	0 / 0	
Gallbladder polyp			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hepatitis alcoholic			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 1	0 / 0	
Ischaemic hepatitis			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Jaundice			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Jaundice cholestatic			

subjects affected / exposed	2 / 10793 (0.02%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Portal vein thrombosis			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Investigations			
Anticoagulation drug level above therapeutic			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Catheterisation cardiac			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Precancerous cells present			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Oxygen saturation decreased			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hepatic enzyme increased			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Heart rate increased			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Fibrin D dimer increased			

subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Injury, poisoning and procedural complications			
Accident			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 1	
Accidental overdose			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Acetabulum fracture			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Acquired encephalocele			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (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	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Ankle fracture			
subjects affected / exposed	3 / 10793 (0.03%)	6 / 21587 (0.03%)	
occurrences causally related to treatment / all	0 / 3	0 / 6	
deaths causally related to treatment / all	0 / 0	0 / 0	
Aortic injury			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Bladder injury			

subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Burns second degree			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cartilage injury			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cervical vertebral fracture			
subjects affected / exposed	1 / 10793 (0.01%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 1	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Chemical peritonitis			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
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	1 / 10793 (0.01%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 1	0 / 2	
deaths causally related to treatment / all	0 / 1	0 / 0	
Comminuted fracture			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Concussion			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Contusion			

subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Craniocerebral injury			
subjects affected / exposed	2 / 10793 (0.02%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 3	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Facial bones fracture			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Fall			
subjects affected / exposed	0 / 10793 (0.00%)	3 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Femoral neck fracture			
subjects affected / exposed	2 / 10793 (0.02%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 2	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Femur fracture			
subjects affected / exposed	6 / 10793 (0.06%)	3 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 6	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Fibula fracture			
subjects affected / exposed	1 / 10793 (0.01%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 1	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Foot fracture			
subjects affected / exposed	2 / 10793 (0.02%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Forearm fracture			

subjects affected / exposed	1 / 10793 (0.01%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Fracture displacement			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Gun shot wound			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 1	
Hand fracture			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Colon injury			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hip fracture			
subjects affected / exposed	4 / 10793 (0.04%)	6 / 21587 (0.03%)	
occurrences causally related to treatment / all	0 / 4	0 / 7	
deaths causally related to treatment / all	0 / 0	0 / 0	
Medication error			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hypobarism			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Ilium fracture			

subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Incarcerated incisional hernia			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Incision site haematoma			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Incisional hernia			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Injury			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (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 / 10793 (0.01%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Joint dislocation			
subjects affected / exposed	1 / 10793 (0.01%)	3 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 1	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Ligament rupture			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Ligament sprain			

subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Limb crushing injury			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Lower limb fracture			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Lumbar vertebral fracture			
subjects affected / exposed	1 / 10793 (0.01%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Humerus fracture			
subjects affected / exposed	1 / 10793 (0.01%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 1	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Multiple injuries			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Multiple fractures			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Overdose			
subjects affected / exposed	1 / 10793 (0.01%)	5 / 21587 (0.02%)	
occurrences causally related to treatment / all	0 / 1	0 / 5	
deaths causally related to treatment / all	0 / 0	0 / 3	
Patella fracture			

subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pelvic fracture			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Periprosthetic fracture			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pneumothorax traumatic			
subjects affected / exposed	0 / 10793 (0.00%)	3 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Post procedural haematuria			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Post procedural swelling			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Post-traumatic pain			
subjects affected / exposed	1 / 10793 (0.01%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Postoperative ileus			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
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 / 10793 (0.00%)	1 / 21587 (0.00%)	
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 / 10793 (0.00%)	1 / 21587 (0.00%)	
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	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Reactive gastropathy			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Rib fracture			
subjects affected / exposed	1 / 10793 (0.01%)	6 / 21587 (0.03%)	
occurrences causally related to treatment / all	0 / 1	0 / 6	
deaths causally related to treatment / all	0 / 0	0 / 0	
Osteochondral fracture			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Road traffic accident			
subjects affected / exposed	2 / 10793 (0.02%)	3 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 2	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 2	
Toxicity to various agents			
subjects affected / exposed	2 / 10793 (0.02%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 1	
deaths causally related to treatment / all	0 / 2	0 / 1	
Soft tissue injury			

subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Spinal compression fracture			
subjects affected / exposed	2 / 10793 (0.02%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 3	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Spinal cord injury			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 1	0 / 0	
Spinal fracture			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Splenic rupture			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Stab wound			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Sternal fracture			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Subdural haematoma			
subjects affected / exposed	1 / 10793 (0.01%)	3 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 1	0 / 4	
deaths causally related to treatment / all	0 / 0	0 / 0	
Subdural haemorrhage			

subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Tendon injury			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Tendon rupture			
subjects affected / exposed	2 / 10793 (0.02%)	4 / 21587 (0.02%)	
occurrences causally related to treatment / all	0 / 2	0 / 4	
deaths causally related to treatment / all	0 / 0	0 / 0	
Thermal burn			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 1	
Tibia fracture			
subjects affected / exposed	2 / 10793 (0.02%)	4 / 21587 (0.02%)	
occurrences causally related to treatment / all	0 / 2	0 / 5	
deaths causally related to treatment / all	0 / 0	0 / 0	
Skin laceration			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Traumatic haemothorax			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Traumatic intracranial haemorrhage			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Traumatic liver injury			

subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Wrist fracture			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Traumatic intracranial haematoma			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
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			
Thalassaemia minor			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cerebral arteriovenous malformation haemorrhagic			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cerebral cavernous malformation			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cerebral palsy			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Haemorrhagic arteriovenous malformation			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	

Hamartoma			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hypertrophic cardiomyopathy			
subjects affected / exposed	1 / 10793 (0.01%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Multiple congenital abnormalities			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Renal aplasia			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cardiac disorders			
Atrial flutter			
subjects affected / exposed	2 / 10793 (0.02%)	1 / 21587 (0.00%)	
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	13 / 10793 (0.12%)	21 / 21587 (0.10%)	
occurrences causally related to treatment / all	0 / 13	0 / 22	
deaths causally related to treatment / all	0 / 0	0 / 0	
Arteriosclerosis coronary artery			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (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	2 / 10793 (0.02%)	1 / 21587 (0.00%)	
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	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Angina unstable			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Angina pectoris			
subjects affected / exposed	1 / 10793 (0.01%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Acute myocardial infarction			
subjects affected / exposed	7 / 10793 (0.06%)	27 / 21587 (0.13%)	
occurrences causally related to treatment / all	0 / 7	0 / 27	
deaths causally related to treatment / all	0 / 1	0 / 1	
Acute left ventricular failure			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Acute coronary syndrome			
subjects affected / exposed	2 / 10793 (0.02%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Atrial tachycardia			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Chronic left ventricular failure			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cardiomyopathy			

subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cardio-respiratory arrest			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 2	
Cardiac tamponade			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cardiac failure congestive			
subjects affected / exposed	6 / 10793 (0.06%)	9 / 21587 (0.04%)	
occurrences causally related to treatment / all	0 / 9	0 / 11	
deaths causally related to treatment / all	0 / 1	0 / 1	
Cardiac failure acute			
subjects affected / exposed	0 / 10793 (0.00%)	4 / 21587 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 5	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cardiac failure			
subjects affected / exposed	0 / 10793 (0.00%)	3 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cardiac disorder			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 1	
Cardiac arrest			
subjects affected / exposed	4 / 10793 (0.04%)	5 / 21587 (0.02%)	
occurrences causally related to treatment / all	0 / 4	0 / 5	
deaths causally related to treatment / all	0 / 3	0 / 3	
Bundle branch block right			

subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Brugada syndrome			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Bradycardia			
subjects affected / exposed	2 / 10793 (0.02%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Atrioventricular block second degree			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Atrioventricular block complete			
subjects affected / exposed	3 / 10793 (0.03%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 3	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Atrioventricular block			
subjects affected / exposed	2 / 10793 (0.02%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cardiopulmonary failure			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Congestive cardiomyopathy			
subjects affected / exposed	1 / 10793 (0.01%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pericardial effusion			

subjects affected / exposed	0 / 10793 (0.00%)	3 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Coronary artery dissection			
subjects affected / exposed	1 / 10793 (0.01%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Coronary artery occlusion			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Coronary artery stenosis			
subjects affected / exposed	1 / 10793 (0.01%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 1	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hypertensive heart disease			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 1	
Ischaemic cardiomyopathy			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Mitral valve incompetence			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Mitral valve prolapse			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Myocardial infarction			

subjects affected / exposed	10 / 10793 (0.09%)	12 / 21587 (0.06%)	
occurrences causally related to treatment / all	0 / 10	1 / 12	
deaths causally related to treatment / all	0 / 2	0 / 3	
Myocardial ischaemia			
subjects affected / exposed	2 / 10793 (0.02%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Myocarditis			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Myxomatous mitral valve degeneration			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Non-obstructive cardiomyopathy			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Palpitations			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Coronary artery disease			
subjects affected / exposed	5 / 10793 (0.05%)	25 / 21587 (0.12%)	
occurrences causally related to treatment / all	0 / 7	0 / 26	
deaths causally related to treatment / all	0 / 0	0 / 1	
Pericarditis			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Sinus bradycardia			

subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
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	1 / 10793 (0.01%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Supraventricular tachycardia			
subjects affected / exposed	2 / 10793 (0.02%)	4 / 21587 (0.02%)	
occurrences causally related to treatment / all	0 / 2	0 / 4	
deaths causally related to treatment / all	0 / 0	0 / 0	
Tachycardia			
subjects affected / exposed	1 / 10793 (0.01%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Torsade de pointes			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Ventricular fibrillation			
subjects affected / exposed	0 / 10793 (0.00%)	3 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Ventricular tachycardia			
subjects affected / exposed	1 / 10793 (0.01%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Right ventricular failure			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Nervous system disorders			
Cerebral haemorrhage			

subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Alcoholic seizure			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Amnesia			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Bulbar palsy			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Carotid artery dissection			
subjects affected / exposed	2 / 10793 (0.02%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 3	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Carotid artery stenosis			
subjects affected / exposed	3 / 10793 (0.03%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 3	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cerebellar stroke			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cerebral artery occlusion			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Focal dyscognitive seizures			

subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Embolic stroke			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cerebral infarction			
subjects affected / exposed	1 / 10793 (0.01%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cerebral venous thrombosis			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cerebrovascular accident			
subjects affected / exposed	3 / 10793 (0.03%)	6 / 21587 (0.03%)	
occurrences causally related to treatment / all	0 / 3	0 / 6	
deaths causally related to treatment / all	0 / 0	0 / 0	
Chronic inflammatory demyelinating polyradiculoneuropathy			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cognitive disorder			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Corticobasal degeneration			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Delayed myelination			

subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Dementia			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Dementia Alzheimer's type			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 1	
Demyelinating polyneuropathy			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Dizziness			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Dysarthria			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Generalised tonic-clonic seizure			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Haemorrhage intracranial			
subjects affected / exposed	1 / 10793 (0.01%)	3 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 1	0 / 3	
deaths causally related to treatment / all	0 / 1	0 / 0	
Metabolic encephalopathy			

subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Headache			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hemianopia			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hemianopia homonymous			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
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 / 10793 (0.00%)	3 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hydrocephalus			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hypoaesthesia			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hypoxic-ischaemic encephalopathy			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Internal capsule infarction			

subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
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 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Intraventricular haemorrhage			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Ischaemic cerebral infarction			
subjects affected / exposed	1 / 10793 (0.01%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Ischaemic stroke			
subjects affected / exposed	7 / 10793 (0.06%)	15 / 21587 (0.07%)	
occurrences causally related to treatment / all	0 / 8	0 / 15	
deaths causally related to treatment / all	0 / 0	0 / 0	
Loss of consciousness			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
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	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Lumbosacral radiculopathy			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Haemorrhagic transformation stroke			

subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 1	0 / 0	
Migraine			
subjects affected / exposed	1 / 10793 (0.01%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Seizure			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Myasthenia gravis			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Myelomalacia			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Myelopathy			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Myoclonic epilepsy			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Neurodegenerative disorder			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 1	0 / 0	
Neuropathy peripheral			

subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Normal pressure hydrocephalus			
subjects affected / exposed	1 / 10793 (0.01%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Paraesthesia			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	1 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Parkinson's disease			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Peripheral sensory neuropathy			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Post stroke seizure			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Presyncope			
subjects affected / exposed	1 / 10793 (0.01%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Radiculopathy			
subjects affected / exposed	1 / 10793 (0.01%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Sciatica			

subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Multiple sclerosis			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Sensory loss			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Subarachnoid haemorrhage			
subjects affected / exposed	1 / 10793 (0.01%)	3 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 1	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Syncope			
subjects affected / exposed	4 / 10793 (0.04%)	8 / 21587 (0.04%)	
occurrences causally related to treatment / all	0 / 4	0 / 8	
deaths causally related to treatment / all	0 / 0	0 / 0	
Thalamic infarction			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Thrombotic stroke			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Toxic encephalopathy			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Transient global amnesia			

subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Transient ischaemic attack			
subjects affected / exposed	8 / 10793 (0.07%)	10 / 21587 (0.05%)	
occurrences causally related to treatment / all	0 / 8	0 / 10	
deaths causally related to treatment / all	0 / 0	0 / 0	
Spinal cord compression			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Blood and lymphatic system disorders			
Anaemia			
subjects affected / exposed	0 / 10793 (0.00%)	6 / 21587 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 6	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hypercoagulation			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Iron deficiency anaemia			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Neutropenia			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Splenic cyst			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Splenic haematoma			

subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Splenic infarction			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Thrombocytopenia			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Blood loss anaemia			
subjects affected / exposed	3 / 10793 (0.03%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 3	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Ear and labyrinth disorders			
Deafness bilateral			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Deafness			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Deafness unilateral			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Neurosensory hypoacusis			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Vertigo			

subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Vertigo positional			
subjects affected / exposed	0 / 10793 (0.00%)	3 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Eye disorders			
Amaurosis fugax			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Endocrine ophthalmopathy			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Optic ischaemic neuropathy			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	1 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Retinal artery occlusion			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Retinal detachment			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Retinal vein occlusion			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Gastrointestinal disorders			

Diverticular perforation			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Abdominal hernia			
subjects affected / exposed	0 / 10793 (0.00%)	3 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 4	
deaths causally related to treatment / all	0 / 0	0 / 0	
Abdominal pain lower			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Abdominal pain upper			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Abdominal strangulated hernia			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Abdominal wall haematoma			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Aphthous ulcer			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
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	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Colitis			

subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Colitis ischaemic			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Constipation			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Diarrhoea			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Diverticulum intestinal			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Incarcerated inguinal hernia			
subjects affected / exposed	2 / 10793 (0.02%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Duodenal perforation			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Duodenal ulcer haemorrhage			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Erosive oesophagitis			

subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Femoral hernia incarcerated			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Food poisoning			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Gastric fistula			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Gastric ulcer haemorrhage			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Gastric ulcer perforation			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Gastrointestinal haemorrhage			
subjects affected / exposed	2 / 10793 (0.02%)	5 / 21587 (0.02%)	
occurrences causally related to treatment / all	0 / 2	0 / 5	
deaths causally related to treatment / all	0 / 0	0 / 0	
Gastrointestinal necrosis			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Gastrooesophageal reflux disease			

subjects affected / exposed	1 / 10793 (0.01%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Haemorrhoids thrombosed			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hiatus hernia			
subjects affected / exposed	0 / 10793 (0.00%)	4 / 21587 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 4	
deaths causally related to treatment / all	0 / 0	0 / 0	
Ileus			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Diverticulum intestinal haemorrhagic			
subjects affected / exposed	0 / 10793 (0.00%)	3 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Mesenteric vein thrombosis			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Lower gastrointestinal haemorrhage			
subjects affected / exposed	1 / 10793 (0.01%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Oesophageal ulcer haemorrhage			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Oesophageal varices haemorrhage			

subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Oesophagitis			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Oesophagitis haemorrhagic			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pancreatic failure			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 1	
Pancreatitis			
subjects affected / exposed	2 / 10793 (0.02%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 2	0 / 2	
deaths causally related to treatment / all	0 / 1	0 / 0	
Pancreatitis acute			
subjects affected / exposed	1 / 10793 (0.01%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pancreatitis necrotising			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Inguinal hernia			
subjects affected / exposed	1 / 10793 (0.01%)	3 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 1	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Internal hernia			

subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Intestinal ischaemia			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Intestinal obstruction			
subjects affected / exposed	1 / 10793 (0.01%)	9 / 21587 (0.04%)	
occurrences causally related to treatment / all	0 / 1	0 / 9	
deaths causally related to treatment / all	0 / 0	0 / 0	
Intestinal perforation			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Large intestine perforation			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Obstructive pancreatitis			
subjects affected / exposed	1 / 10793 (0.01%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 1	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pancreatolithiasis			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pharyngo-oesophageal diverticulum			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Rectal haemorrhage			

subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Small intestinal obstruction			
subjects affected / exposed	5 / 10793 (0.05%)	12 / 21587 (0.06%)	
occurrences causally related to treatment / all	0 / 5	0 / 13	
deaths causally related to treatment / all	0 / 0	0 / 0	
Small intestinal perforation			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Umbilical hernia			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Upper gastrointestinal haemorrhage			
subjects affected / exposed	1 / 10793 (0.01%)	3 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 1	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Volvulus			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
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 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Peptic ulcer haemorrhage			
subjects affected / exposed	1 / 10793 (0.01%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Skin and subcutaneous tissue disorders			
Petechiae			

subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Angioedema			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Dermatitis			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
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	1 / 10793 (0.01%)	0 / 21587 (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			
Nephrolithiasis			
subjects affected / exposed	4 / 10793 (0.04%)	6 / 21587 (0.03%)	
occurrences causally related to treatment / all	0 / 4	0 / 6	
deaths causally related to treatment / all	0 / 0	0 / 0	
Acute kidney injury			
subjects affected / exposed	6 / 10793 (0.06%)	13 / 21587 (0.06%)	
occurrences causally related to treatment / all	0 / 7	0 / 13	
deaths causally related to treatment / all	0 / 0	0 / 0	
Bladder outlet obstruction			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Bladder perforation			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Chronic kidney disease			

subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Haematuria			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hydronephrosis			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Nephropathy toxic			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pelvi-ureteric obstruction			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Renal aneurysm			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Renal failure			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Renal mass			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Renal tubular necrosis			

subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Ureterolithiasis			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Urethral caruncle			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Urinary retention			
subjects affected / exposed	2 / 10793 (0.02%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Urinary tract obstruction			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Nephrotic syndrome			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Endocrine disorders			
Goitre			
subjects affected / exposed	2 / 10793 (0.02%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hypothyroidism			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Secondary adrenocortical insufficiency			

subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Thyroid mass			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Thyroiditis subacute			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Thyrotoxic crisis			
subjects affected / exposed	2 / 10793 (0.02%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Musculoskeletal and connective tissue disorders			
Arthralgia			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Rhabdomyolysis			
subjects affected / exposed	2 / 10793 (0.02%)	5 / 21587 (0.02%)	
occurrences causally related to treatment / all	0 / 2	0 / 5	
deaths causally related to treatment / all	0 / 0	0 / 0	
Back pain			
subjects affected / exposed	1 / 10793 (0.01%)	4 / 21587 (0.02%)	
occurrences causally related to treatment / all	0 / 1	0 / 4	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cervical spinal stenosis			
subjects affected / exposed	2 / 10793 (0.02%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 2	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Costochondritis			

subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Haemarthrosis			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Haematoma muscle			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Intervertebral disc degeneration			
subjects affected / exposed	0 / 10793 (0.00%)	3 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 4	
deaths causally related to treatment / all	0 / 0	0 / 0	
Intervertebral disc protrusion			
subjects affected / exposed	4 / 10793 (0.04%)	5 / 21587 (0.02%)	
occurrences causally related to treatment / all	0 / 5	0 / 7	
deaths causally related to treatment / all	0 / 0	0 / 0	
Limb mass			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Lumbar spinal stenosis			
subjects affected / exposed	2 / 10793 (0.02%)	6 / 21587 (0.03%)	
occurrences causally related to treatment / all	0 / 2	0 / 6	
deaths causally related to treatment / all	0 / 0	0 / 0	
Muscle spasms			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Musculoskeletal chest pain			

subjects affected / exposed	1 / 10793 (0.01%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Neck pain			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Osteoarthritis			
subjects affected / exposed	10 / 10793 (0.09%)	14 / 21587 (0.06%)	
occurrences causally related to treatment / all	0 / 13	0 / 16	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pain in extremity			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Arthritis			
subjects affected / exposed	0 / 10793 (0.00%)	5 / 21587 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 7	
deaths causally related to treatment / all	0 / 0	0 / 0	
Spinal stenosis			
subjects affected / exposed	3 / 10793 (0.03%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 3	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Spinal pain			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (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	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Vertebral foraminal stenosis			

subjects affected / exposed	1 / 10793 (0.01%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Rheumatoid arthritis			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Rotator cuff syndrome			
subjects affected / exposed	2 / 10793 (0.02%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Scoliosis			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Spinal osteoarthritis			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Spinal synovial cyst			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Infections and infestations			
Abdominal sepsis			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Abdominal abscess			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Abdominal wall abscess			

subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Abscess limb			
subjects affected / exposed	0 / 10793 (0.00%)	4 / 21587 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 4	
deaths causally related to treatment / all	0 / 0	0 / 0	
Chronic sinusitis			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cholecystitis infective			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cellulitis			
subjects affected / exposed	7 / 10793 (0.06%)	11 / 21587 (0.05%)	
occurrences causally related to treatment / all	0 / 8	0 / 12	
deaths causally related to treatment / all	0 / 0	0 / 0	
Campylobacter colitis			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
COVID-19 pneumonia			
subjects affected / exposed	21 / 10793 (0.19%)	29 / 21587 (0.13%)	
occurrences causally related to treatment / all	0 / 23	0 / 34	
deaths causally related to treatment / all	0 / 3	0 / 2	
COVID-19			
subjects affected / exposed	16 / 10793 (0.15%)	20 / 21587 (0.09%)	
occurrences causally related to treatment / all	0 / 17	0 / 23	
deaths causally related to treatment / all	0 / 2	0 / 1	
Bursitis infective			

subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Balanitis candida			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Bacterial pyelonephritis			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Bacteraemia			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Arthritis bacterial			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Appendicitis perforated			
subjects affected / exposed	2 / 10793 (0.02%)	4 / 21587 (0.02%)	
occurrences causally related to treatment / all	0 / 2	0 / 4	
deaths causally related to treatment / all	0 / 0	0 / 0	
Appendicitis			
subjects affected / exposed	11 / 10793 (0.10%)	29 / 21587 (0.13%)	
occurrences causally related to treatment / all	0 / 11	0 / 31	
deaths causally related to treatment / all	0 / 0	0 / 0	
Anal abscess			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Encephalitis			

subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Diverticulitis intestinal perforated			
subjects affected / exposed	0 / 10793 (0.00%)	3 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 4	
deaths causally related to treatment / all	0 / 0	0 / 0	
Diverticulitis			
subjects affected / exposed	1 / 10793 (0.01%)	7 / 21587 (0.03%)	
occurrences causally related to treatment / all	0 / 1	0 / 9	
deaths causally related to treatment / all	0 / 0	0 / 0	
Device related sepsis			
subjects affected / exposed	0 / 10793 (0.00%)	3 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Device related infection			
subjects affected / exposed	3 / 10793 (0.03%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 3	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cystitis			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Enterococcal bacteraemia			
subjects affected / exposed	0 / 10793 (0.00%)	3 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 4	
deaths causally related to treatment / all	0 / 0	0 / 0	
Complicated appendicitis			
subjects affected / exposed	2 / 10793 (0.02%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Colonic abscess			

subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Coccidioidomycosis			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Clostridium difficile infection			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
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 / 10793 (0.02%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 1	
deaths causally related to treatment / all	0 / 1	0 / 0	
Enterobacter sepsis			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Coronavirus infection			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Enterococcal sepsis			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Influenza			
subjects affected / exposed	1 / 10793 (0.01%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Infective tenosynovitis			

subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Infected skin ulcer			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Giardiasis			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Gastrointestinal infection			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
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	1 / 10793 (0.01%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Gastroenteritis shigella			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Gastroenteritis			
subjects affected / exposed	2 / 10793 (0.02%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Gangrene			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Focal peritonitis			

subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Extradural abscess			
subjects affected / exposed	0 / 10793 (0.00%)	3 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Escherichia urinary tract infection			
subjects affected / exposed	1 / 10793 (0.01%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Escherichia pyelonephritis			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Escherichia bacteraemia			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (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 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Peritonitis			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Peritonsillar abscess			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Periorbital cellulitis			

subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pelvic abscess			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Parainfluenzae virus infection			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Otitis externa			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Osteomyelitis			
subjects affected / exposed	5 / 10793 (0.05%)	5 / 21587 (0.02%)	
occurrences causally related to treatment / all	0 / 5	0 / 6	
deaths causally related to treatment / all	0 / 0	0 / 0	
Oropharyngeal candidiasis			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Orchitis			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Monkeypox			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Localised infection			

subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Large intestine infection			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Labyrinthitis			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Kidney infection			
subjects affected / exposed	1 / 10793 (0.01%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Intervertebral discitis			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Perirectal abscess			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pneumonia bacterial			
subjects affected / exposed	0 / 10793 (0.00%)	5 / 21587 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 5	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pneumonia klebsiella			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pneumonia legionella			

subjects affected / exposed	1 / 10793 (0.01%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pneumonia streptococcal			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Post procedural infection			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Postoperative abscess			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Postoperative wound infection			
subjects affected / exposed	1 / 10793 (0.01%)	3 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 1	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pseudomonal sepsis			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Psoas abscess			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pyelonephritis			
subjects affected / exposed	0 / 10793 (0.00%)	5 / 21587 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 5	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pyelonephritis acute			

subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Renal cyst infection			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Respiratory syncytial virus infection			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
SARS-CoV-2 sepsis			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Salmonellosis			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pharyngitis			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pneumonia			
subjects affected / exposed	8 / 10793 (0.07%)	32 / 21587 (0.15%)	
occurrences causally related to treatment / all	0 / 9	0 / 36	
deaths causally related to treatment / all	0 / 0	0 / 1	
Scrotal cellulitis			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Vascular device infection			

subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Sepsis syndrome			
subjects affected / exposed	1 / 10793 (0.01%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Septic shock			
subjects affected / exposed	2 / 10793 (0.02%)	5 / 21587 (0.02%)	
occurrences causally related to treatment / all	0 / 2	0 / 5	
deaths causally related to treatment / all	0 / 1	0 / 0	
Sinusitis bacterial			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Staphylococcal bacteraemia			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Staphylococcal infection			
subjects affected / exposed	2 / 10793 (0.02%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 2	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Staphylococcal sepsis			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Streptococcal bacteraemia			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Suspected COVID-19			

subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Toxic shock syndrome staphylococcal			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 1	
Tubo-ovarian abscess			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
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	2 / 10793 (0.02%)	8 / 21587 (0.04%)	
occurrences causally related to treatment / all	0 / 2	0 / 9	
deaths causally related to treatment / all	0 / 0	0 / 0	
Urinary tract infection bacterial			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 1	
Urinary tract infection enterococcal			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Urosepsis			
subjects affected / exposed	0 / 10793 (0.00%)	3 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Sepsis			
subjects affected / exposed	3 / 10793 (0.03%)	11 / 21587 (0.05%)	
occurrences causally related to treatment / all	0 / 3	0 / 11	
deaths causally related to treatment / all	0 / 0	0 / 0	
Viral infection			

subjects affected / exposed	1 / 10793 (0.01%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Viral pericarditis			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
West Nile viral infection			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Metabolism and nutrition disorders			
Dehydration			
subjects affected / exposed	1 / 10793 (0.01%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 1	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Failure to thrive			
subjects affected / exposed	0 / 10793 (0.00%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Gout			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hypercalcaemia			
subjects affected / exposed	0 / 10793 (0.00%)	4 / 21587 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 4	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hyperglycaemia			
subjects affected / exposed	2 / 10793 (0.02%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hyperglycaemic hyperosmolar nonketotic syndrome			

subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hyperkalaemia			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hypoglycaemia			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hypokalaemia			
subjects affected / exposed	2 / 10793 (0.02%)	2 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 2	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hypomagnesaemia			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hyponatraemia			
subjects affected / exposed	4 / 10793 (0.04%)	3 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 4	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hypovolaemia			
subjects affected / exposed	1 / 10793 (0.01%)	0 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Obesity			
subjects affected / exposed	0 / 10793 (0.00%)	1 / 21587 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Type 2 diabetes mellitus			

subjects affected / exposed	1 / 10793 (0.01%)	3 / 21587 (0.01%)	
occurrences causally related to treatment / all	0 / 1	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Diabetic ketoacidosis			
subjects affected / exposed	4 / 10793 (0.04%)	7 / 21587 (0.03%)	
occurrences causally related to treatment / all	0 / 4	0 / 8	
deaths causally related to treatment / all	0 / 1	0 / 0	

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

Non-serious adverse events	Placebo	AZD1222	
Total subjects affected by non-serious adverse events			
subjects affected / exposed	4234 / 10793 (39.23%)	11846 / 21587 (54.88%)	
Nervous system disorders			
Headache			
subjects affected / exposed	996 / 10793 (9.23%)	3511 / 21587 (16.26%)	
occurrences (all)	1125	4106	
General disorders and administration site conditions			
Pyrexia			
subjects affected / exposed	73 / 10793 (0.68%)	1174 / 21587 (5.44%)	
occurrences (all)	78	1209	
Fatigue			
subjects affected / exposed	853 / 10793 (7.90%)	2836 / 21587 (13.14%)	
occurrences (all)	942	3250	
Injection site pain			
subjects affected / exposed	532 / 10793 (4.93%)	3624 / 21587 (16.79%)	
occurrences (all)	595	4363	
Pain			
subjects affected / exposed	264 / 10793 (2.45%)	1857 / 21587 (8.60%)	
occurrences (all)	283	2023	
Chills			
subjects affected / exposed	248 / 10793 (2.30%)	2208 / 21587 (10.23%)	
occurrences (all)	264	2366	

Musculoskeletal and connective tissue disorders			
Myalgia			
subjects affected / exposed	278 / 10793 (2.58%)	1622 / 21587 (7.51%)	
occurrences (all)	296	1736	
Infections and infestations			
COVID-19			
subjects affected / exposed	2715 / 10793 (25.16%)	6207 / 21587 (28.75%)	
occurrences (all)	2921	6717	

More information

Substantial protocol amendments (globally)

Were there any global substantial amendments to the protocol? Yes

Date	Amendment
26 October 2020	Revised important potential risks. Increased the sample size from 30000 to 40000 participants. Added a description of the independent neurological AESI Expert Committee. Expanded the AESI list and specified that AESIs were reported to the Sponsor within one day of becoming aware of the event. New section for evaluation of potential neurological AESIs.
14 December 2020	Revised the first secondary endpoint. Identified the 4 key secondary endpoints. Added an immunogenicity cohort of approximately 300 participants. Reduced the sample size from 40000 back to the original number of 30000. Added text to indicate that experimental vaccinations, other than AZD1222, for prevention of SARS-CoV-2 or COVID-19 are prohibited. Also, noted that participants who choose to receive an authorized COVID-19 vaccine should inform the Investigator so it can be properly documented, and they should be encouraged to continue study conduct. Specified the procedures for unblinding in the event a study participant is contacted about receiving an authorized COVID-19 vaccine. Specified that if the primary endpoint results of the interim or primary analysis are statistically significant, an additional analysis with 5% alpha was performed once all participants had completed the first year of follow up. Revised to indicate how the primary efficacy endpoints and 4 key secondary endpoints were assessed.
19 February 2021	Removed “and Adolescents” from the title and removed text from the study rationale regarding adolescents. Upgraded exploratory endpoint "To estimate the efficacy of AZD1222 compared to saline placebo for the prevention of COVID-19 following the first dose" to secondary endpoint. Removed all references to Part 1 and Part 2 (Parts 1 and 2 were introduced in Amendment 5; an amendment which was not implemented).
29 March 2021	Revised to reflect the addition of an unblinded phase with an open-label dosing option (Part B). The original randomized, double-blind part of the study was labeled as Part A. Added additional exploratory objectives to assess the long-term safety and tolerability of 2 IM doses of AZD1222.

Notes:

Interruptions (globally)

Were there any global interruptions to the trial? No

Limitations and caveats

None reported