Clinical trial results:
A Randomized, Observer-blind, Active Comparator-controlled, Multicenter, Phase 3 Study to Assess the Efficacy, Safety, and Immunogenicity of a Plant-derived Quadrivalent VLP Influenza Vaccine in Adults 65 Years of Age and Older

Summary

EudraCT number	2018-001894-26
Trial protocol	DE FI
Global end of trial date	16 July 2019

Results information

Result version number	v1 (current)
This version publication date	29 September 2023
First version publication date	29 September 2023

Trial information

Trial identification

Sponsor protocol code	CP-PRO-QVLP-014

Additional study identifiers

ISRCTN number	-
ClinicalTrials.gov id (NCT number)	NCT03739112
WHO universal trial number (UTN)	-

Notes:

Sponsors

Sponsor organisation name	Medicago
Sponsor organisation address	1020 route de l'Église, bureau 600, Québec, Canada,
Public contact	Medical Director, Medicago, +1 418658-9393, clinicaltrialinquiries@medicago.com
Scientific contact	Medical Director, Medicago, +1 418658-9393, clinicaltrialinquiries@medicago.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	16 July 2019
Is this the analysis of the primary completion data?	No

Global end of trial reached?	Yes
Global end of trial date	16 July 2019
Was the trial ended prematurely?	No

Notes:

General information about the trial

Main objective of the trial:

- To evaluate the efficacy, relative to an active comparator, of a single 30 microgram (µg)/strain dose of the Quadrivalent Virus-like Particle (VLP) Influenza Vaccine, against protocol-defined influenza-like illness (ILI) caused by any influenza viral type/subtype (reverse transcription polymerase chain reaction [RT-PCR]).

Protection of trial subjects:

This study was conducted in accordance with the current International Council for Harmonisation (ICH) guidance, Good Clinical Practice (GCP) as established by the ICH (ICH E6 GCP), the European Union Clinical Trials Directive 2001 / 20 / EC, United States (US) 21 Code of Federal Regulations dealing with clinical studies, applicable federal, state, and/or local laws and regulations in the countries where the clinical study was conducted, clinical study contractual obligations, and the principles enunciated in the World Medical Association Declaration of Helsinki.

The Investigator or designee fully informed the subject of the risks and requirements of the study and, during the study, subjects were given any new information that could have affected their decision to continue participation. Subjects were told that their consent to participate in the study was voluntary and that it could be withdrawn at any time with no reason given and without penalty or loss of benefits to which they would otherwise be entitled. Only subjects who were fully able to understand the risks, benefits, and potential adverse events (AEs) of the study, and who provided their consent voluntarily were enrolled. The Investigator or designee answered all questions prior to requesting the subject's signature on the informed consent form (ICF). Subjects had sufficient time to consider the risks and benefits associated with participation in the study prior to signing the ICF. Each subject signed the ICF containing appropriate study and study drug information and was provided a copy of the ICF.

Background therapy: -

Evidence for comparator: -

Actual start date of recruitment	18 September 2018
Long term follow-up planned	No
Independent data monitoring committee (IDMC) involvement?	No

Notes:

Population of trial subjects

Subjects enrolled per country

Country: Number of subjects enrolled	Finland: 1329
Country: Number of subjects enrolled	Germany: 2683
Country: Number of subjects enrolled	Thailand: 537
Country: Number of subjects enrolled	Canada: 2517
Country: Number of subjects enrolled	United States: 5728
Worldwide total number of subjects	12794
EEA total number of subjects	4012

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)	0
From 65 to 84 years	12419
85 years and over	375

Subject disposition

Recruitment

Recruitment details:

Participants were randomized in a 1:1 ratio to receive the Quadrivalent virus-like particle (VLP) Influenza Vaccine at a dose of 30 µg/strain or the active comparator.

Pre-assignment

Screening details:

Participants aged 65 years or older with no acute or evolving medical problems were assessed.

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, Carer, Assessor

Arms

Are arms mutually exclusive?	Yes
Arm title	Quadrivalent (30 µg) VLP Vaccine

Arm description:

Participants received one intramuscular (IM) injection of 0.5 mL of 30 µg/strain of the Quadrivalent VLP Influenza Vaccine on Day 0.

Arm type	Experimental
Investigational medicinal product name	Quadrivalent VLP Influenza Vaccine
Investigational medicinal product code	
Other name	
Pharmaceutical forms	Injection
Routes of administration	Intramuscular use

Dosage and administration details:

Single dose of a 30 µg/strain of Quadrivalent VLP Vaccine.

Arm title	Active Comparator (Fluarix)

Arm description:

Participants received one IM injection of 0.5 milliliter (mL) of 15 µg/strain of the Fluarix Quadrivalent® comparator vaccine on Day 0.

Arm type	Active comparator
Investigational medicinal product name	Fluarix
Investigational medicinal product code	
Other name	
Pharmaceutical forms	Injection
Routes of administration	Intramuscular use

Dosage and administration details:

Single dose of a 15 µg/strain of Fluarix Quadrivalent® Comparator Vaccine.

Number of subjects in period 1	Quadrivalent (30 µg) VLP Vaccine	Active Comparator (Fluarix)
Started	6396	6398
Completed	6196	6222
Not completed	200	176
Adverse event, serious fatal	12	16
Consent withdrawn by subject	38	19
Physician decision	1	-
Site went on hold	65	67
Adverse event, non-fatal	2	2
Randomized but not vaccinated	31	25
Other than specified	8	5
Lost to follow-up	31	33
Protocol deviation	12	9

Baseline characteristics

Reporting groups	
Reporting group title	Quadrivalent (30 µg) VLP Vaccine
Reporting group description:	
Participants received one intramuscular (IM) injection of 0.5 mL of 30 µg/strain of the Quadrivalent VLP Influenza Vaccine on Day 0.	
Reporting group title	Active Comparator (Fluarix)
Reporting group description:	
Participants received one IM injection of 0.5 milliliter (mL) of 15 µg/strain of the Fluarix Quadrivalent® comparator vaccine on Day 0.	

Reporting group values	Quadrivalent (30 µg) VLP Vaccine	Active Comparator (Fluarix)	Total
Number of subjects	6396	6398	12794
Age categorical Units: Subjects			
Age continuous			
Baseline characteristics are summarized for the SAS.			
Units: years			
arithmetic mean	72.2	72.2	
standard deviation	± 5.67	± 5.7	-
Gender categorical Units: Subjects			
Female	3577	3575	7152
Male	2819	2823	5642
Ethnicity Units: Subjects			
Hispanic or Latino	202	198	400
Not Hispanic or Latino	6179	6190	12369
Unknown or Not Reported	15	10	25
Race Units: Subjects			
American Indian or Alaska Native	19	13	32
Asian	308	300	608
Native Hawaiian or Other Pacific Islander	5	5	10
Black or African American	251	266	517
White	5805	5803	11608
More than one race	6	10	16
Unknown or Not Reported	2	1	3

End points

End points reporting groups

Reporting group title	Quadrivalent (30 µg) VLP Vaccine
Reporting group description:	
Participants received one intramuscular (IM) injection of 0.5 mL of 30 µg/strain of the Quadrivalent VLP Influenza Vaccine on Day 0.	
Reporting group title	Active Comparator (Fluarix)
Reporting group description:	
Participants received one IM injection of 0.5 milliliter (mL) of 15 µg/strain of the Fluarix Quadrivalent® comparator vaccine on Day 0.	

Primary: Number of occurrences of protocol-defined Influenza-like Illness (ILI) due to any laboratory-confirmed Influenza strains

End point title	Number of occurrences of protocol-defined Influenza-like Illness (ILI) due to any laboratory-confirmed Influenza strains
End point description:	
Occurrences of laboratory-confirmed ILI caused by any influenza viral strains was measured by reverse transcriptase polymerase chain reaction (RT-PCR). A participant was considered to have protocol-defined ILI if the participant met at least one of the following pre-defined respiratory symptoms: sore throat, cough, sputum production, wheezing, or difficulty breathing and at least one of the following systemic symptoms: fever (defined as a temperature > 37.2 °C or > 99.0 °F), chills, tiredness, headache, or myalgia. The number of laboratory-confirmed ILI cases caused by any influenza strains are reported. The Per protocol (PP) set consisted of the participants who participated in the study until at least the end of the peak period or for at least five months or until the end of the surveillance period.	
End point type	Primary
End point timeframe:	
Day 14 (post vaccination) up to 9 months	

End point values	Quadrivalent (30 µg) VLP Vaccine	Active Comparator (Fluarix)		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	5996	6026		
Units: Number of cases				
number (not applicable)	118	130		

Statistical analyses

Statistical analysis title	Statistical analysis 1
Statistical analysis description:	
Vaccine efficacy (VE) of VLP vaccine versus Fluarix = (1-ARVv/ARVc) x 100% where ARVv = attack rate in participants vaccinated with the Quadrivalent VLP Influenza vaccine and ARVc = attack rate in participants vaccinated with an active Fluarix.	
Comparison groups	Quadrivalent (30 µg) VLP Vaccine v Active Comparator (Fluarix)

Number of subjects included in analysis	12022
Analysis specification	Pre-specified
Analysis type	non-inferiority[1]
Parameter estimate	percent VE
Point estimate	8.8
Confidence interval	
level	95 %
sides	2-sided
lower limit	-16.7
upper limit	28.7

Notes:

[1] - Non-inferiority was concluded if the lower limit of the two-sided 95% confidence interval (CI) for relative VE was > -20%.

Secondary: Number of occurrences of laboratory confirmed protocol-defined Influenza-like Illness (ILI) caused by vaccine-matched Influenza strains

End point title	Number of occurrences of laboratory confirmed protocol-defined Influenza-like Illness (ILI) caused by vaccine-matched Influenza strains

End point description:

Occurrences of protocol-defined ILI due to laboratory-confirmed influenza caused by influenza viral types/subtypes that were matched (and/or antigenically similar) to the strains covered in the vaccine formulation was measured by sequential RT-PCR & serotyping. The vaccine-matched strains included: homologous A/Michigan/45/2015 [H1N1], homologous A/Singapore/INFIMH-16-0019/2016 [H3N2], homologous B/Colorado/06/2017 and homologous B/Phuket/3073/2013) covered in the vaccine formulation. A participant was considered to have protocol-defined ILI if the participant met at least one of the following pre-defined respiratory symptoms: sore throat, cough, sputum production, wheezing, or difficulty breathing and at least one of the following systemic symptoms: fever (defined as a temperature > 37.2 °C or > 99.0 °F), chills, tiredness, headache, or myalgia. The number of laboratory-confirmed ILI cases caused by vaccine-matched influenza strains (all matched strains) are reported. PP set.

End point type	Secondary

End point timeframe:

Day 14 (post vaccination) up to 9 months

End point values	Quadrivalent (30 µg) VLP Vaccine	Active Comparator (Fluarix)		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	5996	6026		
Units: Number of cases				
number (not applicable)	42	46		

Statistical analyses

No statistical analyses for this end point

Secondary: Number of occurrences of protocol-defined respiratory illness due to any laboratory confirmed Influenza strains

End point title	Number of occurrences of protocol-defined respiratory illness due to any laboratory confirmed Influenza strains

End point description:

Occurrences of protocol-defined respiratory illness due to laboratory-confirmed influenza strain (matched, mismatched, and un-typed) was measured by sequential RT-PCR. A protocol-defined respiratory illness was determined by the occurrence of at least 1 of the following respiratory symptoms: sneezing, stuffy nose, sore throat, cough, sputum production, wheezing, or difficulty breathing. The number of protocol-defined respiratory illness cases caused by any laboratory-confirmed influenza strain (matched, mismatched, and un-typed) are reported. PP set.

End point type	Secondary

End point timeframe:

Day 14 (post vaccination) up to 9 months

End point values	Quadrivalent (30 µg) VLP Vaccine	Active Comparator (Fluarix)		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	5996	6026		
Units: Number of cases				
number (not applicable)	148	167		

Statistical analyses

No statistical analyses for this end point

Secondary: Number of occurrences of protocol-defined respiratory illness vaccine caused by vaccine-matched Influenza strains

End point title	Number of occurrences of protocol-defined respiratory illness vaccine caused by vaccine-matched Influenza strains

End point description:

Occurrences of protocol-defined respiratory illness vaccine caused by vaccine-matched influenza strains was measured by sequential RT-PCR & serotyping. The vaccine-matched strains included: homologous A/Michigan/45/2015 [H1N1], homologous A/Singapore/INFIMH-16-0019/2016 [H3N2], homologous B/Colorado/06/2017 and homologous B/Phuket/3073/2013. The protocol-defined respiratory illness was determined by the occurrence of at least 1 of the following respiratory symptoms: sneezing, stuffy nose, sore throat, cough, sputum production, wheezing, or difficulty breathing. The number of protocol-defined respiratory illness cases caused by one or more vaccine-matched strains are reported. PP set.

End point type	Secondary

End point timeframe:

Day 14 (post vaccination) up to 9 months

End point values	Quadrivalent (30 µg) VLP Vaccine	Active Comparator (Fluarix)		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	5996	6026		
Units: Number of cases				
number (not applicable)	50	55		

Statistical analyses

No statistical analyses for this end point

Secondary: Number of occurrences of protocol-defined ILI

End point title	Number of occurrences of protocol-defined ILI

End point description:

Occurrences of protocol-defined ILI that were confirmed or not by laboratory testing were assessed. A participant was considered to have protocol-defined ILI if the participant met at least one of the following pre-defined respiratory symptoms: sore throat, cough, sputum production, wheezing, or difficulty breathing and at least one of the following systemic symptoms: fever (defined as a temperature > 37.2 °C or > 99.0 °F), chills, tiredness, headache or myalgia. The number of protocol-defined ILI cases (confirmed or not) are reported. PP set.

End point type	Secondary

End point timeframe:

Day 14 (post vaccination) up to 9 months

End point values	Quadrivalent (30 µg) VLP Vaccine	Active Comparator (Fluarix)		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	5996	6026		
Units: Number of cases				
number (not applicable)	1249	1243		

Statistical analyses

No statistical analyses for this end point

Secondary: Number of participants with at least one immediate complaints

End point title	Number of participants with at least one immediate complaints

End point description:

Immediate complaints were defined as any solicited local or systemic reactions. Solicited local reactions included: erythema, swelling, and pain at the injection site) and solicited systemic reactions included: fever, headache, fatigue, muscle aches, joint aches, chills, a feeling of general discomfort, swelling in the axilla, and swelling in the neck. Safety Analysis Set (SAS) was defined as all participants who received either the Quadrivalent VLP Influenza Vaccine or the active comparator. Participants that were non-compliant to protocol/Good Clinical Practice (GCP), as per investigator, were excluded from analysis.

End point type	Secondary

End point timeframe:

15 minutes post vaccination

End point values	Quadrivalent (30 µg) VLP Vaccine	Active Comparator (Fluarix)		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	6352	6366		
Units: Participants				
number (not applicable)	272	184		

Statistical analyses

No statistical analyses for this end point

Secondary: Number of participants with ≥ Severe solicited local and systemic reactions

End point title	Number of participants with ≥ Severe solicited local and systemic reactions

End point description:

Participants were monitored for both solicited local reactions (erythema, swelling, and pain at the injection site) and solicited systemic reactions (fever, headache, fatigue, muscle aches, joint aches, chills, a feeling of general discomfort, swelling in the axilla, and swelling in the neck). The intensity of the solicited reactions was graded as mild (1)-easily tolerated and does not interfere with usual activity; moderate (2)-interferes with daily activity, but the participant is still able to function; severe (3)-incapacitating and the participant is unable to work or complete usual activity or potentially life threatening; (4)-likely to be life-threatening if not treated in a timely manner, according to the Food and Drug Administration (FDA) Guidance for Industry. ≥ Severe events included severe and potentially life-threatening events. Any ≥severe solicited reactions are reported.

End point type	Secondary

End point timeframe:

Day 0 (post-vaccination) to Day 7

End point values	Quadrivalent (30 µg) VLP Vaccine	Active Comparator (Fluarix)		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	6352[2]	6366[3]		
Units: Participants				
number (not applicable)	47	62		

Notes:

[2] - SAS. Participants non-compliant to protocol/GCP, as per investigator, were excluded from analysis.

[3] - SAS. Participants non-compliant to protocol/GCP, as per investigator, were excluded from analysis.

Statistical analyses

No statistical analyses for this end point

Secondary: Number of participants with ≥ Severe related solicited reactions

End point title	Number of participants with ≥ Severe related solicited

	reactions

End point description:

Participants were monitored for both solicited local reactions (erythema, swelling, & pain at the injection site) & solicited systemic reactions (fever, headache, fatigue, muscle aches, joint aches, chills, a feeling of general discomfort, swelling in axilla, & swelling in neck). The intensity of solicited reactions was graded as mild (1)-easily tolerated and does not interfere with usual activity; moderate (2)-interferes with daily activity, but the participant is still able to function; severe (3)-incapacitating and the participant is unable to work or complete usual activity or potentially life threatening; (4)-likely to be life-threatening if not treated in a timely manner, according to the FDA Guidance for Industry. ≥ Severe events included severe and potentially life-threatening events. Any ≥severe related (possibly related, probably related, and definitely related to the study treatments [as defined by investigator]) solicited events are reported.

End point type	Secondary

End point timeframe:

Day 0 (post-vaccination) to Day 7

End point values	Quadrivalent (30 µg) VLP Vaccine	Active Comparator (Fluarix)		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	6352[4]	6366[5]		
Units: Participants				
number (not applicable)	28	51		

Notes:

[4] - SAS. Participants non-compliant to protocol/GCP, as per investigator, were excluded from analysis.

[5] - SAS. Participants non-compliant to protocol/GCP, as per investigator, were excluded from analysis.

Statistical analyses

No statistical analyses for this end point

Secondary: Number of participants with unsolicited Treatment-Emergent Adverse Events (TEAEs)

End point title	Number of participants with unsolicited Treatment-Emergent Adverse Events (TEAEs)

End point description:

Participants were monitored for unsolicited TEAEs (e.g., nasopharyngitis, upper respiratory tract infection, headache, and pain). An adverse event (AE) or adverse experience was defined as any untoward medical occurrence in a participant or clinical investigation participant who received study drug, with or without a causal relationship with the treatment. An AE was considered treatment-emergent if it began on or after the date and time of Study Day 0 vaccination. SAS. Participants that were non-compliant to protocol/GCP, as per investigator, were excluded from analysis.

End point type	Secondary

End point timeframe:

Day 0 (post-vaccination) to Day 21

End point values	Quadrivalent (30 µg) VLP Vaccine	Active Comparator (Fluarix)		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	6352	6366		
Units: Participants				
number (not applicable)	849	824		

Statistical analyses

No statistical analyses for this end point

Secondary: Number of participants with ≥ Severe unsolicited TEAEs

End point title	Number of participants with ≥ Severe unsolicited TEAEs

End point description:

Participants were monitored for unsolicited TEAEs (e.g., nasopharyngitis, upper respiratory tract infection, headache, and pain). AE: any untoward medical occurrence in a participant or clinical investigation participant who received study drug, with or without a causal relationship with the treatment. An AE was considered treatment-emergent if it began on or after the date and time of Study Day 0 vaccination. The intensity of the solicited reactions was graded as mild (1)-easily tolerated and does not interfere with usual activity; moderate (2)-interferes with daily activity, but the participant is still able to function; severe (3)-incapacitating and the participant is unable to work or complete usual activity or potentially life threatening; (4)-likely to be life-threatening if not treated in a timely manner, according to the FDA Guidance for Industry. ≥ severe events included severe and potentially life-threatening events. Any ≥ severe unsolicited reactions are reported.

End point type	Secondary

End point timeframe:

Day 0 (post-vaccination) to Day 21

End point values	Quadrivalent (30 µg) VLP Vaccine	Active Comparator (Fluarix)		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	6352[6]	6366[7]		
Units: Participants				
number (not applicable)	33	34		

Notes:

[6] - SAS. Participants non-compliant to protocol/GCP, as per investigator, were excluded from analysis.

[7] - SAS. Participants non-compliant to protocol/GCP, as per investigator, were excluded from analysis.

Statistical analyses

No statistical analyses for this end point

Secondary: Number of participants with ≥ Severe related unsolicited reactions

End point title	Number of participants with ≥ Severe related unsolicited reactions

End point description:

Participants were monitored for unsolicited TEAEs (e.g., nasopharyngitis, upper respiratory tract infection, headache, & pain). AE: any untoward medical occurrence in a participant who received study drug, with or without a causal relationship with treatment. An AE was considered treatment-emergent if

it began on or after date & time of Study Day 0 vaccination. Intensity of solicited reactions was graded as mild (1) easily tolerated & not interfere with usual activity; moderate (2) interferes with daily activity, but participant still able to function; severe (3) incapacitating & participant unable to work/complete usual activity/ potentially life threatening; (4) likely to be life-threatening if not treated in a timely manner, according to FDA Guidance for Industry. ≥ severe events included severe & potentially life-threatening events. Any ≥severe related (possibly, probably, & definitely related to study treatments [as defined by investigator]) unsolicited events are reported.

End point type	Secondary

End point timeframe:

Day 0 (post-vaccination) to Day 21

End point values	Quadrivalent (30 µg) VLP Vaccine	Active Comparator (Fluarix)		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	6352[8]	6366[9]		
Units: Participants				
number (not applicable)	1	3		

Notes:

[8] - SAS. Participants non-compliant to protocol/GCP, as per investigator, were excluded from analysis.

[9] - SAS. Participants non-compliant to protocol/GCP, as per investigator, were excluded from analysis.

Statistical analyses

No statistical analyses for this end point

Secondary: Number of participants with at least one Serious Adverse Event (SAE)

End point title	Number of participants with at least one Serious Adverse Event (SAE)

End point description:

An AE was any untoward medical occurrence in a participant who received study drug, with or without a causal relationship with treatment. An SAE was an AE that resulted in death, was life threatening, resulted in a persistent or significant disability or incapacity, resulted in or prolonged an existing hospitalization, was a congenital anomaly or birth defect, or was another important medical event. SAS. Participants that were non-compliant to protocol/GCP, as per investigator, were excluded from analysis.

End point type	Secondary

End point timeframe:

Day 0 to 9 months

End point values	Quadrivalent (30 µg) VLP Vaccine	Active Comparator (Fluarix)		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	6352	6366		
Units: Participants				
number (not applicable)	263	266		

Statistical analyses

No statistical analyses for this end point

Secondary: Number of occurrences of death

End point title	Number of occurrences of death

End point description:

The number of participants who died during the study was assessed. SAS. Participants that were non-compliant to protocol/GCP, as per investigator, were excluded from analysis.

End point type	Secondary

End point timeframe:

Day 0 up to 9 months

End point values	Quadrivalent (30 µg) VLP Vaccine	Active Comparator (Fluarix)		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	6352	6366		
Units: Participants				
number (not applicable)	12	17		

Statistical analyses

No statistical analyses for this end point

Secondary: Number of participants who withdrew due to an AE

End point title	Number of participants who withdrew due to an AE

End point description:

An AE or adverse experience was defined as any untoward medical occurrence in a participant or clinical investigation participant who received study drug, with or without a causal relationship with the treatment. An AE can be any favorable and unintended sign, symptom, or disease temporally associated with the use of a medicinal product, whether or not related to a medicinal product. SAS. Participants that were non-compliant to protocol/GCP, as per investigator, were excluded from analysis.

End point type	Secondary

End point timeframe:

Day 0 up to 9 months

End point values	Quadrivalent (30 µg) VLP Vaccine	Active Comparator (Fluarix)		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	6352	6366		
Units: Participants				
number (not applicable)	15	20		

Statistical analyses

No statistical analyses for this end point

Secondary: Number of participants with at least one New Onset Chronic Diseases (NOCDs)

End point title	Number of participants with at least one New Onset Chronic Diseases (NOCDs)

End point description:

All NOCDs that may plausibly have an allergic, autoimmune or inflammatory component were reported. Plausibility should be interpreted broadly; however, the only clear exceptions were degenerative conditions such as osteoarthritis, age-related physiologic changes and life-style diseases. In this context, most cancers, cardiac conditions and kidney diseases should be reported. NOCDs were collected from vaccination on Day 0 to the end of the surveillance period. SAS. Participants that were non-compliant to protocol/GCP, as per investigator, were excluded from analysis.

End point type	Secondary

End point timeframe:

Day 0 up to 9 months

End point values	Quadrivalent (30 µg) VLP Vaccine	Active Comparator (Fluarix)		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	6352	6366		
Units: Participants				
number (not applicable)	36	23		

Statistical analyses

No statistical analyses for this end point

Secondary: Geometric mean titers (GMTs) of Hemagglutination Inhibition (HI) antibody response for each homologous and heterologous Influenza strains

End point title	Geometric mean titers (GMTs) of Hemagglutination Inhibition (HI) antibody response for each homologous and heterologous Influenza strains

End point description:

GMTs were measured using a HI assay for the homologous strains: H1N1 = A/Michigan/45/2015; H3N2 = A/Singapore/INFIMH-16-0019/2016; B/Colorado = B/Colorado/06/2017; B/Phuket = B/Phuket/3073/2013, and the heterologous strains: H1N1 = A/Brisbane/59/2007; H3N2 = A/Uruguay/716/2007; B/Malaysia = B/Malaysia/2506/2004; B/Florida = B/Florida/4/2006. The Immunogenicity Per Protocol (IPP) set consisted of a subset of participants who participated in the immunogenicity portion of the study and who had a Day 21 immunogenicity sample collection.

End point type	Secondary

End point timeframe:
Baseline (Day 0), Day 21

End point values	Quadrivalent (30 µg) VLP Vaccine	Active Comparator (Fluarix)		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	206	199		
Units: Titers				
geometric mean (confidence interval 95%)				
Homologous Strain (H1N1): Day 0	43.8 (36.3 to 52.7)	36.3 (30.0 to 43.9)		
Homologous Strain (H1N1): Day 21	88.9 (73.7 to 107.2)	146.8 (121.4 to 177.7)		
Homologous Strain (H3N2): Day 0	31.1 (25.5 to 37.8)	30.9 (25.3 to 37.8)		
Homologous Strain (H3N2): Day 21	79.5 (65.2 to 97.1)	90.2 (73.7 to 110.5)		
Homologous Strain (B/Colorado): Day 0	15.5 (13.3 to 18.1)	13.2 (11.3 to 15.5)		
Homologous Strain (B/Colorado): Day 21	23.0 (19.3 to 27.4)	51.7 (43.3 to 61.8)		
Homologous Strain (B/Phuket): Day 0	24.4 (20.8 to 28.7)	22.6 (19.2 to 26.6)		
Homologous Strain (B/Phuket): Day 21	45.0 (37.8 to 53.5)	75.5 (63.3 to 90.1)		
Heterologous Strain (H1N1): Day 0	15.5 (13.6 to 17.6)	13.5 (11.8 to 15.4)		
Heterologous Strain (H1N1): Day 21	16.6 (14.3 to 19.2)	32.4 (28.0 to 37.6)		
Heterologous Strain (H3N2): Day 0	19.5 (16.4 to 23.3)	22.6 (18.9 to 27.1)		
Heterologous Strain (H3N2): Day 21	32.4 (27.1 to 38.6)	46.6 (38.9 to 55.8)		
Heterologous Strain (B/Malaysia): Day 0	12.6 (11.0 to 14.5)	11.4 (10.0 to 13.1)		
Heterologous Strain (B/Malaysia): Day 21	15.9 (13.6 to 18.7)	34.8 (29.5 to 41.0)		
Heterologous Strain (B/Florida): Day 0	22.5 (19.2 to 26.4)	20.6 (17.5 to 24.2)		
Heterologous Strain (B/Florida): Day 21	38.0 (32.2 to 44.8)	72.1 (61.0 to 85.2)		

Statistical analyses

No statistical analyses for this end point

Secondary: Percentage of participants with seroconversion measured by HI antibody response for each homologous and heterologous strain

End point title	Percentage of participants with seroconversion measured by HI antibody response for each homologous and heterologous strain

End point description:

The percentage of participants in a given treatment group with either a ≥ 4-fold increase in reciprocal HI titers between Day 0 and Day 21 or a rise of undetectable HI titer (i.e. < 10) pre-vaccination (Day 0) to an HI titer of ≥ 40 on Day 21 was measured using an HI assay for the homologous strains: H1N1 = A/Michigan/45/2015; H3N2 = A/Singapore/INFIMH-16-0019/2016; B/Colorado = B/Colorado/06/2017; B/Phuket = B/Phuket/3073/2013, and the heterologous strains: H1N1 = A/Brisbane/59/2007; H3N2 = A/Uruguay/716/2007; B/Malaysia = B/Malaysia/2506/2004; B/Florida = B/Florida/4/2006. IPP set.

End point type	Secondary

End point timeframe:

Day 0 (pre-vaccination) to Day 21

End point values	Quadrivalent (30 µg) VLP Vaccine	Active Comparator (Fluarix)		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	206	199		
Units: Percentage of participants				
number (confidence interval 95%)				
Homologous Strain (H1N1)	15.5 (10.9 to 21.2)	40.7 (33.8 to 47.9)		
Homologous Strain (H3N2)	25.2 (19.5 to 31.7)	28.1 (22.0 to 34.9)		
Homologous Strain (B/Colorado)	10.2 (6.4 to 15.2)	38.2 (31.4 to 45.3)		
Homologous Strain (B/Phuket)	16.5 (11.7 to 22.3)	34.7 (28.1 to 41.7)		
Heterologous Strain (H1N1)	0.5 (0.0 to 2.7)	24.1 (18.4 to 30.7)		
Heterologous Strain (H3N2)	10.7 (6.8 to 15.7)	14.6 (10.0 to 20.3)		
Heterologous Strain (B/Malaysia)	5.3 (2.7 to 9.4)	30.7 (24.3 to 37.6)		
Heterologous Strain (B/Florida)	12.1 (8.0 to 17.4)	37.7 (30.9 to 44.8)		

Statistical analyses

No statistical analyses for this end point

Secondary: Percentage of participants with seroprotection measured by HI antibody response for each homologous and heterologous strain

End point title	Percentage of participants with seroprotection measured by HI antibody response for each homologous and heterologous strain

End point description:

The percentage of participants in a given treatment group attaining a reciprocal HI titer of ≥ 40 on Day 21 (the percentage of vaccine recipients with a serum HI titer of at least 1:40 following vaccination) was measured using an HI assay for homologous strains: H1N1 = A/Michigan/45/2015; H3N2 = A/Singapore/INFIMH-16-0019/2016; B/Colorado = B/Colorado/06/2017; B/Phuket = B/Phuket/3073/2013, and the heterologous strains: H1N1 = A/Brisbane/59/2007; H3N2 = A/Uruguay/716/2007; B/Malaysia = B/Malaysia/2506/2004; B/Florida = B/Florida/4/2006. IPP set.

End point type	Secondary

End point timeframe:

Baseline (Day 0), Day 21

End point values	Quadrivalent (30 µg) VLP Vaccine	Active Comparator (Fluarix)		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	206	199		
Units: Percentage of participants				
number (confidence interval 95%)				
Homologous Strain (H1N1): Day 0	57.3 (50.2 to 64.1)	52.8 (45.6 to 59.9)		
Homologous Strain (H1N1): Day 21	77.2 (70.8 to 82.7)	87.9 (82.6 to 92.1)		
Homologous Strain (H3N2): Day 0	48.5 (41.5 to 55.6)	52.8 (45.6 to 59.9)		
Homologous Strain (H3N2): Day 21	72.3 (65.7 to 78.3)	78.4 (72.0 to 83.9)		
Homologous Strain (B/Colorado): Day 0	28.6 (22.6 to 35.3)	22.1 (16.5 to 28.5)		
Homologous Strain (B/Colorado): Day 21	40.3 (33.5 to 47.3)	62.3 (55.2 to 69.1)		
Homologous Strain (B/Phuket): Day 0	46.6 (39.6 to 53.7)	38.7 (31.9 to 45.8)		
Homologous Strain (B/Phuket): Day 21	64.1 (57.1 to 70.6)	79.4 (73.1 to 84.8)		
Heterologous Strain (H1N1): Day 0	21.8 (16.4 to 28.1)	18.1 (13.0 to 24.2)		
Heterologous Strain (H1N1): Day 21	22.8 (17.3 to 29.2)	49.2 (42.1 to 56.4)		
Heterologous Strain (H3N2): Day 0	35.4 (28.9 to 42.4)	40.2 (33.3 to 47.4)		
Heterologous Strain (H3N2): Day 21	50.0 (43.0 to 57.0)	63.8 (56.7 to 70.5)		
Heterologous Strain (B/Malaysia): Day 0	22.8 (17.3 to 29.2)	17.6 (12.6 to 23.6)		
Heterologous Strain (B/Malaysia): Day 21	30.1 (23.9 to 36.9)	53.8 (46.6 to 60.8)		
Heterologous Strain (B/Florida): Day 0	37.4 (30.8 to 44.4)	31.2 (24.8 to 38.1)		
Heterologous Strain (B/Florida): Day 21	58.7 (51.7 to 65.5)	76.9 (70.4 to 82.6)		

Statistical analyses

No statistical analyses for this end point

Secondary: Geometric mean fold rise (GMFR) of HI antibody response for each homologous and heterologous strain

End point title	Geometric mean fold rise (GMFR) of HI antibody response for each homologous and heterologous strain

End point description:

GMFR, the geometric mean of the ratio of GMTs (Day 21/Day 0), was measured using an HI assay

homologous strains: H1N1 = A/Michigan/45/2015; H3N2 = A/Singapore/INFIMH-16-0019/2016; B/Colorado = B/Colorado/06/2017; B/Phuket = B/Phuket/3073/2013, and the heterologous strains: H1N1 = A/Brisbane/59/2007; H3N2 = A/Uruguay/716/2007; B/Malaysia = B/Malaysia/2506/2004; B/Florida = B/Florida/4/2006. IPP set.

End point type	Secondary
End point timeframe:	
Baseline (Day 0), Day 21	

End point values	Quadrivalent (30 µg) VLP Vaccine	Active Comparator (Fluarix)		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	206	199		
Units: Ratio				
geometric mean (confidence interval 95%)				
Homologous Strain (H1N1)	2.1 (1.8 to 2.4)	3.9 (3.4 to 4.5)		
Homologous Strain (H3N2)	2.6 (2.2 to 3.0)	2.9 (2.5 to 3.4)		
Homologous Strain (B/Colorado)	1.5 (1.3 to 1.8)	3.8 (3.3 to 4.4)		
Homologous Strain (B/Phuket)	1.9 (1.6 to 2.1)	3.3 (2.9 to 3.8)		
Heterologous Strain (H1N1)	1.1 (1.0 to 1.2)	2.4 (2.2 to 2.6)		
Heterologous Strain (H3N2)	1.6 (1.5 to 1.8)	2.1 (1.9 to 2.3)		
Heterologous Strain (B/Malaysia)	1.3 (1.1 to 1.4)	3.0 (2.7 to 3.4)		
Heterologous Strain (B/Florida)	1.7 (1.5 to 1.9)	3.4 (3.0 to 3.9)		

Statistical analyses

No statistical analyses for this end point

Secondary: GMTS of Microneutralization (MN) antibody response for each homologous strain

End point title	GMTS of Microneutralization (MN) antibody response for each homologous strain

End point description:

GMTs were measured using an MN assay for homologous strains:H1N1 = A/Michigan/45/2015; H3N2 = A/Singapore/INFIMH-16-0019/2016; B/Colorado = B/Colorado/06/2017; B/Phuket = B/Phuket/3073/2013. IPP set.

End point type	Secondary
End point timeframe:	
Baseline (Day 0), Day 21	

End point values	Quadrivalent (30 µg) VLP Vaccine	Active Comparator (Fluarix)		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	206	199		
Units: Titers				
geometric mean (confidence interval 95%)				
H1N1: Day 0	501.4 (420.6 to 597.8)	511.2 (427.5 to 611.3)		
H1N1: Day 21	1120.7 (926.3 to 1355.9)	1807.0 (1488.6 to 2193.6)		
H3N2: Day 0	227.4 (194.9 to 265.3)	240.1 (205.3 to 280.8)		
H3N2: Day 21	558.5 (461.2 to 676.3)	437.0 (359.7 to 531.0)		
B/Colorado: Day 0	62.7 (54.1 to 72.6)	55.1 (47.5 to 64.0)		
B/Colorado: Day 21	124.1 (105.4 to 146.2)	193.1 (163.5 to 228.1)		
B/Phuket: Day 0	50.5 (44.4 to 57.6)	45.8 (40.1 to 52.3)		
B/Phuket: Day 21	98.2 (85.7 to 112.6)	143.4 (143.4 to 164.7)		

Statistical analyses

No statistical analyses for this end point

Secondary: Percentage of participants with seroconversion measured by MN antibody response for each homologous strain

End point title	Percentage of participants with seroconversion measured by MN antibody response for each homologous strain

End point description:

The percentage of participants in a given treatment group with either a ≥ 4-fold increase in reciprocal MN titers between Day 0 and Day 21 or a rise of undetectable MN titer (i.e. 7.1) pre-vaccination (Day 0) to an MN titer of ≥ 28.3 at Day 21 post-vaccination were measured using an MN assay for homologous strains: H1N1 = A/Michigan/45/2015; H3N2 = A/Singapore/INFIMH-16-0019/2016; B/Colorado = B/Colorado/06/2017; B/Phuket = B/Phuket/3073/2013. IPP set.

End point type	Secondary

End point timeframe:

Day 0 (pre-vaccination) to Day 21

End point values	Quadrivalent (30 µg) VLP Vaccine	Active Comparator (Fluarix)		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	206	199		
Units: Percentage of participants				
number (confidence interval 95%)				
H1N1	25.7 (19.9 to 32.3)	43.2 (36.2 to 50.4)		

H3N2	36.4 (29.8 to 43.4)	26.6 (20.6 to 33.3)		
B/Colorado	19.9 (14.7 to 26.0)	43.7 (36.7 to 50.9)		
B/Phuket	19.4 (14.2 to 25.5)	41.2 (34.3 to 48.4)		

Statistical analyses

No statistical analyses for this end point

Secondary: GMFR of MN antibody response for each homologous strain

End point title	GMFR of MN antibody response for each homologous strain

End point description:

GMFR, the geometric mean of the ratio of GMTs (Day 21/Day 0), was measured using an MN assay for H1N1 = A/Michigan/45/2015; H3N2 = A/Singapore/INFIMH-16-0019/2016; B/Colorado = B/Colorado/06/2017; B/Phuket = B/Phuket/3073/2013. IPP set.

End point type	Secondary

End point timeframe:

Baseline (Day 0), Day 21

End point values	Quadrivalent (30 µg) VLP Vaccine	Active Comparator (Fluarix)		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	206	199		
Units: Ratio				
geometric mean (confidence interval 95%)				
H1N1	2.2 (1.9 to 2.6)	3.5 (3.0 to 4.1)		
H3N2	2.4 (2.1 to 2.8)	1.8 (1.6 to 2.1)		
B/Colorado	2.0 (1.8 to 2.3)	3.4 (3.0 to 3.9)		
B/Phuket	2.0 (1.8 to 2.2)	3.1 (2.8 to 3.4)		

Statistical analyses

No statistical analyses for this end point

Secondary: Geometric Mean Areas (GMA) of Single Radial Hemolysis (SRH) antibody response for each homologous strain

End point title	Geometric Mean Areas (GMA) of Single Radial Hemolysis (SRH) antibody response for each homologous strain

End point description:

GMA was measured using an SRH assay for homologous strains: H1N1 = A/Michigan/45/2015; H3N2 = A/Singapore/INFIMH-16-0019/2016; B/Colorado = B/Colorado/06/2017; B/Phuket = B/Phuket/3073/2013. IPP set.

End point type	Secondary

End point timeframe:

Baseline (Day 0), Day 21

End point values	Quadrivalent (30 µg) VLP Vaccine	Active Comparator (Fluarix)		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	206	199		
Units: mm^2				
geometric mean (confidence interval 95%)				
H1N1: Day 0	23.8 (20.4 to 27.7)	20.7 (17.7 to 24.2)		
H1N1: Day 21	38.7 (34.7 to 43.1)	48.9 (43.8 to 54.6)		
H3N2: Day 0	6.3 (5.1 to 7.9)	5.1 (4.1 to 6.3)		
H3N2: Day 21	21.9 (17.2 to 28.0)	13.2 (10.3 to 17.0)		
B/Colorado: Day 0	33.7 (28.1 to 40.3)	29.2 (24.3 to 35.1)		
B/Colorado: Day 21	54.1 (47.3 to 61.8)	61.0 (53.3 to 69.9)		
B/Phuket: Day 0	21.5 (17.7 to 26.0)	20.4 (16.8 to 24.8)		
B/Phuket: Day 21	38.8 (33.5 to 44.9)	50.2 (43.3 to 58.3)		

Statistical analyses

No statistical analyses for this end point

Secondary: Percentage of participants with seroconversion measured by SRH antibody response for each homologous strain

End point title	Percentage of participants with seroconversion measured by SRH antibody response for each homologous strain

End point description:

The percentage of participants in a given treatment group showing at least 50 % increase in GMA between Days 0 and 21 were measured using an SRH assay for homologous strains: H1N1 = A/Michigan/45/2015; H3N2 = A/Singapore/INFIMH-16-0019/2016; B/Colorado = B/Colorado/06/2017; B/Phuket = B/Phuket/3073/2013. IPP set.

End point type	Secondary

End point timeframe:

Day 0 (pre-vaccination) to Day 21

End point values	Quadrivalent (30 µg) VLP Vaccine	Active Comparator (Fluarix)		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	206	199		
Units: Percentage of participants				
number (confidence interval 95%)				
H1N1	34.0 (27.5 to 40.9)	55.3 (48.1 to 62.3)		
H3N2	45.1 (38.2 to 52.2)	34.7 (28.1 to 41.7)		
B/Colorado	32.0 (25.7 to 38.9)	49.7 (42.6 to 56.9)		
B/Phuket	35.9 (29.4 to 42.9)	52.3 (45.1 to 59.4)		

Statistical analyses

No statistical analyses for this end point

Secondary: Percentage of participants with seroprotection measured by SRH antibody response for each homologous strain

End point title	Percentage of participants with seroprotection measured by SRH antibody response for each homologous strain

End point description:

The percentage of participants in a given treatment group attaining an area ≥ 25 mm^2 following vaccination (Day 21) were measured using an SRH assay for homologous strains: H1N1 = A/Michigan/45/2015; H3N2 = A/Singapore/INFIMH-16-0019/2016; B/Colorado = B/Colorado/06/2017; B/Phuket = B/Phuket/3073/2013. IPP set.

End point type	Secondary

End point timeframe:

Baseline (Day 0), Day 21

End point values	Quadrivalent (30 µg) VLP Vaccine	Active Comparator (Fluarix)		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	206	199		
Units: Percentage of participants				
number (confidence interval 95%)				
H1N1: Day 0	56.8 (49.7 to 63.7)	51.8 (44.6 to 58.9)		
H1N1: Day 21	77.7 (71.4 to 83.2)	89.4 (84.3 to 93.3)		
H3N2: Day 0	27.7 (21.7 to 34.3)	23.6 (17.9 to 30.1)		
H3N2: Day 21	63.6 (56.6 to 70.2)	51.8 (44.6 to 58.9)		
B/Colorado: Day 0	71.8 (65.2 to 77.9)	69.8 (63.0 to 76.1)		
B/Colorado: Day 21	87.9 (82.6 to 92.0)	89.4 (84.3 to 93.3)		

B/Phuket: Day 0	63.6 (56.6 to 70.2)	60.8 (53.7 to 67.6)		
B/Phuket: Day 21	76.2 (69.8 to 81.9)	87.9 (82.6 to 92.1)		

Statistical analyses

No statistical analyses for this end point

Secondary: GMFR of SRH antibody response for each homologous strain

End point title	GMFR of SRH antibody response for each homologous strain

End point description:

GMFR, the geometric mean of the ratio of GMTs (Day 21/Day 0), was measured using an SRH assay for homologous strains: H1N1 = A/Michigan/45/2015; H3N2 = A/Singapore/INFIMH-16-0019/2016; B/Colorado = B/Colorado/06/2017; B/Phuket = B/Phuket/3073/2013. IPP set.

End point type	Secondary

End point timeframe:

Baseline (Day 0), Day 21

End point values	Quadrivalent (30 µg) VLP Vaccine	Active Comparator (Fluarix)		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	206	199		
Units: Ratio				
geometric mean (confidence interval 95%)				
H1N1	1.7 (1.6 to 1.8)	2.3 (2.1 to 2.5)		
H3N2	3.6 (2.9 to 4.4)	2.5 (2.0 to 3.1)		
B/Colorado	1.7 (1.5 to 1.9)	2.0 (1.8 to 2.2)		
B/Phuket	1.8 (1.6 to 2.1)	2.4 (2.2 to 2.7)		

Statistical analyses

No statistical analyses for this end point

Secondary: Number of participants with at least one solicited local and systemic reactions

End point title	Number of participants with at least one solicited local and systemic reactions

End point description:

Participants were monitored for both solicited local reactions (erythema, swelling, and pain at the injection site) and solicited systemic reactions (fever, headache, fatigue, muscle aches, joint aches, chills, a feeling of general discomfort, swelling in the axilla, and swelling in the neck) from the time of vaccination through Day 7. Any solicited local or systemic immediate complaint was also included. SAS. Participants that were non-compliant to protocol/GCP, as per investigator, were excluded from analysis.

End point type	Secondary

End point timeframe:
Day 0 (post-vaccination) to Day 7

End point values	Quadrivalent (30 µg) VLP Vaccine	Active Comparator (Fluarix)		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	6352	6366		
Units: Participants				
number (not applicable)				
Solicited Local Reactions	1954	1460		
Solicited Systemic Reactions	1682	1497		

Statistical analyses

No statistical analyses for this end point

Adverse events

Adverse events information	
Timeframe for reporting adverse events:	
Day 0 (post-vaccination) up to ~9 months	
Adverse event reporting additional description:	
SAS. Participants that were non-compliant to protocol/GCP, as per investigator, were excluded from analysis.	
Assessment type	Systematic

Dictionary used	
Dictionary name	MedDRA
Dictionary version	21.0

Reporting groups	
Reporting group title	Quadrivalent (30 µg) VLP Vaccine
Reporting group description:	
Participants received one IM injection of 0.5 mL of 30 µg/strain of the Quadrivalent VLP Influenza Vaccine on Day 0.	
Reporting group title	Active Comparator (Fluarix)
Reporting group description:	
Participants received one IM injection of 0.5 mL of 15 µg/strain of the Fluarix Quadrivalent® comparator vaccine on Day 0.	

Serious adverse events	Quadrivalent (30 µg) VLP Vaccine	Active Comparator (Fluarix)	
Total subjects affected by serious adverse events			
subjects affected / exposed	263 / 6352 (4.14%)	266 / 6366 (4.18%)	
number of deaths (all causes)	12	17	
number of deaths resulting from adverse events			
Neoplasms benign, malignant and unspecified (incl cysts and polyps)			
Acral lentiginous melanoma			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Adenocarcinoma			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Adenocarcinoma of colon			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	

Basal cell carcinoma			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
B-cell lymphoma			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Benign neoplasm of bladder			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Biliary neoplasm			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Bladder cancer			
subjects affected / exposed	1 / 6352 (0.02%)	2 / 6366 (0.03%)	
occurrences causally related to treatment / all	0 / 1	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Breast cancer			
subjects affected / exposed	1 / 6352 (0.02%)	5 / 6366 (0.08%)	
occurrences causally related to treatment / all	0 / 1	0 / 5	
deaths causally related to treatment / all	0 / 0	0 / 0	
Bronchial neoplasm			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Chronic lymphocytic leukaemia			
subjects affected / exposed	2 / 6352 (0.03%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Colon adenoma			

subjects affected / exposed	0 / 6352 (0.00%)	2 / 6366 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Colon cancer			
subjects affected / exposed	1 / 6352 (0.02%)	2 / 6366 (0.03%)	
occurrences causally related to treatment / all	0 / 1	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Endometrial cancer stage I			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Female reproductive neoplasm			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hepatic cancer metastatic			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 1	
Hepatocellular carcinoma			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Intraductal papillary mucinous neoplasm			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Invasive ductal breast carcinoma			
subjects affected / exposed	1 / 6352 (0.02%)	2 / 6366 (0.03%)	
occurrences causally related to treatment / all	0 / 1	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Laryngeal cancer			

subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Lung adenocarcinoma			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Lung neoplasm malignant			
subjects affected / exposed	4 / 6352 (0.06%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 4	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Lung squamous cell carcinoma stage III			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Lymphoplasmacytoid lymphoma/immunocytoma recurrent			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Malignant melanoma			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Malignant melanoma stage II			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Mesothelioma			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Metastatic malignant melanoma			

subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 1	
Neoplasm malignant			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Oesophageal adenocarcinoma			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pancreatic carcinoma			
subjects affected / exposed	1 / 6352 (0.02%)	2 / 6366 (0.03%)	
occurrences causally related to treatment / all	0 / 1	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Papillary thyroid cancer			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Parathyroid tumour benign			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
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 / 6352 (0.02%)	2 / 6366 (0.03%)	
occurrences causally related to treatment / all	0 / 1	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Prostate cancer			
subjects affected / exposed	3 / 6352 (0.05%)	6 / 6366 (0.09%)	
occurrences causally related to treatment / all	0 / 3	0 / 6	
deaths causally related to treatment / all	0 / 0	0 / 0	
Prostate cancer metastatic			

subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Rectal cancer			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Renal neoplasm			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
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 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Squamous cell carcinoma of lung			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
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 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Thyroid cancer recurrent			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Tongue neoplasm malignant stage unspecified			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Tonsil cancer			

subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Triple negative breast cancer			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Waldenstrom's macroglobulinaemia			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Vascular disorders			
Accelerated hypertension			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Angiodysplasia			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Aortic aneurysm			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Aortic dissection			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Aortic stenosis			
subjects affected / exposed	2 / 6352 (0.03%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Circulatory collapse			

subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Deep vein thrombosis			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Extremity necrosis			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
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 / 6352 (0.00%)	2 / 6366 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hypertensive crisis			
subjects affected / exposed	3 / 6352 (0.05%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 3	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hypertensive emergency			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hypotension			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 1	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Orthostatic hypotension			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Peripheral arterial occlusive disease			

subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Peripheral vascular disorder			
subjects affected / exposed	2 / 6352 (0.03%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Varicose vein			
subjects affected / exposed	0 / 6352 (0.00%)	2 / 6366 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Surgical and medical procedures			
Atrial appendage closure			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
General disorders and administration site conditions			
Chest pain			
subjects affected / exposed	3 / 6352 (0.05%)	4 / 6366 (0.06%)	
occurrences causally related to treatment / all	0 / 3	0 / 4	
deaths causally related to treatment / all	0 / 0	0 / 0	
Death			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 1	0 / 1	
Incarcerated hernia			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Malaise			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (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	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pain			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pyrexia			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Surgical failure			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Vascular stent stenosis			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Immune system disorders			
Anaphylactic reaction			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	1 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Drug hypersensitivity			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Reproductive system and breast disorders			
Benign prostatic hyperplasia			
subjects affected / exposed	1 / 6352 (0.02%)	2 / 6366 (0.03%)	
occurrences causally related to treatment / all	0 / 1	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	

Endometriosis			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Rectocele			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Uterine polyp			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Uterine prolapse			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Uterovaginal prolapse			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
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 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Vaginal prolapse			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Varicocele			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Respiratory, thoracic and mediastinal disorders			

Acute pulmonary oedema			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Acute respiratory distress syndrome			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Acute respiratory failure			
subjects affected / exposed	1 / 6352 (0.02%)	2 / 6366 (0.03%)	
occurrences causally related to treatment / all	0 / 1	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Asthma			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Bronchospasm			
subjects affected / exposed	2 / 6352 (0.03%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 2	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Chronic obstructive pulmonary disease			
subjects affected / exposed	5 / 6352 (0.08%)	6 / 6366 (0.09%)	
occurrences causally related to treatment / all	0 / 7	0 / 6	
deaths causally related to treatment / all	0 / 0	0 / 0	
Dyspnoea			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Dyspnoea exertional			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Emphysema			

subjects affected / exposed	2 / 6352 (0.03%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Haemoptysis			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pneumothorax			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pulmonary embolism			
subjects affected / exposed	4 / 6352 (0.06%)	6 / 6366 (0.09%)	
occurrences causally related to treatment / all	0 / 4	0 / 6	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pulmonary fibrosis			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 2	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pulmonary oedema			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Respiratory failure			
subjects affected / exposed	0 / 6352 (0.00%)	3 / 6366 (0.05%)	
occurrences causally related to treatment / all	0 / 0	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 2	
Psychiatric disorders			
Anxiety			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Chronic idiopathic pain syndrome			

subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Completed suicide			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 1	
Confusional state			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Delirium			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Disorientation			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Mental status changes			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Mood disorder due to a general medical condition			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Suicidal ideation			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Product issues			
Device dislocation			

subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hepatobiliary disorders			
Bile duct stenosis			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Bile duct stone			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cholecystitis			
subjects affected / exposed	0 / 6352 (0.00%)	2 / 6366 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cholecystitis acute			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cholecystitis chronic			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cholelithiasis			
subjects affected / exposed	2 / 6352 (0.03%)	3 / 6366 (0.05%)	
occurrences causally related to treatment / all	0 / 2	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cholestasis			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hepatic cirrhosis			

subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Sphincter of Oddi dysfunction			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Investigations			
Fibrin D dimer increased			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Injury, poisoning and procedural complications			
Acetabulum fracture			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Animal attack			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
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	2 / 6352 (0.03%)	4 / 6366 (0.06%)	
occurrences causally related to treatment / all	0 / 2	0 / 4	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cervical vertebral fracture			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 1	
Concussion			
subjects affected / exposed	0 / 6352 (0.00%)	2 / 6366 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	

Contusion			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Craniocerebral injury			
subjects affected / exposed	0 / 6352 (0.00%)	2 / 6366 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Fall			
subjects affected / exposed	1 / 6352 (0.02%)	3 / 6366 (0.05%)	
occurrences causally related to treatment / all	0 / 1	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 1	
Fascial rupture			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Femoral neck fracture			
subjects affected / exposed	0 / 6352 (0.00%)	2 / 6366 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Femur fracture			
subjects affected / exposed	4 / 6352 (0.06%)	2 / 6366 (0.03%)	
occurrences causally related to treatment / all	0 / 4	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Forearm fracture			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Foreign body aspiration			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Fractured coccyx			

subjects affected / exposed	2 / 6352 (0.03%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Fractured sacrum			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
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	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
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	3 / 6352 (0.05%)	2 / 6366 (0.03%)	
occurrences causally related to treatment / all	0 / 3	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Lacrimal structure injury			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Limb injury			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Lumbar vertebral fracture			
subjects affected / exposed	2 / 6352 (0.03%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 2	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Meniscus injury			
subjects affected / exposed	2 / 6352 (0.03%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pelvic fracture			

subjects affected / exposed	2 / 6352 (0.03%)	2 / 6366 (0.03%)	
occurrences causally related to treatment / all	0 / 2	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Postoperative ileus			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Procedural pain			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pubis fracture			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
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 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Rib fracture			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Road traffic accident			
subjects affected / exposed	2 / 6352 (0.03%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 2	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Spinal compression fracture			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Spinal fracture			

subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (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 / 6352 (0.02%)	2 / 6366 (0.03%)	
occurrences causally related to treatment / all	0 / 1	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Subdural haematoma			
subjects affected / exposed	2 / 6352 (0.03%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 2	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Tendon rupture			
subjects affected / exposed	2 / 6352 (0.03%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 2	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Tibia fracture			
subjects affected / exposed	1 / 6352 (0.02%)	3 / 6366 (0.05%)	
occurrences causally related to treatment / all	0 / 1	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Upper limb fracture			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Urinary retention postoperative			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
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	2 / 6352 (0.03%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Congenital, familial and genetic disorders			
Arteriovenous malformation			

subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hamartoma			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Syringomyelia			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cardiac disorders			
Acute coronary syndrome			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Acute myocardial infarction			
subjects affected / exposed	6 / 6352 (0.09%)	8 / 6366 (0.13%)	
occurrences causally related to treatment / all	0 / 6	0 / 8	
deaths causally related to treatment / all	0 / 1	0 / 0	
Angina pectoris			
subjects affected / exposed	3 / 6352 (0.05%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 3	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Angina unstable			
subjects affected / exposed	2 / 6352 (0.03%)	0 / 6366 (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 / 6352 (0.03%)	2 / 6366 (0.03%)	
occurrences causally related to treatment / all	0 / 2	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 1	
Arteriosclerosis coronary artery			

subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Atrial fibrillation			
subjects affected / exposed	11 / 6352 (0.17%)	12 / 6366 (0.19%)	
occurrences causally related to treatment / all	0 / 12	0 / 13	
deaths causally related to treatment / all	0 / 0	0 / 0	
Atrial flutter			
subjects affected / exposed	1 / 6352 (0.02%)	2 / 6366 (0.03%)	
occurrences causally related to treatment / all	0 / 1	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Bradycardia			
subjects affected / exposed	0 / 6352 (0.00%)	2 / 6366 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cardiac arrest			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 1	
Cardiac failure			
subjects affected / exposed	0 / 6352 (0.00%)	3 / 6366 (0.05%)	
occurrences causally related to treatment / all	0 / 0	0 / 5	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cardiac failure chronic			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 1	0 / 0	
Cardiac failure congestive			
subjects affected / exposed	5 / 6352 (0.08%)	2 / 6366 (0.03%)	
occurrences causally related to treatment / all	0 / 5	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 1	
Cardio-respiratory arrest			

subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 1	
Coronary artery disease			
subjects affected / exposed	5 / 6352 (0.08%)	3 / 6366 (0.05%)	
occurrences causally related to treatment / all	0 / 6	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Coronary artery occlusion			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Coronary artery stenosis			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Myocardial infarction			
subjects affected / exposed	5 / 6352 (0.08%)	3 / 6366 (0.05%)	
occurrences causally related to treatment / all	0 / 5	0 / 3	
deaths causally related to treatment / all	0 / 2	0 / 0	
Pericardial effusion			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pericarditis			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pulseless electrical activity			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 1	
Sinus arrest			

subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Sinus bradycardia			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Sinus node dysfunction			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Tachyarrhythmia			
subjects affected / exposed	0 / 6352 (0.00%)	2 / 6366 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Nervous system disorders			
Carotid artery stenosis			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cerebellar stroke			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cerebral haemorrhage			
subjects affected / exposed	1 / 6352 (0.02%)	2 / 6366 (0.03%)	
occurrences causally related to treatment / all	0 / 1	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cerebral infarction			
subjects affected / exposed	2 / 6352 (0.03%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cerebrovascular accident			

subjects affected / exposed	7 / 6352 (0.11%)	9 / 6366 (0.14%)	
occurrences causally related to treatment / all	0 / 7	0 / 9	
deaths causally related to treatment / all	0 / 1	0 / 2	
Chronic inflammatory demyelinating polyradiculoneuropathy			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Dementia			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Dementia Alzheimer's type			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 1	0 / 0	
Diabetic neuropathy			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
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 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hepatic encephalopathy			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hydrocephalus			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Ischaemic stroke			

subjects affected / exposed	0 / 6352 (0.00%)	2 / 6366 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
IVth nerve paralysis			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Loss of consciousness			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Neuropathy peripheral			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Parkinsonism			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Parkinson's disease			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Presyncope			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Sciatica			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Seizure			

subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Spondylitic myelopathy			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Syncope			
subjects affected / exposed	2 / 6352 (0.03%)	5 / 6366 (0.08%)	
occurrences causally related to treatment / all	0 / 2	0 / 5	
deaths causally related to treatment / all	0 / 0	0 / 0	
Transient global amnesia			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Transient ischaemic attack			
subjects affected / exposed	6 / 6352 (0.09%)	3 / 6366 (0.05%)	
occurrences causally related to treatment / all	0 / 6	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Vertebrobasilar insufficiency			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Blood and lymphatic system disorders			
Anaemia			
subjects affected / exposed	2 / 6352 (0.03%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Febrile neutropenia			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Haemorrhagic anaemia			

subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Iron deficiency anaemia			
subjects affected / exposed	0 / 6352 (0.00%)	2 / 6366 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Microcytic anaemia			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Splenic haematoma			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Ear and labyrinth disorders			
Deafness			
subjects affected / exposed	2 / 6352 (0.03%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Vertigo			
subjects affected / exposed	2 / 6352 (0.03%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 2	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Vertigo positional			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Vestibular disorder			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Eye disorders			

Amaurosis fugax			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Corneal degeneration			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Dacryostenosis acquired			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Gastrointestinal disorders			
Colitis			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Diverticulum			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Duodenal ulcer			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Gastric haemorrhage			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Gastric ulcer			
subjects affected / exposed	2 / 6352 (0.03%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Gastric ulcer haemorrhage			

subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Gastric ulcer perforation			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Gastritis			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Gastrointestinal haemorrhage			
subjects affected / exposed	3 / 6352 (0.05%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 3	0 / 0	
deaths causally related to treatment / all	0 / 1	0 / 0	
Gastrointestinal inflammation			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
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	2 / 6352 (0.03%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 2	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hiatus hernia			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Ileus			
subjects affected / exposed	2 / 6352 (0.03%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 2	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Incarcerated inguinal hernia			

subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Inguinal hernia			
subjects affected / exposed	3 / 6352 (0.05%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 3	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Intestinal obstruction			
subjects affected / exposed	0 / 6352 (0.00%)	2 / 6366 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Mesenteric arterial occlusion			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pancreatitis			
subjects affected / exposed	2 / 6352 (0.03%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 2	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pancreatitis acute			
subjects affected / exposed	0 / 6352 (0.00%)	3 / 6366 (0.05%)	
occurrences causally related to treatment / all	0 / 0	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Portal hypertensive gastropathy			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Proctitis ulcerative			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (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	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Rectal haemorrhage			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Small intestinal obstruction			
subjects affected / exposed	4 / 6352 (0.06%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 4	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Subileus			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Skin and subcutaneous tissue disorders			
Angioedema			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Renal and urinary disorders			
Acute kidney injury			
subjects affected / exposed	1 / 6352 (0.02%)	3 / 6366 (0.05%)	
occurrences causally related to treatment / all	0 / 1	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Bladder prolapse			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Glomerulonephritis			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Haematuria			

subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Nephrolithiasis			
subjects affected / exposed	2 / 6352 (0.03%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 2	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Renal colic			
subjects affected / exposed	1 / 6352 (0.02%)	2 / 6366 (0.03%)	
occurrences causally related to treatment / all	0 / 1	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Renal failure			
subjects affected / exposed	2 / 6352 (0.03%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Ureterolithiasis			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Urinary incontinence			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Endocrine disorders			
Goitre			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Musculoskeletal and connective tissue disorders			
Arthralgia			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	

Back pain			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Camptocormia			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Exostosis			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Haemarthrosis			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
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	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Intervertebral disc protrusion			
subjects affected / exposed	2 / 6352 (0.03%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 2	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Lateral patellar compression syndrome			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Lumbar spinal stenosis			
subjects affected / exposed	1 / 6352 (0.02%)	2 / 6366 (0.03%)	
occurrences causally related to treatment / all	0 / 1	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Musculoskeletal chest pain			

subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Musculoskeletal pain			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Osteitis			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Osteoarthritis			
subjects affected / exposed	9 / 6352 (0.14%)	9 / 6366 (0.14%)	
occurrences causally related to treatment / all	0 / 9	0 / 9	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pain in extremity			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pathological fracture			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Polymyalgia rheumatica			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Spinal column stenosis			
subjects affected / exposed	1 / 6352 (0.02%)	3 / 6366 (0.05%)	
occurrences causally related to treatment / all	0 / 1	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Spinal osteoarthritis			

subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Spondylolisthesis			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Synovial cyst			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Systemic scleroderma			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Infections and infestations			
Abscess			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Appendicitis			
subjects affected / exposed	0 / 6352 (0.00%)	2 / 6366 (0.03%)	
occurrences causally related to treatment / all	0 / 0	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Appendicitis perforated			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Bronchitis			
subjects affected / exposed	1 / 6352 (0.02%)	4 / 6366 (0.06%)	
occurrences causally related to treatment / all	0 / 1	0 / 4	
deaths causally related to treatment / all	0 / 0	0 / 0	
Bursitis infective			

subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cellulitis			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Diverticulitis			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Empyema			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Endocarditis			
subjects affected / exposed	1 / 6352 (0.02%)	2 / 6366 (0.03%)	
occurrences causally related to treatment / all	0 / 1	0 / 2	
deaths causally related to treatment / all	0 / 1	0 / 0	
Erysipelas			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Gastroenteritis			
subjects affected / exposed	2 / 6352 (0.03%)	2 / 6366 (0.03%)	
occurrences causally related to treatment / all	0 / 2	0 / 2	
deaths causally related to treatment / all	0 / 0	0 / 0	
Gastroenteritis Escherichia coli			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
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 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
H1N1 influenza			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Herpes zoster			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Infective exacerbation of chronic obstructive airways disease			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Influenza			
subjects affected / exposed	3 / 6352 (0.05%)	4 / 6366 (0.06%)	
occurrences causally related to treatment / all	0 / 3	0 / 4	
deaths causally related to treatment / all	0 / 0	0 / 0	
Localised infection			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Necrotising fasciitis staphylococcal			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Orchitis			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Osteomyelitis			

subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Osteomyelitis bacterial			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pharyngeal abscess			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pneumonia			
subjects affected / exposed	20 / 6352 (0.31%)	11 / 6366 (0.17%)	
occurrences causally related to treatment / all	0 / 20	0 / 12	
deaths causally related to treatment / all	0 / 1	0 / 1	
Pneumonia legionella			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pneumonia viral			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Postoperative wound infection			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pseudomonal sepsis			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pyelonephritis			

subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (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 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Sepsis			
subjects affected / exposed	4 / 6352 (0.06%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 4	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Sialoadenitis			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Sinusitis			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Staphylococcal infection			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Streptococcal bacteraemia			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
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	3 / 6352 (0.05%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 3	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Urosepsis			

subjects affected / exposed	3 / 6352 (0.05%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 3	0 / 0	
deaths causally related to treatment / all	0 / 1	0 / 0	
Metabolism and nutrition disorders			
Dehydration			
subjects affected / exposed	2 / 6352 (0.03%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 2	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Diabetes mellitus			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Gout			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hyperkalaemia			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hypoglycaemia			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hypokalaemia			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (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	1 / 6352 (0.02%)	3 / 6366 (0.05%)	
occurrences causally related to treatment / all	0 / 1	0 / 3	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hypovolaemia			

subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Type 2 diabetes mellitus			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences causally related to treatment / all	0 / 1	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	

Frequency threshold for reporting non-serious adverse events: 0 %

Non-serious adverse events	Quadrivalent (30 µg) VLP Vaccine	Active Comparator (Fluarix)	
Total subjects affected by non-serious adverse events			
subjects affected / exposed	88 / 6352 (1.39%)	138 / 6366 (2.17%)	
Neoplasms benign, malignant and unspecified (incl cysts and polyps)			
Adenocarcinoma of colon			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Basal cell carcinoma			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Benign neoplasm of thyroid gland			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Breast cancer stage I			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Chronic lymphocytic leukaemia			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Eye naevus			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Haemangioma of liver			

subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Malignant melanoma			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Squamous cell carcinoma			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	5	0	
Squamous cell carcinoma of skin			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Vascular disorders			
Aneurysm			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Aortic arteriosclerosis			
subjects affected / exposed	0 / 6352 (0.00%)	3 / 6366 (0.05%)	
occurrences (all)	0	3	
Aortic dilatation			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Aortic disorder			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Aortic stenosis			
subjects affected / exposed	2 / 6352 (0.03%)	0 / 6366 (0.00%)	
occurrences (all)	2	0	
Deep vein thrombosis			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Flushing			
subjects affected / exposed	1 / 6352 (0.02%)	3 / 6366 (0.05%)	
occurrences (all)	1	4	
Haematoma			
subjects affected / exposed	0 / 6352 (0.00%)	2 / 6366 (0.03%)	
occurrences (all)	0	2	

Hot flush			
subjects affected / exposed	2 / 6352 (0.03%)	0 / 6366 (0.00%)	
occurrences (all)	2	0	
Hypertension			
subjects affected / exposed	35 / 6352 (0.55%)	27 / 6366 (0.42%)	
occurrences (all)	35	27	
Hypertensive crisis			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Hypotension			
subjects affected / exposed	0 / 6352 (0.00%)	2 / 6366 (0.03%)	
occurrences (all)	0	2	
Microangiopathy			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Peripheral arterial occlusive disease			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Peripheral coldness			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences (all)	1	1	
Peripheral vascular disorder			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences (all)	1	1	
Thrombosis			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Varicose vein			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences (all)	1	1	
General disorders and administration site conditions			
Asthenia			
subjects affected / exposed	4 / 6352 (0.06%)	0 / 6366 (0.00%)	
occurrences (all)	4	0	
Axillary pain			

subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Chest discomfort			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Chest pain			
subjects affected / exposed	3 / 6352 (0.05%)	5 / 6366 (0.08%)	
occurrences (all)	3	5	
Chills			
subjects affected / exposed	15 / 6352 (0.24%)	15 / 6366 (0.24%)	
occurrences (all)	15	15	
Discomfort			
subjects affected / exposed	2 / 6352 (0.03%)	2 / 6366 (0.03%)	
occurrences (all)	2	2	
Face oedema			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Facial pain			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Fatigue			
subjects affected / exposed	26 / 6352 (0.41%)	32 / 6366 (0.50%)	
occurrences (all)	27	35	
Feeling hot			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Gait disturbance			
subjects affected / exposed	0 / 6352 (0.00%)	2 / 6366 (0.03%)	
occurrences (all)	0	2	
General physical health deterioration			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Hypothermia			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Induration			

subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Inflammation			
subjects affected / exposed	0 / 6352 (0.00%)	2 / 6366 (0.03%)	
occurrences (all)	0	2	
Influenza like illness			
subjects affected / exposed	8 / 6352 (0.13%)	11 / 6366 (0.17%)	
occurrences (all)	8	11	
Injected limb mobility decreased			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Injection site bruising			
subjects affected / exposed	2 / 6352 (0.03%)	4 / 6366 (0.06%)	
occurrences (all)	2	4	
Injection site erythema			
subjects affected / exposed	2 / 6352 (0.03%)	3 / 6366 (0.05%)	
occurrences (all)	2	3	
Injection site haematoma			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences (all)	1	1	
Injection site induration			
subjects affected / exposed	2 / 6352 (0.03%)	0 / 6366 (0.00%)	
occurrences (all)	2	0	
Injection site pain			
subjects affected / exposed	12 / 6352 (0.19%)	7 / 6366 (0.11%)	
occurrences (all)	12	7	
Injection site pruritus			
subjects affected / exposed	3 / 6352 (0.05%)	4 / 6366 (0.06%)	
occurrences (all)	3	4	
Injection site reaction			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Injection site swelling			
subjects affected / exposed	3 / 6352 (0.05%)	5 / 6366 (0.08%)	
occurrences (all)	3	5	
Injection site warmth			

subjects affected / exposed	4 / 6352 (0.06%)	2 / 6366 (0.03%)	
occurrences (all)	4	2	
Malaise			
subjects affected / exposed	11 / 6352 (0.17%)	6 / 6366 (0.09%)	
occurrences (all)	11	6	
Mass			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Oedema peripheral			
subjects affected / exposed	1 / 6352 (0.02%)	3 / 6366 (0.05%)	
occurrences (all)	1	3	
Pain			
subjects affected / exposed	4 / 6352 (0.06%)	2 / 6366 (0.03%)	
occurrences (all)	4	2	
Peripheral swelling			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences (all)	1	1	
Pyrexia			
subjects affected / exposed	9 / 6352 (0.14%)	15 / 6366 (0.24%)	
occurrences (all)	9	16	
Swelling			
subjects affected / exposed	5 / 6352 (0.08%)	6 / 6366 (0.09%)	
occurrences (all)	5	6	
Tenderness			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Vaccination site haematoma			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Vaccination site hypoaesthesia			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Vaccination site induration			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Vaccination site pain			

subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences (all)	1	1	
Vaccination site pruritus			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Immune system disorders			
Drug hypersensitivity			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Seasonal allergy			
subjects affected / exposed	2 / 6352 (0.03%)	3 / 6366 (0.05%)	
occurrences (all)	2	3	
Reproductive system and breast disorders			
Benign prostatic hyperplasia			
subjects affected / exposed	4 / 6352 (0.06%)	7 / 6366 (0.11%)	
occurrences (all)	4	7	
Erectile dysfunction			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences (all)	1	1	
Ovarian cyst			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Prostatomegaly			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Vulvovaginal pruritus			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Respiratory, thoracic and mediastinal disorders			
Allergic cough			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Asthma			
subjects affected / exposed	3 / 6352 (0.05%)	0 / 6366 (0.00%)	
occurrences (all)	3	0	
Atelectasis			

subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)		
occurrences (all)	0	1		
Chronic obstructive pulmonary disease				
subjects affected / exposed	8 / 6352 (0.13%)	5 / 6366 (0.08%)		
occurrences (all)	8	5		
Cough				
subjects affected / exposed	42 / 6352 (0.66%)	46 / 6366 (0.72%)		
occurrences (all)	42	47		
Dry throat				
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)		
occurrences (all)	1	0		
Dysphonia				
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)		
occurrences (all)	1	1		
Dyspnoea				
subjects affected / exposed	4 / 6352 (0.06%)	7 / 6366 (0.11%)		
occurrences (all)	4	7		
Dyspnoea exertional				
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)		
occurrences (all)	1	0		
Emphysema				
subjects affected / exposed	3 / 6352 (0.05%)	2 / 6366 (0.03%)		
occurrences (all)	3	2		
Epistaxis				
subjects affected / exposed	4 / 6352 (0.06%)	1 / 6366 (0.02%)		
occurrences (all)	4	1		
Nasal congestion				
subjects affected / exposed	29 / 6352 (0.46%)	30 / 6366 (0.47%)		
occurrences (all)	31	32		
Nasal discomfort				
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)		
occurrences (all)	0	1		
Oropharyngeal pain				
subjects affected / exposed	47 / 6352 (0.74%)	41 / 6366 (0.64%)		
occurrences (all)	47	42		

Pharyngeal oedema			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Pharyngeal paraesthesia			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Productive cough			
subjects affected / exposed	10 / 6352 (0.16%)	21 / 6366 (0.33%)	
occurrences (all)	10	22	
Pulmonary hypertension			
subjects affected / exposed	2 / 6352 (0.03%)	3 / 6366 (0.05%)	
occurrences (all)	2	3	
Pulmonary mass			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Pulmonary oedema			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Respiratory disorder			
subjects affected / exposed	16 / 6352 (0.25%)	17 / 6366 (0.27%)	
occurrences (all)	16	17	
Respiratory symptom			
subjects affected / exposed	0 / 6352 (0.00%)	2 / 6366 (0.03%)	
occurrences (all)	0	2	
Respiratory tract congestion			
subjects affected / exposed	4 / 6352 (0.06%)	0 / 6366 (0.00%)	
occurrences (all)	4	0	
Rhinitis allergic			
subjects affected / exposed	0 / 6352 (0.00%)	3 / 6366 (0.05%)	
occurrences (all)	0	3	
Rhinorrhoea			
subjects affected / exposed	41 / 6352 (0.65%)	56 / 6366 (0.88%)	
occurrences (all)	42	56	
Sinus congestion			
subjects affected / exposed	2 / 6352 (0.03%)	1 / 6366 (0.02%)	
occurrences (all)	2	1	

Sinus pain			
subjects affected / exposed	1 / 6352 (0.02%)	2 / 6366 (0.03%)	
occurrences (all)	1	2	
Sleep apnoea syndrome			
subjects affected / exposed	2 / 6352 (0.03%)	1 / 6366 (0.02%)	
occurrences (all)	2	1	
Sneezing			
subjects affected / exposed	31 / 6352 (0.49%)	25 / 6366 (0.39%)	
occurrences (all)	33	27	
Throat irritation			
subjects affected / exposed	2 / 6352 (0.03%)	2 / 6366 (0.03%)	
occurrences (all)	2	2	
Throat tightness			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Tonsillar cyst			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Tonsillar hypertrophy			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Upper respiratory tract congestion			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Upper-airway cough syndrome			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences (all)	1	1	
Wheezing			
subjects affected / exposed	4 / 6352 (0.06%)	1 / 6366 (0.02%)	
occurrences (all)	4	1	
Psychiatric disorders			
Abnormal dreams			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Aggression			

subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Anxiety			
subjects affected / exposed	5 / 6352 (0.08%)	1 / 6366 (0.02%)	
occurrences (all)	5	1	
Confusional state			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Depression			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences (all)	1	1	
Fear			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Initial insomnia			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Insomnia			
subjects affected / exposed	4 / 6352 (0.06%)	2 / 6366 (0.03%)	
occurrences (all)	4	2	
Listless			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Nervousness			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences (all)	1	1	
Restlessnes			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences (all)	1	2	
Sleep disorder			
subjects affected / exposed	2 / 6352 (0.03%)	0 / 6366 (0.00%)	
occurrences (all)	3	0	
Investigations			
Blood cholesterol increased			
subjects affected / exposed	0 / 6352 (0.00%)	3 / 6366 (0.05%)	
occurrences (all)	0	3	

Blood pressure increased			
subjects affected / exposed	13 / 6352 (0.20%)	9 / 6366 (0.14%)	
occurrences (all)	13	9	
Blood thyroid stimulating hormone increased			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Body temperature increased			
subjects affected / exposed	0 / 6352 (0.00%)	2 / 6366 (0.03%)	
occurrences (all)	0	2	
Cardiac imaging procedure abnormal			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Hepatic enzyme increased			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Thyroid hormones decreased			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Injury, poisoning and procedural complications			
Anaemia postoperative			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Animal bite			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Arthropod bite			
subjects affected / exposed	1 / 6352 (0.02%)	4 / 6366 (0.06%)	
occurrences (all)	1	4	
Arthropod sting			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences (all)	1	1	
Back injury			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Chest injury			

subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Contusion			
subjects affected / exposed	2 / 6352 (0.03%)	2 / 6366 (0.03%)	
occurrences (all)	2	3	
Epicondylitis			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Eye injury			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Face injury			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Facial bones fracture			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Fall			
subjects affected / exposed	1 / 6352 (0.02%)	7 / 6366 (0.11%)	
occurrences (all)	1	7	
Head injury			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Joint injury			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Laceration			
subjects affected / exposed	3 / 6352 (0.05%)	4 / 6366 (0.06%)	
occurrences (all)	3	4	
Ligament sprain			
subjects affected / exposed	3 / 6352 (0.05%)	0 / 6366 (0.00%)	
occurrences (all)	3	0	
Limb injury			
subjects affected / exposed	2 / 6352 (0.03%)	0 / 6366 (0.00%)	
occurrences (all)	2	0	
Meniscus injury			

subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Muscle contusion			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Muscle rupture			
subjects affected / exposed	2 / 6352 (0.03%)	0 / 6366 (0.00%)	
occurrences (all)	2	0	
Muscle strain			
subjects affected / exposed	5 / 6352 (0.08%)	3 / 6366 (0.05%)	
occurrences (all)	5	3	
Post-traumatic pain			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Procedural pain			
subjects affected / exposed	3 / 6352 (0.05%)	5 / 6366 (0.08%)	
occurrences (all)	3	6	
Pubis fracture			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Rib fracture			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences (all)	1	1	
Scratch			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Skin abrasion			
subjects affected / exposed	0 / 6352 (0.00%)	3 / 6366 (0.05%)	
occurrences (all)	0	3	
Spinal compression fracture			
subjects affected / exposed	1 / 6352 (0.02%)	2 / 6366 (0.03%)	
occurrences (all)	1	2	
Sunburn			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Synovial rupture			

subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Tendon rupture			
subjects affected / exposed	2 / 6352 (0.03%)	0 / 6366 (0.00%)	
occurrences (all)	3	0	
Tibia fracture			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Tongue injury			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Toxicity to various agents			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Wound			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences (all)	1	1	
Wound complication			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Wrist fracture			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Bone contusion			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Cardiac disorders			
Acute left ventricular failure			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences (all)	1	1	
Angina pectoris			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Aortic valve incompetence			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences (all)	1	1	

Aortic valve stenosis			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Arrhythmia			
subjects affected / exposed	2 / 6352 (0.03%)	2 / 6366 (0.03%)	
occurrences (all)	2	2	
Arteriosclerosis coronary artery			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Atrial fibrillation			
subjects affected / exposed	11 / 6352 (0.17%)	11 / 6366 (0.17%)	
occurrences (all)	11	11	
Atrial flutter			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences (all)	1	1	
Bradycardia			
subjects affected / exposed	3 / 6352 (0.05%)	0 / 6366 (0.00%)	
occurrences (all)	3	0	
Bundle branch block bilateral			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Cardiac dysfunction			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Cardiac failure congestive			
subjects affected / exposed	1 / 6352 (0.02%)	2 / 6366 (0.03%)	
occurrences (all)	1	2	
Cardiomegaly			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Cardiomyopathy			
subjects affected / exposed	0 / 6352 (0.00%)	2 / 6366 (0.03%)	
occurrences (all)	0	2	
Cardiovascular disorder			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	

Coronary artery disease			
subjects affected / exposed	3 / 6352 (0.05%)	5 / 6366 (0.08%)	
occurrences (all)	3	5	
Coronary artery stenosis			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Dilatation atrial			
subjects affected / exposed	0 / 6352 (0.00%)	2 / 6366 (0.03%)	
occurrences (all)	0	2	
Extrasystoles			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Hypertensive heart disease			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Left atrial dilatation			
subjects affected / exposed	0 / 6352 (0.00%)	2 / 6366 (0.03%)	
occurrences (all)	0	2	
Left ventricular dysfunction			
subjects affected / exposed	0 / 6352 (0.00%)	2 / 6366 (0.03%)	
occurrences (all)	0	2	
Left ventricular hypertrophy			
subjects affected / exposed	0 / 6352 (0.00%)	5 / 6366 (0.08%)	
occurrences (all)	0	5	
Mitral valve calcification			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Mitral valve incompetence			
subjects affected / exposed	3 / 6352 (0.05%)	4 / 6366 (0.06%)	
occurrences (all)	3	4	
Palpitations			
subjects affected / exposed	0 / 6352 (0.00%)	2 / 6366 (0.03%)	
occurrences (all)	0	2	
Pulmonary valve incompetence			
subjects affected / exposed	0 / 6352 (0.00%)	2 / 6366 (0.03%)	
occurrences (all)	0	2	

Right atrial dilatation			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Sinus node dysfunction			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Tachycardia			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences (all)	1	1	
Tricuspid valve incompetence			
subjects affected / exposed	1 / 6352 (0.02%)	5 / 6366 (0.08%)	
occurrences (all)	1	5	
Nervous system disorders			
Carotid artery stenosis			
subjects affected / exposed	2 / 6352 (0.03%)	1 / 6366 (0.02%)	
occurrences (all)	2	1	
Carpal tunnel syndrome			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences (all)	1	1	
Cerebral atrophy			
subjects affected / exposed	0 / 6352 (0.00%)	3 / 6366 (0.05%)	
occurrences (all)	0	3	
Cerebral microangiopathy			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Complex regional pain syndrome			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Dementia			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Dementia Alzheimer's type			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences (all)	1	1	
Dizziness			

subjects affected / exposed	19 / 6352 (0.30%)	24 / 6366 (0.38%)	
occurrences (all)	20	25	
Dysgeusia			
subjects affected / exposed	1 / 6352 (0.02%)	2 / 6366 (0.03%)	
occurrences (all)	1	2	
Facial paralysis			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Head discomfort			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Headache			
subjects affected / exposed	61 / 6352 (0.96%)	66 / 6366 (1.04%)	
occurrences (all)	73	71	
Hypoaesthesia			
subjects affected / exposed	2 / 6352 (0.03%)	1 / 6366 (0.02%)	
occurrences (all)	2	1	
Hypogeusia			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Memory impairment			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Migraine			
subjects affected / exposed	2 / 6352 (0.03%)	3 / 6366 (0.05%)	
occurrences (all)	2	3	
Movement disorder			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Myasthenia gravis			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Neuralgia			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences (all)	1	1	
Neuropathy peripheral			

subjects affected / exposed	0 / 6352 (0.00%)	2 / 6366 (0.03%)	
occurrences (all)	0	3	
Paraesthesia			
subjects affected / exposed	3 / 6352 (0.05%)	0 / 6366 (0.00%)	
occurrences (all)	3	0	
Parkinson's disease			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences (all)	1	1	
Polyneuropathy			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Posterior cortical atrophy			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Presyncope			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Radiculopathy			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Restless legs syndrome			
subjects affected / exposed	2 / 6352 (0.03%)	3 / 6366 (0.05%)	
occurrences (all)	2	3	
Sciatica			
subjects affected / exposed	2 / 6352 (0.03%)	2 / 6366 (0.03%)	
occurrences (all)	2	2	
Sinus headache			
subjects affected / exposed	2 / 6352 (0.03%)	2 / 6366 (0.03%)	
occurrences (all)	2	2	
Somnolence			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Syncope			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Tension headache			

subjects affected / exposed	2 / 6352 (0.03%)	0 / 6366 (0.00%)	
occurrences (all)	2	0	
Trigeminal neuralgia			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
VIth nerve paralysis			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Visual field defect			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences (all)	1	3	
White matter lesion			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Blood and lymphatic system disorders			
Anaemia			
subjects affected / exposed	3 / 6352 (0.05%)	2 / 6366 (0.03%)	
occurrences (all)	3	2	
Increased tendency to bruise			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Iron deficiency anemia			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Lymph node pain			
subjects affected / exposed	2 / 6352 (0.03%)	0 / 6366 (0.00%)	
occurrences (all)	2	0	
Lymphadenopathy			
subjects affected / exposed	3 / 6352 (0.05%)	1 / 6366 (0.02%)	
occurrences (all)	3	1	
Ear and labyrinth disorders			
Cerumen impaction			
subjects affected / exposed	2 / 6352 (0.03%)	0 / 6366 (0.00%)	
occurrences (all)	2	0	
Ear discomfort			

subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences (all)	1	1	
Ear pain			
subjects affected / exposed	7 / 6352 (0.11%)	7 / 6366 (0.11%)	
occurrences (all)	7	7	
Tinnitus			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Vertigo			
subjects affected / exposed	3 / 6352 (0.05%)	4 / 6366 (0.06%)	
occurrences (all)	3	4	
Vertigo positional			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Eye disorders			
Astigmatism			
subjects affected / exposed	0 / 6352 (0.00%)	2 / 6366 (0.03%)	
occurrences (all)	0	2	
Binocular eye movement disorder			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Blepharitis			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Blepharochalasis			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Blindness transient			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Cataract			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Chalazion			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	

Conjunctival haemorrhage			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Corneal warpage			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Dry age-related macular degeneration			
subjects affected / exposed	2 / 6352 (0.03%)	0 / 6366 (0.00%)	
occurrences (all)	2	0	
Dry eye			
subjects affected / exposed	2 / 6352 (0.03%)	1 / 6366 (0.02%)	
occurrences (all)	2	1	
Eye discharge			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Eye haemorrhage			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Eye inflammation			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Eye irritation			
subjects affected / exposed	2 / 6352 (0.03%)	0 / 6366 (0.00%)	
occurrences (all)	2	0	
Eye pain			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Eye pruritus			
subjects affected / exposed	0 / 6352 (0.00%)	2 / 6366 (0.03%)	
occurrences (all)	0	2	
Eye swelling			
subjects affected / exposed	0 / 6352 (0.00%)	2 / 6366 (0.03%)	
occurrences (all)	0	2	
Eyelid irritation			

subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)		
occurrences (all)	0	1		
Eyelid margin crusting				
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)		
occurrences (all)	0	1		
Eyelid oedema				
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)		
occurrences (all)	0	1		
Eyelid pain				
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)		
occurrences (all)	1	0		
Glaucoma				
subjects affected / exposed	2 / 6352 (0.03%)	1 / 6366 (0.02%)		
occurrences (all)	2	1		
Hyalosis asteroid				
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)		
occurrences (all)	0	1		
Keratitis				
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)		
occurrences (all)	0	1		
Lacrimation increased				
subjects affected / exposed	1 / 6352 (0.02%)	2 / 6366 (0.03%)		
occurrences (all)	1	2		
Macular degeneration				
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)		
occurrences (all)	0	1		
Macular fibrosis				
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)		
occurrences (all)	1	0		
Neovascular age-related macular degeneration				
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)		
occurrences (all)	1	0		
Ocular hyperaemia				
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)		
occurrences (all)	1	1		

Ocular rosacea			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Vision blurred			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences (all)	1	1	
Visual impairment			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Vitreous haemorrhage			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Gastrointestinal disorders			
Abdominal discomfort			
subjects affected / exposed	3 / 6352 (0.05%)	0 / 6366 (0.00%)	
occurrences (all)	3	0	
Abdominal distension			
subjects affected / exposed	2 / 6352 (0.03%)	1 / 6366 (0.02%)	
occurrences (all)	2	1	
Abdominal hernia			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences (all)	1	1	
Abdominal pain			
subjects affected / exposed	8 / 6352 (0.13%)	7 / 6366 (0.11%)	
occurrences (all)	8	7	
Abdominal pain lower			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Abdominal pain upper			
subjects affected / exposed	2 / 6352 (0.03%)	8 / 6366 (0.13%)	
occurrences (all)	2	10	
Anal incontinence			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Aphthous ulcer			

subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences (all)	1	1	
Barrett's oesophagus			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences (all)	1	1	
Cheilitis			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Chronic gastritis			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Constipation			
subjects affected / exposed	4 / 6352 (0.06%)	6 / 6366 (0.09%)	
occurrences (all)	4	6	
Diarrhoea			
subjects affected / exposed	50 / 6352 (0.79%)	53 / 6366 (0.83%)	
occurrences (all)	53	56	
Diverticulum			
subjects affected / exposed	1 / 6352 (0.02%)	3 / 6366 (0.05%)	
occurrences (all)	1	3	
Diverticulum intestinal			
subjects affected / exposed	1 / 6352 (0.02%)	2 / 6366 (0.03%)	
occurrences (all)	1	2	
Dry mouth			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Dyspepsia			
subjects affected / exposed	2 / 6352 (0.03%)	3 / 6366 (0.05%)	
occurrences (all)	2	3	
Dysphagia			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Enlarged uvula			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Faeces soft			

subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Flatulence			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Food poisoning			
subjects affected / exposed	1 / 6352 (0.02%)	2 / 6366 (0.03%)	
occurrences (all)	1	2	
Gastric ulcer			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Gastritis			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences (all)	1	1	
Gastrooesophageal reflux disease			
subjects affected / exposed	3 / 6352 (0.05%)	5 / 6366 (0.08%)	
occurrences (all)	3	5	
Haemorrhoidal haemorrhage			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Haemorrhoids			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Hiatus hernia			
subjects affected / exposed	2 / 6352 (0.03%)	2 / 6366 (0.03%)	
occurrences (all)	2	2	
Irritable bowel syndrome			
subjects affected / exposed	0 / 6352 (0.00%)	2 / 6366 (0.03%)	
occurrences (all)	0	2	
Large intestine polyp			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Lip erythema			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	2	
Lip pruritus			

subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	2	
Lip swelling			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Lip ulceration			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Malocclusion			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Nausea			
subjects affected / exposed	27 / 6352 (0.43%)	30 / 6366 (0.47%)	
occurrences (all)	27	32	
Oesophageal food impaction			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Oesophageal motility disorder			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Oesophagitis			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences (all)	1	1	
Oral pruritus			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Palatal oedema			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Rectal haemorrhage			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Salivary gland calculus			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Swollen tongue			

subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Tongue blistering			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Tooth impacted			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Tooth loss			
subjects affected / exposed	2 / 6352 (0.03%)	0 / 6366 (0.00%)	
occurrences (all)	2	0	
Toothache			
subjects affected / exposed	8 / 6352 (0.13%)	6 / 6366 (0.09%)	
occurrences (all)	8	6	
Vomiting			
subjects affected / exposed	13 / 6352 (0.20%)	13 / 6366 (0.20%)	
occurrences (all)	13	14	
Hepatobiliary disorders			
Biliary colic			
subjects affected / exposed	2 / 6352 (0.03%)	0 / 6366 (0.00%)	
occurrences (all)	2	0	
Cholangitis			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Cholangitis acute			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Cholecystitis acute			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Hepatic cirrhosis			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Hepatic steatosis			
subjects affected / exposed	0 / 6352 (0.00%)	2 / 6366 (0.03%)	
occurrences (all)	0	2	

Portal vein thrombosis			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Steatohepatitis			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Skin and subcutaneous tissue disorders			
Acne			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Blood blister			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Dermatitis			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Dermatitis allergic			
subjects affected / exposed	1 / 6352 (0.02%)	2 / 6366 (0.03%)	
occurrences (all)	1	2	
Dermatitis contact			
subjects affected / exposed	2 / 6352 (0.03%)	2 / 6366 (0.03%)	
occurrences (all)	2	2	
Diabetic foot			
subjects affected / exposed	2 / 6352 (0.03%)	0 / 6366 (0.00%)	
occurrences (all)	2	0	
Eczema			
subjects affected / exposed	3 / 6352 (0.05%)	3 / 6366 (0.05%)	
occurrences (all)	3	3	
Erythema			
subjects affected / exposed	1 / 6352 (0.02%)	5 / 6366 (0.08%)	
occurrences (all)	1	5	
Hyperhidrosis			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences (all)	1	1	
Ingrowing nail			

subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Ingrown hair			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Nail bed disorder			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Night sweats			
subjects affected / exposed	1 / 6352 (0.02%)	2 / 6366 (0.03%)	
occurrences (all)	1	2	
Pruritus			
subjects affected / exposed	9 / 6352 (0.14%)	8 / 6366 (0.13%)	
occurrences (all)	10	9	
Pruritus generalised			
subjects affected / exposed	3 / 6352 (0.05%)	3 / 6366 (0.05%)	
occurrences (all)	3	4	
Rash			
subjects affected / exposed	12 / 6352 (0.19%)	8 / 6366 (0.13%)	
occurrences (all)	12	8	
Rash generalised			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Skin discolouration			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Skin ulcer			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Stasis dermatitis			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Urticaria			
subjects affected / exposed	2 / 6352 (0.03%)	2 / 6366 (0.03%)	
occurrences (all)	2	2	
Renal and urinary disorders			

Acute kidney injury			
subjects affected / exposed	2 / 6352 (0.03%)	0 / 6366 (0.00%)	
occurrences (all)	2	0	
Bladder discomfort			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Calculus urinary			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Chronic kidney disease			
subjects affected / exposed	3 / 6352 (0.05%)	4 / 6366 (0.06%)	
occurrences (all)	3	4	
Hypertensive nephropathy			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Hypertonic bladder			
subjects affected / exposed	2 / 6352 (0.03%)	0 / 6366 (0.00%)	
occurrences (all)	2	0	
Micturition urgency			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Nephrolithiasis			
subjects affected / exposed	1 / 6352 (0.02%)	2 / 6366 (0.03%)	
occurrences (all)	1	2	
Nephrotic syndrome			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Pollakiuria			
subjects affected / exposed	0 / 6352 (0.00%)	3 / 6366 (0.05%)	
occurrences (all)	0	3	
Renal cyst			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Renal disorder			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	

Urinary incontinence			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Urinary tract obstruction			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Lower urinary tract symptoms			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Endocrine disorders			
Goitre			
subjects affected / exposed	0 / 6352 (0.00%)	2 / 6366 (0.03%)	
occurrences (all)	0	2	
Hyperthyroidism			
subjects affected / exposed	2 / 6352 (0.03%)	1 / 6366 (0.02%)	
occurrences (all)	2	1	
Hypothyroidism			
subjects affected / exposed	3 / 6352 (0.05%)	4 / 6366 (0.06%)	
occurrences (all)	3	4	
Musculoskeletal and connective tissue disorders			
Arthralgia			
subjects affected / exposed	25 / 6352 (0.39%)	30 / 6366 (0.47%)	
occurrences (all)	25	30	
Arthritis			
subjects affected / exposed	4 / 6352 (0.06%)	1 / 6366 (0.02%)	
occurrences (all)	4	1	
Back pain			
subjects affected / exposed	20 / 6352 (0.31%)	26 / 6366 (0.41%)	
occurrences (all)	21	27	
Bone pain			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Bursitis			
subjects affected / exposed	3 / 6352 (0.05%)	3 / 6366 (0.05%)	
occurrences (all)	3	3	
Coccydynia			

subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Dupuytren's contracture			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Facet joint syndrome			
subjects affected / exposed	0 / 6352 (0.00%)	2 / 6366 (0.03%)	
occurrences (all)	0	2	
Flank pain			
subjects affected / exposed	2 / 6352 (0.03%)	0 / 6366 (0.00%)	
occurrences (all)	2	0	
Groin pain			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Intervertebral disc degeneration			
subjects affected / exposed	1 / 6352 (0.02%)	2 / 6366 (0.03%)	
occurrences (all)	1	2	
Intervertebral disc protrusion			
subjects affected / exposed	3 / 6352 (0.05%)	0 / 6366 (0.00%)	
occurrences (all)	3	0	
Joint swelling			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences (all)	1	1	
Limb discomfort			
subjects affected / exposed	2 / 6352 (0.03%)	1 / 6366 (0.02%)	
occurrences (all)	2	1	
Lumbar spinal stenosis			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Muscle spasms			
subjects affected / exposed	9 / 6352 (0.14%)	3 / 6366 (0.05%)	
occurrences (all)	9	3	
Muscle tightness			
subjects affected / exposed	3 / 6352 (0.05%)	1 / 6366 (0.02%)	
occurrences (all)	3	1	
Muscular weakness			

subjects affected / exposed	2 / 6352 (0.03%)	0 / 6366 (0.00%)	
occurrences (all)	2	0	
Musculoskeletal chest pain			
subjects affected / exposed	1 / 6352 (0.02%)	3 / 6366 (0.05%)	
occurrences (all)	1	3	
Musculoskeletal pain			
subjects affected / exposed	11 / 6352 (0.17%)	12 / 6366 (0.19%)	
occurrences (all)	11	12	
Musculoskeletal stiffness			
subjects affected / exposed	1 / 6352 (0.02%)	5 / 6366 (0.08%)	
occurrences (all)	1	5	
Myalgia			
subjects affected / exposed	32 / 6352 (0.50%)	35 / 6366 (0.55%)	
occurrences (all)	34	35	
Myosclerosis			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Neck mass			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Neck pain			
subjects affected / exposed	10 / 6352 (0.16%)	10 / 6366 (0.16%)	
occurrences (all)	11	10	
Nuchal rigidity			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Osteoarthritis			
subjects affected / exposed	7 / 6352 (0.11%)	12 / 6366 (0.19%)	
occurrences (all)	7	12	
Osteopenia			
subjects affected / exposed	2 / 6352 (0.03%)	2 / 6366 (0.03%)	
occurrences (all)	2	2	
Osteoporosis			
subjects affected / exposed	6 / 6352 (0.09%)	4 / 6366 (0.06%)	
occurrences (all)	6	4	
Osteosclerosis			

subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Pain in extremity			
subjects affected / exposed	14 / 6352 (0.22%)	11 / 6366 (0.17%)	
occurrences (all)	14	11	
Periarthritis			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Plantar fasciitis			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences (all)	1	1	
Polyarthritis			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Polymyalgia rheumatica			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Rheumatoid arthritis			
subjects affected / exposed	2 / 6352 (0.03%)	1 / 6366 (0.02%)	
occurrences (all)	2	1	
Rotator cuff syndrome			
subjects affected / exposed	0 / 6352 (0.00%)	2 / 6366 (0.03%)	
occurrences (all)	0	2	
Spinal column stenosis			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Spinal osteoarthritis			
subjects affected / exposed	3 / 6352 (0.05%)	1 / 6366 (0.02%)	
occurrences (all)	3	1	
Spondylitis			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Spondylolisthesis			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Synovial cyst			

subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Temporomandibular joint syndrome			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Tendonitis			
subjects affected / exposed	2 / 6352 (0.03%)	0 / 6366 (0.00%)	
occurrences (all)	2	0	
Torticollis			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Vertebral foraminal stenosis			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Infections and infestations			
Acute sinusitis			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences (all)	1	1	
Adenovirus infection			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Borrelia infection			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Bronchitis			
subjects affected / exposed	12 / 6352 (0.19%)	8 / 6366 (0.13%)	
occurrences (all)	12	8	
Cellulitis			
subjects affected / exposed	1 / 6352 (0.02%)	3 / 6366 (0.05%)	
occurrences (all)	1	3	
Conjunctivitis			
subjects affected / exposed	2 / 6352 (0.03%)	3 / 6366 (0.05%)	
occurrences (all)	2	3	
Cystitis			
subjects affected / exposed	2 / 6352 (0.03%)	6 / 6366 (0.09%)	
occurrences (all)	2	6	

Cystitis escherichia			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Diverticulitis			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Ear infection			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Erysipelas			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences (all)	1	1	
Eye infection			
subjects affected / exposed	1 / 6352 (0.02%)	2 / 6366 (0.03%)	
occurrences (all)	1	2	
Eye infection bacterial			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Folliculitis			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Gastroenteritis			
subjects affected / exposed	6 / 6352 (0.09%)	8 / 6366 (0.13%)	
occurrences (all)	6	8	
Gastroenteritis viral			
subjects affected / exposed	3 / 6352 (0.05%)	1 / 6366 (0.02%)	
occurrences (all)	3	1	
Gastrointestinal infection			
subjects affected / exposed	1 / 6352 (0.02%)	2 / 6366 (0.03%)	
occurrences (all)	1	2	
Gastrointestinal viral infection			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Genital infection fungal			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	

Gingivitis			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences (all)	1	1	
Haemorrhagic fever with renal syndrome			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Helicobacter infection			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Hepatitis C			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Herpes simplex			
subjects affected / exposed	1 / 6352 (0.02%)	3 / 6366 (0.05%)	
occurrences (all)	1	3	
Herpes zoster			
subjects affected / exposed	5 / 6352 (0.08%)	5 / 6366 (0.08%)	
occurrences (all)	5	5	
Hordeolum			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Infected bite			
subjects affected / exposed	0 / 6352 (0.00%)	2 / 6366 (0.03%)	
occurrences (all)	0	2	
Infection			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Influenza			
subjects affected / exposed	3 / 6352 (0.05%)	13 / 6366 (0.20%)	
occurrences (all)	3	14	
Laryngitis			
subjects affected / exposed	2 / 6352 (0.03%)	0 / 6366 (0.00%)	
occurrences (all)	2	0	
Localised infection			

subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Lower respiratory tract infection			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences (all)	1	1	
Nasal herpes			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Nasopharyngitis			
subjects affected / exposed	88 / 6352 (1.39%)	76 / 6366 (1.19%)	
occurrences (all)	88	77	
Onychomycosis			
subjects affected / exposed	1 / 6352 (0.02%)	2 / 6366 (0.03%)	
occurrences (all)	1	2	
Oral candidiasis			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Oral herpes			
subjects affected / exposed	5 / 6352 (0.08%)	3 / 6366 (0.05%)	
occurrences (all)	5	3	
Otitis media			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Periodontitis			
subjects affected / exposed	2 / 6352 (0.03%)	1 / 6366 (0.02%)	
occurrences (all)	2	1	
Pertussis			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Pharyngitis			
subjects affected / exposed	4 / 6352 (0.06%)	0 / 6366 (0.00%)	
occurrences (all)	4	0	
Pilonidal cyst			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Pneumonia			

subjects affected / exposed	1 / 6352 (0.02%)	4 / 6366 (0.06%)	
occurrences (all)	1	4	
Respiratory tract infection			
subjects affected / exposed	8 / 6352 (0.13%)	5 / 6366 (0.08%)	
occurrences (all)	8	5	
Respiratory tract infection viral			
subjects affected / exposed	2 / 6352 (0.03%)	0 / 6366 (0.00%)	
occurrences (all)	2	0	
Rhinitis			
subjects affected / exposed	9 / 6352 (0.14%)	6 / 6366 (0.09%)	
occurrences (all)	9	6	
Root canal infection			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Sinusitis			
subjects affected / exposed	13 / 6352 (0.20%)	9 / 6366 (0.14%)	
occurrences (all)	13	9	
Sinusitis bacterial			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Streptococcal infection			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Tinea infection			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Tooth abscess			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences (all)	1	1	
Tooth infection			
subjects affected / exposed	2 / 6352 (0.03%)	1 / 6366 (0.02%)	
occurrences (all)	2	1	
Upper respiratory tract infection			
subjects affected / exposed	52 / 6352 (0.82%)	47 / 6366 (0.74%)	
occurrences (all)	53	48	
Urinary tract infection			

subjects affected / exposed	15 / 6352 (0.24%)	13 / 6366 (0.20%)	
occurrences (all)	15	13	
Vaginal infection			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Varicella zoster virus infection			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Viral infection			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Wound infection			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Metabolism and nutrition disorders			
Dehydration			
subjects affected / exposed	3 / 6352 (0.05%)	0 / 6366 (0.00%)	
occurrences (all)	3	0	
Dyslipidaemia			
subjects affected / exposed	3 / 6352 (0.05%)	2 / 6366 (0.03%)	
occurrences (all)	3	2	
Glucose tolerance impaired			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences (all)	1	1	
Gout			
subjects affected / exposed	0 / 6352 (0.00%)	5 / 6366 (0.08%)	
occurrences (all)	0	5	
Hypercalcaemia			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Hypercholesterolaemia			
subjects affected / exposed	8 / 6352 (0.13%)	5 / 6366 (0.08%)	
occurrences (all)	8	5	
Hyperglycaemia			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences (all)	1	1	

Hyperlipidaemia			
subjects affected / exposed	5 / 6352 (0.08%)	2 / 6366 (0.03%)	
occurrences (all)	5	2	
Hypoglycaemia			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	3	
Hypokalaemia			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Iron deficiency			
subjects affected / exposed	2 / 6352 (0.03%)	0 / 6366 (0.00%)	
occurrences (all)	2	0	
Ketoacidosis			
subjects affected / exposed	1 / 6352 (0.02%)	0 / 6366 (0.00%)	
occurrences (all)	1	0	
Lactose intolerance			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Lipid metabolism disorder			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences (all)	1	1	
Type 2 diabetes mellitus			
subjects affected / exposed	4 / 6352 (0.06%)	8 / 6366 (0.13%)	
occurrences (all)	4	8	
Vitamin B12 deficiency			
subjects affected / exposed	0 / 6352 (0.00%)	1 / 6366 (0.02%)	
occurrences (all)	0	1	
Vitamin D deficiency			
subjects affected / exposed	1 / 6352 (0.02%)	1 / 6366 (0.02%)	
occurrences (all)	1	1	

More information

Substantial protocol amendments (globally)

Were there any global substantial amendments to the protocol? Yes

Date	Amendment
24 July 2018	The purpose of Amendment No. 01 was to add precision to Protocol version 1.1 around key elements of the protocol: • Clarification of the testing to be applied for primary, secondary, and exploratory outcomes; • Clarification of the rationale for comparator vaccine choice; • Clarification of the statistical analysis plan; • Specification of the needle length required for obese subjects (body mass index greater than 30 kg/m^2).

Notes:

Interruptions (globally)

Were there any global interruptions to the trial? No

Limitations and caveats

None reported