Clinical trial results:
A Phase 2, Randomized, Multicenter, Placebo-Controlled, Double-Blind, Parallel-Group Study, with an Open-Label Extension to Evaluate the Efficacy, Safety, and Pharmacokinetics of E5501 in Subjects with Chronic Hepatitis C Virus Related Thrombocytopenia who are Potential Candidates for Antiviral Treatment

Summary

EudraCT number	2010-024479-20
Trial protocol	DE BG
Global end of trial date	01 May 2014

Results information

Result version number	v2 (current)
This version publication date	25 March 2016
First version publication date	05 August 2015
Version creation reason	• Correction of full data set Results are being revised due to training issue with our staff and to reconcile the results to ensure consistency with ClinicalTrials.gov results.

Trial information

Trial identification

Sponsor protocol code	E5501-G000-203

Additional study identifiers

ISRCTN number	-
ClinicalTrials.gov id (NCT number)	NCT01355289
WHO universal trial number (UTN)	-

Notes:

Sponsors

Sponsor organisation name	Eisai
Sponsor organisation address	155 Tice Boulevard, Woodcliff Lake, United States, 07677
Public contact	Medical Information, Eisai Limited, +44 08456761400, LmedInfo@eisai.net
Scientific contact	Medical Information, Eisai Limited, +44 08456761400, LmedInfo@eisai.net

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	01 May 2014
Is this the analysis of the primary completion data?	No

Global end of trial reached?	Yes
Global end of trial date	01 May 2014
Was the trial ended prematurely?	No

Notes:

General information about the trial

Main objective of the trial:

To evaluate the efficacy of E5501 by measuring platelet response in subjects with chronic hepatitis C virus (HCV)-related thrombocytopenia who require antiviral treatment

Protection of trial subjects:

This study was conducted in accordance with standard operating procedures (SOPs) of the sponsor (or designee), which are designed to ensure adherence to Good Clinical Practice (GCP) guidelines as required by the following:
- Principles of the World Medical Association Declaration of Helsinki (World Medical Association, 2008)
- International Conference on Harmonisation (ICH) E6 Guideline for GCP (CPMP/ICH/135/95) of the European Agency for the Evaluation of Medicinal
Products, Committee for Proprietary Medicinal Products, International Conference on Harmonisation of Pharmaceuticals for Human Use
- Title 21 of the United States (US) Code of Federal Regulations (US 21 CFR) regarding clinical studies, including Part 50 and Part 56 concerning informed subject consent and Institutional Review Board (IRB) regulations and applicable sections of US 21 CFR Part 312
- European Good Clinical Practice Directive 2005/28/EC and Clinical Trial Directive 2001/20/EC for studies conducted within any European Union (EU) country. All suspected unexpected serious adverse reactions were reported, as required, to the Competent Authorities of all involved EU member states.
- Article 14, Paragraph 3, and Article 80-2 of the Pharmaceutical Affairs Law (Law No. 145, 1960) for studies conducted in Japan, in addition to Japan’s GCP Subject Information and Informed Consent.

Background therapy: -

Evidence for comparator: -

Actual start date of recruitment	10 November 2011
Long term follow-up planned	No
Independent data monitoring committee (IDMC) involvement?	Yes

Notes:

Population of trial subjects

Subjects enrolled per country

Country: Number of subjects enrolled	Bulgaria: 3
Country: Number of subjects enrolled	France: 3
Country: Number of subjects enrolled	Germany: 5
Country: Number of subjects enrolled	Israel: 9
Country: Number of subjects enrolled	United States: 45

Worldwide total number of subjects	65
EEA total number of subjects	11

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)	60
From 65 to 84 years	5
85 years and over	0

Subject disposition

Recruitment

Recruitment details: -

Pre-assignment

Screening details:

The Screening Period encompassed 14 days ±7 days. Prerandomization assessments took place in all participants who had provided informed consent.

Period 1

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

Arms

Are arms mutually exclusive?	Yes
Arm title	Placebo (Core Study)

Arm description:

Placebo, was administered orally, once daily for up to 21 days.

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

Dosage and administration details:

Once daily.

Arm title	Avatrombopag 10 mg (Core Study)

Arm description:

Avatrombopag 10 mg, was administered orally, once daily, preferably with food for up to 21 days.

Arm type	Active comparator
Investigational medicinal product name	Avatrombopag
Investigational medicinal product code	E5501
Other name	
Pharmaceutical forms	Tablet
Routes of administration	Oral use

Dosage and administration details:

10 mg, tablet form, orally, once daily.

Arm title	Avatrombopag 20 mg (Core Study)

Arm description:

Avatrombopag 20 mg, was administered orally, once daily, preferably with food for up to 21 days.

Arm type	Active comparator
Investigational medicinal product name	Avatrombopag
Investigational medicinal product code	E5501
Other name	
Pharmaceutical forms	Tablet
Routes of administration	Oral use

Dosage and administration details:
20 mg, tablet form, orally, once daily.

Arm title	Avatrombopag 30 mg (Core Study)

Arm description:
Avatrombopag 30 mg, was administered orally, once daily, preferably with food for up to 21 days.

Arm type	Active comparator
Investigational medicinal product name	Avatrombopag
Investigational medicinal product code	E5501
Other name	
Pharmaceutical forms	Tablet
Routes of administration	Oral use

Dosage and administration details:
30 mg, tablet form, orally, once daily.

Number of subjects in period 1	Placebo (Core Study)	Avatrombopag 10 mg (Core Study)	Avatrombopag 20 mg (Core Study)
Started	17	16	18
Completed	16	16	18
Not completed	1	0	0
Adverse event, non-fatal	1	-	-
Not specified	-	-	-
Inadequate therapeutic effect	-	-	-

Number of subjects in period 1	Avatrombopag 30 mg (Core Study)
Started	14
Completed	12
Not completed	2
Adverse event, non-fatal	-
Not specified	1
Inadequate therapeutic effect	1

Period 2

Period 2 title	Open Label Extension (OLE)
Is this the baseline period?	No
Allocation method	Non-randomised - controlled
Blinding used	Not blinded

Arms

Arm title	Avatrombopag (Open Label Extension)

Arm description:

Avatrombopag was initiated at a dose of 20 mg, once daily in the open label extension (OLE) period. The avatrombopag dose was titrated up or down in accordance with their individual response within the range of a minimum of 5 mg and a maximum of 50 mg for up to 48 weeks.

Arm type	Experimental
Investigational medicinal product name	Avatrombopag
Investigational medicinal product code	E5501
Other name	
Pharmaceutical forms	Tablet
Routes of administration	Oral use

Dosage and administration details:

5-50 mg, tablet form, orally, once daily.

Number of subjects in period 2	Avatrombopag (Open Label Extension)
Started	62
Completed	28
Not completed	34
Adverse event, non-fatal	1
Not specified	3
Lost to follow-up	1
Lack of efficacy	29

Baseline characteristics

Reporting groups	
Reporting group title	Placebo (Core Study)
Reporting group description:	
Placebo, was administered orally, once daily for up to 21 days.	
Reporting group title	Avatrombopag 10 mg (Core Study)
Reporting group description:	
Avatrombopag 10 mg, was administered orally, once daily, preferably with food for up to 21 days.	
Reporting group title	Avatrombopag 20 mg (Core Study)
Reporting group description:	
Avatrombopag 20 mg, was administered orally, once daily, preferably with food for up to 21 days.	
Reporting group title	Avatrombopag 30 mg (Core Study)
Reporting group description:	
Avatrombopag 30 mg, was administered orally, once daily, preferably with food for up to 21 days.	

Reporting group values	Placebo (Core Study)	Avatrombopag 10 mg (Core Study)	Avatrombopag 20 mg (Core Study)
Number of subjects	17	16	18
Age categorical Units: Subjects			
In utero Preterm newborn infants (gestational age < 37 wks) Newborns (0-27 days) Infants and toddlers (28 days-23 months) Children (2-11 years) Adolescents (12-17 years) Adults (18-64 years) From 65-84 years 85 years and over			
Age continuous Units: years			
arithmetic mean	50.2	55.3	54.9
standard deviation	± 7.96	± 8.06	± 7.38
Gender categorical Units: Subjects			
Female	3	4	5
Male	14	12	13

Reporting group values	Avatrombopag 30 mg (Core Study)	Total	
Number of subjects	14	65	
Age categorical Units: Subjects			
In utero		0	
Preterm newborn infants (gestational age < 37 wks)		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 65-84 years		0	
85 years and over		0	
Age continuous			
Units: years			
arithmetic mean	53.6		
standard deviation	± 7.26	-	
Gender categorical			
Units: Subjects			
Female	5	17	
Male	9	48	

End points

End points reporting groups

Reporting group title	Placebo (Core Study)
Reporting group description:	
Placebo, was administered orally, once daily for up to 21 days.	
Reporting group title	Avatrombopag 10 mg (Core Study)
Reporting group description:	
Avatrombopag 10 mg, was administered orally, once daily, preferably with food for up to 21 days.	
Reporting group title	Avatrombopag 20 mg (Core Study)
Reporting group description:	
Avatrombopag 20 mg, was administered orally, once daily, preferably with food for up to 21 days.	
Reporting group title	Avatrombopag 30 mg (Core Study)
Reporting group description:	
Avatrombopag 30 mg, was administered orally, once daily, preferably with food for up to 21 days.	
Reporting group title	Avatrombopag (Open Label Extension)
Reporting group description:	
Avatrombopag was initiated at a dose of 20 mg, once daily in the open label extension (OLE) period. The avatrombopag dose was titrated up or down in accordance with their individual response within the range of a minimum of 5 mg and a maximum of 50 mg for up to 48 weeks.	
Subject analysis set title	Full Analysis Set
Subject analysis set type	Full analysis
Subject analysis set description:	
The full analysis set was the group of randomized participants.	

Primary: Number of participants who achieved Platelet Response (greater than or equal to 100 x 10^9/L) by Day 21 of Treatment Period A1 of Core Study

End point title	Number of participants who achieved Platelet Response (greater than or equal to 100 x 10^9/L) by Day 21 of Treatment Period A1 of Core Study
End point description:	
A responder was defined as a participant having a platelet count of greater than or equal to 100x10^9/L by Day 21 starting from an average baseline platelet count of greater than 20 x 10^9/L to less than or equal to 70 x 10^9/L.	
End point type	Primary
End point timeframe:	
Baseline to Day 21	

End point values	Placebo (Core Study)	Avatrombopag 10 mg (Core Study)	Avatrombopag 20 mg (Core Study)	Avatrombopag 30 mg (Core Study)
Subject group type	Reporting group	Reporting group	Reporting group	Reporting group
Number of subjects analysed	17	16	18	14
Units: Participants				
number (not applicable)				
Yes	1	6	12	9
No	16	10	6	5

Statistical analyses

Statistical analysis title	Difference of response rate (10 mg) vs placebo
Comparison groups	Avatrombopag 10 mg (Core Study) v Placebo (Core Study)
Number of subjects included in analysis	33
Analysis specification	Pre-specified
Analysis type	superiority
P-value	= 0.0236
Method	Cochran-Mantel-Haenszel
Parameter estimate	Median difference (final values)
Point estimate	31.62
Confidence interval	
level	95 %
sides	2-sided
lower limit	5.39
upper limit	57.84

Statistical analysis title	Difference of response rate (20 mg) vs placebo
Comparison groups	Placebo (Core Study) v Avatrombopag 20 mg (Core Study)
Number of subjects included in analysis	35
Analysis specification	Pre-specified
Analysis type	superiority
P-value	= 0.0003
Method	Cochran-Mantel-Haenszel
Parameter estimate	Median difference (final values)
Point estimate	60.78
Confidence interval	
level	95 %
sides	2-sided
lower limit	36.3
upper limit	85.27

Statistical analysis title	Difference of response rate (30 mg) vs placebo
Comparison groups	Placebo (Core Study) v Avatrombopag 30 mg (Core Study)

Number of subjects included in analysis	31
Analysis specification	Pre-specified
Analysis type	superiority
P-value	= 0.0003
Method	Cochran-Mantel-Haenszel
Parameter estimate	Median difference (final values)
Point estimate	58.4
Confidence interval	
level	95 %
sides	2-sided
lower limit	30.92
upper limit	85.88

Secondary: Change from Baseline of Local Platelet Count by Visit during Treatment Period A1 of Core Study

End point title	Change from Baseline of Local Platelet Count by Visit during Treatment Period A1 of Core Study

End point description:

Missing platelet counts were imputed using last observation carried forward (LOCF) approach for subjects who achieved platelet response at prior visits.

End point type	Secondary

End point timeframe:

Day 7 and Day 14

End point values	Placebo (Core Study)	Avatrombopag 10 mg (Core Study)	Avatrombopag 20 mg (Core Study)	Avatrombopag 30 mg (Core Study)
Subject group type	Reporting group	Reporting group	Reporting group	Reporting group
Number of subjects analysed	17	16	18	14
Units: x10^9/L				
arithmetic mean (standard deviation)				
Day 7	-0.1 (± 7.15)	19.8 (± 17.59)	26.5 (± 22.06)	30.9 (± 37.65)
Day 14	-0.2 (± 13.79)	29.2 (± 18.32)	57.2 (± 31.39)	55.4 (± 37.47)

Statistical analyses

No statistical analyses for this end point

Secondary: Number of Participants who achieved Platelet Count greater than 30 X 10^9/L from Baseline to Day 21 during Treatment Period A1 of Core Study

End point title	Number of Participants who achieved Platelet Count greater than 30 X 10^9/L from Baseline to Day 21 during Treatment Period A1 of Core Study

End point description:

Blood draws were taken to monitor platelet counts.

End point type	Secondary

End point timeframe:

Baseline to Day 21

End point values	Placebo (Core Study)	Avatrombopag 10 mg (Core Study)	Avatrombopag 20 mg (Core Study)	Avatrombopag 30 mg (Core Study)
Subject group type	Reporting group	Reporting group	Reporting group	Reporting group
Number of subjects analysed	17	16	18	14
Units: Participants				
number (not applicable)				
Yes	1	9	16	11
No	16	7	2	3

Statistical analyses

No statistical analyses for this end point

Secondary: Number of Participants Who Initiated Antiviral Treatment by Day 21 of Period A1 of Core Study

End point title	Number of Participants Who Initiated Antiviral Treatment by Day 21 of Period A1 of Core Study

End point description:

Blood draws were taken to monitor platelet counts during the first 21 days of study treatment. When a platelet count of greater than or equal to 100 X 10^9/L was attained, antiviral treatment was initiated.

End point type	Secondary

End point timeframe:

Baseline to Day 21

End point values	Placebo (Core Study)	Avatrombopag 10 mg (Core Study)	Avatrombopag 20 mg (Core Study)	Avatrombopag 30 mg (Core Study)
Subject group type	Reporting group	Reporting group	Reporting group	Reporting group
Number of subjects analysed	17	16	18	14
Units: Participants				
number (not applicable)				
Yes	1	6	13	9
No	16	10	5	5

Statistical analyses

No statistical analyses for this end point

Adverse events

Adverse events information

Timeframe for reporting adverse events:

For each participant, treatment emergent adverse events were collected from the first day of administration of study drug up to 30 days after the last dose of study drug or up to approximately 2.5 years.

Adverse event reporting additional description:

Safety Analysis Set was used which combines data of the Core and Extension Phase and includes subjects who had at least 1 dose of avatrombopag. Placebo treatment arm was excluded since not part of study design for the Extension Phase.

Assessment type	Systematic

Dictionary used

Dictionary name	MedDRA
Dictionary version	16.0

Reporting groups

Reporting group title	Avatrombopag (Open Label Extension)

Reporting group description:

During the core study, participants were administered a fixed dose of Avatrombopag 10mg, 20mg, or 30mg,orally, once daily, preferably with food for upto 21 days. The participants initiated Open label extension with Avontrombopag 20 mg, once daily. The avotrambopag dose was titrated up or down in accordance with their individual response, within the range of a minimum of 5mg and a maximum of 50 mg for up to 48 weeks.

Reporting group title	Placebo (Core Study)

Reporting group description:

Placebo, was given orally for up to 21 days once daily.

Reporting group title	Avatrombopag 10 mg (Core Study)

Reporting group description:

Avatrombopag 10 mg, was administered orally, once daily, preferably with food for up to 21 days.

Reporting group title	Avatrombopag 20 mg (Core Study)

Reporting group description:

Avatrombopag 20 mg, was administered orally, once daily, preferably with food for up to 21 days.

Reporting group title	Avatrombopag 30 mg (Core Study)

Reporting group description:

Avatrombopag 30 mg, was administered orally, once daily, preferably with food for up to 21 days.

Serious adverse events	Avatrombopag (Open Label Extension)	Placebo (Core Study)	Avatrombopag 10 mg (Core Study)
Total subjects affected by serious adverse events			
subjects affected / exposed	13 / 64 (20.31%)	0 / 17 (0.00%)	1 / 16 (6.25%)
number of deaths (all causes)	1	0	0
number of deaths resulting from adverse events	0	0	0
Nervous system disorders			
Paraesthesia			

subjects affected / exposed	1 / 64 (1.56%)	0 / 17 (0.00%)	0 / 16 (0.00%)
occurrences causally related to treatment / all	1 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Blood and lymphatic system disorders			
Anaemia			
subjects affected / exposed	2 / 64 (3.13%)	0 / 17 (0.00%)	0 / 16 (0.00%)
occurrences causally related to treatment / all	0 / 2	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Leukopenia			
subjects affected / exposed	1 / 64 (1.56%)	0 / 17 (0.00%)	0 / 16 (0.00%)
occurrences causally related to treatment / all	0 / 2	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Neutropenia			
subjects affected / exposed	2 / 64 (3.13%)	0 / 17 (0.00%)	0 / 16 (0.00%)
occurrences causally related to treatment / all	0 / 2	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pancytopenia			
subjects affected / exposed	1 / 64 (1.56%)	0 / 17 (0.00%)	0 / 16 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Thrombocytopenia			
subjects affected / exposed	2 / 64 (3.13%)	0 / 17 (0.00%)	1 / 16 (6.25%)
occurrences causally related to treatment / all	0 / 2	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Gastrointestinal disorders			
Abdominal pain			
subjects affected / exposed	1 / 64 (1.56%)	0 / 17 (0.00%)	0 / 16 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hepatobiliary disorders			
Hepatic mass			
subjects affected / exposed	1 / 64 (1.56%)	0 / 17 (0.00%)	0 / 16 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0

Renal and urinary disorders			
Haematuria			
subjects affected / exposed	1 / 64 (1.56%)	0 / 17 (0.00%)	0 / 16 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Renal failure acute			
subjects affected / exposed	1 / 64 (1.56%)	0 / 17 (0.00%)	0 / 16 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Infections and infestations			
Gastroenteritis			
subjects affected / exposed	1 / 64 (1.56%)	0 / 17 (0.00%)	0 / 16 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Urinary tract infection			
subjects affected / exposed	1 / 64 (1.56%)	0 / 17 (0.00%)	0 / 16 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Metabolism and nutrition disorders			
Hyperglycaemia			
subjects affected / exposed	1 / 64 (1.56%)	0 / 17 (0.00%)	0 / 16 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hyperuricaemia			
subjects affected / exposed	1 / 64 (1.56%)	0 / 17 (0.00%)	0 / 16 (0.00%)
occurrences causally related to treatment / all	1 / 1	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0

Serious adverse events	Avatrombopag 20 mg (Core Study)	Avatrombopag 30 mg (Core Study)	
Total subjects affected by serious adverse events			
subjects affected / exposed	0 / 18 (0.00%)	1 / 14 (7.14%)	
number of deaths (all causes)	0	0	
number of deaths resulting from adverse events	0	0	
Nervous system disorders			
Paraesthesia			

subjects affected / exposed	0 / 18 (0.00%)	1 / 14 (7.14%)	
occurrences causally related to treatment / all	0 / 0	1 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	
Blood and lymphatic system disorders			
Anaemia			
subjects affected / exposed	0 / 18 (0.00%)	0 / 14 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Leukopenia			
subjects affected / exposed	0 / 18 (0.00%)	0 / 14 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Neutropenia			
subjects affected / exposed	0 / 18 (0.00%)	0 / 14 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Pancytopenia			
subjects affected / exposed	0 / 18 (0.00%)	0 / 14 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Thrombocytopenia			
subjects affected / exposed	0 / 18 (0.00%)	0 / 14 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Gastrointestinal disorders			
Abdominal pain			
subjects affected / exposed	0 / 18 (0.00%)	0 / 14 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hepatobiliary disorders			
Hepatic mass			
subjects affected / exposed	0 / 18 (0.00%)	0 / 14 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	

Renal and urinary disorders			
Haematuria			
subjects affected / exposed	0 / 18 (0.00%)	0 / 14 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Renal failure acute			
subjects affected / exposed	0 / 18 (0.00%)	0 / 14 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Infections and infestations			
Gastroenteritis			
subjects affected / exposed	0 / 18 (0.00%)	0 / 14 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Urinary tract infection			
subjects affected / exposed	0 / 18 (0.00%)	0 / 14 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Metabolism and nutrition disorders			
Hyperglycaemia			
subjects affected / exposed	0 / 18 (0.00%)	0 / 14 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hyperuricaemia			
subjects affected / exposed	0 / 18 (0.00%)	0 / 14 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	

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

Non-serious adverse events	Avatrombopag (Open Label Extension)	Placebo (Core Study)	Avatrombopag 10 mg (Core Study)
Total subjects affected by non-serious adverse events			
subjects affected / exposed	55 / 64 (85.94%)	6 / 17 (35.29%)	11 / 16 (68.75%)
Vascular disorders			

Hot flush			
subjects affected / exposed	0 / 64 (0.00%)	0 / 17 (0.00%)	1 / 16 (6.25%)
occurrences (all)	0	0	1
Pallor			
subjects affected / exposed	0 / 64 (0.00%)	0 / 17 (0.00%)	0 / 16 (0.00%)
occurrences (all)	0	0	0
General disorders and administration site conditions			
Asthenia			
subjects affected / exposed	8 / 64 (12.50%)	0 / 17 (0.00%)	1 / 16 (6.25%)
occurrences (all)	9	0	2
Chills			
subjects affected / exposed	13 / 64 (20.31%)	1 / 17 (5.88%)	3 / 16 (18.75%)
occurrences (all)	14	1	3
Fatigue			
subjects affected / exposed	16 / 64 (25.00%)	0 / 17 (0.00%)	1 / 16 (6.25%)
occurrences (all)	17	0	1
Influenza like illness			
subjects affected / exposed	11 / 64 (17.19%)	2 / 17 (11.76%)	1 / 16 (6.25%)
occurrences (all)	13	2	1
Injection site erythema			
subjects affected / exposed	6 / 64 (9.38%)	0 / 17 (0.00%)	1 / 16 (6.25%)
occurrences (all)	7	0	1
Irritability			
subjects affected / exposed	7 / 64 (10.94%)	0 / 17 (0.00%)	0 / 16 (0.00%)
occurrences (all)	7	0	0
Oedema peripheral			
subjects affected / exposed	6 / 64 (9.38%)	0 / 17 (0.00%)	0 / 16 (0.00%)
occurrences (all)	6	0	0
Pyrexia			
subjects affected / exposed	9 / 64 (14.06%)	1 / 17 (5.88%)	0 / 16 (0.00%)
occurrences (all)	13	1	0
Chest discomfort			
subjects affected / exposed	0 / 64 (0.00%)	0 / 17 (0.00%)	1 / 16 (6.25%)
occurrences (all)	0	0	1
Feeling cold			

subjects affected / exposed	0 / 64 (0.00%)	0 / 17 (0.00%)	0 / 16 (0.00%)
occurrences (all)	0	0	0
Feeling of body temperature change			
subjects affected / exposed	0 / 64 (0.00%)	0 / 17 (0.00%)	0 / 16 (0.00%)
occurrences (all)	0	0	0
Injection site pruritus			
subjects affected / exposed	0 / 64 (0.00%)	0 / 17 (0.00%)	0 / 16 (0.00%)
occurrences (all)	0	0	0
Pain			
subjects affected / exposed	0 / 64 (0.00%)	0 / 17 (0.00%)	0 / 16 (0.00%)
occurrences (all)	0	0	0
Reproductive system and breast disorders			
Spontaneous penile erection			
subjects affected / exposed	0 / 64 (0.00%)	0 / 17 (0.00%)	1 / 16 (6.25%)
occurrences (all)	0	0	1
Respiratory, thoracic and mediastinal disorders			
Cough			
subjects affected / exposed	9 / 64 (14.06%)	0 / 17 (0.00%)	0 / 16 (0.00%)
occurrences (all)	10	0	0
Dyspnoea			
subjects affected / exposed	4 / 64 (6.25%)	0 / 17 (0.00%)	0 / 16 (0.00%)
occurrences (all)	4	0	0
Dyspnoea exertional			
subjects affected / exposed	4 / 64 (6.25%)	0 / 17 (0.00%)	1 / 16 (6.25%)
occurrences (all)	5	0	1
Epistaxis			
subjects affected / exposed	6 / 64 (9.38%)	0 / 17 (0.00%)	1 / 16 (6.25%)
occurrences (all)	7	0	1
Productive cough			
subjects affected / exposed	0 / 64 (0.00%)	0 / 17 (0.00%)	0 / 16 (0.00%)
occurrences (all)	0	0	0
Respiratory tract congestion			
subjects affected / exposed	0 / 64 (0.00%)	0 / 17 (0.00%)	0 / 16 (0.00%)
occurrences (all)	0	0	0
Upper-airway cough syndrome			

subjects affected / exposed	0 / 64 (0.00%)	0 / 17 (0.00%)	0 / 16 (0.00%)
occurrences (all)	0	0	0
Psychiatric disorders			
Depression			
subjects affected / exposed	5 / 64 (7.81%)	0 / 17 (0.00%)	0 / 16 (0.00%)
occurrences (all)	6	0	0
Insomnia			
subjects affected / exposed	13 / 64 (20.31%)	1 / 17 (5.88%)	1 / 16 (6.25%)
occurrences (all)	13	1	1
Anxiety			
subjects affected / exposed	0 / 64 (0.00%)	1 / 17 (5.88%)	1 / 16 (6.25%)
occurrences (all)	0	1	1
Food aversion			
subjects affected / exposed	0 / 64 (0.00%)	0 / 17 (0.00%)	1 / 16 (6.25%)
occurrences (all)	0	0	0
Mood altered			
subjects affected / exposed	0 / 64 (0.00%)	0 / 17 (0.00%)	0 / 16 (0.00%)
occurrences (all)	0	0	0
Sleep disorder			
subjects affected / exposed	0 / 64 (0.00%)	0 / 17 (0.00%)	1 / 16 (6.25%)
occurrences (all)	0	0	1
Investigations			
Staphylococcus test positive			
subjects affected / exposed	0 / 64 (0.00%)	0 / 17 (0.00%)	0 / 16 (0.00%)
occurrences (all)	0	0	0
Weight decreased			
subjects affected / exposed	0 / 64 (0.00%)	1 / 17 (5.88%)	0 / 16 (0.00%)
occurrences (all)	0	1	0
Injury, poisoning and procedural complications			
Burns first degree			
subjects affected / exposed	0 / 64 (0.00%)	0 / 17 (0.00%)	1 / 16 (6.25%)
occurrences (all)	0	0	1
Tooth fracture			
subjects affected / exposed	0 / 64 (0.00%)	0 / 17 (0.00%)	0 / 16 (0.00%)
occurrences (all)	0	0	0
Congenital, familial and genetic disorders			

Congenital lymphoedema			
subjects affected / exposed	0 / 64 (0.00%)	0 / 17 (0.00%)	1 / 16 (6.25%)
occurrences (all)	0	0	1
Cardiac disorders			
Palpitations			
subjects affected / exposed	0 / 64 (0.00%)	1 / 17 (5.88%)	0 / 16 (0.00%)
occurrences (all)	0	1	0
Tachycardia			
subjects affected / exposed	0 / 64 (0.00%)	0 / 17 (0.00%)	0 / 16 (0.00%)
occurrences (all)	0	0	0
Nervous system disorders			
Dizziness			
subjects affected / exposed	4 / 64 (6.25%)	1 / 17 (5.88%)	0 / 16 (0.00%)
occurrences (all)	4	1	0
Headache			
subjects affected / exposed	12 / 64 (18.75%)	1 / 17 (5.88%)	0 / 16 (0.00%)
occurrences (all)	16	1	0
Disturbance in attention			
subjects affected / exposed	0 / 64 (0.00%)	0 / 17 (0.00%)	1 / 16 (6.25%)
occurrences (all)	0	0	1
Dysgeusia			
subjects affected / exposed	0 / 64 (0.00%)	0 / 17 (0.00%)	0 / 16 (0.00%)
occurrences (all)	0	0	0
Hypoaesthesia			
subjects affected / exposed	0 / 64 (0.00%)	0 / 17 (0.00%)	0 / 16 (0.00%)
occurrences (all)	0	0	0
Mental impairment			
subjects affected / exposed	0 / 64 (0.00%)	0 / 17 (0.00%)	0 / 16 (0.00%)
occurrences (all)	0	0	0
Sciatica			
subjects affected / exposed	0 / 64 (0.00%)	0 / 17 (0.00%)	0 / 16 (0.00%)
occurrences (all)	0	0	0
Somnolence			
subjects affected / exposed	0 / 64 (0.00%)	0 / 17 (0.00%)	0 / 16 (0.00%)
occurrences (all)	0	0	0
Syncope			

subjects affected / exposed	0 / 64 (0.00%)	0 / 17 (0.00%)	0 / 16 (0.00%)
occurrences (all)	0	0	0
Blood and lymphatic system disorders			
Anaemia			
subjects affected / exposed	21 / 64 (32.81%)	0 / 17 (0.00%)	2 / 16 (12.50%)
occurrences (all)	26	0	3
Leukopenia			
subjects affected / exposed	14 / 64 (21.88%)	0 / 17 (0.00%)	1 / 16 (6.25%)
occurrences (all)	14	0	2
Lymphopenia			
subjects affected / exposed	5 / 64 (7.81%)	0 / 17 (0.00%)	0 / 16 (0.00%)
occurrences (all)	6	0	0
Neutropenia			
subjects affected / exposed	20 / 64 (31.25%)	0 / 17 (0.00%)	2 / 16 (12.50%)
occurrences (all)	24	0	2
Ear and labyrinth disorders			
Vertigo			
subjects affected / exposed	0 / 64 (0.00%)	0 / 17 (0.00%)	1 / 16 (6.25%)
occurrences (all)	0	0	1
Gastrointestinal disorders			
Abdominal pain			
subjects affected / exposed	6 / 64 (9.38%)	0 / 17 (0.00%)	0 / 16 (0.00%)
occurrences (all)	6	0	0
Abdominal pain upper			
subjects affected / exposed	5 / 64 (7.81%)	1 / 17 (5.88%)	1 / 16 (6.25%)
occurrences (all)	5	1	1
Ascites			
subjects affected / exposed	6 / 64 (9.38%)	0 / 17 (0.00%)	0 / 16 (0.00%)
occurrences (all)	6	0	0
Diarrhoea			
subjects affected / exposed	11 / 64 (17.19%)	0 / 17 (0.00%)	1 / 16 (6.25%)
occurrences (all)	12	0	1
Dyspepsia			
subjects affected / exposed	4 / 64 (6.25%)	0 / 17 (0.00%)	2 / 16 (12.50%)
occurrences (all)	4	0	2
Haemorrhoids			

subjects affected / exposed	4 / 64 (6.25%)	0 / 17 (0.00%)	0 / 16 (0.00%)
occurrences (all)	6	0	0
Nausea			
subjects affected / exposed	20 / 64 (31.25%)	0 / 17 (0.00%)	3 / 16 (18.75%)
occurrences (all)	22	0	3
Vomiting			
subjects affected / exposed	8 / 64 (12.50%)	0 / 17 (0.00%)	0 / 16 (0.00%)
occurrences (all)	10	0	0
Abdominal distension			
subjects affected / exposed	0 / 64 (0.00%)	1 / 17 (5.88%)	0 / 16 (0.00%)
occurrences (all)	0	1	0
Anal pruritus			
subjects affected / exposed	0 / 64 (0.00%)	0 / 17 (0.00%)	0 / 16 (0.00%)
occurrences (all)	0	0	0
Aphthous stomatitis			
subjects affected / exposed	0 / 64 (0.00%)	1 / 17 (5.88%)	0 / 16 (0.00%)
occurrences (all)	0	1	0
Constipation			
subjects affected / exposed	0 / 64 (0.00%)	1 / 17 (5.88%)	1 / 16 (6.25%)
occurrences (all)	0	1	1
Flatulence			
subjects affected / exposed	0 / 64 (0.00%)	1 / 17 (5.88%)	1 / 16 (6.25%)
occurrences (all)	0	1	1
Proctalgia			
subjects affected / exposed	0 / 64 (0.00%)	0 / 17 (0.00%)	0 / 16 (0.00%)
occurrences (all)	0	0	0
Hepatobiliary disorders			
Liver tenderness			
subjects affected / exposed	0 / 64 (0.00%)	0 / 17 (0.00%)	1 / 16 (6.25%)
occurrences (all)	0	0	1
Skin and subcutaneous tissue disorders			
Pruritus			
subjects affected / exposed	15 / 64 (23.44%)	0 / 17 (0.00%)	4 / 16 (25.00%)
occurrences (all)	15	0	4
Rash			

subjects affected / exposed	12 / 64 (18.75%)	2 / 17 (11.76%)	1 / 16 (6.25%)
occurrences (all)	14	2	1
Rash macular			
subjects affected / exposed	0 / 64 (0.00%)	0 / 17 (0.00%)	0 / 16 (0.00%)
occurrences (all)	0	0	0
Renal and urinary disorders			
Haematuria			
subjects affected / exposed	0 / 64 (0.00%)	0 / 17 (0.00%)	1 / 16 (6.25%)
occurrences (all)	0	0	1
Musculoskeletal and connective tissue disorders			
Arthralgia			
subjects affected / exposed	0 / 64 (0.00%)	0 / 17 (0.00%)	0 / 16 (0.00%)
occurrences (all)	0	0	0
Muscle tightness			
subjects affected / exposed	0 / 64 (0.00%)	1 / 17 (5.88%)	0 / 16 (0.00%)
occurrences (all)	0	1	0
Muscular weakness			
subjects affected / exposed	0 / 64 (0.00%)	0 / 17 (0.00%)	0 / 16 (0.00%)
occurrences (all)	0	0	0
Infections and infestations			
Nasopharyngitis			
subjects affected / exposed	4 / 64 (6.25%)	0 / 17 (0.00%)	0 / 16 (0.00%)
occurrences (all)	4	0	0
Upper respiratory tract infection			
subjects affected / exposed	4 / 64 (6.25%)	0 / 17 (0.00%)	2 / 16 (12.50%)
occurrences (all)	5	0	2
Abscess limb			
subjects affected / exposed	0 / 64 (0.00%)	0 / 17 (0.00%)	0 / 16 (0.00%)
occurrences (all)	0	0	0
Cellulitis			
subjects affected / exposed	0 / 64 (0.00%)	0 / 17 (0.00%)	0 / 16 (0.00%)
occurrences (all)	0	0	0
Gastroenteritis			
subjects affected / exposed	0 / 64 (0.00%)	0 / 17 (0.00%)	0 / 16 (0.00%)
occurrences (all)	0	0	0
Metabolism and nutrition disorders			

Hyperuricaemia			
subjects affected / exposed	4 / 64 (6.25%)	1 / 17 (5.88%)	0 / 16 (0.00%)
occurrences (all)	5	1	0
Hypertriglyceridaemia			
subjects affected / exposed	0 / 64 (0.00%)	0 / 17 (0.00%)	0 / 16 (0.00%)
occurrences (all)	0	0	0
Hypokalaemia			
subjects affected / exposed	0 / 64 (0.00%)	1 / 17 (5.88%)	0 / 16 (0.00%)
occurrences (all)	0	1	0

Non-serious adverse events	Avatrombopag 20 mg (Core Study)	Avatrombopag 30 mg (Core Study)	
Total subjects affected by non-serious adverse events			
subjects affected / exposed	12 / 18 (66.67%)	11 / 14 (78.57%)	
Vascular disorders			
Hot flush			
subjects affected / exposed	0 / 18 (0.00%)	0 / 14 (0.00%)	
occurrences (all)	0	0	
Pallor			
subjects affected / exposed	1 / 18 (5.56%)	0 / 14 (0.00%)	
occurrences (all)	1	0	
General disorders and administration site conditions			
Asthenia			
subjects affected / exposed	2 / 18 (11.11%)	1 / 14 (7.14%)	
occurrences (all)	2	1	
Chills			
subjects affected / exposed	3 / 18 (16.67%)	1 / 14 (7.14%)	
occurrences (all)	3	2	
Fatigue			
subjects affected / exposed	3 / 18 (16.67%)	2 / 14 (14.29%)	
occurrences (all)	4	2	
Influenza like illness			
subjects affected / exposed	4 / 18 (22.22%)	3 / 14 (21.43%)	
occurrences (all)	4	3	
Injection site erythema			
subjects affected / exposed	1 / 18 (5.56%)	0 / 14 (0.00%)	
occurrences (all)	1	0	

Irritability			
subjects affected / exposed	1 / 18 (5.56%)	1 / 14 (7.14%)	
occurrences (all)	1	1	
Oedema peripheral			
subjects affected / exposed	1 / 18 (5.56%)	0 / 14 (0.00%)	
occurrences (all)	1	0	
Pyrexia			
subjects affected / exposed	2 / 18 (11.11%)	2 / 14 (14.29%)	
occurrences (all)	2	2	
Chest discomfort			
subjects affected / exposed	0 / 18 (0.00%)	0 / 14 (0.00%)	
occurrences (all)	0	0	
Feeling cold			
subjects affected / exposed	1 / 18 (5.56%)	0 / 14 (0.00%)	
occurrences (all)	1	0	
Feeling of body temperature change			
subjects affected / exposed	0 / 18 (0.00%)	1 / 14 (7.14%)	
occurrences (all)	0	1	
Injection site pruritus			
subjects affected / exposed	1 / 18 (5.56%)	0 / 14 (0.00%)	
occurrences (all)	1	0	
Pain			
subjects affected / exposed	1 / 18 (5.56%)	0 / 14 (0.00%)	
occurrences (all)	1	0	
Reproductive system and breast disorders			
Spontaneous penile erection			
subjects affected / exposed	0 / 18 (0.00%)	0 / 14 (0.00%)	
occurrences (all)	0	0	
Respiratory, thoracic and mediastinal disorders			
Cough			
subjects affected / exposed	2 / 18 (11.11%)	3 / 14 (21.43%)	
occurrences (all)	2	3	
Dyspnoea			
subjects affected / exposed	2 / 18 (11.11%)	0 / 14 (0.00%)	
occurrences (all)	2	0	
Dyspnoea exertional			

subjects affected / exposed	1 / 18 (5.56%)	1 / 14 (7.14%)	
occurrences (all)	1	1	
Epistaxis			
subjects affected / exposed	0 / 18 (0.00%)	2 / 14 (14.29%)	
occurrences (all)	0	2	
Productive cough			
subjects affected / exposed	1 / 18 (5.56%)	1 / 14 (7.14%)	
occurrences (all)	1	1	
Respiratory tract congestion			
subjects affected / exposed	1 / 18 (5.56%)	0 / 14 (0.00%)	
occurrences (all)	1	0	
Upper-airway cough syndrome			
subjects affected / exposed	0 / 18 (0.00%)	1 / 14 (7.14%)	
occurrences (all)	0	1	
Psychiatric disorders			
Depression			
subjects affected / exposed	0 / 18 (0.00%)	1 / 14 (7.14%)	
occurrences (all)	0	1	
Insomnia			
subjects affected / exposed	0 / 18 (0.00%)	3 / 14 (21.43%)	
occurrences (all)	0	3	
Anxiety			
subjects affected / exposed	1 / 18 (5.56%)	1 / 14 (7.14%)	
occurrences (all)	1	1	
Food aversion			
subjects affected / exposed	0 / 18 (0.00%)	0 / 14 (0.00%)	
occurrences (all)	0	0	
Mood altered			
subjects affected / exposed	0 / 18 (0.00%)	1 / 14 (7.14%)	
occurrences (all)	0	2	
Sleep disorder			
subjects affected / exposed	0 / 18 (0.00%)	0 / 14 (0.00%)	
occurrences (all)	0	0	
Investigations			
Staphylococcus test positive			

subjects affected / exposed	1 / 18 (5.56%)	0 / 14 (0.00%)	
occurrences (all)	1	0	
Weight decreased			
subjects affected / exposed	0 / 18 (0.00%)	0 / 14 (0.00%)	
occurrences (all)	0	0	
Injury, poisoning and procedural complications			
Burns first degree			
subjects affected / exposed	0 / 18 (0.00%)	0 / 14 (0.00%)	
occurrences (all)	0	0	
Tooth fracture			
subjects affected / exposed	0 / 18 (0.00%)	1 / 14 (7.14%)	
occurrences (all)	0	1	
Congenital, familial and genetic disorders			
Congenital lymphoedema			
subjects affected / exposed	0 / 18 (0.00%)	0 / 14 (0.00%)	
occurrences (all)	0	0	
Cardiac disorders			
Palpitations			
subjects affected / exposed	0 / 18 (0.00%)	0 / 14 (0.00%)	
occurrences (all)	0	0	
Tachycardia			
subjects affected / exposed	0 / 18 (0.00%)	1 / 14 (7.14%)	
occurrences (all)	0	1	
Nervous system disorders			
Dizziness			
subjects affected / exposed	0 / 18 (0.00%)	1 / 14 (7.14%)	
occurrences (all)	0	1	
Headache			
subjects affected / exposed	3 / 18 (16.67%)	2 / 14 (14.29%)	
occurrences (all)	3	3	
Disturbance in attention			
subjects affected / exposed	0 / 18 (0.00%)	0 / 14 (0.00%)	
occurrences (all)	0	0	
Dysgeusia			
subjects affected / exposed	0 / 18 (0.00%)	1 / 14 (7.14%)	
occurrences (all)	0	1	

Hypoaesthesia			
subjects affected / exposed	1 / 18 (5.56%)	1 / 14 (7.14%)	
occurrences (all)	1	1	
Mental impairment			
subjects affected / exposed	0 / 18 (0.00%)	1 / 14 (7.14%)	
occurrences (all)	0	1	
Sciatica			
subjects affected / exposed	0 / 18 (0.00%)	1 / 14 (7.14%)	
occurrences (all)	0	1	
Somnolence			
subjects affected / exposed	1 / 18 (5.56%)	0 / 14 (0.00%)	
occurrences (all)	1	0	
Syncope			
subjects affected / exposed	1 / 18 (5.56%)	0 / 14 (0.00%)	
occurrences (all)	1	0	
Blood and lymphatic system disorders			
Anaemia			
subjects affected / exposed	2 / 18 (11.11%)	1 / 14 (7.14%)	
occurrences (all)	2	1	
Leukopenia			
subjects affected / exposed	3 / 18 (16.67%)	1 / 14 (7.14%)	
occurrences (all)	3	1	
Lymphopenia			
subjects affected / exposed	0 / 18 (0.00%)	1 / 14 (7.14%)	
occurrences (all)	0	1	
Neutropenia			
subjects affected / exposed	4 / 18 (22.22%)	2 / 14 (14.29%)	
occurrences (all)	4	2	
Ear and labyrinth disorders			
Vertigo			
subjects affected / exposed	0 / 18 (0.00%)	0 / 14 (0.00%)	
occurrences (all)	0	0	
Gastrointestinal disorders			
Abdominal pain			
subjects affected / exposed	0 / 18 (0.00%)	0 / 14 (0.00%)	
occurrences (all)	0	0	
Abdominal pain upper			

subjects affected / exposed	1 / 18 (5.56%)	0 / 14 (0.00%)	
occurrences (all)	1	0	
Ascites			
subjects affected / exposed	0 / 18 (0.00%)	1 / 14 (7.14%)	
occurrences (all)	0	1	
Diarrhoea			
subjects affected / exposed	2 / 18 (11.11%)	2 / 14 (14.29%)	
occurrences (all)	2	2	
Dyspepsia			
subjects affected / exposed	0 / 18 (0.00%)	0 / 14 (0.00%)	
occurrences (all)	0	0	
Haemorrhoids			
subjects affected / exposed	1 / 18 (5.56%)	1 / 14 (7.14%)	
occurrences (all)	1	1	
Nausea			
subjects affected / exposed	4 / 18 (22.22%)	2 / 14 (14.29%)	
occurrences (all)	4	2	
Vomiting			
subjects affected / exposed	1 / 18 (5.56%)	0 / 14 (0.00%)	
occurrences (all)	1	0	
Abdominal distension			
subjects affected / exposed	0 / 18 (0.00%)	0 / 14 (0.00%)	
occurrences (all)	0	0	
Anal pruritus			
subjects affected / exposed	2 / 18 (11.11%)	0 / 14 (0.00%)	
occurrences (all)	2	0	
Aphthous stomatitis			
subjects affected / exposed	0 / 18 (0.00%)	0 / 14 (0.00%)	
occurrences (all)	0	0	
Constipation			
subjects affected / exposed	1 / 18 (5.56%)	0 / 14 (0.00%)	
occurrences (all)	1	0	
Flatulence			
subjects affected / exposed	1 / 18 (5.56%)	0 / 14 (0.00%)	
occurrences (all)	1	0	
Proctalgia			

subjects affected / exposed	1 / 18 (5.56%)	0 / 14 (0.00%)	
occurrences (all)	1	0	
Hepatobiliary disorders			
Liver tenderness			
subjects affected / exposed	0 / 18 (0.00%)	0 / 14 (0.00%)	
occurrences (all)	0	0	
Skin and subcutaneous tissue disorders			
Pruritus			
subjects affected / exposed	0 / 18 (0.00%)	2 / 14 (14.29%)	
occurrences (all)	0	2	
Rash			
subjects affected / exposed	2 / 18 (11.11%)	1 / 14 (7.14%)	
occurrences (all)	2	1	
Rash macular			
subjects affected / exposed	0 / 18 (0.00%)	1 / 14 (7.14%)	
occurrences (all)	0	2	
Renal and urinary disorders			
Haematuria			
subjects affected / exposed	0 / 18 (0.00%)	0 / 14 (0.00%)	
occurrences (all)	0	0	
Musculoskeletal and connective tissue disorders			
Arthralgia			
subjects affected / exposed	0 / 18 (0.00%)	1 / 14 (7.14%)	
occurrences (all)	0	1	
Muscle tightness			
subjects affected / exposed	0 / 18 (0.00%)	0 / 14 (0.00%)	
occurrences (all)	0	0	
Muscular weakness			
subjects affected / exposed	0 / 18 (0.00%)	1 / 14 (7.14%)	
occurrences (all)	0	1	
Infections and infestations			
Nasopharyngitis			
subjects affected / exposed	1 / 18 (5.56%)	1 / 14 (7.14%)	
occurrences (all)	1	1	
Upper respiratory tract infection			

subjects affected / exposed	0 / 18 (0.00%)	1 / 14 (7.14%)	
occurrences (all)	0	1	
Abscess limb			
subjects affected / exposed	0 / 18 (0.00%)	1 / 14 (7.14%)	
occurrences (all)	0	1	
Cellulitis			
subjects affected / exposed	0 / 18 (0.00%)	1 / 14 (7.14%)	
occurrences (all)	0	1	
Gastroenteritis			
subjects affected / exposed	0 / 18 (0.00%)	1 / 14 (7.14%)	
occurrences (all)	0	1	
Metabolism and nutrition disorders			
Hyperuricaemia			
subjects affected / exposed	1 / 18 (5.56%)	1 / 14 (7.14%)	
occurrences (all)	1	1	
Hypertriglyceridaemia			
subjects affected / exposed	1 / 18 (5.56%)	0 / 14 (0.00%)	
occurrences (all)	1	0	
Hypokalaemia			
subjects affected / exposed	0 / 18 (0.00%)	0 / 14 (0.00%)	
occurrences (all)	0	0	

More information

Substantial protocol amendments (globally)

Were there any global substantial amendments to the protocol? No

Interruptions (globally)

Were there any global interruptions to the trial? No

Limitations and caveats

None reported