Clinical trial results:
A Phase 2b Randomized, Double-Blind, Placebo-Controlled, Parallel Group, DoseRanging Study to Assess the Efficacy, Safety, and Tolerability of Vupanorsen (PF07285557) in Statin-Treated Subjects with Dyslipidemia

Summary

EudraCT number	2020-002796-35
Trial protocol	PL
Global end of trial date	06 December 2021

Results information

Result version number	v1 (current)
This version publication date	15 December 2022
First version publication date	15 December 2022

Trial information

Trial identification

Sponsor protocol code	C4491011

Additional study identifiers

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

Notes:

Sponsors

Sponsor organisation name	Pfizer Inc.
Sponsor organisation address	235 E 42nd Street, New York, United States, NY 10017
Public contact	Pfizer ClinicalTrials.gov Call Center, Pfizer Inc., 001 8007181021, ClinicalTrials.gov_Inquiries@pfizer.com
Scientific contact	Pfizer ClinicalTrials.gov Call Center, Pfizer Inc., 001 8007181021, ClinicalTrials.gov_Inquiries@pfizer.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	28 June 2022
Is this the analysis of the primary completion data?	No

Global end of trial reached?	Yes
Global end of trial date	06 December 2021
Was the trial ended prematurely?	No

Notes:

General information about the trial

Main objective of the trial:

To estimate the effects of multiple dose levels and regimens of vupanorsen compared to placebo on non-high-density lipoprotein cholesterol (non-HDL-C).

Protection of trial subjects:

The study was in compliance with the ethical principles derived from the Declaration of Helsinki and in compliance with all International Council for Harmonisation (ICH) Good Clinical Practice (GCP) Guidelines. All the local regulatory requirements pertinent to safety of trial subjects were followed.

Background therapy: -

Evidence for comparator: -

Actual start date of recruitment	28 September 2020
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	Canada: 126
Country: Number of subjects enrolled	Poland: 33
Country: Number of subjects enrolled	United States: 127
Worldwide total number of subjects	286
EEA total number of subjects	33

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)	151
From 65 to 84 years	134

85 years and over	1

Subject disposition

Recruitment

Recruitment details:

Adult subjects aged greater than equal to (>=) 40 years with dyslipidemia on a stable dose of statin (with or without ezetimibe) were included in the study. The study was conducted across 3 countries.

Pre-assignment

Screening details:

727 subjects signed the informed consent form (ICF). 391 subjects were screen failures who did not meet criteria and were not enrolled. 336 subjects were enrolled into the study of which 50 subjects were not randomised and 286 subjects were assigned to a study treatment.

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	Investigator, Subject

Arms

Are arms mutually exclusive?	Yes
Arm title	Placebo

Arm description:

Subjects were randomised to receive vupanorsen (PF-07285557) matched placebo subcutaneous (SC) injection. Single or double administration was given at every 2 or 4 weeks (Q2W or Q4W) to match active treatment groups. Administration locations were upper arm, thigh, or abdomen quadrant, as preferred by the subject. Treatment duration was up to 24 weeks. Subjects were followed up to 12 weeks after last dose of study intervention.

Arm type	Placebo
Investigational medicinal product name	Placebo
Investigational medicinal product code	
Other name	
Pharmaceutical forms	Solution for injection in pre-filled syringe
Routes of administration	Subcutaneous use

Dosage and administration details:

Subjects received vupanorsen matched placebo SC injection administered Q2W or Q4W up to 24 weeks.

Arm title	Vupanorsen: 80 mg Q4W

Arm description:

Subjects were randomised to receive vupanorsen 80 mg single SC injection at Q4W. Administration locations were upper arm, thigh, or abdomen quadrant, as preferred by the subject. Treatment duration was up to 24 weeks. Subjects were followed up to 12 weeks after last dose of study intervention.

Arm type	Experimental
Investigational medicinal product name	Vupanorsen
Investigational medicinal product code	PF-07285557
Other name	
Pharmaceutical forms	Solution for injection in pre-filled syringe
Routes of administration	Subcutaneous use

Dosage and administration details:

Subjects received vupanorsen 80 mg single SC injection administered Q4W for 24 weeks.

Arm title	Vupanorsen: 60 mg Q2W

Arm description:

Subjects were randomised to receive vupanorsen 60 mg single SC injection at Q2W. Administration locations were upper arm, thigh, or abdomen quadrant, as preferred by the subject. Treatment duration

was up to 24 weeks. Subjects were followed up to 12 weeks after last dose of study intervention.

Arm type	Experimental
Investigational medicinal product name	Vupanorsen
Investigational medicinal product code	PF-07285557
Other name	
Pharmaceutical forms	Solution for injection in pre-filled syringe
Routes of administration	Subcutaneous use

Dosage and administration details:

Subjects received vupanorsen 60 mg single SC injection administered Q2W for 24 weeks.

Arm title	Vupanorsen: 120 mg Q4W

Arm description:

Subjects were randomised to receive vupanorsen 60 mg double SC injection (120 mg total) in different locations at Q4W. Administration locations were upper arm, thigh, or abdomen quadrant, as preferred by the subject. Treatment duration was up to 24 weeks. Subjects were followed up to 12 weeks after last dose of study intervention.

Arm type	Experimental
Investigational medicinal product name	Vupanorsen
Investigational medicinal product code	PF-07285557
Other name	
Pharmaceutical forms	Solution for injection in pre-filled syringe
Routes of administration	Subcutaneous use

Dosage and administration details:

Subjects received vupanorsen 60 mg double SC injection (120 mg in total) administered Q4W for 24 weeks.

Arm title	Vupanorsen: 80 mg Q2W

Arm description:

Subjects were randomised to receive vupanorsen 80 mg single SC injection at Q2W. Administration locations were upper arm, thigh, or abdomen quadrant, as preferred by the subject. Treatment duration was up to 24 weeks. Subjects were followed up to 12 weeks after last dose of study intervention.

Arm type	Experimental
Investigational medicinal product name	Vupanorsen
Investigational medicinal product code	PF-07285557
Other name	
Pharmaceutical forms	Solution for injection in pre-filled syringe
Routes of administration	Subcutaneous use

Dosage and administration details:

Subjects received vupanorsen 80 mg single SC injection administered Q2W for 24 weeks.

Arm title	Vupanorsen: 160 mg Q4W

Arm description:

Subjects were randomised to receive vupanorsen 80 mg double SC injection (160 mg total) in different locations at Q4W. Administration locations were upper arm, thigh, or abdomen quadrant, as preferred by the subject. Treatment duration was up to 24 weeks. Subjects were followed up to 12 weeks after last dose of study intervention.

Arm type	Experimental
Investigational medicinal product name	Vupanorsen
Investigational medicinal product code	PF-07285557
Other name	
Pharmaceutical forms	Solution for injection in pre-filled syringe
Routes of administration	Subcutaneous use

Dosage and administration details:

Subjects received vupanorsen 80 mg double SC injection (160 mg in total) administered Q4W for 24 weeks.

Arm title	Vupanorsen: 120 mg Q2W

Arm description:

Subjects were randomised to receive vupanorsen 60 mg double SC injection (120 mg in total) in different locations at Q2W. Administration locations were upper arm, thigh, or abdomen quadrant, as preferred by the subject. Treatment duration was up to 24 weeks. Subjects were followed up to 12 weeks after last dose of study intervention.

Arm type	Experimental
Investigational medicinal product name	Vupanorsen
Investigational medicinal product code	PF-07285557
Other name	
Pharmaceutical forms	Solution for injection in pre-filled syringe
Routes of administration	Subcutaneous use

Dosage and administration details:

Subjects received vupanorsen 60 mg double SC injection (120 mg in total) administered Q2W for 24 weeks.

Arm title	Vupanorsen: 160 mg Q2W

Arm description:

Subjects were randomised to receive vupanorsen 80 mg double SC injection (160 mg in total) in different locations at Q2W. Administration locations were upper arm, thigh, or abdomen quadrant, as preferred by the subject. Treatment duration was up to 24 weeks. Subjects were followed up to 12 weeks after last dose of study intervention.

Arm type	Experimental
Investigational medicinal product name	Vupanorsen
Investigational medicinal product code	PF-07285557
Other name	
Pharmaceutical forms	Solution for injection in pre-filled syringe
Routes of administration	Subcutaneous use

Dosage and administration details:

Subjects received vupanorsen 80 mg double SC injection (160 mg in total) administered Q2W for 24 weeks.

Number of subjects in period 1	Placebo	Vupanorsen: 80 mg Q4W	Vupanorsen: 60 mg Q2W
Started	44	23	24
Completed	44	22	23
Not completed	0	1	1
Consent withdrawn by subject	-	-	-
Adverse event, non-fatal	-	-	1
Unspecified	-	1	-

Number of subjects in period 1	Vupanorsen: 120 mg Q4W	Vupanorsen: 80 mg Q2W	Vupanorsen: 160 mg Q4W
Started	23	45	45
Completed	21	43	45
Not completed	2	2	0
Consent withdrawn by subject	-	1	-
Adverse event, non-fatal	-	1	-
Unspecified	2	-	-

Number of subjects in period 1	Vupanorsen: 120 mg Q2W	Vupanorsen: 160 mg Q2W

Started	46	36
Completed	46	36
Not completed	0	0
Consent withdrawn by subject	-	-
Adverse event, non-fatal	-	-
Unspecified	-	-

Baseline characteristics

Reporting groups	
Reporting group title	Placebo
Reporting group description:	
Subjects were randomised to receive vupanorsen (PF-07285557) matched placebo subcutaneous (SC) injection. Single or double administration was given at every 2 or 4 weeks (Q2W or Q4W) to match active treatment groups. Administration locations were upper arm, thigh, or abdomen quadrant, as preferred by the subject. Treatment duration was up to 24 weeks. Subjects were followed up to 12 weeks after last dose of study intervention.	
Reporting group title	Vupanorsen: 80 mg Q4W
Reporting group description:	
Subjects were randomised to receive vupanorsen 80 mg single SC injection at Q4W. Administration locations were upper arm, thigh, or abdomen quadrant, as preferred by the subject. Treatment duration was up to 24 weeks. Subjects were followed up to 12 weeks after last dose of study intervention.	
Reporting group title	Vupanorsen: 60 mg Q2W
Reporting group description:	
Subjects were randomised to receive vupanorsen 60 mg single SC injection at Q2W. Administration locations were upper arm, thigh, or abdomen quadrant, as preferred by the subject. Treatment duration was up to 24 weeks. Subjects were followed up to 12 weeks after last dose of study intervention.	
Reporting group title	Vupanorsen: 120 mg Q4W
Reporting group description:	
Subjects were randomised to receive vupanorsen 60 mg double SC injection (120 mg total) in different locations at Q4W. Administration locations were upper arm, thigh, or abdomen quadrant, as preferred by the subject. Treatment duration was up to 24 weeks. Subjects were followed up to 12 weeks after last dose of study intervention.	
Reporting group title	Vupanorsen: 80 mg Q2W
Reporting group description:	
Subjects were randomised to receive vupanorsen 80 mg single SC injection at Q2W. Administration locations were upper arm, thigh, or abdomen quadrant, as preferred by the subject. Treatment duration was up to 24 weeks. Subjects were followed up to 12 weeks after last dose of study intervention.	
Reporting group title	Vupanorsen: 160 mg Q4W
Reporting group description:	
Subjects were randomised to receive vupanorsen 80 mg double SC injection (160 mg total) in different locations at Q4W. Administration locations were upper arm, thigh, or abdomen quadrant, as preferred by the subject. Treatment duration was up to 24 weeks. Subjects were followed up to 12 weeks after last dose of study intervention.	
Reporting group title	Vupanorsen: 120 mg Q2W
Reporting group description:	
Subjects were randomised to receive vupanorsen 60 mg double SC injection (120 mg in total) in different locations at Q2W. Administration locations were upper arm, thigh, or abdomen quadrant, as preferred by the subject. Treatment duration was up to 24 weeks. Subjects were followed up to 12 weeks after last dose of study intervention.	
Reporting group title	Vupanorsen: 160 mg Q2W
Reporting group description:	
Subjects were randomised to receive vupanorsen 80 mg double SC injection (160 mg in total) in different locations at Q2W. Administration locations were upper arm, thigh, or abdomen quadrant, as preferred by the subject. Treatment duration was up to 24 weeks. Subjects were followed up to 12 weeks after last dose of study intervention.	

Reporting group values	Placebo	Vupanorsen: 80 mg Q4W	Vupanorsen: 60 mg Q2W
Number of subjects	44	23	24
Age categorical Units: Subjects			
In utero	0	0	0

Preterm newborn infants (gestational age < 37 wks)	0	0	0
Newborns (0-27 days)	0	0	0
Infants and toddlers (28 days-23 months)	0	0	0
Children (2-11 years)	0	0	0
Adolescents (12-17 years)	0	0	0
Adults (18-64 years)	23	11	11
From 65-84 years	21	12	12
85 years and over	0	0	1
Age Continuous			
Units: Years			
arithmetic mean	64.23	65.78	64.21
standard deviation	± 8.09	± 7.27	± 9.92
Sex: Female, Male			
Units: Subjects			
Female	17	13	7
Male	27	10	17
Race			
Units: Subjects			
Asian	4	2	2
Black or African American	0	1	1
White	38	20	21
Unknown or Not Reported	2	0	0
Ethnicity			
Units: Subjects			
Hispanic or Latino	5	2	1
Not Hispanic or Latino	39	21	23

Reporting group values	Vupanorsen: 120 mg Q4W	Vupanorsen: 80 mg Q2W	Vupanorsen: 160 mg Q4W
Number of subjects	23	45	45
Age categorical			
Units: Subjects			
In utero	0	0	0
Preterm newborn infants (gestational age < 37 wks)	0	0	0
Newborns (0-27 days)	0	0	0
Infants and toddlers (28 days-23 months)	0	0	0
Children (2-11 years)	0	0	0
Adolescents (12-17 years)	0	0	0
Adults (18-64 years)	13	23	23
From 65-84 years	10	22	22
85 years and over	0	0	0
Age Continuous			
Units: Years			
arithmetic mean	61.04	63.38	63.09
standard deviation	± 9.31	± 8.41	± 8.80
Sex: Female, Male			
Units: Subjects			
Female	9	24	17
Male	14	21	28

Race Units: Subjects			
Asian	1	3	0
Black or African American	0	4	1
White	21	38	44
Unknown or Not Reported	1	0	0
Ethnicity Units: Subjects			
Hispanic or Latino	6	4	6
Not Hispanic or Latino	17	41	39

Reporting group values	Vupanorsen: 120 mg Q2W	Vupanorsen: 160 mg Q2W	Total
Number of subjects	46	36	286
Age categorical Units: Subjects			
In utero	0	0	0
Preterm newborn infants (gestational age < 37 wks)	0	0	0
Newborns (0-27 days)	0	0	0
Infants and toddlers (28 days-23 months)	0	0	0
Children (2-11 years)	0	0	0
Adolescents (12-17 years)	0	0	0
Adults (18-64 years)	30	17	151
From 65-84 years	16	19	134
85 years and over	0	0	1
Age Continuous Units: Years			
arithmetic mean	62.74	64.47	
standard deviation	± 8.64	± 7.74	-
Sex: Female, Male Units: Subjects			
Female	20	19	126
Male	26	17	160
Race Units: Subjects			
Asian	4	4	20
Black or African American	4	1	12
White	37	31	250
Unknown or Not Reported	1	0	4
Ethnicity Units: Subjects			
Hispanic or Latino	2	1	27
Not Hispanic or Latino	44	35	259

End points

End points reporting groups

Reporting group title	Placebo

Reporting group description:

Subjects were randomised to receive vupanorsen (PF-07285557) matched placebo subcutaneous (SC) injection. Single or double administration was given at every 2 or 4 weeks (Q2W or Q4W) to match active treatment groups. Administration locations were upper arm, thigh, or abdomen quadrant, as preferred by the subject. Treatment duration was up to 24 weeks. Subjects were followed up to 12 weeks after last dose of study intervention.

Reporting group title	Vupanorsen: 80 mg Q4W

Reporting group description:

Subjects were randomised to receive vupanorsen 80 mg single SC injection at Q4W. Administration locations were upper arm, thigh, or abdomen quadrant, as preferred by the subject. Treatment duration was up to 24 weeks. Subjects were followed up to 12 weeks after last dose of study intervention.

Reporting group title	Vupanorsen: 60 mg Q2W

Reporting group description:

Subjects were randomised to receive vupanorsen 60 mg single SC injection at Q2W. Administration locations were upper arm, thigh, or abdomen quadrant, as preferred by the subject. Treatment duration was up to 24 weeks. Subjects were followed up to 12 weeks after last dose of study intervention.

Reporting group title	Vupanorsen: 120 mg Q4W

Reporting group description:

Subjects were randomised to receive vupanorsen 60 mg double SC injection (120 mg total) in different locations at Q4W. Administration locations were upper arm, thigh, or abdomen quadrant, as preferred by the subject. Treatment duration was up to 24 weeks. Subjects were followed up to 12 weeks after last dose of study intervention.

Reporting group title	Vupanorsen: 80 mg Q2W

Reporting group description:

Subjects were randomised to receive vupanorsen 80 mg single SC injection at Q2W. Administration locations were upper arm, thigh, or abdomen quadrant, as preferred by the subject. Treatment duration was up to 24 weeks. Subjects were followed up to 12 weeks after last dose of study intervention.

Reporting group title	Vupanorsen: 160 mg Q4W

Reporting group description:

Subjects were randomised to receive vupanorsen 80 mg double SC injection (160 mg total) in different locations at Q4W. Administration locations were upper arm, thigh, or abdomen quadrant, as preferred by the subject. Treatment duration was up to 24 weeks. Subjects were followed up to 12 weeks after last dose of study intervention.

Reporting group title	Vupanorsen: 120 mg Q2W

Reporting group description:

Subjects were randomised to receive vupanorsen 60 mg double SC injection (120 mg in total) in different locations at Q2W. Administration locations were upper arm, thigh, or abdomen quadrant, as preferred by the subject. Treatment duration was up to 24 weeks. Subjects were followed up to 12 weeks after last dose of study intervention.

Reporting group title	Vupanorsen: 160 mg Q2W

Reporting group description:

Subjects were randomised to receive vupanorsen 80 mg double SC injection (160 mg in total) in different locations at Q2W. Administration locations were upper arm, thigh, or abdomen quadrant, as preferred by the subject. Treatment duration was up to 24 weeks. Subjects were followed up to 12 weeks after last dose of study intervention.

Primary: Percent Change From Baseline in Non-High-Density Lipoprotein-Cholesterol (non-HDL-C) at Week 24

End point title	Percent Change From Baseline in Non-High-Density Lipoprotein-Cholesterol (non-HDL-C) at Week 24

End point description:

Fasting was required at least 10 hours before blood sample collection. Baseline was calculated using the average of all values obtained at Screening and on Day 1 prior to dosing. Full analysis set primary (FAS_primary) included all subjects randomised to study intervention and who took at least 1 dose of study intervention, had a baseline measurement and at least one post-baseline measurement with all observations that occurred after discontinuation of treatment or after initiation of severe hypertriglyceridaemia excluded.

End point type	Primary

End point timeframe:

Baseline, Week 24

End point values	Placebo	Vupanorsen: 80 mg Q4W	Vupanorsen: 60 mg Q2W	Vupanorsen: 120 mg Q4W
Subject group type	Reporting group	Reporting group	Reporting group	Reporting group
Number of subjects analysed	44	23	23	23
Units: Percent change				
least squares mean (standard error)	-1.1 (± 2.76)	-23.5 (± 4.08)	-23.2 (± 4.02)	-25.3 (± 4.23)

End point values	Vupanorsen: 80 mg Q2W	Vupanorsen: 160 mg Q4W	Vupanorsen: 120 mg Q2W	Vupanorsen: 160 mg Q2W
Subject group type	Reporting group	Reporting group	Reporting group	Reporting group
Number of subjects analysed	42	45	46	35
Units: Percent change				
least squares mean (standard error)	-28.8 (± 3.02)	-27.8 (± 2.88)	-25.8 (± 2.84)	-27.6 (± 3.57)

Statistical analyses

Statistical analysis title	Placebo versus Vupanorsen: 80 mg Q4W

Statistical analysis description:

Mixed Model Analysis (MMRM) with baseline value, treatment, visit, interaction term of treatment by visit as fixed effects and unstructured covariance structure was used.

Comparison groups	Placebo v Vupanorsen: 80 mg Q4W
Number of subjects included in analysis	67
Analysis specification	Pre-specified
Analysis type	other
P-value	< 0.001
Method	Mixed models analysis
Parameter estimate	Least Square (LS) Mean difference
Point estimate	-22.4
Confidence interval	
level	95 %
sides	2-sided
lower limit	-32.1
upper limit	-12.7
Variability estimate	Standard error of the mean
Dispersion value	4.93

Statistical analysis title	Placebo versus Vupanorsen: 60 mg Q2W
Statistical analysis description: MMRM with baseline value, treatment, visit, interaction term of treatment by visit as fixed effects and unstructured covariance structure was used.	
Comparison groups	Placebo v Vupanorsen: 60 mg Q2W
Number of subjects included in analysis	67
Analysis specification	Pre-specified
Analysis type	other
P-value	< 0.001
Method	Mixed models analysis
Parameter estimate	LS Mean difference
Point estimate	-22
Confidence interval	
level	95 %
sides	2-sided
lower limit	-31.7
upper limit	-12.4
Variability estimate	Standard error of the mean
Dispersion value	4.88

Statistical analysis title	Placebo versus Vupanorsen: 120 mg Q4W
Statistical analysis description: MMRM with baseline value, treatment, visit, interaction term of treatment by visit as fixed effects and unstructured covariance structure was used.	
Comparison groups	Placebo v Vupanorsen: 120 mg Q4W
Number of subjects included in analysis	67
Analysis specification	Pre-specified
Analysis type	other
P-value	< 0.001
Method	Mixed models analysis
Parameter estimate	LS Mean difference
Point estimate	-24.1
Confidence interval	
level	95 %
sides	2-sided
lower limit	-34.1
upper limit	-14.2
Variability estimate	Standard error of the mean
Dispersion value	5.05

Statistical analysis title	Placebo versus Vupanorsen: 80 mg Q2W
Statistical analysis description: MMRM with baseline value, treatment, visit, interaction term of treatment by visit as fixed effects and unstructured covariance structure was used.	

Comparison groups	Placebo v Vupanorsen: 80 mg Q2W
Number of subjects included in analysis	86
Analysis specification	Pre-specified
Analysis type	other
P-value	< 0.001
Method	Mixed models analysis
Parameter estimate	LS Mean difference
Point estimate	-27.7
Confidence interval	
level	95 %
sides	2-sided
lower limit	-35.7
upper limit	-19.6
Variability estimate	Standard error of the mean
Dispersion value	4.09

Statistical analysis title	Placebo versus Vupanorsen: 160 mg Q4W

Statistical analysis description:

MMRM with baseline value, treatment, visit, interaction term of treatment by visit as fixed effects and unstructured covariance structure was used.

Comparison groups	Placebo v Vupanorsen: 160 mg Q4W
Number of subjects included in analysis	89
Analysis specification	Pre-specified
Analysis type	other
P-value	< 0.001
Method	Mixed models analysis
Parameter estimate	LS Mean difference
Point estimate	-26.6
Confidence interval	
level	95 %
sides	2-sided
lower limit	-34.5
upper limit	-18.8
Variability estimate	Standard error of the mean
Dispersion value	3.98

Statistical analysis title	Placebo versus Vupanorsen: 120 mg Q2W

Statistical analysis description:

MMRM with baseline value, treatment, visit, interaction term of treatment by visit as fixed effects and unstructured covariance structure was used.

Comparison groups	Placebo v Vupanorsen: 120 mg Q2W

Number of subjects included in analysis	90
Analysis specification	Pre-specified
Analysis type	other
P-value	< 0.001
Method	Mixed models analysis
Parameter estimate	LS Mean difference
Point estimate	-24.7
Confidence interval	
level	95 %
sides	2-sided
lower limit	-32.5
upper limit	-16.9
Variability estimate	Standard error of the mean
Dispersion value	3.96

Statistical analysis title	Placebo versus Vupanorsen: 160 mg Q2W

Statistical analysis description:

MMRM with baseline value, treatment, visit, interaction term of treatment by visit as fixed effects and unstructured covariance structure was used.

Comparison groups	Placebo v Vupanorsen: 160 mg Q2W
Number of subjects included in analysis	79
Analysis specification	Pre-specified
Analysis type	other
P-value	< 0.001
Method	Mixed models analysis
Parameter estimate	LS Mean difference
Point estimate	-26.5
Confidence interval	
level	95 %
sides	2-sided
lower limit	-35.4
upper limit	-17.6
Variability estimate	Standard error of the mean
Dispersion value	4.51

Secondary: Percent Change From Baseline in Triglyceride (TG), Apolipoprotein B (ApoB), Low-Density Lipoprotein-Cholesterol (LDL-C), and non-HDL-C at Week 16

End point title	Percent Change From Baseline in Triglyceride (TG), Apolipoprotein B (ApoB), Low-Density Lipoprotein-Cholesterol (LDL-C), and non-HDL-C at Week 16

End point description:

Blood samples were collected from subjects in a fasted state for the measurement of TG, ApoB, HDL-C and LDL-C. Fasting was required at least 10 hours before blood sample collection. Non-HDL-C was calculated as total cholesterol minus HDL cholesterol. Baseline was calculated using the average of all values obtained at Screening and on Day 1 prior to dosing. FAS included all subjects randomised to study intervention and who took at least 1 dose of study intervention and had a baseline measurement and at least one post-baseline measurement. Here, "n" signifies subjects evaluable for specific rows.

End point type	Secondary

End point timeframe:
Baseline, Week 16

End point values	Placebo	Vupanorsen: 80 mg Q4W	Vupanorsen: 60 mg Q2W	Vupanorsen: 120 mg Q4W
Subject group type	Reporting group	Reporting group	Reporting group	Reporting group
Number of subjects analysed	44	23	24	23
Units: Percent change				
arithmetic mean (standard deviation)				
TG (n=43,22,23,21,40,43,44,35)	-2.53 (± 31.247)	-42.17 (± 28.081)	-44.71 (± 22.725)	-39.57 (± 32.546)
ApoB (n=41,21,22,22,40,44,43,34)	0.66 (± 20.372)	-12.12 (± 15.839)	-12.30 (± 12.604)	-10.41 (± 17.067)
LDL-C (n=41,22,22,20,40,42,44,35)	-1.42 (± 24.092)	-10.40 (± 31.199)	-12.61 (± 21.904)	-2.09 (± 26.921)
Non-HDL-C (n=43,22,23,21,40,43,44,35)	-3.25 (± 21.631)	-21.65 (± 24.347)	-24.71 (± 16.678)	-20.06 (± 16.698)

End point values	Vupanorsen: 80 mg Q2W	Vupanorsen: 160 mg Q4W	Vupanorsen: 120 mg Q2W	Vupanorsen: 160 mg Q2W
Subject group type	Reporting group	Reporting group	Reporting group	Reporting group
Number of subjects analysed	43	45	46	36
Units: Percent change				
arithmetic mean (standard deviation)				
TG (n=43,22,23,21,40,43,44,35)	-49.21 (± 21.049)	-40.33 (± 21.311)	-50.55 (± 20.234)	-55.76 (± 14.625)
ApoB (n=41,21,22,22,40,44,43,34)	-10.40 (± 14.690)	-10.42 (± 19.332)	-11.74 (± 15.560)	-6.76 (± 18.890)
LDL-C (n=41,22,22,20,40,42,44,35)	-11.38 (± 22.001)	-10.88 (± 26.640)	-13.78 (± 19.861)	-6.30 (± 29.825)
Non-HDL-C (n=43,22,23,21,40,43,44,35)	-24.79 (± 18.266)	-21.68 (± 23.898)	-27.23 (± 12.799)	-22.44 (± 24.145)

Statistical analyses

No statistical analyses for this end point

Secondary: Percent Change From Baseline in TG, ApoB, and LDL-C at Week 24

End point title	Percent Change From Baseline in TG, ApoB, and LDL-C at Week 24

End point description:

Fasting was required for all lipid measures at least 10 hours before blood sample collection. Baseline was calculated using the average of all values obtained at Screening and on Day 1 prior to dosing.
FAS_primary included all subjects randomised to study intervention and who took at least 1 dose of study intervention, had a baseline measurement and at least one post-baseline measurement with all observations that occurred after discontinuation of treatment or after initiation of severe hypertriglyceridaemia excluded. Here, "n" signifies subjects evaluable for specific rows.

End point type	Secondary

End point timeframe:
Baseline, Week 24

End point values	Placebo	Vupanorsen: 80 mg Q4W	Vupanorsen: 60 mg Q2W	Vupanorsen: 120 mg Q4W
Subject group type	Reporting group	Reporting group	Reporting group	Reporting group
Number of subjects analysed	44	23	23	23
Units: Percent change				
least squares mean (standard error)				
TG (n=44,23,23,23,42,45,46,35)	-1.8 (± 3.71)	-45.8 (± 5.53)	-45.6 (± 5.50)	-43.1 (± 5.76)
ApoB (n=42,22,22,23,41,45,44,35)	0.3 (± 2.46)	-14.8 (± 3.60)	-10.3 (± 3.57)	-11.2 (± 3.76)
LDL-C (n=43,23,22,22,42,43,46,35)	-1.2 (± 3.69)	-11.2 (± 5.41)	-9.1 (± 5.52)	-12.7 (± 5.63)

End point values	Vupanorsen: 80 mg Q2W	Vupanorsen: 160 mg Q4W	Vupanorsen: 120 mg Q2W	Vupanorsen: 160 mg Q2W
Subject group type	Reporting group	Reporting group	Reporting group	Reporting group
Number of subjects analysed	42	45	46	35
Units: Percent change				
least squares mean (standard error)				
TG (n=44,23,23,23,42,45,46,35)	-52.3 (± 4.10)	-47.7 (± 3.90)	-52.5 (± 3.85)	-58.6 (± 4.90)
ApoB (n=42,22,22,23,41,45,44,35)	-12.2 (± 2.68)	-12.2 (± 2.55)	-5.6 (± 2.57)	-8.1 (± 3.17)
LDL-C (n=43,23,22,22,42,43,46,35)	-17.3 (± 4.00)	-15.7 (± 3.92)	-9.1 (± 3.77)	-10.2 (± 4.74)

Statistical analyses

Statistical analysis title	Placebo versus Vupanorsen: 80 mg Q4W - TG

Statistical analysis description:

MMRM with baseline value, treatment, visit, interaction term of treatment by visit as fixed effects and unstructured covariance structure was used.

Comparison groups	Placebo v Vupanorsen: 80 mg Q4W
Number of subjects included in analysis	67
Analysis specification	Pre-specified
Analysis type	other
P-value	< 0.001
Method	Mixed models analysis
Parameter estimate	LS Mean difference
Point estimate	-44
Confidence interval	
level	95 %
sides	2-sided
lower limit	-57.1
upper limit	-30.8
Variability estimate	Standard error of the mean
Dispersion value	6.66

Statistical analysis title	Placebo versus Vupanorsen: 60 mg Q2W - TG

Statistical analysis description:

MMRM with baseline value, treatment, visit, interaction term of treatment by visit as fixed effects and unstructured covariance structure was used.

Comparison groups	Placebo v Vupanorsen: 60 mg Q2W
Number of subjects included in analysis	67
Analysis specification	Pre-specified
Analysis type	other
P-value	< 0.001
Method	Mixed models analysis
Parameter estimate	LS Mean difference
Point estimate	-43.8
Confidence interval	
level	95 %
sides	2-sided
lower limit	-56.9
upper limit	-30.7
Variability estimate	Standard error of the mean
Dispersion value	6.64

Statistical analysis title	Placebo versus Vupanorsen: 120 mg Q4W - TG

Statistical analysis description:

MMRM with baseline value, treatment, visit, interaction term of treatment by visit as fixed effects and unstructured covariance structure was used.

Comparison groups	Placebo v Vupanorsen: 120 mg Q4W
Number of subjects included in analysis	67
Analysis specification	Pre-specified
Analysis type	other
P-value	< 0.001
Method	Mixed models analysis
Parameter estimate	LS Mean difference
Point estimate	-41.3
Confidence interval	
level	95 %
sides	2-sided
lower limit	-54.8
upper limit	-27.8
Variability estimate	Standard error of the mean
Dispersion value	6.85

Statistical analysis title	Placebo versus Vupanorsen: 80 mg Q2W - TG

Statistical analysis description:

MMRM with baseline value, treatment, visit, interaction term of treatment by visit as fixed effects and unstructured covariance structure was used.

Comparison groups	Placebo v Vupanorsen: 80 mg Q2W
Number of subjects included in analysis	86
Analysis specification	Pre-specified
Analysis type	other
P-value	< 0.001
Method	Mixed models analysis
Parameter estimate	LS Mean difference
Point estimate	-50.5
Confidence interval	
level	95 %
sides	2-sided
lower limit	-61.4
upper limit	-39.6
Variability estimate	Standard error of the mean
Dispersion value	5.54

Statistical analysis title	Placebo versus Vupanorsen: 160 mg Q4W - TG

Statistical analysis description:

MMRM with baseline value, treatment, visit, interaction term of treatment by visit as fixed effects and unstructured covariance structure was used.

Comparison groups	Placebo v Vupanorsen: 160 mg Q4W
Number of subjects included in analysis	89
Analysis specification	Pre-specified
Analysis type	other
P-value	< 0.001
Method	Mixed models analysis
Parameter estimate	LS Mean difference
Point estimate	-45.9
Confidence interval	
level	95 %
sides	2-sided
lower limit	-56.5
upper limit	-35.2
Variability estimate	Standard error of the mean
Dispersion value	5.38

Statistical analysis title	Placebo versus Vupanorsen: 120 mg Q2W - TG

Statistical analysis description:

MMRM with baseline value, treatment, visit, interaction term of treatment by visit as fixed effects and unstructured covariance structure was used.

Comparison groups	Placebo v Vupanorsen: 120 mg Q2W

Number of subjects included in analysis	90
Analysis specification	Pre-specified
Analysis type	other
P-value	< 0.001
Method	Mixed models analysis
Parameter estimate	LS Mean difference
Point estimate	-50.7
Confidence interval	
level	95 %
sides	2-sided
lower limit	-61.2
upper limit	-40.1
Variability estimate	Standard error of the mean
Dispersion value	5.35

Statistical analysis title	Placebo versus Vupanorsen: 160 mg Q2W - TG

Statistical analysis description:

MMRM with baseline value, treatment, visit, interaction term of treatment by visit as fixed effects and unstructured covariance structure was used.

Comparison groups	Placebo v Vupanorsen: 160 mg Q2W
Number of subjects included in analysis	79
Analysis specification	Pre-specified
Analysis type	other
P-value	< 0.001
Method	Mixed models analysis
Parameter estimate	LS Mean difference
Point estimate	-56.8
Confidence interval	
level	95 %
sides	2-sided
lower limit	-68.9
upper limit	-44.7
Variability estimate	Standard error of the mean
Dispersion value	6.14

Statistical analysis title	Placebo versus Vupanorsen: 80 mg Q4W - ApoB

Statistical analysis description:

MMRM with baseline value, treatment, visit, interaction term of treatment by visit as fixed effects and unstructured covariance structure was used.

Comparison groups	Placebo v Vupanorsen: 80 mg Q4W
Number of subjects included in analysis	67
Analysis specification	Pre-specified
Analysis type	other
P-value	< 0.001
Method	Mixed models analysis
Parameter estimate	LS Mean difference
Point estimate	-15.1

Confidence interval	
level	95 %
sides	2-sided
lower limit	-23.7
upper limit	-6.5
Variability estimate	Standard error of the mean
Dispersion value	4.36

Statistical analysis title	Placebo versus Vupanorsen: 60 mg Q2W - ApoB

Statistical analysis description:

MMRM with baseline value, treatment, visit, interaction term of treatment by visit as fixed effects and unstructured covariance structure was used.

Comparison groups	Placebo v Vupanorsen: 60 mg Q2W
Number of subjects included in analysis	67
Analysis specification	Pre-specified
Analysis type	other
P-value	= 0.015
Method	Mixed models analysis
Parameter estimate	LS Mean difference
Point estimate	-10.6
Confidence interval	
level	95 %
sides	2-sided
lower limit	-19.2
upper limit	-2.1
Variability estimate	Standard error of the mean
Dispersion value	4.34

Statistical analysis title	Placebo versus Vupanorsen: 120 mg Q4W - ApoB

Statistical analysis description:

MMRM with baseline value, treatment, visit, interaction term of treatment by visit as fixed effects and unstructured covariance structure was used.

Comparison groups	Placebo v Vupanorsen: 120 mg Q4W
Number of subjects included in analysis	67
Analysis specification	Pre-specified
Analysis type	other
P-value	= 0.011
Method	Mixed models analysis
Parameter estimate	LS Mean difference
Point estimate	-11.5
Confidence interval	
level	95 %
sides	2-sided
lower limit	-20.3
upper limit	-2.7
Variability estimate	Standard error of the mean
Dispersion value	4.49

Statistical analysis title	Placebo versus Vupanorsen: 80 mg Q2W - ApoB
Statistical analysis description: MMRM with baseline value, treatment, visit, interaction term of treatment by visit as fixed effects and unstructured covariance structure was used.	
Comparison groups	Placebo v Vupanorsen: 80 mg Q2W
Number of subjects included in analysis	86
Analysis specification	Pre-specified
Analysis type	other
P-value	< 0.001
Method	Mixed models analysis
Parameter estimate	LS Mean difference
Point estimate	-12.5
Confidence interval	
level	95 %
sides	2-sided
lower limit	-19.7
upper limit	-5.3
Variability estimate	Standard error of the mean
Dispersion value	3.64

Statistical analysis title	Placebo versus Vupanorsen: 160 mg Q4W - ApoB
Statistical analysis description: MMRM with baseline value, treatment, visit, interaction term of treatment by visit as fixed effects and unstructured covariance structure was used.	
Comparison groups	Placebo v Vupanorsen: 160 mg Q4W
Number of subjects included in analysis	89
Analysis specification	Pre-specified
Analysis type	other
P-value	< 0.001
Method	Mixed models analysis
Parameter estimate	LS Mean difference
Point estimate	-12.6
Confidence interval	
level	95 %
sides	2-sided
lower limit	-19.5
upper limit	-5.6
Variability estimate	Standard error of the mean
Dispersion value	3.54

Statistical analysis title	Placebo versus Vupanorsen: 120 mg Q2W - ApoB
Statistical analysis description: MMRM with baseline value, treatment, visit, interaction term of treatment by visit as fixed effects and unstructured covariance structure was used.	

Comparison groups	Placebo v Vupanorsen: 120 mg Q2W
Number of subjects included in analysis	90
Analysis specification	Pre-specified
Analysis type	other
P-value	= 0.095
Method	Mixed models analysis
Parameter estimate	LS Mean difference
Point estimate	-6
Confidence interval	
level	95 %
sides	2-sided
lower limit	-13
upper limit	1
Variability estimate	Standard error of the mean
Dispersion value	3.56

Statistical analysis title	Placebo versus Vupanorsen: 160 mg Q2W - ApoB

Statistical analysis description:

MMRM with baseline value, treatment, visit, interaction term of treatment by visit as fixed effects and unstructured covariance structure was used.

Comparison groups	Placebo v Vupanorsen: 160 mg Q2W
Number of subjects included in analysis	79
Analysis specification	Pre-specified
Analysis type	other
P-value	= 0.036
Method	Mixed models analysis
Parameter estimate	LS Mean difference
Point estimate	-8.5
Confidence interval	
level	95 %
sides	2-sided
lower limit	-16.4
upper limit	-0.6
Variability estimate	Standard error of the mean
Dispersion value	4.01

Statistical analysis title	Placebo versus Vupanorsen: 80 mg Q4W - LDL-C

Statistical analysis description:

MMRM with baseline value, treatment, visit, interaction term of treatment by visit as fixed effects and unstructured covariance structure was used.

Comparison groups	Placebo v Vupanorsen: 80 mg Q4W

Number of subjects included in analysis	67
Analysis specification	Pre-specified
Analysis type	other
P-value	= 0.129
Method	Mixed models analysis
Parameter estimate	LS Mean difference
Point estimate	-10
Confidence interval	
level	95 %
sides	2-sided
lower limit	-22.9
upper limit	2.9
Variability estimate	Standard error of the mean
Dispersion value	6.55

Statistical analysis title	Placebo versus Vupanorsen: 60 mg Q2W - LDL-C
Statistical analysis description:	
MMRM with baseline value, treatment, visit, interaction term of treatment by visit as fixed effects and unstructured covariance structure was used.	
Comparison groups	Placebo v Vupanorsen: 60 mg Q2W
Number of subjects included in analysis	67
Analysis specification	Pre-specified
Analysis type	other
P-value	= 0.238
Method	Mixed models analysis
Parameter estimate	LS Mean difference
Point estimate	-7.9
Confidence interval	
level	95 %
sides	2-sided
lower limit	-21
upper limit	5.2
Variability estimate	Standard error of the mean
Dispersion value	6.65

Statistical analysis title	Placebo versus Vupanorsen: 120 mg Q4W - LDL-C
Statistical analysis description:	
MMRM with baseline value, treatment, visit, interaction term of treatment by visit as fixed effects and unstructured covariance structure was used.	
Comparison groups	Placebo v Vupanorsen: 120 mg Q4W
Number of subjects included in analysis	67
Analysis specification	Pre-specified
Analysis type	other
P-value	= 0.09
Method	Mixed models analysis
Parameter estimate	LS Mean difference
Point estimate	-11.4

Confidence interval	
level	95 %
sides	2-sided
lower limit	-24.7
upper limit	1.8
Variability estimate	Standard error of the mean
Dispersion value	6.73

Statistical analysis title	Placebo versus Vupanorsen: 160 mg Q4W - LDL-C
Statistical analysis description:	
MMRM with baseline value, treatment, visit, interaction term of treatment by visit as fixed effects and unstructured covariance structure was used.	
Comparison groups	Placebo v Vupanorsen: 160 mg Q4W
Number of subjects included in analysis	89
Analysis specification	Pre-specified
Analysis type	other
P-value	= 0.008
Method	Mixed models analysis
Parameter estimate	LS Mean difference
Point estimate	-14.5
Confidence interval	
level	95 %
sides	2-sided
lower limit	-25.1
upper limit	-3.9
Variability estimate	Standard error of the mean
Dispersion value	5.38

Statistical analysis title	Placebo versus Vupanorsen: 80 mg Q2W - LDL-C
Statistical analysis description:	
MMRM with baseline value, treatment, visit, interaction term of treatment by visit as fixed effects and unstructured covariance structure was used.	
Comparison groups	Placebo v Vupanorsen: 80 mg Q2W
Number of subjects included in analysis	86
Analysis specification	Pre-specified
Analysis type	other
P-value	= 0.004
Method	Mixed models analysis
Parameter estimate	LS Mean difference
Point estimate	-16
Confidence interval	
level	95 %
sides	2-sided
lower limit	-26.7
upper limit	-5.3
Variability estimate	Standard error of the mean
Dispersion value	5.44

Statistical analysis title	Placebo versus Vupanorsen: 120 mg Q2W - LDL-C
Statistical analysis description: MMRM with baseline value, treatment, visit, interaction term of treatment by visit as fixed effects and unstructured covariance structure was used.	
Comparison groups	Placebo v Vupanorsen: 120 mg Q2W
Number of subjects included in analysis	90
Analysis specification	Pre-specified
Analysis type	other
P-value	= 0.136
Method	Mixed models analysis
Parameter estimate	LS Mean difference
Point estimate	-7.9
Confidence interval	
level	95 %
sides	2-sided
lower limit	-18.3
upper limit	2.5
Variability estimate	Standard error of the mean
Dispersion value	5.28

Statistical analysis title	Placebo versus Vupanorsen: 160 mg Q2W - LDL-C
Statistical analysis description: MMRM with baseline value, treatment, visit, interaction term of treatment by visit as fixed effects and unstructured covariance structure was used.	
Comparison groups	Placebo v Vupanorsen: 160 mg Q2W
Number of subjects included in analysis	79
Analysis specification	Pre-specified
Analysis type	other
P-value	= 0.138
Method	Mixed models analysis
Parameter estimate	LS Mean difference
Point estimate	-9
Confidence interval	
level	95 %
sides	2-sided
lower limit	-20.8
upper limit	2.9
Variability estimate	Standard error of the mean
Dispersion value	6.01

Secondary: Percent Change From Baseline in Angiopoietin-like Protein 3 (ANGPTL3) at Week 16

End point title	Percent Change From Baseline in Angiopoietin-like Protein 3 (ANGPTL3) at Week 16

End point description:

ANGPTL3 is a protein primarily synthesized and secreted by the liver and is a member of the angiopoietin-like family of proteins. Blood samples were collected from subjects in a fasted state for the measurement of ANGPTL3. Fasting was required at least 10 hours before blood sample collection. Baseline was calculated using the average of all values obtained at Screening and on Day 1 prior to dosing. FAS included all subjects randomised to study intervention and who took at least 1 dose of study intervention and had a baseline measurement and at least one post-baseline measurement. Here, "Number of Subjects Analyzed" signifies subjects evaluable for this endpoint.

End point type	Secondary

End point timeframe:

Baseline, Week 16

End point values	Placebo	Vupanorsen: 80 mg Q4W	Vupanorsen: 60 mg Q2W	Vupanorsen: 120 mg Q4W
Subject group type	Reporting group	Reporting group	Reporting group	Reporting group
Number of subjects analysed	41	21	21	21
Units: Percent change				
arithmetic mean (standard deviation)	9.14 (± 36.729)	-51.12 (± 30.444)	-63.61 (± 16.934)	-58.19 (± 22.175)

End point values	Vupanorsen: 80 mg Q2W	Vupanorsen: 160 mg Q4W	Vupanorsen: 120 mg Q2W	Vupanorsen: 160 mg Q2W
Subject group type	Reporting group	Reporting group	Reporting group	Reporting group
Number of subjects analysed	39	44	41	32
Units: Percent change				
arithmetic mean (standard deviation)	-65.56 (± 21.908)	-60.35 (± 21.040)	-77.55 (± 15.469)	-75.14 (± 23.549)

Statistical analyses

No statistical analyses for this end point

Secondary: Percent Change From Baseline in ANGPTL3 at Week 24

End point title	Percent Change From Baseline in ANGPTL3 at Week 24

End point description:

ANGPTL3 is a protein primarily synthesized and secreted by the liver and is a member of the angiopoietin-like family of proteins. Blood samples were collected from subjects in a fasted state for the measurement of ANGPTL3. Fasting was required at least 10 hours before blood sample collection. Baseline was calculated using the average of all values obtained at Screening and on Day 1 prior to dosing. FAS_primary included all subjects randomised to study intervention and who took at least 1 dose of study intervention, had a baseline measurement and at least one post-baseline measurement with all observations that occurred after discontinuation of treatment or after initiation of severe hypertriglyceridaemia excluded. Here, "Number of Subjects Analysed" signifies subjects evaluable for this endpoint.

End point type	Secondary

End point timeframe:

Baseline, Week 24

End point values	Placebo	Vupanorsen: 80 mg Q4W	Vupanorsen: 60 mg Q2W	Vupanorsen: 120 mg Q4W
Subject group type	Reporting group	Reporting group	Reporting group	Reporting group
Number of subjects analysed	42	22	21	22
Units: Percent change				
least squares mean (standard error)	13.3 (± 3.36)	-56.6 (± 4.92)	-66.3 (± 5.01)	-63.8 (± 5.22)

End point values	Vupanorsen: 80 mg Q2W	Vupanorsen: 160 mg Q4W	Vupanorsen: 120 mg Q2W	Vupanorsen: 160 mg Q2W
Subject group type	Reporting group	Reporting group	Reporting group	Reporting group
Number of subjects analysed	40	45	42	33
Units: Percent change				
least squares mean (standard error)	-73.0 (± 3.76)	-67.1 (± 3.46)	-78.9 (± 3.58)	-81.9 (± 4.48)

Statistical analyses

Statistical analysis title	Placebo versus Vupanorsen: 80 mg Q4W
Statistical analysis description:	
MMRM with baseline value, treatment, visit, interaction term of treatment by visit as fixed effects and unstructured covariance structure was used.	
Comparison groups	Placebo v Vupanorsen: 80 mg Q4W
Number of subjects included in analysis	64
Analysis specification	Pre-specified
Analysis type	other
P-value	< 0.001
Method	Mixed models analysis
Parameter estimate	LS Mean difference
Point estimate	-69.9
Confidence interval	
level	95 %
sides	2-sided
lower limit	-81.6
upper limit	-58.1
Variability estimate	Standard error of the mean
Dispersion value	5.97

Statistical analysis title	Placebo versus Vupanorsen: 60 mg Q2W
Statistical analysis description:	
MMRM with baseline value, treatment, visit, interaction term of treatment by visit as fixed effects and unstructured covariance structure was used.	
Comparison groups	Placebo v Vupanorsen: 60 mg Q2W

Number of subjects included in analysis	63
Analysis specification	Pre-specified
Analysis type	other
P-value	< 0.001
Method	Mixed models analysis
Parameter estimate	LS Mean difference
Point estimate	-79.6
Confidence interval	
level	95 %
sides	2-sided
lower limit	-91.5
upper limit	-67.7
Variability estimate	Standard error of the mean
Dispersion value	6.03

Statistical analysis title	Placebo versus Vupanorsen: 120 mg Q4W

Statistical analysis description:

MMRM with baseline value, treatment, visit, interaction term of treatment by visit as fixed effects and unstructured covariance structure was used.

Comparison groups	Placebo v Vupanorsen: 120 mg Q4W
Number of subjects included in analysis	64
Analysis specification	Pre-specified
Analysis type	other
P-value	< 0.001
Method	Mixed models analysis
Parameter estimate	LS Mean difference
Point estimate	-77.1
Confidence interval	
level	95 %
sides	2-sided
lower limit	-89.4
upper limit	-64.9
Variability estimate	Standard error of the mean
Dispersion value	6.22

Statistical analysis title	Placebo versus Vupanorsen: 80 mg Q2W

Statistical analysis description:

MMRM with baseline value, treatment, visit, interaction term of treatment by visit as fixed effects and unstructured covariance structure was used.

Comparison groups	Placebo v Vupanorsen: 80 mg Q2W
Number of subjects included in analysis	82
Analysis specification	Pre-specified
Analysis type	other
P-value	< 0.001
Method	Mixed models analysis
Parameter estimate	LS Mean difference
Point estimate	-86.3

Confidence interval	
level	95 %
sides	2-sided
lower limit	-96.2
upper limit	-76.3
Variability estimate	Standard error of the mean
Dispersion value	5.04

Statistical analysis title	Placebo versus Vupanorsen: 160 mg Q4W

Statistical analysis description:

MMRM with baseline value, treatment, visit, interaction term of treatment by visit as fixed effects and unstructured covariance structure was used.

Comparison groups	Placebo v Vupanorsen: 160 mg Q4W
Number of subjects included in analysis	87
Analysis specification	Pre-specified
Analysis type	other
P-value	< 0.001
Method	Mixed models analysis
Parameter estimate	LS Mean difference
Point estimate	-80.4
Confidence interval	
level	95 %
sides	2-sided
lower limit	-89.9
upper limit	-70.9
Variability estimate	Standard error of the mean
Dispersion value	4.82

Statistical analysis title	Placebo versus Vupanorsen: 120 mg Q2W

Statistical analysis description:

MMRM with baseline value, treatment, visit, interaction term of treatment by visit as fixed effects and unstructured covariance structure was used.

Comparison groups	Placebo v Vupanorsen: 120 mg Q2W
Number of subjects included in analysis	84
Analysis specification	Pre-specified
Analysis type	other
P-value	< 0.001
Method	Mixed models analysis
Parameter estimate	LS Mean difference
Point estimate	-92.2
Confidence interval	
level	95 %
sides	2-sided
lower limit	-101.9
upper limit	-82.6
Variability estimate	Standard error of the mean
Dispersion value	4.92

Statistical analysis title	Placebo versus Vupanorsen: 160 mg Q2W
Statistical analysis description: MMRM with baseline value, treatment, visit, interaction term of treatment by visit as fixed effects and unstructured covariance structure was used.	
Comparison groups	Placebo v Vupanorsen: 160 mg Q2W
Number of subjects included in analysis	75
Analysis specification	Pre-specified
Analysis type	other
P-value	< 0.001
Method	Mixed models analysis
Parameter estimate	LS Mean difference
Point estimate	-95.2
Confidence interval	
level	95 %
sides	2-sided
lower limit	-106.2
upper limit	-84.2
Variability estimate	Standard error of the mean
Dispersion value	5.59

Adverse events

Adverse events information

Timeframe for reporting adverse events:

From start of study intervention up to 12 weeks after last dose of study intervention (maximum for 36 weeks)

Adverse event reporting additional description:

Same event may appear as both an AE and SAE. However, what is presented are distinct events. An event may be categorised as serious in 1 subject and as non-serious in another, or a subject may have experienced both a serious and non-serious event.

Assessment type	Non-systematic

Dictionary used

Dictionary name	MedDRA
Dictionary version	24.1

Reporting groups

Reporting group title	Placebo

Reporting group description:

Subjects were randomised to receive vupanorsen (PF-07285557) matched placebo SC injection. Single or double administration was given at Q2W or Q4W to match active treatment groups. Administration locations were upper arm, thigh, or abdomen quadrant, as preferred by the subject. Treatment duration was up to 24 weeks. Subjects were followed up to 12 weeks after last dose of study intervention.

Reporting group title	Vupanorsen: 80 mg Q4W

Reporting group description:

Subjects were randomised to receive vupanorsen 80 mg single SC injection at Q4W. Administration locations were upper arm, thigh, or abdomen quadrant, as preferred by the subject. Treatment duration was up to 24 weeks. Subjects were followed up to 12 weeks after last dose of study intervention.

Reporting group title	Vupanorsen: 60 mg Q2W

Reporting group description:

Subjects were randomised to receive vupanorsen 60 mg single SC injection at Q2W. Administration locations were upper arm, thigh, or abdomen quadrant, as preferred by the subject. Treatment duration was up to 24 weeks. Subjects were followed up to 12 weeks after last dose of study intervention.

Reporting group title	Vupanorsen: 120 mg Q4W

Reporting group description:

Subjects were randomised to receive vupanorsen 60 mg double SC injection (120 mg total) in different locations at Q4W. Administration locations were upper arm, thigh, or abdomen quadrant, as preferred by the subject. Treatment duration was up to 24 weeks. Subjects were followed up to 12 weeks after last dose of study intervention.

Reporting group title	Vupanorsen: 160 mg Q4W

Reporting group description:

Subjects were randomised to receive vupanorsen 80 mg double SC injection (160 mg total) in different locations at Q4W. Administration locations were upper arm, thigh, or abdomen quadrant, as preferred by the subject. Treatment duration was up to 24 weeks. Subjects were followed up to 12 weeks after last dose of study intervention.

Reporting group title	Vupanorsen: 80 mg Q2W

Reporting group description:

Subjects were randomised to receive vupanorsen 80 mg single SC injection at Q2W. Administration locations were upper arm, thigh, or abdomen quadrant, as preferred by the subject. Treatment duration was up to 24 weeks. Subjects were followed up to 12 weeks after last dose of study intervention.

Reporting group title	Vupanorsen: 120 mg Q2W

Reporting group description:

Subjects were randomised to receive vupanorsen 60 mg double SC injection (120 mg in total) in different locations at Q2W. Administration locations were upper arm, thigh, or abdomen quadrant, as preferred by the subject. Treatment duration was up to 24 weeks. Subjects were followed up to 12 weeks after last dose of study intervention.

Reporting group title	Vupanorsen: 160 mg Q2W

Reporting group description:

Subjects were randomised to receive vupanorsen 80 mg double SC injection (160 mg in total) in different locations at Q2W. Administration locations were upper arm, thigh, or abdomen quadrant, as preferred by the subject. Treatment duration was up to 24 weeks. Subjects were followed up to 12 weeks after last dose of study intervention.

Serious adverse events	Placebo	Vupanorsen: 80 mg Q4W	Vupanorsen: 60 mg Q2W
Total subjects affected by serious adverse events			
subjects affected / exposed	4 / 44 (9.09%)	2 / 23 (8.70%)	2 / 24 (8.33%)
number of deaths (all causes)	0	0	0
number of deaths resulting from adverse events			
Investigations			
SARS-CoV-2 test positive			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Neoplasms benign, malignant and unspecified (incl cysts and polyps)			
Acoustic neuroma			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Basal cell carcinoma			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	1 / 24 (4.17%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Bladder cancer			
subjects affected / exposed	1 / 44 (2.27%)	0 / 23 (0.00%)	0 / 24 (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
Vascular disorders			
Thrombosis			
subjects affected / exposed	0 / 44 (0.00%)	1 / 23 (4.35%)	0 / 24 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cardiac disorders			
Acute coronary syndrome			

subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Acute myocardial infarction			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Atrial fibrillation			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Nervous system disorders			
Ischaemic stroke			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Seizure			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
General disorders and administration site conditions			
Chest pain			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hepatobiliary disorders			
Cholangitis			
subjects affected / exposed	1 / 44 (2.27%)	0 / 23 (0.00%)	0 / 24 (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
Psychiatric disorders			
Depression			

subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Musculoskeletal and connective tissue disorders			
Back pain			
subjects affected / exposed	0 / 44 (0.00%)	1 / 23 (4.35%)	0 / 24 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Osteoarthritis			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Spinal stenosis			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Infections and infestations			
Gastroenteritis			
subjects affected / exposed	0 / 44 (0.00%)	1 / 23 (4.35%)	0 / 24 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Urinary tract infection			
subjects affected / exposed	1 / 44 (2.27%)	0 / 23 (0.00%)	1 / 24 (4.17%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Metabolism and nutrition disorders			
Dehydration			
subjects affected / exposed	1 / 44 (2.27%)	0 / 23 (0.00%)	0 / 24 (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

Serious adverse events	Vupanorsen: 120 mg Q4W	Vupanorsen: 160 mg Q4W	Vupanorsen: 80 mg Q2W
Total subjects affected by serious adverse events			
subjects affected / exposed	0 / 23 (0.00%)	1 / 45 (2.22%)	6 / 45 (13.33%)

number of deaths (all causes)	0	0	0
number of deaths resulting from adverse events			
Investigations			
SARS-CoV-2 test positive			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Neoplasms benign, malignant and unspecified (incl cysts and polyps)			
Acoustic neuroma			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	1 / 45 (2.22%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Basal cell carcinoma			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Bladder cancer			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Vascular disorders			
Thrombosis			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cardiac disorders			
Acute coronary syndrome			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Acute myocardial infarction			
subjects affected / exposed	0 / 23 (0.00%)	1 / 45 (2.22%)	0 / 45 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Atrial fibrillation			

subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	1 / 45 (2.22%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Nervous system disorders			
Ischaemic stroke			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Seizure			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	1 / 45 (2.22%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
General disorders and administration site conditions			
Chest pain			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	1 / 45 (2.22%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Hepatobiliary disorders			
Cholangitis			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Psychiatric disorders			
Depression			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Musculoskeletal and connective tissue disorders			
Back pain			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Osteoarthritis			

subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	1 / 45 (2.22%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Spinal stenosis			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	1 / 45 (2.22%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Infections and infestations			
Gastroenteritis			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Urinary tract infection			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Metabolism and nutrition disorders			
Dehydration			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0

Serious adverse events	Vupanorsen: 120 mg Q2W	Vupanorsen: 160 mg Q2W	
Total subjects affected by serious adverse events			
subjects affected / exposed	3 / 46 (6.52%)	1 / 36 (2.78%)	
number of deaths (all causes)	0	0	
number of deaths resulting from adverse events			
Investigations			
SARS-CoV-2 test positive			
subjects affected / exposed	1 / 46 (2.17%)	0 / 36 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Neoplasms benign, malignant and unspecified (incl cysts and polyps)			
Acoustic neuroma			

subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Basal cell carcinoma			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Bladder cancer			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Vascular disorders			
Thrombosis			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Cardiac disorders			
Acute coronary syndrome			
subjects affected / exposed	1 / 46 (2.17%)	0 / 36 (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	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Atrial fibrillation			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Nervous system disorders			
Ischaemic stroke			
subjects affected / exposed	0 / 46 (0.00%)	1 / 36 (2.78%)	
occurrences causally related to treatment / all	0 / 0	0 / 1	
deaths causally related to treatment / all	0 / 0	0 / 0	

Seizure			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
General disorders and administration site conditions			
Chest pain			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Hepatobiliary disorders			
Cholangitis			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Psychiatric disorders			
Depression			
subjects affected / exposed	1 / 46 (2.17%)	0 / 36 (0.00%)	
occurrences causally related to treatment / all	0 / 1	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Musculoskeletal and connective tissue disorders			
Back pain			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Osteoarthritis			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences causally related to treatment / all	0 / 0	0 / 0	
deaths causally related to treatment / all	0 / 0	0 / 0	
Spinal stenosis			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (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 / 46 (0.00%)	0 / 36 (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 / 46 (0.00%)	0 / 36 (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			
Dehydration			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (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: 2 %

Non-serious adverse events	Placebo	Vupanorsen: 80 mg Q4W	Vupanorsen: 60 mg Q2W
Total subjects affected by non-serious adverse events			
subjects affected / exposed	31 / 44 (70.45%)	15 / 23 (65.22%)	17 / 24 (70.83%)
Neoplasms benign, malignant and unspecified (incl cysts and polyps)			
Basal cell carcinoma			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Vascular disorders			
Deep vein thrombosis			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Hot flush			
subjects affected / exposed	1 / 44 (2.27%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	1	0	0
Hypertension			
subjects affected / exposed	3 / 44 (6.82%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	3	0	0
Hypotension			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	1 / 24 (4.17%)
occurrences (all)	0	0	1

Peripheral coldness			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
General disorders and administration site conditions			
Asthenia			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Chest discomfort			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Chest pain			
subjects affected / exposed	1 / 44 (2.27%)	0 / 23 (0.00%)	1 / 24 (4.17%)
occurrences (all)	1	0	2
Chills			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Fatigue			
subjects affected / exposed	2 / 44 (4.55%)	1 / 23 (4.35%)	1 / 24 (4.17%)
occurrences (all)	2	1	1
Hunger			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Injection site bruising			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Injection site dermatitis			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Injection site erythema			
subjects affected / exposed	0 / 44 (0.00%)	1 / 23 (4.35%)	1 / 24 (4.17%)
occurrences (all)	0	2	1
Injection site pain			
subjects affected / exposed	1 / 44 (2.27%)	1 / 23 (4.35%)	0 / 24 (0.00%)
occurrences (all)	1	1	0
Injection site rash			

subjects affected / exposed	0 / 44 (0.00%)	1 / 23 (4.35%)	1 / 24 (4.17%)
occurrences (all)	0	2	1
Injection site reaction			
subjects affected / exposed	0 / 44 (0.00%)	1 / 23 (4.35%)	3 / 24 (12.50%)
occurrences (all)	0	2	7
Injection site recall reaction			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	2 / 24 (8.33%)
occurrences (all)	0	0	2
Injection site vesicles			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Malaise			
subjects affected / exposed	1 / 44 (2.27%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	1	0	0
Non-cardiac chest pain			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Oedema peripheral			
subjects affected / exposed	1 / 44 (2.27%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	1	0	0
Pain			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	1 / 24 (4.17%)
occurrences (all)	0	0	1
Peripheral swelling			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Pyrexia			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Therapeutic response unexpected			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Thirst			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Vessel puncture site bruise			

subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Immune system disorders			
Hypersensitivity			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Seasonal allergy			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Reproductive system and breast disorders			
Abnormal uterine bleeding			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Atrophic vulvovaginitis			
subjects affected / exposed	0 / 44 (0.00%)	1 / 23 (4.35%)	0 / 24 (0.00%)
occurrences (all)	0	1	0
Erectile dysfunction			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Genital atrophy			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	1 / 24 (4.17%)
occurrences (all)	0	0	1
Prostatitis			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Vulvovaginal dryness			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Respiratory, thoracic and mediastinal disorders			
Cough			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Dyspnoea			
subjects affected / exposed	2 / 44 (4.55%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	2	0	0
Epistaxis			

subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Haemoptysis			
subjects affected / exposed	1 / 44 (2.27%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	1	0	0
Nasal congestion			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Productive cough			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Pulmonary mass			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	1 / 24 (4.17%)
occurrences (all)	0	0	1
Rhinitis allergic			
subjects affected / exposed	2 / 44 (4.55%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	2	0	0
Sinus pain			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Sputum increased			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Psychiatric disorders			
Adjustment disorder			
subjects affected / exposed	0 / 44 (0.00%)	1 / 23 (4.35%)	0 / 24 (0.00%)
occurrences (all)	0	1	0
Anxiety			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Depression			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Insomnia			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0

Major depression			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Investigations			
Activated partial thromboplastin time prolonged			
subjects affected / exposed	1 / 44 (2.27%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	1	0	0
Alanine aminotransferase abnormal			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Alanine aminotransferase increased			
subjects affected / exposed	2 / 44 (4.55%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	2	0	0
Albumin urine present			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Aspartate aminotransferase increased			
subjects affected / exposed	1 / 44 (2.27%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	1	0	0
Blood calcium increased			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Blood glucose increased			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Blood potassium abnormal			
subjects affected / exposed	1 / 44 (2.27%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	1	0	0
Blood pressure decreased			
subjects affected / exposed	1 / 44 (2.27%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	1	0	0
Blood pressure increased			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Body temperature increased			

subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Creatinine renal clearance abnormal			
subjects affected / exposed	1 / 44 (2.27%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	1	0	0
Electrocardiogram QRS complex prolonged			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Electrocardiogram QT prolonged			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Electrocardiogram abnormal			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	1 / 24 (4.17%)
occurrences (all)	0	0	1
Haematocrit increased			
subjects affected / exposed	0 / 44 (0.00%)	1 / 23 (4.35%)	0 / 24 (0.00%)
occurrences (all)	0	1	0
Liver function test abnormal			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Hepatic enzyme increased			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	1 / 24 (4.17%)
occurrences (all)	0	0	1
Liver function test increased			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Magnetic resonance imaging spinal abnormal			
subjects affected / exposed	1 / 44 (2.27%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	1	0	0
Mean cell haemoglobin concentration decreased			
subjects affected / exposed	0 / 44 (0.00%)	1 / 23 (4.35%)	0 / 24 (0.00%)
occurrences (all)	0	1	0
Mean cell volume increased			

subjects affected / exposed	0 / 44 (0.00%)	1 / 23 (4.35%)	0 / 24 (0.00%)
occurrences (all)	0	1	0
Urine albumin/creatinine ratio abnormal			
subjects affected / exposed	1 / 44 (2.27%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	1	0	0
Urine albumin/creatinine ratio increased			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Urine analysis abnormal			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	2 / 24 (8.33%)
occurrences (all)	0	0	2
Weight decreased			
subjects affected / exposed	1 / 44 (2.27%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	1	0	0
Transaminases increased			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Injury, poisoning and procedural complications			
Animal bite			
subjects affected / exposed	1 / 44 (2.27%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	1	0	0
Arthropod bite			
subjects affected / exposed	1 / 44 (2.27%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	1	0	0
Back injury			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Epicondylitis			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Fall			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	1 / 24 (4.17%)
occurrences (all)	0	0	1
Foot fracture			

subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Head injury			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Injection related reaction			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Ligament sprain			
subjects affected / exposed	1 / 44 (2.27%)	0 / 23 (0.00%)	1 / 24 (4.17%)
occurrences (all)	1	0	1
Muscle rupture			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	1 / 24 (4.17%)
occurrences (all)	0	0	1
Post procedural discomfort			
subjects affected / exposed	1 / 44 (2.27%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	1	0	0
Procedural pain			
subjects affected / exposed	0 / 44 (0.00%)	1 / 23 (4.35%)	0 / 24 (0.00%)
occurrences (all)	0	1	0
Skin abrasion			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	1 / 24 (4.17%)
occurrences (all)	0	0	1
Vaccination complication			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Cardiac disorders			
Atrial fibrillation			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Brugada syndrome			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	1 / 24 (4.17%)
occurrences (all)	0	0	1
Myocardial injury			
subjects affected / exposed	1 / 44 (2.27%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	1	0	0

Palpitations			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Sinus tachycardia			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Nervous system disorders			
Balance disorder			
subjects affected / exposed	0 / 44 (0.00%)	1 / 23 (4.35%)	0 / 24 (0.00%)
occurrences (all)	0	1	0
Carpal tunnel syndrome			
subjects affected / exposed	1 / 44 (2.27%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	1	0	0
Dizziness			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Headache			
subjects affected / exposed	2 / 44 (4.55%)	1 / 23 (4.35%)	1 / 24 (4.17%)
occurrences (all)	2	1	1
Hypoaesthesia			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Migraine			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Neuropathy peripheral			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Paraesthesia			
subjects affected / exposed	0 / 44 (0.00%)	1 / 23 (4.35%)	0 / 24 (0.00%)
occurrences (all)	0	1	0
Radiculopathy			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Sciatica			

subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Sensory disturbance			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Transient ischaemic attack			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Blood and lymphatic system disorders			
Anaemia			
subjects affected / exposed	1 / 44 (2.27%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	1	0	0
Eosinophilia			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Ear and labyrinth disorders			
Ear pain			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Vertigo			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Eye disorders			
Cataract			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Conjunctivitis allergic			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Ulcerative keratitis			
subjects affected / exposed	1 / 44 (2.27%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	1	0	0
Vitreous floaters			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Gastrointestinal disorders			

Abdominal discomfort			
subjects affected / exposed	1 / 44 (2.27%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	1	0	0
Abdominal distension			
subjects affected / exposed	0 / 44 (0.00%)	1 / 23 (4.35%)	0 / 24 (0.00%)
occurrences (all)	0	1	0
Abdominal pain			
subjects affected / exposed	1 / 44 (2.27%)	1 / 23 (4.35%)	0 / 24 (0.00%)
occurrences (all)	1	1	0
Abdominal pain upper			
subjects affected / exposed	2 / 44 (4.55%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	2	0	0
Acid peptic disease			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	1 / 24 (4.17%)
occurrences (all)	0	0	1
Constipation			
subjects affected / exposed	1 / 44 (2.27%)	1 / 23 (4.35%)	0 / 24 (0.00%)
occurrences (all)	1	1	0
Crohn's disease			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Dental necrosis			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Diarrhoea			
subjects affected / exposed	3 / 44 (6.82%)	1 / 23 (4.35%)	0 / 24 (0.00%)
occurrences (all)	3	1	0
Diverticulum			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Dyspepsia			
subjects affected / exposed	0 / 44 (0.00%)	1 / 23 (4.35%)	0 / 24 (0.00%)
occurrences (all)	0	1	0
Faeces soft			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0

Gastrooesophageal reflux disease			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Haematochezia			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Nausea			
subjects affected / exposed	2 / 44 (4.55%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	3	0	0
Oesophageal pain			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Rectal haemorrhage			
subjects affected / exposed	0 / 44 (0.00%)	1 / 23 (4.35%)	0 / 24 (0.00%)
occurrences (all)	0	1	0
Toothache			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Vomiting			
subjects affected / exposed	1 / 44 (2.27%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	2	0	0
Hepatobiliary disorders			
Cholelithiasis			
subjects affected / exposed	1 / 44 (2.27%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	1	0	0
Drug-induced liver injury			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Hepatic steatosis			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Nonalcoholic fatty liver disease			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Skin and subcutaneous tissue disorders			

Actinic keratosis			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Alopecia			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Dermatitis			
subjects affected / exposed	0 / 44 (0.00%)	1 / 23 (4.35%)	0 / 24 (0.00%)
occurrences (all)	0	1	0
Dermatitis allergic			
subjects affected / exposed	2 / 44 (4.55%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	2	0	0
Dermatitis contact			
subjects affected / exposed	1 / 44 (2.27%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	1	0	0
Erythema			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Hyperhidrosis			
subjects affected / exposed	0 / 44 (0.00%)	1 / 23 (4.35%)	0 / 24 (0.00%)
occurrences (all)	0	2	0
Miliaria			
subjects affected / exposed	0 / 44 (0.00%)	1 / 23 (4.35%)	0 / 24 (0.00%)
occurrences (all)	0	1	0
Petechiae			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Pruritus			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Rash			
subjects affected / exposed	0 / 44 (0.00%)	1 / 23 (4.35%)	0 / 24 (0.00%)
occurrences (all)	0	1	0
Rosacea			
subjects affected / exposed	1 / 44 (2.27%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	1	0	0

Skin discolouration			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Skin lesion			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Skin mass			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Renal and urinary disorders			
Acute kidney injury			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	2 / 24 (8.33%)
occurrences (all)	0	0	2
Calculus bladder			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Chronic kidney disease			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Dysuria			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Haematuria			
subjects affected / exposed	1 / 44 (2.27%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	1	0	0
Microalbuminuria			
subjects affected / exposed	1 / 44 (2.27%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	1	0	0
Nephrolithiasis			
subjects affected / exposed	1 / 44 (2.27%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	1	0	0
Pollakiuria			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Proteinuria			

subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Renal cyst			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Renal failure			
subjects affected / exposed	1 / 44 (2.27%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	1	0	0
Urge incontinence			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Urinary incontinence			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Musculoskeletal and connective tissue disorders			
Arthralgia			
subjects affected / exposed	2 / 44 (4.55%)	1 / 23 (4.35%)	1 / 24 (4.17%)
occurrences (all)	2	2	1
Arthritis			
subjects affected / exposed	1 / 44 (2.27%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	1	0	0
Bone cyst			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Bursitis			
subjects affected / exposed	1 / 44 (2.27%)	1 / 23 (4.35%)	0 / 24 (0.00%)
occurrences (all)	1	1	0
Flank pain			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Intervertebral disc degeneration			
subjects affected / exposed	1 / 44 (2.27%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	1	0	0
Intervertebral disc disorder			

subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	1 / 24 (4.17%)
occurrences (all)	0	0	1
Joint stiffness			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Joint swelling			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	1 / 24 (4.17%)
occurrences (all)	0	0	1
Muscle fatigue			
subjects affected / exposed	1 / 44 (2.27%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	1	0	0
Muscle spasms			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Muscular weakness			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Musculoskeletal chest pain			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	1 / 24 (4.17%)
occurrences (all)	0	0	1
Musculoskeletal pain			
subjects affected / exposed	1 / 44 (2.27%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	1	0	0
Myalgia			
subjects affected / exposed	1 / 44 (2.27%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	1	0	0
Neck pain			
subjects affected / exposed	1 / 44 (2.27%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	1	0	0
Osteoarthritis			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	2 / 24 (8.33%)
occurrences (all)	0	0	2
Pain in extremity			
subjects affected / exposed	1 / 44 (2.27%)	0 / 23 (0.00%)	3 / 24 (12.50%)
occurrences (all)	1	0	4
Plantar fascial fibromatosis			

subjects affected / exposed	0 / 44 (0.00%)	1 / 23 (4.35%)	0 / 24 (0.00%)
occurrences (all)	0	1	0
Rotator cuff syndrome			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	1 / 24 (4.17%)
occurrences (all)	0	0	1
Scoliosis			
subjects affected / exposed	1 / 44 (2.27%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	1	0	0
Spinal pain			
subjects affected / exposed	1 / 44 (2.27%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	1	0	0
Temporomandibular joint syndrome			
subjects affected / exposed	1 / 44 (2.27%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	1	0	0
Tendonitis			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	1 / 24 (4.17%)
occurrences (all)	0	0	1
Tenosynovitis			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Back pain			
subjects affected / exposed	2 / 44 (4.55%)	0 / 23 (0.00%)	2 / 24 (8.33%)
occurrences (all)	2	0	2
Infections and infestations			
Acute sinusitis			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Bronchitis			
subjects affected / exposed	2 / 44 (4.55%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	2	0	0
COVID-19			
subjects affected / exposed	1 / 44 (2.27%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	1	0	0
Cellulitis			
subjects affected / exposed	2 / 44 (4.55%)	1 / 23 (4.35%)	0 / 24 (0.00%)
occurrences (all)	2	1	0

Conjunctivitis			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	1 / 24 (4.17%)
occurrences (all)	0	0	1
Cystitis			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Fungal infection			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Fungal skin infection			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Gastroenteritis			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Gastroenteritis viral			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Gastrointestinal bacterial infection			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Helicobacter infection			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Herpes zoster			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Infection parasitic			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Injection site cellulitis			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Kidney infection			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0

Nasopharyngitis			
subjects affected / exposed	0 / 44 (0.00%)	1 / 23 (4.35%)	0 / 24 (0.00%)
occurrences (all)	0	1	0
Oral candidiasis			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Otitis externa			
subjects affected / exposed	1 / 44 (2.27%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	1	0	0
Pharyngitis			
subjects affected / exposed	1 / 44 (2.27%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	1	0	0
Sinusitis			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Tinea cruris			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Tooth abscess			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Tooth infection			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Upper respiratory tract infection			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	1 / 24 (4.17%)
occurrences (all)	0	0	1
Urinary tract infection			
subjects affected / exposed	3 / 44 (6.82%)	1 / 23 (4.35%)	1 / 24 (4.17%)
occurrences (all)	3	1	1
Viral infection			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Vulvovaginal candidiasis			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0

Vulvovaginal mycotic infection			
subjects affected / exposed	0 / 44 (0.00%)	1 / 23 (4.35%)	0 / 24 (0.00%)
occurrences (all)	0	1	0
Metabolism and nutrition disorders			
Decreased appetite			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Dehydration			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	1 / 24 (4.17%)
occurrences (all)	0	0	1
Diabetes mellitus			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Diabetes mellitus inadequate control			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Glucose tolerance impaired			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Gout			
subjects affected / exposed	1 / 44 (2.27%)	0 / 23 (0.00%)	2 / 24 (8.33%)
occurrences (all)	1	0	2
Hyperglycaemia			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Hyperkalaemia			
subjects affected / exposed	1 / 44 (2.27%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	1	0	0
Hypoglycaemia			
subjects affected / exposed	0 / 44 (0.00%)	1 / 23 (4.35%)	0 / 24 (0.00%)
occurrences (all)	0	1	0
Hypokalaemia			
subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Increased appetite			

subjects affected / exposed	0 / 44 (0.00%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	0	0	0
Type 2 diabetes mellitus			
subjects affected / exposed	1 / 44 (2.27%)	0 / 23 (0.00%)	0 / 24 (0.00%)
occurrences (all)	1	0	0

Non-serious adverse events	Vupanorsen: 120 mg Q4W	Vupanorsen: 160 mg Q4W	Vupanorsen: 80 mg Q2W
Total subjects affected by non-serious adverse events			
subjects affected / exposed	12 / 23 (52.17%)	28 / 45 (62.22%)	30 / 45 (66.67%)
Neoplasms benign, malignant and unspecified (incl cysts and polyps)			
Basal cell carcinoma			
subjects affected / exposed	0 / 23 (0.00%)	1 / 45 (2.22%)	0 / 45 (0.00%)
occurrences (all)	0	1	0
Vascular disorders			
Deep vein thrombosis			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Hot flush			
subjects affected / exposed	1 / 23 (4.35%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	1	0	0
Hypertension			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Hypotension			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Peripheral coldness			
subjects affected / exposed	0 / 23 (0.00%)	1 / 45 (2.22%)	0 / 45 (0.00%)
occurrences (all)	0	1	0
General disorders and administration site conditions			
Asthenia			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	1 / 45 (2.22%)
occurrences (all)	0	0	1
Chest discomfort			
subjects affected / exposed	1 / 23 (4.35%)	0 / 45 (0.00%)	1 / 45 (2.22%)
occurrences (all)	1	0	1

Chest pain			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Chills			
subjects affected / exposed	0 / 23 (0.00%)	1 / 45 (2.22%)	0 / 45 (0.00%)
occurrences (all)	0	1	0
Fatigue			
subjects affected / exposed	0 / 23 (0.00%)	2 / 45 (4.44%)	1 / 45 (2.22%)
occurrences (all)	0	2	2
Hunger			
subjects affected / exposed	0 / 23 (0.00%)	1 / 45 (2.22%)	0 / 45 (0.00%)
occurrences (all)	0	1	0
Injection site bruising			
subjects affected / exposed	0 / 23 (0.00%)	1 / 45 (2.22%)	0 / 45 (0.00%)
occurrences (all)	0	1	0
Injection site dermatitis			
subjects affected / exposed	1 / 23 (4.35%)	1 / 45 (2.22%)	0 / 45 (0.00%)
occurrences (all)	3	4	0
Injection site erythema			
subjects affected / exposed	2 / 23 (8.70%)	1 / 45 (2.22%)	2 / 45 (4.44%)
occurrences (all)	2	2	3
Injection site pain			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Injection site rash			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Injection site reaction			
subjects affected / exposed	4 / 23 (17.39%)	4 / 45 (8.89%)	4 / 45 (8.89%)
occurrences (all)	10	7	5
Injection site recall reaction			
subjects affected / exposed	1 / 23 (4.35%)	0 / 45 (0.00%)	2 / 45 (4.44%)
occurrences (all)	1	0	3
Injection site vesicles			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	1 / 45 (2.22%)
occurrences (all)	0	0	1

Malaise			
subjects affected / exposed	1 / 23 (4.35%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	1	0	0
Non-cardiac chest pain			
subjects affected / exposed	1 / 23 (4.35%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	1	0	0
Oedema peripheral			
subjects affected / exposed	0 / 23 (0.00%)	1 / 45 (2.22%)	2 / 45 (4.44%)
occurrences (all)	0	1	2
Pain			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Peripheral swelling			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	2 / 45 (4.44%)
occurrences (all)	0	0	2
Pyrexia			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	2 / 45 (4.44%)
occurrences (all)	0	0	2
Therapeutic response unexpected			
subjects affected / exposed	0 / 23 (0.00%)	1 / 45 (2.22%)	0 / 45 (0.00%)
occurrences (all)	0	1	0
Thirst			
subjects affected / exposed	1 / 23 (4.35%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	1	0	0
Vessel puncture site bruise			
subjects affected / exposed	0 / 23 (0.00%)	1 / 45 (2.22%)	1 / 45 (2.22%)
occurrences (all)	0	1	1
Immune system disorders			
Hypersensitivity			
subjects affected / exposed	0 / 23 (0.00%)	1 / 45 (2.22%)	0 / 45 (0.00%)
occurrences (all)	0	1	0
Seasonal allergy			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Reproductive system and breast disorders			

Abnormal uterine bleeding			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Atrophic vulvovaginitis			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Erectile dysfunction			
subjects affected / exposed	0 / 23 (0.00%)	1 / 45 (2.22%)	0 / 45 (0.00%)
occurrences (all)	0	1	0
Genital atrophy			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Prostatitis			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Vulvovaginal dryness			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	1 / 45 (2.22%)
occurrences (all)	0	0	1
Respiratory, thoracic and mediastinal disorders			
Cough			
subjects affected / exposed	1 / 23 (4.35%)	0 / 45 (0.00%)	1 / 45 (2.22%)
occurrences (all)	1	0	1
Dyspnoea			
subjects affected / exposed	0 / 23 (0.00%)	1 / 45 (2.22%)	1 / 45 (2.22%)
occurrences (all)	0	1	1
Epistaxis			
subjects affected / exposed	1 / 23 (4.35%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	1	0	0
Haemoptysis			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Nasal congestion			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Productive cough			

subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	1 / 45 (2.22%)
occurrences (all)	0	0	1
Pulmonary mass			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Rhinitis allergic			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Sinus pain			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Sputum increased			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	1 / 45 (2.22%)
occurrences (all)	0	0	1
Psychiatric disorders			
Adjustment disorder			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Anxiety			
subjects affected / exposed	2 / 23 (8.70%)	2 / 45 (4.44%)	1 / 45 (2.22%)
occurrences (all)	2	2	1
Depression			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Insomnia			
subjects affected / exposed	0 / 23 (0.00%)	1 / 45 (2.22%)	0 / 45 (0.00%)
occurrences (all)	0	1	0
Major depression			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	1 / 45 (2.22%)
occurrences (all)	0	0	1
Investigations			
Activated partial thromboplastin time prolonged			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	1 / 45 (2.22%)
occurrences (all)	0	0	1
Alanine aminotransferase abnormal			

subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Alanine aminotransferase increased			
subjects affected / exposed	0 / 23 (0.00%)	1 / 45 (2.22%)	1 / 45 (2.22%)
occurrences (all)	0	1	1
Albumin urine present			
subjects affected / exposed	0 / 23 (0.00%)	1 / 45 (2.22%)	0 / 45 (0.00%)
occurrences (all)	0	1	0
Aspartate aminotransferase increased			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Blood calcium increased			
subjects affected / exposed	0 / 23 (0.00%)	1 / 45 (2.22%)	0 / 45 (0.00%)
occurrences (all)	0	1	0
Blood glucose increased			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Blood potassium abnormal			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Blood pressure decreased			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Blood pressure increased			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	1 / 45 (2.22%)
occurrences (all)	0	0	1
Body temperature increased			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Creatinine renal clearance abnormal			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Electrocardiogram QRS complex prolonged			

subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	1 / 45 (2.22%)
occurrences (all)	0	0	1
Electrocardiogram QT prolonged			
subjects affected / exposed	0 / 23 (0.00%)	1 / 45 (2.22%)	1 / 45 (2.22%)
occurrences (all)	0	1	1
Electrocardiogram abnormal			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Haematocrit increased			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Liver function test abnormal			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Hepatic enzyme increased			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Liver function test increased			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	1 / 45 (2.22%)
occurrences (all)	0	0	1
Magnetic resonance imaging spinal abnormal			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Mean cell haemoglobin concentration decreased			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Mean cell volume increased			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Urine albumin/creatinine ratio abnormal			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Urine albumin/creatinine ratio increased			

subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Urine analysis abnormal			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Weight decreased			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Transaminases increased			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	2 / 45 (4.44%)
occurrences (all)	0	0	2
Injury, poisoning and procedural complications			
Animal bite			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Arthropod bite			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Back injury			
subjects affected / exposed	1 / 23 (4.35%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	1	0	0
Epicondylitis			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	1 / 45 (2.22%)
occurrences (all)	0	0	1
Fall			
subjects affected / exposed	1 / 23 (4.35%)	0 / 45 (0.00%)	1 / 45 (2.22%)
occurrences (all)	1	0	1
Foot fracture			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Head injury			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	1 / 45 (2.22%)
occurrences (all)	0	0	1
Injection related reaction			

subjects affected / exposed	0 / 23 (0.00%)	1 / 45 (2.22%)	0 / 45 (0.00%)
occurrences (all)	0	1	0
Ligament sprain			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Muscle rupture			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Post procedural discomfort			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Procedural pain			
subjects affected / exposed	0 / 23 (0.00%)	1 / 45 (2.22%)	0 / 45 (0.00%)
occurrences (all)	0	1	0
Skin abrasion			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	1 / 45 (2.22%)
occurrences (all)	0	0	1
Vaccination complication			
subjects affected / exposed	0 / 23 (0.00%)	1 / 45 (2.22%)	0 / 45 (0.00%)
occurrences (all)	0	1	0
Cardiac disorders			
Atrial fibrillation			
subjects affected / exposed	0 / 23 (0.00%)	1 / 45 (2.22%)	0 / 45 (0.00%)
occurrences (all)	0	1	0
Brugada syndrome			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Myocardial injury			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Palpitations			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	2 / 45 (4.44%)
occurrences (all)	0	0	2
Sinus tachycardia			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0

Nervous system disorders			
Balance disorder			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Carpal tunnel syndrome			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	1 / 45 (2.22%)
occurrences (all)	0	0	1
Dizziness			
subjects affected / exposed	0 / 23 (0.00%)	1 / 45 (2.22%)	2 / 45 (4.44%)
occurrences (all)	0	1	2
Headache			
subjects affected / exposed	1 / 23 (4.35%)	3 / 45 (6.67%)	2 / 45 (4.44%)
occurrences (all)	1	4	3
Hypoaesthesia			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	1 / 45 (2.22%)
occurrences (all)	0	0	1
Migraine			
subjects affected / exposed	1 / 23 (4.35%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	1	0	0
Neuropathy peripheral			
subjects affected / exposed	0 / 23 (0.00%)	1 / 45 (2.22%)	0 / 45 (0.00%)
occurrences (all)	0	1	0
Paraesthesia			
subjects affected / exposed	0 / 23 (0.00%)	2 / 45 (4.44%)	0 / 45 (0.00%)
occurrences (all)	0	2	0
Radiculopathy			
subjects affected / exposed	0 / 23 (0.00%)	1 / 45 (2.22%)	0 / 45 (0.00%)
occurrences (all)	0	1	0
Sciatica			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Sensory disturbance			
subjects affected / exposed	0 / 23 (0.00%)	1 / 45 (2.22%)	0 / 45 (0.00%)
occurrences (all)	0	1	0
Transient ischaemic attack			

subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Blood and lymphatic system disorders			
Anaemia			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Eosinophilia			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Ear and labyrinth disorders			
Ear pain			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Vertigo			
subjects affected / exposed	0 / 23 (0.00%)	1 / 45 (2.22%)	0 / 45 (0.00%)
occurrences (all)	0	1	0
Eye disorders			
Cataract			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	1 / 45 (2.22%)
occurrences (all)	0	0	2
Conjunctivitis allergic			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Ulcerative keratitis			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Vitreous floaters			
subjects affected / exposed	0 / 23 (0.00%)	1 / 45 (2.22%)	0 / 45 (0.00%)
occurrences (all)	0	1	0
Gastrointestinal disorders			
Abdominal discomfort			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Abdominal distension			
subjects affected / exposed	0 / 23 (0.00%)	1 / 45 (2.22%)	0 / 45 (0.00%)
occurrences (all)	0	1	0
Abdominal pain			

subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Abdominal pain upper			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	1 / 45 (2.22%)
occurrences (all)	0	0	1
Acid peptic disease			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Constipation			
subjects affected / exposed	1 / 23 (4.35%)	1 / 45 (2.22%)	1 / 45 (2.22%)
occurrences (all)	1	1	1
Crohn's disease			
subjects affected / exposed	0 / 23 (0.00%)	1 / 45 (2.22%)	0 / 45 (0.00%)
occurrences (all)	0	1	0
Dental necrosis			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	1 / 45 (2.22%)
occurrences (all)	0	0	1
Diarrhoea			
subjects affected / exposed	0 / 23 (0.00%)	1 / 45 (2.22%)	1 / 45 (2.22%)
occurrences (all)	0	2	1
Diverticulum			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Dyspepsia			
subjects affected / exposed	1 / 23 (4.35%)	1 / 45 (2.22%)	0 / 45 (0.00%)
occurrences (all)	1	1	0
Faeces soft			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Gastrooesophageal reflux disease			
subjects affected / exposed	0 / 23 (0.00%)	2 / 45 (4.44%)	0 / 45 (0.00%)
occurrences (all)	0	2	0
Haematochezia			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	1 / 45 (2.22%)
occurrences (all)	0	0	1
Nausea			

subjects affected / exposed	1 / 23 (4.35%)	1 / 45 (2.22%)	2 / 45 (4.44%)
occurrences (all)	1	1	2
Oesophageal pain			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Rectal haemorrhage			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Toothache			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Vomiting			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	1 / 45 (2.22%)
occurrences (all)	0	0	2
Hepatobiliary disorders			
Cholelithiasis			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Drug-induced liver injury			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	1 / 45 (2.22%)
occurrences (all)	0	0	1
Hepatic steatosis			
subjects affected / exposed	0 / 23 (0.00%)	1 / 45 (2.22%)	0 / 45 (0.00%)
occurrences (all)	0	1	0
Nonalcoholic fatty liver disease			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	1 / 45 (2.22%)
occurrences (all)	0	0	1
Skin and subcutaneous tissue disorders			
Actinic keratosis			
subjects affected / exposed	0 / 23 (0.00%)	1 / 45 (2.22%)	0 / 45 (0.00%)
occurrences (all)	0	1	0
Alopecia			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	1 / 45 (2.22%)
occurrences (all)	0	0	1
Dermatitis			

subjects affected / exposed	0 / 23 (0.00%)	1 / 45 (2.22%)	0 / 45 (0.00%)
occurrences (all)	0	1	0
Dermatitis allergic			
subjects affected / exposed	0 / 23 (0.00%)	1 / 45 (2.22%)	0 / 45 (0.00%)
occurrences (all)	0	2	0
Dermatitis contact			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Erythema			
subjects affected / exposed	1 / 23 (4.35%)	0 / 45 (0.00%)	1 / 45 (2.22%)
occurrences (all)	1	0	1
Hyperhidrosis			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Miliaria			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Petechiae			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Pruritus			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	1 / 45 (2.22%)
occurrences (all)	0	0	1
Rash			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	2 / 45 (4.44%)
occurrences (all)	0	0	3
Rosacea			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Skin discolouration			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Skin lesion			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	2 / 45 (4.44%)
occurrences (all)	0	0	2
Skin mass			

subjects affected / exposed	1 / 23 (4.35%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	1	0	0
Renal and urinary disorders			
Acute kidney injury			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Calculus bladder			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Chronic kidney disease			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Dysuria			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Haematuria			
subjects affected / exposed	1 / 23 (4.35%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	1	0	0
Microalbuminuria			
subjects affected / exposed	0 / 23 (0.00%)	1 / 45 (2.22%)	0 / 45 (0.00%)
occurrences (all)	0	1	0
Nephrolithiasis			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	1 / 45 (2.22%)
occurrences (all)	0	0	1
Pollakiuria			
subjects affected / exposed	1 / 23 (4.35%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	1	0	0
Proteinuria			
subjects affected / exposed	0 / 23 (0.00%)	2 / 45 (4.44%)	0 / 45 (0.00%)
occurrences (all)	0	2	0
Renal cyst			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Renal failure			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0

Urge incontinence			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	1 / 45 (2.22%)
occurrences (all)	0	0	1
Urinary incontinence			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	1 / 45 (2.22%)
occurrences (all)	0	0	1
Musculoskeletal and connective tissue disorders			
Arthralgia			
subjects affected / exposed	0 / 23 (0.00%)	1 / 45 (2.22%)	4 / 45 (8.89%)
occurrences (all)	0	1	4
Arthritis			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Bone cyst			
subjects affected / exposed	0 / 23 (0.00%)	1 / 45 (2.22%)	0 / 45 (0.00%)
occurrences (all)	0	1	0
Bursitis			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Flank pain			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Intervertebral disc degeneration			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Intervertebral disc disorder			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Joint stiffness			
subjects affected / exposed	0 / 23 (0.00%)	1 / 45 (2.22%)	0 / 45 (0.00%)
occurrences (all)	0	1	0
Joint swelling			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Muscle fatigue			

subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Muscle spasms			
subjects affected / exposed	1 / 23 (4.35%)	1 / 45 (2.22%)	0 / 45 (0.00%)
occurrences (all)	1	1	0
Muscular weakness			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Musculoskeletal chest pain			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Musculoskeletal pain			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Myalgia			
subjects affected / exposed	0 / 23 (0.00%)	3 / 45 (6.67%)	2 / 45 (4.44%)
occurrences (all)	0	3	3
Neck pain			
subjects affected / exposed	0 / 23 (0.00%)	1 / 45 (2.22%)	0 / 45 (0.00%)
occurrences (all)	0	1	0
Osteoarthritis			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Pain in extremity			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	3 / 45 (6.67%)
occurrences (all)	0	0	3
Plantar fascial fibromatosis			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Rotator cuff syndrome			
subjects affected / exposed	1 / 23 (4.35%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	1	0	0
Scoliosis			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Spinal pain			

subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Temporomandibular joint syndrome			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Tendonitis			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Tenosynovitis			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Back pain			
subjects affected / exposed	0 / 23 (0.00%)	1 / 45 (2.22%)	6 / 45 (13.33%)
occurrences (all)	0	1	6
Infections and infestations			
Acute sinusitis			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Bronchitis			
subjects affected / exposed	1 / 23 (4.35%)	2 / 45 (4.44%)	0 / 45 (0.00%)
occurrences (all)	5	2	0
COVID-19			
subjects affected / exposed	1 / 23 (4.35%)	0 / 45 (0.00%)	1 / 45 (2.22%)
occurrences (all)	1	0	1
Cellulitis			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Conjunctivitis			
subjects affected / exposed	0 / 23 (0.00%)	1 / 45 (2.22%)	0 / 45 (0.00%)
occurrences (all)	0	1	0
Cystitis			
subjects affected / exposed	0 / 23 (0.00%)	1 / 45 (2.22%)	0 / 45 (0.00%)
occurrences (all)	0	1	0
Fungal infection			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	1 / 45 (2.22%)
occurrences (all)	0	0	1

Fungal skin infection			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	1 / 45 (2.22%)
occurrences (all)	0	0	1
Gastroenteritis			
subjects affected / exposed	1 / 23 (4.35%)	1 / 45 (2.22%)	0 / 45 (0.00%)
occurrences (all)	1	1	0
Gastroenteritis viral			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Gastrointestinal bacterial infection			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	1 / 45 (2.22%)
occurrences (all)	0	0	1
Helicobacter infection			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Herpes zoster			
subjects affected / exposed	1 / 23 (4.35%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	1	0	0
Infection parasitic			
subjects affected / exposed	0 / 23 (0.00%)	1 / 45 (2.22%)	0 / 45 (0.00%)
occurrences (all)	0	1	0
Injection site cellulitis			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Kidney infection			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Nasopharyngitis			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	1 / 45 (2.22%)
occurrences (all)	0	0	1
Oral candidiasis			
subjects affected / exposed	1 / 23 (4.35%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	1	0	0
Otitis externa			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0

Pharyngitis			
subjects affected / exposed	1 / 23 (4.35%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	1	0	0
Sinusitis			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	1 / 45 (2.22%)
occurrences (all)	0	0	1
Tinea cruris			
subjects affected / exposed	1 / 23 (4.35%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	1	0	0
Tooth abscess			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	1 / 45 (2.22%)
occurrences (all)	0	0	1
Tooth infection			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Upper respiratory tract infection			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Urinary tract infection			
subjects affected / exposed	1 / 23 (4.35%)	1 / 45 (2.22%)	3 / 45 (6.67%)
occurrences (all)	1	1	4
Viral infection			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	1 / 45 (2.22%)
occurrences (all)	0	0	1
Vulvovaginal candidiasis			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Vulvovaginal mycotic infection			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Metabolism and nutrition disorders			
Decreased appetite			
subjects affected / exposed	1 / 23 (4.35%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	1	0	0
Dehydration			

subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Diabetes mellitus			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Diabetes mellitus inadequate control			
subjects affected / exposed	0 / 23 (0.00%)	1 / 45 (2.22%)	0 / 45 (0.00%)
occurrences (all)	0	1	0
Glucose tolerance impaired			
subjects affected / exposed	0 / 23 (0.00%)	1 / 45 (2.22%)	0 / 45 (0.00%)
occurrences (all)	0	1	0
Gout			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Hyperglycaemia			
subjects affected / exposed	0 / 23 (0.00%)	1 / 45 (2.22%)	0 / 45 (0.00%)
occurrences (all)	0	1	0
Hyperkalaemia			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	0 / 45 (0.00%)
occurrences (all)	0	0	0
Hypoglycaemia			
subjects affected / exposed	0 / 23 (0.00%)	1 / 45 (2.22%)	0 / 45 (0.00%)
occurrences (all)	0	1	0
Hypokalaemia			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	1 / 45 (2.22%)
occurrences (all)	0	0	1
Increased appetite			
subjects affected / exposed	0 / 23 (0.00%)	0 / 45 (0.00%)	1 / 45 (2.22%)
occurrences (all)	0	0	1
Type 2 diabetes mellitus			
subjects affected / exposed	1 / 23 (4.35%)	1 / 45 (2.22%)	1 / 45 (2.22%)
occurrences (all)	1	1	1

Non-serious adverse events	Vupanorsen: 120 mg Q2W	Vupanorsen: 160 mg Q2W	
Total subjects affected by non-serious adverse events			
subjects affected / exposed	29 / 46 (63.04%)	31 / 36 (86.11%)	

Neoplasms benign, malignant and unspecified (incl cysts and polyps)			
Basal cell carcinoma			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Vascular disorders			
Deep vein thrombosis			
subjects affected / exposed	0 / 46 (0.00%)	1 / 36 (2.78%)	
occurrences (all)	0	1	
Hot flush			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Hypertension			
subjects affected / exposed	0 / 46 (0.00%)	3 / 36 (8.33%)	
occurrences (all)	0	3	
Hypotension			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Peripheral coldness			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
General disorders and administration site conditions			
Asthenia			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Chest discomfort			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Chest pain			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Chills			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Fatigue			
subjects affected / exposed	3 / 46 (6.52%)	0 / 36 (0.00%)	
occurrences (all)	3	0	

Hunger			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Injection site bruising			
subjects affected / exposed	1 / 46 (2.17%)	1 / 36 (2.78%)	
occurrences (all)	1	1	
Injection site dermatitis			
subjects affected / exposed	0 / 46 (0.00%)	1 / 36 (2.78%)	
occurrences (all)	0	1	
Injection site erythema			
subjects affected / exposed	3 / 46 (6.52%)	4 / 36 (11.11%)	
occurrences (all)	5	6	
Injection site pain			
subjects affected / exposed	1 / 46 (2.17%)	0 / 36 (0.00%)	
occurrences (all)	1	0	
Injection site rash			
subjects affected / exposed	1 / 46 (2.17%)	0 / 36 (0.00%)	
occurrences (all)	1	0	
Injection site reaction			
subjects affected / exposed	6 / 46 (13.04%)	8 / 36 (22.22%)	
occurrences (all)	9	27	
Injection site recall reaction			
subjects affected / exposed	5 / 46 (10.87%)	3 / 36 (8.33%)	
occurrences (all)	5	4	
Injection site vesicles			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Malaise			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Non-cardiac chest pain			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Oedema peripheral			
subjects affected / exposed	1 / 46 (2.17%)	0 / 36 (0.00%)	
occurrences (all)	1	0	

Pain			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Peripheral swelling			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Pyrexia			
subjects affected / exposed	0 / 46 (0.00%)	1 / 36 (2.78%)	
occurrences (all)	0	1	
Therapeutic response unexpected			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Thirst			
subjects affected / exposed	0 / 46 (0.00%)	1 / 36 (2.78%)	
occurrences (all)	0	1	
Vessel puncture site bruise			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Immune system disorders			
Hypersensitivity			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Seasonal allergy			
subjects affected / exposed	1 / 46 (2.17%)	0 / 36 (0.00%)	
occurrences (all)	1	0	
Reproductive system and breast disorders			
Abnormal uterine bleeding			
subjects affected / exposed	1 / 46 (2.17%)	0 / 36 (0.00%)	
occurrences (all)	1	0	
Atrophic vulvovaginitis			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Erectile dysfunction			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Genital atrophy			

subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Prostatitis			
subjects affected / exposed	1 / 46 (2.17%)	0 / 36 (0.00%)	
occurrences (all)	1	0	
Vulvovaginal dryness			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Respiratory, thoracic and mediastinal disorders			
Cough			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Dyspnoea			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Epistaxis			
subjects affected / exposed	0 / 46 (0.00%)	1 / 36 (2.78%)	
occurrences (all)	0	1	
Haemoptysis			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Nasal congestion			
subjects affected / exposed	1 / 46 (2.17%)	0 / 36 (0.00%)	
occurrences (all)	1	0	
Productive cough			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Pulmonary mass			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Rhinitis allergic			
subjects affected / exposed	0 / 46 (0.00%)	1 / 36 (2.78%)	
occurrences (all)	0	1	
Sinus pain			

subjects affected / exposed	1 / 46 (2.17%)	0 / 36 (0.00%)	
occurrences (all)	1	0	
Sputum increased			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Psychiatric disorders			
Adjustment disorder			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Anxiety			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Depression			
subjects affected / exposed	0 / 46 (0.00%)	1 / 36 (2.78%)	
occurrences (all)	0	1	
Insomnia			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Major depression			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Investigations			
Activated partial thromboplastin time prolonged			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Alanine aminotransferase abnormal			
subjects affected / exposed	1 / 46 (2.17%)	0 / 36 (0.00%)	
occurrences (all)	1	0	
Alanine aminotransferase increased			
subjects affected / exposed	4 / 46 (8.70%)	9 / 36 (25.00%)	
occurrences (all)	5	10	
Albumin urine present			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Aspartate aminotransferase increased			

subjects affected / exposed	2 / 46 (4.35%)	4 / 36 (11.11%)		
occurrences (all)	2	4		
Blood calcium increased				
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)		
occurrences (all)	0	0		
Blood glucose increased				
subjects affected / exposed	0 / 46 (0.00%)	1 / 36 (2.78%)		
occurrences (all)	0	1		
Blood potassium abnormal				
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)		
occurrences (all)	0	0		
Blood pressure decreased				
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)		
occurrences (all)	0	0		
Blood pressure increased				
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)		
occurrences (all)	0	0		
Body temperature increased				
subjects affected / exposed	0 / 46 (0.00%)	1 / 36 (2.78%)		
occurrences (all)	0	1		
Creatinine renal clearance abnormal				
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)		
occurrences (all)	0	0		
Electrocardiogram QRS complex prolonged				
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)		
occurrences (all)	0	0		
Electrocardiogram QT prolonged				
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)		
occurrences (all)	0	0		
Electrocardiogram abnormal				
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)		
occurrences (all)	0	0		
Haematocrit increased				
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)		
occurrences (all)	0	0		

Liver function test abnormal			
subjects affected / exposed	1 / 46 (2.17%)	0 / 36 (0.00%)	
occurrences (all)	1	0	
Hepatic enzyme increased			
subjects affected / exposed	2 / 46 (4.35%)	3 / 36 (8.33%)	
occurrences (all)	2	3	
Liver function test increased			
subjects affected / exposed	1 / 46 (2.17%)	0 / 36 (0.00%)	
occurrences (all)	1	0	
Magnetic resonance imaging spinal abnormal			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Mean cell haemoglobin concentration decreased			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Mean cell volume increased			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Urine albumin/creatinine ratio abnormal			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Urine albumin/creatinine ratio increased			
subjects affected / exposed	0 / 46 (0.00%)	1 / 36 (2.78%)	
occurrences (all)	0	1	
Urine analysis abnormal			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Weight decreased			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Transaminases increased			
subjects affected / exposed	1 / 46 (2.17%)	6 / 36 (16.67%)	
occurrences (all)	1	6	
Injury, poisoning and procedural complications			

Animal bite				
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)		
occurrences (all)	0	0		
Arthropod bite				
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)		
occurrences (all)	0	0		
Back injury				
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)		
occurrences (all)	0	0		
Epicondylitis				
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)		
occurrences (all)	0	0		
Fall				
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)		
occurrences (all)	0	0		
Foot fracture				
subjects affected / exposed	1 / 46 (2.17%)	0 / 36 (0.00%)		
occurrences (all)	1	0		
Head injury				
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)		
occurrences (all)	0	0		
Injection related reaction				
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)		
occurrences (all)	0	0		
Ligament sprain				
subjects affected / exposed	1 / 46 (2.17%)	0 / 36 (0.00%)		
occurrences (all)	1	0		
Muscle rupture				
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)		
occurrences (all)	0	0		
Post procedural discomfort				
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)		
occurrences (all)	0	0		
Procedural pain				
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)		
occurrences (all)	0	0		

Skin abrasion			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Vaccination complication			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Cardiac disorders			
Atrial fibrillation			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Brugada syndrome			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Myocardial injury			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Palpitations			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Sinus tachycardia			
subjects affected / exposed	0 / 46 (0.00%)	1 / 36 (2.78%)	
occurrences (all)	0	1	
Nervous system disorders			
Balance disorder			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Carpal tunnel syndrome			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Dizziness			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Headache			
subjects affected / exposed	1 / 46 (2.17%)	0 / 36 (0.00%)	
occurrences (all)	4	0	
Hypoaesthesia			

subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Migraine			
subjects affected / exposed	1 / 46 (2.17%)	0 / 36 (0.00%)	
occurrences (all)	1	0	
Neuropathy peripheral			
subjects affected / exposed	1 / 46 (2.17%)	0 / 36 (0.00%)	
occurrences (all)	1	0	
Paraesthesia			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Radiculopathy			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Sciatica			
subjects affected / exposed	1 / 46 (2.17%)	0 / 36 (0.00%)	
occurrences (all)	1	0	
Sensory disturbance			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Transient ischaemic attack			
subjects affected / exposed	0 / 46 (0.00%)	1 / 36 (2.78%)	
occurrences (all)	0	1	
Blood and lymphatic system disorders			
Anaemia			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Eosinophilia			
subjects affected / exposed	0 / 46 (0.00%)	1 / 36 (2.78%)	
occurrences (all)	0	1	
Ear and labyrinth disorders			
Ear pain			
subjects affected / exposed	1 / 46 (2.17%)	0 / 36 (0.00%)	
occurrences (all)	1	0	
Vertigo			

subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Eye disorders			
Cataract			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Conjunctivitis allergic			
subjects affected / exposed	0 / 46 (0.00%)	1 / 36 (2.78%)	
occurrences (all)	0	1	
Ulcerative keratitis			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Vitreous floaters			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Gastrointestinal disorders			
Abdominal discomfort			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Abdominal distension			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Abdominal pain			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Abdominal pain upper			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Acid peptic disease			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Constipation			
subjects affected / exposed	1 / 46 (2.17%)	1 / 36 (2.78%)	
occurrences (all)	1	1	
Crohn's disease			

subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Dental necrosis			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Diarrhoea			
subjects affected / exposed	2 / 46 (4.35%)	2 / 36 (5.56%)	
occurrences (all)	2	3	
Diverticulum			
subjects affected / exposed	1 / 46 (2.17%)	0 / 36 (0.00%)	
occurrences (all)	1	0	
Dyspepsia			
subjects affected / exposed	0 / 46 (0.00%)	1 / 36 (2.78%)	
occurrences (all)	0	1	
Faeces soft			
subjects affected / exposed	1 / 46 (2.17%)	0 / 36 (0.00%)	
occurrences (all)	1	0	
Gastrooesophageal reflux disease			
subjects affected / exposed	0 / 46 (0.00%)	2 / 36 (5.56%)	
occurrences (all)	0	3	
Haematochezia			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Nausea			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Oesophageal pain			
subjects affected / exposed	1 / 46 (2.17%)	0 / 36 (0.00%)	
occurrences (all)	1	0	
Rectal haemorrhage			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Toothache			
subjects affected / exposed	2 / 46 (4.35%)	0 / 36 (0.00%)	
occurrences (all)	2	0	
Vomiting			

subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Hepatobiliary disorders			
Cholelithiasis			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Drug-induced liver injury			
subjects affected / exposed	3 / 46 (6.52%)	0 / 36 (0.00%)	
occurrences (all)	3	0	
Hepatic steatosis			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Nonalcoholic fatty liver disease			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Skin and subcutaneous tissue disorders			
Actinic keratosis			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Alopecia			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Dermatitis			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Dermatitis allergic			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Dermatitis contact			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Erythema			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Hyperhidrosis			

subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Miliaria			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Petechiae			
subjects affected / exposed	0 / 46 (0.00%)	1 / 36 (2.78%)	
occurrences (all)	0	1	
Pruritus			
subjects affected / exposed	1 / 46 (2.17%)	1 / 36 (2.78%)	
occurrences (all)	1	1	
Rash			
subjects affected / exposed	1 / 46 (2.17%)	0 / 36 (0.00%)	
occurrences (all)	1	0	
Rosacea			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Skin discolouration			
subjects affected / exposed	1 / 46 (2.17%)	0 / 36 (0.00%)	
occurrences (all)	1	0	
Skin lesion			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Skin mass			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Renal and urinary disorders			
Acute kidney injury			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Calculus bladder			
subjects affected / exposed	0 / 46 (0.00%)	1 / 36 (2.78%)	
occurrences (all)	0	1	
Chronic kidney disease			
subjects affected / exposed	0 / 46 (0.00%)	1 / 36 (2.78%)	
occurrences (all)	0	1	

Dysuria			
subjects affected / exposed	0 / 46 (0.00%)	1 / 36 (2.78%)	
occurrences (all)	0	1	
Haematuria			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Microalbuminuria			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Nephrolithiasis			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Pollakiuria			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Proteinuria			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Renal cyst			
subjects affected / exposed	0 / 46 (0.00%)	1 / 36 (2.78%)	
occurrences (all)	0	1	
Renal failure			
subjects affected / exposed	0 / 46 (0.00%)	1 / 36 (2.78%)	
occurrences (all)	0	1	
Urge incontinence			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Urinary incontinence			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Musculoskeletal and connective tissue disorders			
Arthralgia			
subjects affected / exposed	3 / 46 (6.52%)	1 / 36 (2.78%)	
occurrences (all)	3	4	
Arthritis			

subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Bone cyst			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Bursitis			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Flank pain			
subjects affected / exposed	0 / 46 (0.00%)	2 / 36 (5.56%)	
occurrences (all)	0	2	
Intervertebral disc degeneration			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Intervertebral disc disorder			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Joint stiffness			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Joint swelling			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Muscle fatigue			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Muscle spasms			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Muscular weakness			
subjects affected / exposed	0 / 46 (0.00%)	3 / 36 (8.33%)	
occurrences (all)	0	3	
Musculoskeletal chest pain			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Musculoskeletal pain			

subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)		
occurrences (all)	0	0		
Myalgia				
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)		
occurrences (all)	0	0		
Neck pain				
subjects affected / exposed	1 / 46 (2.17%)	0 / 36 (0.00%)		
occurrences (all)	1	0		
Osteoarthritis				
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)		
occurrences (all)	0	0		
Pain in extremity				
subjects affected / exposed	2 / 46 (4.35%)	0 / 36 (0.00%)		
occurrences (all)	2	0		
Plantar fascial fibromatosis				
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)		
occurrences (all)	0	0		
Rotator cuff syndrome				
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)		
occurrences (all)	0	0		
Scoliosis				
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)		
occurrences (all)	0	0		
Spinal pain				
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)		
occurrences (all)	0	0		
Temporomandibular joint syndrome				
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)		
occurrences (all)	0	0		
Tendonitis				
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)		
occurrences (all)	0	0		
Tenosynovitis				
subjects affected / exposed	1 / 46 (2.17%)	0 / 36 (0.00%)		
occurrences (all)	1	0		
Back pain				

subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Infections and infestations			
Acute sinusitis			
subjects affected / exposed	1 / 46 (2.17%)	0 / 36 (0.00%)	
occurrences (all)	1	0	
Bronchitis			
subjects affected / exposed	0 / 46 (0.00%)	1 / 36 (2.78%)	
occurrences (all)	0	1	
COVID-19			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Cellulitis			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Conjunctivitis			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Cystitis			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Fungal infection			
subjects affected / exposed	1 / 46 (2.17%)	0 / 36 (0.00%)	
occurrences (all)	1	0	
Fungal skin infection			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Gastroenteritis			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Gastroenteritis viral			
subjects affected / exposed	1 / 46 (2.17%)	0 / 36 (0.00%)	
occurrences (all)	1	0	
Gastrointestinal bacterial infection			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	

Helicobacter infection				
subjects affected / exposed	0 / 46 (0.00%)	1 / 36 (2.78%)		
occurrences (all)	0	1		
Herpes zoster				
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)		
occurrences (all)	0	0		
Infection parasitic				
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)		
occurrences (all)	0	0		
Injection site cellulitis				
subjects affected / exposed	1 / 46 (2.17%)	0 / 36 (0.00%)		
occurrences (all)	1	0		
Kidney infection				
subjects affected / exposed	1 / 46 (2.17%)	0 / 36 (0.00%)		
occurrences (all)	1	0		
Nasopharyngitis				
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)		
occurrences (all)	0	0		
Oral candidiasis				
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)		
occurrences (all)	0	0		
Otitis externa				
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)		
occurrences (all)	0	0		
Pharyngitis				
subjects affected / exposed	1 / 46 (2.17%)	0 / 36 (0.00%)		
occurrences (all)	1	0		
Sinusitis				
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)		
occurrences (all)	0	0		
Tinea cruris				
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)		
occurrences (all)	0	0		
Tooth abscess				
subjects affected / exposed	0 / 46 (0.00%)	1 / 36 (2.78%)		
occurrences (all)	0	1		

Tooth infection			
subjects affected / exposed	2 / 46 (4.35%)	0 / 36 (0.00%)	
occurrences (all)	2	0	
Upper respiratory tract infection			
subjects affected / exposed	0 / 46 (0.00%)	1 / 36 (2.78%)	
occurrences (all)	0	1	
Urinary tract infection			
subjects affected / exposed	1 / 46 (2.17%)	3 / 36 (8.33%)	
occurrences (all)	1	1	
Viral infection			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Vulvovaginal candidiasis			
subjects affected / exposed	0 / 46 (0.00%)	1 / 36 (2.78%)	
occurrences (all)	0	1	
Vulvovaginal mycotic infection			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Metabolism and nutrition disorders			
Decreased appetite			
subjects affected / exposed	1 / 46 (2.17%)	0 / 36 (0.00%)	
occurrences (all)	1	0	
Dehydration			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Diabetes mellitus			
subjects affected / exposed	0 / 46 (0.00%)	1 / 36 (2.78%)	
occurrences (all)	0	1	
Diabetes mellitus inadequate control			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Glucose tolerance impaired			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Gout			

subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Hyperglycaemia			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Hyperkalaemia			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Hypoglycaemia			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Hypokalaemia			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Increased appetite			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	
Type 2 diabetes mellitus			
subjects affected / exposed	0 / 46 (0.00%)	0 / 36 (0.00%)	
occurrences (all)	0	0	

More information

Substantial protocol amendments (globally)

Were there any global substantial amendments to the protocol? Yes

Date	Amendment
18 September 2020	New appendix was added to describe alternative measures that were allowed during a public emergency, such as COVID-19.

Notes:

Interruptions (globally)

Were there any global interruptions to the trial? No

Limitations and caveats

None reported