Clinical trial results:
A Phase 3, Open Label Trial Evaluating the Safety, Immunogenicity and Impact of 13-valent Pneumococcal Conjugate Vaccine (13vPnC) in Alaskan Native Children.

Summary

EudraCT number	2008-003648-12
Trial protocol	Outside EU/EEA
Global end of trial date	10 September 2010

Results information

Result version number	v1 (current)
This version publication date	29 June 2016
First version publication date	02 August 2015

Trial information

Trial identification

Sponsor protocol code	6096A1-3010 (B1851009)

Additional study identifiers

ISRCTN number	-
ClinicalTrials.gov id (NCT number)	NCT00743652
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)	Yes
EMA paediatric investigation plan number(s)	EMEA-000036-PIP01-07
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?	Yes

Notes:

Results analysis stage

Analysis stage	Final
Date of interim/final analysis	17 October 2011
Is this the analysis of the primary completion data?	No

Global end of trial reached?	Yes
Global end of trial date	10 September 2010
Was the trial ended prematurely?	No

Notes:

General information about the trial

Main objective of the trial:

To assess the impact of 13vPnC on the incidence of invasive pneumococcal disease (IPD) in the Yukon Kuskokwim (YK) Delta region due to the 13 vaccine Streptococcus (S.) pneumoniae serotypes.

Protection of trial subjects:

This study was conducted in accordance with the International Conference on Harmonisation (ICH) Guideline for Good Clinical Practice (GCP) and the ethical principles that have their origins in the Declaration of Helsinki. Written informed consent was obtained from all parent(s)/legal guardian(s) of every subject before enrollment in the study and before performance of any study-related procedures.

Background therapy: -

Evidence for comparator: -

Actual start date of recruitment	30 January 2009
Long term follow-up planned	Yes
Long term follow-up rationale	Safety
Long term follow-up duration	6 Months
Independent data monitoring committee (IDMC) involvement?	Yes

Notes:

Population of trial subjects

Subjects enrolled per country

Country: Number of subjects enrolled	United States: 373
Worldwide total number of subjects	373
EEA total number of subjects	0

Notes:

Subjects enrolled per age group

In utero	0
Preterm newborn - gestational age < 37 wk	0
Newborns (0-27 days)	0
Infants and toddlers (28 days-23 months)	294
Children (2-11 years)	79
Adolescents (12-17 years)	0
Adults (18-64 years)	0
From 65 to 84 years	0

85 years and over	0

Subject disposition

Recruitment

Recruitment details:

Blood sample was optional for subjects living in the Bethel area. Active vaccination stopped when 13vPnC was commercially available in Alaska. Study ended after a 6-month follow-up for safety following the last vaccination.

Pre-assignment

Screening details:

Total 373 subjects were enrolled in 24 centres of United States. Study started on 30 Jan 2009 and completed on 10 Sep 2010.

Period 1

Period 1 title	Overall Study (overall period)
Is this the baseline period?	Yes
Allocation method	Not applicable
Blinding used	Not blinded

Arms

Are arms mutually exclusive?	Yes
Arm title	13vPnC Group 1 (3 Doses Infant Series and 1 Toddler Dose)

Arm description:

Subjects 6 weeks to less than (<) 10 months of age with 0 prior doses of Prevnar received 3 doses of 13-valent pneumococcal conjugate vaccine (13vPnC) at least 28 days apart (infant series) and a single dose of 13vPnC at greater than (>) 12 months of age (toddler dose), at least 60 days after last infant dose.

Arm type	Experimental
Investigational medicinal product name	13-valent pneumococcal conjugate vaccine
Investigational medicinal product code	
Other name	
Pharmaceutical forms	Injection
Routes of administration	Intramuscular use

Dosage and administration details:

Subjects 6 weeks to < 10 months of age with 0 prior doses of Prevnar received 3 single intramuscular (IM) 0.5 milliliter (mL) doses of 13vPnC at least 28 days apart (infant series), and a single IM 0.5 mL dose of 13vPnC at > 12 months of age (toddler dose), at least 60 days after last infant dose.

Arm title	13vPnC Group 2 (2 Doses Infant Series and 1 Toddler Dose)

Arm description:

Subjects <12 months of age with 1 prior dose of Prevnar received 2 doses of 13vPnC at least 28 days apart (infant series), and a single dose of 13vPnC at >12 months of age (toddler dose), at least 60 days after last infant dose.

Arm type	Experimental
Investigational medicinal product name	13-valent pneumococcal conjugate vaccine
Investigational medicinal product code	
Other name	
Pharmaceutical forms	Injection
Routes of administration	Intramuscular use

Dosage and administration details:

Subjects <12 months of age with 1 prior dose of Prevnar received 2 single IM 0.5 mL doses of 13vPnC at least 28 days apart (infant series), and a single IM 0.5 mL dose of 13vPnC at >12 months of age (toddler dose), at least 60 days after last infant dose.

Arm title	13vPnC Group 3 (1 Dose Infant Series and 1 Toddler Dose)

Arm description:

Subjects <12 months of age with 2 prior doses of Prevnar received a single dose of 13vPnC (infant

series) and a single dose of 13vPnC at >12 months of age (toddler dose), at least 60 days after last infant dose.

Arm type	Experimental
Investigational medicinal product name	13-valent pneumococcal conjugate vaccine
Investigational medicinal product code	
Other name	
Pharmaceutical forms	Injection
Routes of administration	Intramuscular use

Dosage and administration details:

Subjects <12 months of age with 2 prior doses of Prevnar received a single IM 0.5 mL dose of 13vPnC (infant series) and a single IM 0.5 mL dose of 13vPnC at >12 months of age (toddler dose), at least 60 days after last infant dose.

Arm title	13vPnC Group 4 (2 Catch-Up Doses)

Arm description:

Subjects greater than or equal to (≥) 12 months to <2 years of age received 2 single doses of 13vPnC at least 60 days apart.

Arm type	Experimental
Investigational medicinal product name	13-valent pneumococcal conjugate vaccine
Investigational medicinal product code	
Other name	
Pharmaceutical forms	Injection
Routes of administration	Intramuscular use

Dosage and administration details:

Subjects ≥ 12 months to <2 years of age received 2 single IM 0.5 mL doses of 13vPnC at least 60 days apart.

Arm title	13vPnC Group 5 (1 Catch-Up Dose)

Arm description:

Subjects ≥2 years to <5 years of age received a single dose of 13vPnC.

Arm type	Experimental
Investigational medicinal product name	13-valent pneumococcal conjugate vaccine
Investigational medicinal product code	
Other name	
Pharmaceutical forms	Injection
Routes of administration	Intramuscular use

Dosage and administration details:

Subjects ≥2 years to <5 years of age received a single IM 0.5 mL dose of 13vPnC.

Number of subjects in period 1	13vPnC Group 1 (3 Doses Infant Series and 1 Toddler Dose)	13vPnC Group 2 (2 Doses Infant Series and 1 Toddler Dose)	13vPnC Group 3 (1 Dose Infant Series and 1 Toddler Dose)
Started	151	51	25
Vaccinated Dose 1	151	51	25
Vaccinated Dose 2	112 [1]	41 [2]	0 [3]
Vaccinated Dose 3	74 [4]	0 [5]	0 [6]
Toddler Dose	15 [7]	23 [8]	16 [9]
Completed	141	48	23
Not completed	10	3	2
'Parent/Legal Guardian Request '	3	2	2

'Adverse Event '	3	-	-
Death	2	-	-
Unspecified	1	1	-
Lost to follow-up	1	-	-
'Investigator Request '	-	-	-

Number of subjects in period 1	13vPnC Group 4 (2 Catch-Up Doses)	13vPnC Group 5 (1 Catch-Up Dose)
Started	67	79
Vaccinated Dose 1	66	79
Vaccinated Dose 2	49 [10]	0 [11]
Vaccinated Dose 3	0 [12]	0 [13]
Toddler Dose	0 [14]	0 [15]
Completed	63	79
Not completed	4	0
'Parent/Legal Guardian Request '	2	-
'Adverse Event '	-	-
Death	-	-
Unspecified	-	-
Lost to follow-up	-	-
'Investigator Request '	2	-

Notes:

[1] - The number of subjects at this milestone seems inconsistent with the number of subjects in the arm. It is expected that the number of subjects will be greater than, or equal to the number that completed, minus those who left.
Justification: Subjects who completed the study were not administered with all vaccinations.

[2] - The number of subjects at this milestone seems inconsistent with the number of subjects in the arm. It is expected that the number of subjects will be greater than, or equal to the number that completed, minus those who left.
Justification: Subjects who completed the study were not administered with all vaccinations.

[3] - The number of subjects at this milestone seems inconsistent with the number of subjects in the arm. It is expected that the number of subjects will be greater than, or equal to the number that completed, minus those who left.
Justification: Subjects who completed the study were not administered with all vaccinations.

[4] - The number of subjects at this milestone seems inconsistent with the number of subjects in the arm. It is expected that the number of subjects will be greater than, or equal to the number that completed, minus those who left.
Justification: Subjects who completed the study were not administered with all vaccinations.

[5] - The number of subjects at this milestone seems inconsistent with the number of subjects in the arm. It is expected that the number of subjects will be greater than, or equal to the number that completed, minus those who left.
Justification: Subjects who completed the study were not administered with all vaccinations.

[6] - The number of subjects at this milestone seems inconsistent with the number of subjects in the arm. It is expected that the number of subjects will be greater than, or equal to the number that completed, minus those who left.
Justification: Subjects of 13vPnC Group 3 were not administered with Vaccination Dose 2 and Vaccination Dose 3.

[7] - The number of subjects at this milestone seems inconsistent with the number of subjects in the arm. It is expected that the number of subjects will be greater than, or equal to the number that completed, minus those who left.
Justification: Subjects who completed the study were not administered with all vaccinations.

[8] - The number of subjects at this milestone seems inconsistent with the number of subjects in the arm. It is expected that the number of subjects will be greater than, or equal to the number that completed, minus those who left.
Justification: Subjects who completed the study were not administered with all vaccinations.

[9] - The number of subjects at this milestone seems inconsistent with the number of subjects in the arm. It is expected that the number of subjects will be greater than, or equal to the number that completed, minus those who left.
Justification: Subjects who completed the study were not administered with all vaccinations.

[10] - The number of subjects at this milestone seems inconsistent with the number of subjects in the arm. It is expected that the number of subjects will be greater than, or equal to the number that completed, minus those who left.
Justification: Subjects of 13vPnC Group 4 were not administered with Vaccination Dose 3 and Toddler dose.
[11] - The number of subjects at this milestone seems inconsistent with the number of subjects in the arm. It is expected that the number of subjects will be greater than, or equal to the number that completed, minus those who left.
Justification: Subjects of 13vPnC Group 5 were not administered with Vaccination Dose 2, Vaccination Dose 3 and Toddler dose.
[12] - The number of subjects at this milestone seems inconsistent with the number of subjects in the arm. It is expected that the number of subjects will be greater than, or equal to the number that completed, minus those who left.
Justification: Subjects of 13vPnC Group 4 were not administered with Vaccination Dose 3 and Toddler dose.
[13] - The number of subjects at this milestone seems inconsistent with the number of subjects in the arm. It is expected that the number of subjects will be greater than, or equal to the number that completed, minus those who left.
Justification: Subjects of 13vPnC Group 5 were not administered with Vaccination Dose 2, Vaccination Dose 3 and Toddler dose.
[14] - The number of subjects at this milestone seems inconsistent with the number of subjects in the arm. It is expected that the number of subjects will be greater than, or equal to the number that completed, minus those who left.
Justification: Subjects of 13vPnC Group 4 were not administered with Vaccination Dose 3 and Toddler dose.
[15] - The number of subjects at this milestone seems inconsistent with the number of subjects in the arm. It is expected that the number of subjects will be greater than, or equal to the number that completed, minus those who left.
Justification: Subjects of 13vPnC Group 5 were not administered with Vaccination Dose 2, Vaccination Dose 3 and Toddler dose.

Baseline characteristics

Reporting groups	
Reporting group title	13vPnC Group 1 (3 Doses Infant Series and 1 Toddler Dose)
Reporting group description:	
Subjects 6 weeks to less than (<) 10 months of age with 0 prior doses of Prevnar received 3 doses of 13-valent pneumococcal conjugate vaccine (13vPnC) at least 28 days apart (infant series) and a single dose of 13vPnC at greater than (>) 12 months of age (toddler dose), at least 60 days after last infant dose.	
Reporting group title	13vPnC Group 2 (2 Doses Infant Series and 1 Toddler Dose)
Reporting group description:	
Subjects <12 months of age with 1 prior dose of Prevnar received 2 doses of 13vPnC at least 28 days apart (infant series), and a single dose of 13vPnC at >12 months of age (toddler dose), at least 60 days after last infant dose.	
Reporting group title	13vPnC Group 3 (1 Dose Infant Series and 1 Toddler Dose)
Reporting group description:	
Subjects <12 months of age with 2 prior doses of Prevnar received a single dose of 13vPnC (infant series) and a single dose of 13vPnC at >12 months of age (toddler dose), at least 60 days after last infant dose.	
Reporting group title	13vPnC Group 4 (2 Catch-Up Doses)
Reporting group description:	
Subjects greater than or equal to (≥) 12 months to <2 years of age received 2 single doses of 13vPnC at least 60 days apart.	
Reporting group title	13vPnC Group 5 (1 Catch-Up Dose)
Reporting group description:	
Subjects ≥2 years to <5 years of age received a single dose of 13vPnC.	

Reporting group values	13vPnC Group 1 (3 Doses Infant Series and 1 Toddler Dose)	13vPnC Group 2 (2 Doses Infant Series and 1 Toddler Dose)	13vPnC Group 3 (1 Dose Infant Series and 1 Toddler Dose)
Number of subjects	151	51	25
Age categorical			
Units: Subjects			
Age continuous			
Units: months			
arithmetic mean	1.4	4.4	7.1
standard deviation	± 0.6	± 0.8	± 1.6
Gender categorical			
Units: Subjects			
Female	73	23	11
Male	78	28	14

Reporting group values	13vPnC Group 4 (2 Catch-Up Doses)	13vPnC Group 5 (1 Catch-Up Dose)	Total
Number of subjects	67	79	373
Age categorical			
Units: Subjects			
Age continuous			
Units: months			
arithmetic mean	15.2	40.1	

standard deviation	± 3.4	± 11.1	-

Gender categorical Units: Subjects			
Female	36	47	190
Male	31	32	183

End points

End points reporting groups

Reporting group title	13vPnC Group 1 (3 Doses Infant Series and 1 Toddler Dose)
Reporting group description:	
Subjects 6 weeks to less than (<) 10 months of age with 0 prior doses of Prevnar received 3 doses of 13-valent pneumococcal conjugate vaccine (13vPnC) at least 28 days apart (infant series) and a single dose of 13vPnC at greater than (>) 12 months of age (toddler dose), at least 60 days after last infant dose.	
Reporting group title	13vPnC Group 2 (2 Doses Infant Series and 1 Toddler Dose)
Reporting group description:	
Subjects <12 months of age with 1 prior dose of Prevnar received 2 doses of 13vPnC at least 28 days apart (infant series), and a single dose of 13vPnC at >12 months of age (toddler dose), at least 60 days after last infant dose.	
Reporting group title	13vPnC Group 3 (1 Dose Infant Series and 1 Toddler Dose)
Reporting group description:	
Subjects <12 months of age with 2 prior doses of Prevnar received a single dose of 13vPnC (infant series) and a single dose of 13vPnC at >12 months of age (toddler dose), at least 60 days after last infant dose.	
Reporting group title	13vPnC Group 4 (2 Catch-Up Doses)
Reporting group description:	
Subjects greater than or equal to (≥) 12 months to <2 years of age received 2 single doses of 13vPnC at least 60 days apart.	
Reporting group title	13vPnC Group 5 (1 Catch-Up Dose)
Reporting group description:	
Subjects ≥2 years to <5 years of age received a single dose of 13vPnC.	
Subject analysis set title	13vPnC (All Subjects)
Subject analysis set type	Safety analysis
Subject analysis set description:	
Safety population.	

Primary: Percentage of Subjects Achieving Serotype-Specific Pneumococcal Immunoglobulin G (IgG) Antibody Level ≥0.35 Micrograms Per Milliliter (Mcg/mL) 1 Month After the Infant Series

End point title	Percentage of Subjects Achieving Serotype-Specific Pneumococcal Immunoglobulin G (IgG) Antibody Level ≥0.35 Micrograms Per Milliliter (Mcg/mL) 1 Month After the Infant Series[1][2]
End point description:	
Percentage of subjects in 13vPnC Groups 1, 2 and 3 achieving predefined antibody threshold ≥0.35 mcg/mL along with the corresponding 95% confidence interval (CI) for the 7 common pneumococcal serotypes (serotypes 4, 6B, 9V, 14, 18C, 19F, and 23F) and 6 additional pneumococcal serotypes specific to 13vPnC (serotypes 1, 3, 5, 6A, 7F, and 19A) are presented. Exact 2-sided CI based upon the observed proportion of subjects. Evaluable immunogenicity population: received treatments as randomized at all expected doses, blood drawn within specified timeframes, at least 1 valid and determinate assay result for proposed analysis, and no major protocol violations.	
End point type	Primary
End point timeframe:	
28 to 56 days after vaccination 3 for Group 1, after vaccination 2 for Group 2, and after vaccination 1 for Group 3.	

Notes:

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

[2] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: The endpoint was planned to be assessed for subjects who received infant dose and toddler

dose.

End point values	13vPnC Group 1 (3 Doses Infant Series and 1 Toddler Dose)	13vPnC Group 2 (2 Doses Infant Series and 1 Toddler Dose)	13vPnC Group 3 (1 Dose Infant Series and 1 Toddler Dose)	
Subject group type	Reporting group	Reporting group	Reporting group	
Number of subjects analysed	11[3]	6[4]	3[5]	
Units: Percentage of Subjects				
number (confidence interval 95%)				
Common serotypes - serotype 4	100 (71.5 to 100)	100 (54.1 to 100)	100 (29.2 to 100)	
Common serotypes - serotype 6B	100 (71.5 to 100)	83.3 (35.9 to 99.6)	100 (15.8 to 100)	
Common serotypes - serotype 9V	100 (71.5 to 100)	83.3 (35.9 to 99.6)	100 (29.2 to 100)	
Common serotypes - serotype 14	90.9 (58.7 to 99.8)	100 (54.1 to 100)	100 (15.8 to 100)	
Common serotype - serotype 18C	100 (71.5 to 100)	100 (54.1 to 100)	100 (29.2 to 100)	
Common serotype - serotype 19F	100 (71.5 to 100)	100 (54.1 to 100)	100 (29.2 to 100)	
Common serotype - serotype 23F	90.9 (58.7 to 99.8)	83.3 (35.9 to 99.6)	100 (29.2 to 100)	
Additional serotype - serotype 1	100 (71.5 to 100)	100 (54.1 to 100)	100 (29.2 to 100)	
Additional serotype - serotype 3	90.9 (58.7 to 99.8)	100 (54.1 to 100)	100 (29.2 to 100)	
Additional serotype - serotype 5	100 (71.5 to 100)	100 (54.1 to 100)	100 (29.2 to 100)	
Additional serotype - serotype 6A	90.9 (58.7 to 99.8)	100 (54.1 to 100)	100 (29.2 to 100)	
Additional serotype - serotype 7F	100 (71.5 to 100)	100 (54.1 to 100)	100 (29.2 to 100)	
Additional serotype - serotype 19A	100 (71.5 to 100)	100 (54.1 to 100)	100 (29.2 to 100)	

Notes:

[3] - Number of subjects with a determinate IgG antibody concentration to the given serotype.

[4] - Number of subjects with a determinate IgG antibody concentration to the given serotype.

[5] - Number of subjects with a determinate IgG antibody concentration to the given serotype.

Statistical analyses

No statistical analyses for this end point

Primary: Percentage of Subjects Achieving Serotype-Specific Pneumococcal IgG Antibody Level ≥0.35 Mcg/mL 1 Month After the Toddler Dose

End point title	Percentage of Subjects Achieving Serotype-Specific Pneumococcal IgG Antibody Level ≥0.35 Mcg/mL 1 Month After the Toddler Dose[6][7]

End point description:

Percentage of subjects in 13vPnC Groups 1, 2, and 3 achieving predefined antibody threshold ≥0.35 mcg/mL along with the corresponding 95% CI for the 7 common pneumococcal serotypes (serotypes 4, 6B, 9V, 14, 18C, 19F, and 23F) and 6 additional pneumococcal serotypes specific to 13vPnC (serotypes 1, 3, 5, 6A, 7F, and 19A) are presented. Exact 2-sided CI based upon the observed proportion of subjects. Evaluable immunogenicity population.

End point type	Primary

End point timeframe:

28 to 56 days after vaccination 4 for Group 1, after vaccination 3 for Group 2, and after vaccination 2 for Group 3.

Notes:

[6] - No statistical analyses have been specified for this primary end point. It is expected there is at least one statistical analysis for each primary end point.
Justification: Only descriptive data was planned to be reported for this endpoint.

[7] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: The endpoint was planned to be assessed for subjects who received infant dose and toddler dose.

End point values	13vPnC Group 1 (3 Doses Infant Series and 1 Toddler Dose)	13vPnC Group 2 (2 Doses Infant Series and 1 Toddler Dose)	13vPnC Group 3 (1 Dose Infant Series and 1 Toddler Dose)	
Subject group type	Reporting group	Reporting group	Reporting group	
Number of subjects analysed	2[8]	6[9]	2[10]	
Units: Percentage of Subjects				
number (confidence interval 95%)				
Common serotypes - serotype 4	100 (15.8 to 100)	100 (54.1 to 100)	100 (15.8 to 100)	
Common serotypes - serotype 6B	100 (15.8 to 100)	100 (54.1 to 100)	100 (15.8 to 100)	
Common serotypes - serotype 9V	100 (15.8 to 100)	100 (54.1 to 100)	100 (15.8 to 100)	
Common serotypes - serotype 14	100 (15.8 to 100)	100 (54.1 to 100)	100 (15.8 to 100)	
Common serotype - serotype 18C	100 (15.8 to 100)	100 (54.1 to 100)	100 (15.8 to 100)	
Common serotype - serotype 19F	100 (15.8 to 100)	100 (54.1 to 100)	100 (15.8 to 100)	
Common serotype - serotype 23F	100 (15.8 to 100)	100 (54.1 to 100)	100 (15.8 to 100)	
Additional serotype - serotype 1	100 (15.8 to 100)	100 (54.1 to 100)	100 (15.8 to 100)	
Additional serotype - serotype 3	100 (15.8 to 100)	100 (54.1 to 100)	100 (15.8 to 100)	
Additional serotype - serotype 5	100 (15.8 to 100)	100 (54.1 to 100)	100 (15.8 to 100)	
Additional serotype - serotype 6A	100 (15.8 to 100)	100 (54.1 to 100)	100 (15.8 to 100)	
Additional serotype - serotype 7F	100 (15.8 to 100)	100 (54.1 to 100)	100 (15.8 to 100)	
Additional serotype - serotype 19A	100 (15.8 to 100)	100 (54.1 to 100)	100 (15.8 to 100)	

Notes:

[8] - N=number of subjects with a determinate IgG antibody concentration to the given serotype.

[9] - N=number of subjects with a determinate IgG antibody concentration to the given serotype.

[10] - N=number of subjects with a determinate IgG antibody concentration to the given serotype.

Statistical analyses

No statistical analyses for this end point

Primary: Percentage of Subjects Achieving Serotype-specific Pneumococcal IgG Antibody Level ≥0.35 Mcg/mL 1 Month After the Relevant Catch-Up Dose

End point title	Percentage of Subjects Achieving Serotype-specific Pneumococcal IgG Antibody Level ≥0.35 Mcg/mL 1 Month After the Relevant Catch-Up Dose[11][12]

End point description:

Percentage of subjects in 13vPnC Groups 4 and 5 achieving predefined antibody threshold ≥0.35 mcg/mL along with the corresponding 95% CI for the 7 common pneumococcal serotypes (serotypes 4, 6B, 9V, 14, 18C, 19F, and 23F) and 6 additional pneumococcal serotypes specific to 13vPnC (serotypes 1, 3, 5, 6A, 7F, and 19A) are presented. Exact 2-sided CI based upon the observed proportion of subjects. Evaluable immunogenicity population.

End point type	Primary

End point timeframe:

28 to 56 days after vaccination 2 for Group 4, and after the single vaccination in Group 5.

Notes:

[11] - No statistical analyses have been specified for this primary end point. It is expected there is at least one statistical analysis for each primary end point.
Justification: Only descriptive data was planned to be reported for this endpoint.

[12] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: The endpoint was planned to be assessed for subjects who received catch-up dose.

End point values	13vPnC Group 4 (2 Catch-Up Doses)	13vPnC Group 5 (1 Catch-Up Dose)		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	4[13]	9[14]		
Units: Percentage of Subjects				
number (confidence interval 95%)				
Common serotypes - serotype 4	100 (39.8 to 100)	100 (66.4 to 100)		
Common serotypes - serotype 6B	100 (39.8 to 100)	100 (66.4 to 100)		
Common serotypes - serotype 9V	100 (39.8 to 100)	100 (66.4 to 100)		
Common serotypes - serotype 14	100 (39.8 to 100)	100 (66.4 to 100)		
Common serotype - serotype 18C	100 (39.8 to 100)	100 (66.4 to 100)		
Common serotype - serotype 19F	100 (39.8 to 100)	100 (66.4 to 100)		
Common serotype - serotype 23F	100 (39.8 to 100)	100 (66.4 to 100)		
Additional serotype - serotype 1	100 (39.8 to 100)	100 (66.4 to 100)		
Additional serotype - serotype 3	100 (39.8 to 100)	100 (66.4 to 100)		
Additional serotype - serotype 5	100 (39.8 to 100)	100 (66.4 to 100)		
Additional serotype - serotype 6A	100 (39.8 to 100)	100 (66.4 to 100)		
Additional serotype - serotype 7F	100 (39.8 to 100)	100 (66.4 to 100)		
Additional serotype - serotype 19A	100 (39.8 to 100)	100 (66.4 to 100)		

Notes:

[13] - N=number of subjects with a determinate IgG antibody concentration to the given serotype.

[14] - N=number of subjects with a determinate IgG antibody concentration to the given serotype.

Statistical analyses

No statistical analyses for this end point

Primary: Number of Cases of Invasive Pneumococcal Disease (IPD) in Subjects Less Than 5 Years of Age Due to Any Serotype Contained in 13vPnC

End point title	Number of Cases of Invasive Pneumococcal Disease (IPD) in Subjects Less Than 5 Years of Age Due to Any Serotype Contained in 13vPnC[15]

End point description:

In order to assess the impact of 13vPnC on the incidence of IPD in the YK Delta region, the Centers for Disease Control and Prevention (CDC) Arctic Investigation Program (AIP) accessed IPD data through evaluation of ongoing statewide IPD surveillance in Alaska. The CDC's AIP followed IPD (including serotype and vaccination history) to show whether identified cases of IPD received Prevnar, 13vPnC, or both. These data were combined with statewide data and used to identify the overall trend in IPD in the YK Delta region after introduction of 13vPnC. Safety population.

End point type	Primary

End point timeframe:

Baseline to 6 months after last vaccination

Notes:

[15] - No statistical analyses have been specified for this primary end point. It is expected there is at least one statistical analysis for each primary end point.
Justification: Only descriptive data was planned to be reported for this end point.

End point values	13vPnC (All Subjects)			
Subject group type	Subject analysis set			
Number of subjects analysed	373			
Units: Subjects				
number (not applicable)	0			

Statistical analyses

No statistical analyses for this end point

Secondary: Percentage of Subjects Achieving Serum IgG Antibody Level ≥0.35 Mcg/mL Prior to Vaccination With 13vPnC (Groups 4 and 5 Only)

End point title	Percentage of Subjects Achieving Serum IgG Antibody Level ≥0.35 Mcg/mL Prior to Vaccination With 13vPnC (Groups 4 and 5 Only)[16]

End point description:

Percentage of subjects in 13vPnC Groups 4 and 5 achieving predefined antibody threshold ≥0.35 mcg/mL along with the corresponding 95% CI for the 7 common pneumococcal serotypes (serotypes 4, 6B, 9V, 14, 18C, 19F, and 23F) and 6 additional pneumococcal serotypes specific to 13vPnC (serotypes 1, 3, 5, 6A, 7F, and 19A) are presented. Exact 2-sided CI based upon the observed proportion of subjects. Evaluable immunogenicity population.

End point type	Secondary

End point timeframe:

28 to 56 days before vaccination 2 for Group 4, and before the single vaccination in Group 5.

Notes:

[16] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: The endpoint was planned to be assessed for subjects who received catch-up dose.

End point values	13vPnC Group 4 (2 Catch-Up Doses)	13vPnC Group 5 (1 Catch-Up Dose)		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	4[17]	9[18]		
Units: Percentage of Subjects				
number (confidence interval 95%)				
Common serotypes - serotype 4	50 (6.8 to 93.2)	50 (15.7 to 84.3)		
Common serotypes - serotype 6B	100 (39.8 to 100)	88.9 (51.8 to 99.7)		
Common serotypes - serotype 9V	100 (39.8 to 100)	66.7 (29.9 to 92.5)		
Common serotypes - serotype 14	100 (29.2 to 100)	88.9 (51.8 to 99.7)		
Common serotype - serotype 18C	75 (19.4 to 99.4)	33.3 (7.5 to 70.1)		
Common serotype - serotype 19F	100 (39.8 to 100)	88.9 (51.8 to 99.7)		
Common serotype - serotype 23F	100 (39.8 to 100)	88.9 (51.8 to 99.7)		
Additional serotype - serotype 1	0 (0 to 60.2)	12.5 (0.3 to 52.7)		
Additional serotype - serotype 3	0 (0 to 60.2)	37.5 (8.5 to 75.5)		
Additional serotype - serotype 5	50 (6.8 to 93.2)	100 (66.4 to 100)		
Additional serotype - serotype 6A	100 (39.8 to 100)	88.9 (51.8 to 99.7)		
Additional serotype - serotype 7F	0 (0 to 60.2)	33.3 (7.5 to 70.1)		
Additional serotype - serotype 19A	100 (39.8 to 100)	88.9 (51.8 to 99.7)		

Notes:

[17] - N=number of subjects with a determinate IgG antibody concentration to the given serotype.

[18] - N=number of subjects with a determinate IgG antibody concentration to the given serotype.

Statistical analyses

No statistical analyses for this end point

Secondary: Percentage of Subjects Achieving Serotype-specific Pneumococcal IgG Antibody Level ≥1.0 Mcg/mL 1 Month After the Infant Series

End point title	Percentage of Subjects Achieving Serotype-specific Pneumococcal IgG Antibody Level ≥1.0 Mcg/mL 1 Month After the Infant Series[19]

End point description:

Percentage of subjects in 13vPnC Groups 1, 2, and 3 achieving predefined antibody threshold ≥1.0 mcg/mL along with the corresponding 95% CI for the 7 common pneumococcal serotypes (serotypes 4, 6B, 9V, 14, 18C, 19F, and 23F) and 6 additional pneumococcal serotypes specific to 13vPnC (serotypes 1, 3, 5, 6A, 7F, and 19A) are presented. Exact 2-sided CI based upon the observed proportion of subjects. Evaluable immunogenicity population.

End point type	Secondary

End point timeframe:

28 to 56 days after vaccination 3 for Group 1, after vaccination 2 for Group 2, and after vaccination 1 for Group 3.

Notes:

[19] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.

Justification: The endpoint was planned to be assessed for subjects who received infant dose and toddler dose.

End point values	13vPnC Group 1 (3 Doses Infant Series and 1 Toddler Dose)	13vPnC Group 2 (2 Doses Infant Series and 1 Toddler Dose)	13vPnC Group 3 (1 Dose Infant Series and 1 Toddler Dose)	
Subject group type	Reporting group	Reporting group	Reporting group	
Number of subjects analysed	11[20]	6[21]	3[22]	
Units: Percentage of Subjects				
number (confidence interval 95%)				
Common serotypes - serotype 4	90.9 (58.7 to 99.8)	100 (54.1 to 100)	100 (29.2 to 100)	
Common serotypes - serotype 6B	72.7 (39 to 94)	83.3 (35.9 to 99.6)	100 (15.8 to 100)	
Common serotypes - serotype 9V	81.8 (48.2 to 97.7)	66.7 (22.3 to 95.7)	66.7 (9.4 to 99.2)	
Common serotypes - serotype 14	90.9 (58.7 to 99.8)	83.3 (35.9 to 99.6)	100 (15.8 to 100)	
Common serotype - serotype 18C	63.6 (30.8 to 89.1)	83.3 (35.9 to 99.6)	100 (29.2 to 100)	
Common serotype - serotype 19F	100 (71.5 to 100)	66.7 (22.3 to 95.7)	100 (29.2 to 100)	
Common serotype - serotype 23F	63.6 (30.8 to 89.1)	50 (11.8 to 88.2)	100 (29.2 to 100)	
Additional serotype - serotype 1	81.8 (48.2 to 97.7)	83.3 (35.9 to 99.6)	100 (29.2 to 100)	
Additional serotype - serotype 3	54.5 (23.4 to 83.3)	66.7 (22.3 to 95.7)	100 (29.2 to 100)	
Additional serotype - serotype 5	81.8 (48.2 to 97.7)	83.3 (35.9 to 99.6)	66.7 (9.4 to 99.2)	
Additional serotype - serotype 6A	72.7 (39 to 94)	83.3 (35.9 to 99.6)	66.7 (9.4 to 99.2)	
Additional serotype - serotype 7F	100 (71.5 to 100)	83.3 (35.9 to 99.6)	100 (29.2 to 100)	
Additional serotype - serotype 19A	72.7 (39 to 100)	50 (11.8 to 88.2)	33.3 (0.8 to 90.6)	

Notes:

[20] - N=number of subjects with a determinate IgG antibody concentration to the given serotype.

[21] - N=number of subjects with a determinate IgG antibody concentration to the given serotype.

[22] - N=number of subjects with a determinate IgG antibody concentration to the given serotype.

Statistical analyses

No statistical analyses for this end point

Secondary: Percentage of Subjects Achieving Serotype-specific Pneumococcal IgG Antibody Level ≥1.0 Mcg/mL 1 Month After the Toddler Dose

End point title	Percentage of Subjects Achieving Serotype-specific Pneumococcal IgG Antibody Level ≥1.0 Mcg/mL 1 Month After the Toddler Dose[23]

End point description:

Percentage of subjects in 13vPnC Groups 1, 2, and 3 achieving predefined antibody threshold ≥1.0 mcg/mL along with the corresponding 95% CI for the 7 common pneumococcal serotypes (serotypes 4, 6B, 9V, 14, 18C, 19F, and 23F) and 6 additional pneumococcal serotypes specific to 13vPnC (serotypes

1, 3, 5, 6A, 7F, and 19A) are presented. Exact 2-sided CI based upon the observed proportion of subjects. Evaluable immunogenicity population.

End point type	Secondary

End point timeframe:

28 to 56 days after vaccination 4 for Group 1, after vaccination 3 for Group 2, and after vaccination 2 for Group 3

Notes:

[23] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.

Justification: The endpoint was planned to be assessed for subjects who received infant dose and toddler dose.

End point values	13vPnC Group 1 (3 Doses Infant Series and 1 Toddler Dose)	13vPnC Group 2 (2 Doses Infant Series and 1 Toddler Dose)	13vPnC Group 3 (1 Dose Infant Series and 1 Toddler Dose)	
Subject group type	Reporting group	Reporting group	Reporting group	
Number of subjects analysed	2[24]	6[25]	2[26]	
Units: Percentage of Subjects				
number (confidence interval 95%)				
Common serotypes - serotype 4	100 (15.8 to 100)	100 (54.1 to 100)	50 (1.3 to 98.7)	
Common serotypes - serotype 6B	100 (15.8 to 100)	100 (54.1 to 100)	100 (15.8 to 100)	
Common serotypes - serotype 9V	100 (15.8 to 100)	100 (54.1 to 100)	50 (1.3 to 98.7)	
Common serotypes - serotype 14	100 (15.8 to 100)	100 (54.1 to 100)	100 (15.8 to 100)	
Common serotype - serotype 18C	100 (15.8 to 100)	83.3 (35.9 to 99.6)	100 (15.8 to 100)	
Common serotype - serotype 19F	100 (15.8 to 100)	83.3 (35.9 to 99.6)	100 (15.8 to 100)	
Common serotype - serotype 23F	100 (15.8 to 100)	83.3 (35.9 to 99.6)	100 (15.8 to 100)	
Additional serotype - serotype 1	100 (15.8 to 100)	83.3 (35.9 to 99.6)	100 (15.8 to 100)	
Additional serotype - serotype 3	50 (1.3 to 98.7)	66.7 (22.3 to 95.7)	100 (15.8 to 100)	
Additional serotype - serotype 5	100 (15.8 to 100)	100 (54.1 to 100)	100 (15.8 to 100)	
Additional serotype - serotype 6A	100 (15.8 to 100)	100 (54.1 to 100)	100 (15.8 to 100)	
Additional serotype - serotype 7F	100 (15.8 to 100)	100 (54.1 to 100)	100 (15.8 to 100)	
Additional serotype - serotype 19A	100 (15.8 to 100)	100 (54.1 to 100)	100 (15.8 to 100)	

Notes:

[24] - N=number of subjects with a determinate IgG antibody concentration to the given serotype.

[25] - N=number of subjects with a determinate IgG antibody concentration to the given serotype.

[26] - N=number of subjects with a determinate IgG antibody concentration to the given serotype.

Statistical analyses

No statistical analyses for this end point

Secondary: Percentage of Subjects Achieving Serotype-specific Pneumococcal IgG Antibody Level ≥1.0 Mcg/mL 1 Month After the Relevant Catch-up Dose

End point title	Percentage of Subjects Achieving Serotype-specific Pneumococcal IgG Antibody Level ≥1.0 Mcg/mL 1 Month After the Relevant Catch-up Dose[27]

End point description:

Percentage of subjects in 13vPnC Groups 4 and 5 achieving predefined antibody threshold ≥1.0 mcg/mL along with the corresponding 95% CI for the 7 common pneumococcal serotypes (serotypes 4, 6B, 9V, 14, 18C, 19F, and 23F) and 6 additional pneumococcal serotypes specific to 13vPnC (serotypes 1, 3, 5, 6A, 7F, and 19A) are presented. Exact 2-sided CI based upon the observed proportion of subjects. Evaluable immunogenicity population.

End point type	Secondary

End point timeframe:

28 to 56 days after vaccination 2 for Group 4, and after the single vaccination in Group 5

Notes:

[27] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: The endpoint was planned to be assessed for subjects who received catch-up dose.

End point values	13vPnC Group 4 (2 Catch-Up Doses)	13vPnC Group 5 (1 Catch-Up Dose)		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	4[28]	9[29]		
Units: Percentage of Subjects				
number (confidence interval 95%)				
Common serotypes - serotype 4	100 (39.8 to 100)	100 (66.4 to 100)		
Common serotypes - serotype 6B	100 (39.8 to 100)	100 (66.4 to 100)		
Common serotypes - serotype 9V	100 (39.8 to 100)	100 (66.4 to 100)		
Common serotypes - serotype 14	100 (39.8 to 100)	100 (66.4 to 100)		
Common serotype - serotype 18C	100 (39.8 to 100)	100 (66.4 to 100)		
Common serotype - serotype 19F	100 (39.8 to 100)	100 (66.4 to 100)		
Common serotype - serotype 23F	100 (39.8 to 100)	100 (66.4 to 100)		
Additional serotype - serotype 1	100 (39.8 to 100)	66.7 (29.9 to 92.5)		
Additional serotype - serotype 3	100 (39.8 to 100)	77.8 (40 to 97.2)		
Additional serotype - serotype 5	100 (39.8 to 100)	88.9 (51.8 to 99.7)		
Additional serotype - serotype 6A	100 (39.8 to 100)	88.9 (51.8 to 99.7)		
Additional serotype - serotype 7F	100 (39.8 to 100)	100 (66.4 to 100)		
Additional serotype - serotype 19A	100 (39.8 to 100)	100 (66.4 to 100)		

Notes:

[28] - N=number of subjects with a determinate IgG antibody concentration to the given serotype.

[29] - N=number of subjects with a determinate IgG antibody concentration to the given serotype.

Statistical analyses

No statistical analyses for this end point

Secondary: Percentage of Subjects Reporting Pre-Specified Local Reactions: Catch-up Dose 1

End point title	Percentage of Subjects Reporting Pre-Specified Local Reactions: Catch-up Dose 1[30]

End point description:

Local reactions were reported by the parent/legal guardian using a diary card. Tenderness was scaled as Any (tenderness present); Significant (present and interfered with limb movement). Redness and swelling were scaled as Any (redness or swelling present); Mild (0.5 centimeters [cm] to 2.0 cm); Moderate (2.5 to 7.0 cm); Severe (> 7.0 cm). Subjects may have been represented in more than 1 category. Safety Population: all subjects who received at least 1 dose of 13vPnC; n=number of subjects with specific characteristics.

End point type	Secondary

End point timeframe:

Day 1 through Day 7 after vaccination 1 for Group 4 and after the single vaccination in Group 5.

Notes:

[30] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: The endpoint was planned to be assessed for subjects who received catch-up dose.

End point values	13vPnC Group 4 (2 Catch-Up Doses)	13vPnC Group 5 (1 Catch-Up Dose)		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	65[31]	77[32]		
Units: Percentage of Subjects				
number (not applicable)				
Tenderness: Any (n=65,77)	32.3	48.1		
Tenderness: Significant (n=63,77)	3.2	9.1		
Swelling: Any (n=64,77)	7.8	7.8		
Swelling: Mild (n=62,76)	3.2	2.6		
Swelling: Moderate (n=63,77)	3.2	6.5		
Swelling: Severe (n=63,77)	1.6	2.6		
Redness: Any (n=65,77)	15.4	13		
Redness: Mild (n=63,77)	11.1	11.7		
Redness: Moderate (n=64,77)	4.7	5.2		
Redness: Severe (n=63,77)	1.6	2.6		

Notes:

[31] - N=number of subjects reporting yes for at least 1 day or no for all days.

[32] - N=number of subjects reporting yes for at least 1 day or no for all days.

Statistical analyses

No statistical analyses for this end point

Secondary: Percentage of Subjects Reporting Pre-Specified Local Reactions: Catch-up Dose 2

End point title	Percentage of Subjects Reporting Pre-Specified Local Reactions: Catch-up Dose 2[33]

End point description:

Local reactions were reported by the parent/legal guardian using a diary card. Tenderness was scaled as Any (tenderness present); Significant (present and interfered with limb movement). Redness and swelling were scaled as Any (redness or swelling present); Mild (0.5 centimeters [cm] to 2.0 cm); Moderate (2.5 to 7.0 cm); Severe (> 7.0 cm). Subjects may have been represented in more than 1 category. Safety Population; n=number of subjects with specific characteristics.

End point type	Secondary

End point timeframe:

Day 1 through Day 7 after vaccination 2 for Group 4

Notes:

[33] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: The endpoint was planned to be assessed for subjects who received catch-up dose.

End point values	13vPnC Group 4 (2 Catch-Up Doses)			
Subject group type	Reporting group			
Number of subjects analysed	49[34]			
Units: Percentage of Subjects				
number (not applicable)				
Tenderness: Any (n=49)	36.7			
Tenderness: Significant (n=49)	10.2			
Swelling: Any (n=48)	10.4			
Swelling: Mild (n=47)	8.5			
Swelling: Moderate (n=47)	0			
Swelling: Severe (n=47)	0			
Redness: Any (n=49)	6.1			
Redness: Mild (n=48)	4.2			
Redness: Moderate (n=48)	0			
Redness: Severe (n=48)	0			

Notes:

[34] - N=number of subjects reporting yes for at least 1 day or no for all days.

Statistical analyses

No statistical analyses for this end point

Secondary: Percentage of Subjects Reporting Pre-Specified Systemic Events: Catch-up Dose 1

End point title	Percentage of Subjects Reporting Pre-Specified Systemic Events: Catch-up Dose 1[35]

End point description:

Systemic events (any fever 38 degrees Celsius [C] or higher, decreased appetite, irritability, increased sleep, decreased sleep, hives [urticaria], and use of antipyretic medication) were reported using a diary card. Subjects may have been represented in more than 1 category. Subjects may have been represented in more than 1 category. Safety Population, n=number of subjects with specific characteristics.

End point type	Secondary

End point timeframe:

Day 1 through Day 7 after vaccination 1 for Group 4 and after the single vaccination in Group 5

Notes:

[35] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: The endpoint was planned to be assessed for subjects who received catch-up dose.

End point values	13vPnC Group 4 (2 Catch-Up Doses)	13vPnC Group 5 (1 Catch-Up Dose)		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	66[36]	77[37]		
Units: Percentage of Subjects				
number (not applicable)				
Fever ≥38 degrees C but ≤39 degrees C (n=35,47)	42.9	27.7		
Fever >39 degrees C but ≤40 degrees C (n=29,43)	20.7	11.6		
Fever >40 degrees C (n=28,40)	7.1	0		
Decreased appetite (n=66,77)	31.8	24.7		
Irritability (n=66,77)	53	41.6		
Increased sleep (n=66,77)	33.3	22.1		
Decreased sleep (n=66,77)	24.2	11.7		
Hives (urticaria) (n=66,77)	3	2.6		

Notes:

[36] - N=number of subjects reporting yes for at least 1 day or no for all days.

[37] - N=number of subjects reporting yes for at least 1 day or no for all days.

Statistical analyses

No statistical analyses for this end point

Secondary: Percentage of Subjects Reporting Pre-Specified Systemic Events: Catch-up Dose 2

End point title	Percentage of Subjects Reporting Pre-Specified Systemic Events: Catch-up Dose 2[38]

End point description:

Systemic events (any fever 38 degrees C or higher, decreased appetite, irritability, increased sleep, decreased sleep, hives [urticaria], and use of antipyretic medication) were reported using a diary card. Subjects may have been represented in more than 1 category. Safety Population; n=number of subjects with specific characteristics.

End point type	Secondary

End point timeframe:

Day 1 through Day 7 after vaccination 2 for Group 4

Notes:

[38] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: The endpoint was planned to be assessed for subjects who received catch-up dose.

End point values	13vPnC Group 4 (2 Catch-Up Doses)			
Subject group type	Reporting group			
Number of subjects analysed	49[39]			
Units: Percentage of Subjects				
number (not applicable)				
Fever ≥38 degrees C but ≤39 degrees C (n=26)	23.1			
Fever >39 degrees C but ≤40 degrees C (n=22)	9.1			
Fever >40 degrees C (n=21)	0			
Decreased appetite (n=49)	22.4			

Irritability (n=49)	57.1			
Increased sleep (n=49)	26.5			
Decreased sleep (n=49)	4.1			
Hives (urticaria) (n=49)	2			

Notes:

[39] - N=number of subjects reporting yes for at least 1 day or no for all days.

Statistical analyses

No statistical analyses for this end point

Other pre-specified: Percentage of Subjects Achieving Serotype-Specific Pneumococcal IgG Antibody Level ≥0.15 Mcg/mL 1 Month After the Infant Series

End point title	Percentage of Subjects Achieving Serotype-Specific Pneumococcal IgG Antibody Level ≥0.15 Mcg/mL 1 Month After the Infant Series[40]

End point description:

Percentage of subjects in 13vPnC Groups 1, 2, and 3 achieving predefined antibody threshold ≥0.15 mcg/mL along with the corresponding 95% CI for the 7 common pneumococcal serotypes (serotypes 4, 6B, 9V, 14, 18C, 19F, and 23F) and 6 additional pneumococcal serotypes specific to 13vPnC (serotypes 1, 3, 5, 6A, 7F, and 19A) are presented. Exact 2-sided CI based upon the observed proportion of subjects. Evaluable immunogenicity population.

End point type	Other pre-specified

End point timeframe:

28 to 56 days after vaccination 3 for Group 1, after vaccination 2 for Group 2, and after vaccination 1 for Group 3

Notes:

[40] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: The endpoint was planned to be assessed for subjects who received infant dose and toddler dose.

End point values	13vPnC Group 1 (3 Doses Infant Series and 1 Toddler Dose)	13vPnC Group 2 (2 Doses Infant Series and 1 Toddler Dose)	13vPnC Group 3 (1 Dose Infant Series and 1 Toddler Dose)	
Subject group type	Reporting group	Reporting group	Reporting group	
Number of subjects analysed	11[41]	6[42]	3[43]	
Units: Percentage of Subjects				
number (confidence interval 95%)				
Common serotypes - serotype 4	100 (71.5 to 100)	100 (54.1 to 100)	100 (29.2 to 100)	
Common serotypes - serotype 6B	100 (71.5 to 100)	100 (54.1 to 100)	100 (15.8 to 100)	
Common serotypes - serotype 9V	100 (71.5 to 100)	100 (54.1 to 100)	100 (29.2 to 100)	
Common serotypes - serotype 14	90.9 (58.7 to 99.8)	100 (54.1 to 100)	100 (15.8 to 100)	
Common serotype - serotype 18C	100 (71.5 to 100)	100 (54.1 to 100)	100 (29.2 to 100)	
Common serotype - serotype 19F	100 (71.5 to 100)	100 (54.1 to 100)	100 (29.2 to 100)	
Common serotype - serotype 23F	100 (71.5 to 100)	100 (54.1 to 100)	100 (29.2 to 100)	
Additional serotype - serotype 1	100 (71.5 to 100)	100 (54.1 to 100)	100 (29.2 to 100)	

Additional serotype - serotype 3	100 (71.5 to 100)	100 (54.1 to 100)	100 (29.2 to 100)	
Additional serotype - serotype 5	100 (71.5 to 100)	100 (54.1 to 100)	100 (29.2 to 100)	
Additional serotype - serotype 6A	100 (71.5 to 100)	100 (54.1 to 100)	100 (29.2 to 100)	
Additional serotype - serotype 7F	100 (71.5 to 100)	100 (54.1 to 100)	100 (29.2 to 100)	
Additional serotype - serotype 19A	100 (71.5 to 100)	100 (54.1 to 100)	100 (29.2 to 100)	

Notes:

[41] - N=number of subjects with a determinate IgG antibody concentration to the given serotype.

[42] - N=number of subjects with a determinate IgG antibody concentration to the given serotype.

[43] - N=number of subjects with a determinate IgG antibody concentration to the given serotype.

Statistical analyses

No statistical analyses for this end point

Other pre-specified: Percentage of Subjects Achieving Serotype-specific Pneumococcal IgG Antibody Level ≥0.15 Mcg/mL 1 Month After the Toddler Dose

End point title	Percentage of Subjects Achieving Serotype-specific Pneumococcal IgG Antibody Level ≥0.15 Mcg/mL 1 Month After the Toddler Dose[44]

End point description:

Percentage of subjects in 13vPnC Groups 1, 2, and 3 achieving predefined antibody threshold ≥0.15 mcg/mL along with the corresponding 95% CI for the 7 common pneumococcal serotypes (serotypes 4, 6B, 9V, 14, 18C, 19F, and 23F) and 6 additional pneumococcal serotypes specific to 13vPnC (serotypes 1, 3, 5, 6A, 7F, and 19A) are presented. Exact 2-sided CI based upon the observed proportion of subjects. Evaluable immunogenicity population.

End point type	Other pre-specified

End point timeframe:

28 to 56 days after vaccination 4 for Group 1, after vaccination 3 for Group 2, and after vaccination 2 for Group 3.

Notes:

[44] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: The endpoint was planned to be assessed for subjects who received infant dose and toddler dose.

End point values	13vPnC Group 1 (3 Doses Infant Series and 1 Toddler Dose)	13vPnC Group 2 (2 Doses Infant Series and 1 Toddler Dose)	13vPnC Group 3 (1 Dose Infant Series and 1 Toddler Dose)	
Subject group type	Reporting group	Reporting group	Reporting group	
Number of subjects analysed	2[45]	6[46]	2[47]	
Units: Percentage of Subjects				
number (confidence interval 95%)				
Common serotypes - serotype 4	100 (15.8 to 100)	100 (54.1 to 100)	100 (15.8 to 100)	
Common serotypes - serotype 6B	100 (15.8 to 100)	100 (54.1 to 100)	100 (15.8 to 100)	
Common serotypes - serotype 9V	100 (15.8 to 100)	100 (54.1 to 100)	100 (15.8 to 100)	
Common serotypes - serotype 14	100 (15.8 to 100)	100 (54.1 to 100)	100 (15.8 to 100)	
Common serotype - serotype 18C	100 (15.8 to 100)	100 (54.1 to 100)	100 (15.8 to 100)	

Common serotype - serotype 19F	100 (15.8 to 100)	100 (54.1 to 100)	100 (15.8 to 100)	
Common serotype - serotype 23F	100 (15.8 to 100)	100 (54.1 to 100)	100 (15.8 to 100)	
Additional serotype - serotype 1	100 (15.8 to 100)	100 (54.1 to 100)	100 (15.8 to 100)	
Additional serotype - serotype 3	100 (15.8 to 100)	100 (54.1 to 100)	100 (15.8 to 100)	
Additional serotype - serotype 5	100 (15.8 to 100)	100 (54.1 to 100)	100 (15.8 to 100)	
Additional serotype - serotype 6A	100 (15.8 to 100)	100 (54.1 to 100)	100 (15.8 to 100)	
Additional serotype - serotype 7F	100 (15.8 to 100)	100 (54.1 to 100)	100 (15.8 to 100)	
Additional serotype - serotype 19A	100 (15.8 to 100)	100 (54.1 to 100)	100 (15.8 to 100)	

Notes:

[45] - N=number of subjects with a determinate IgG antibody concentration to the given serotype.

[46] - N=number of subjects with a determinate IgG antibody concentration to the given serotype.

[47] - N=number of subjects with a determinate IgG antibody concentration to the given serotype.

Statistical analyses

No statistical analyses for this end point

Other pre-specified: Percentage of Subjects Achieving Serotype-specific Pneumococcal IgG Antibody Level ≥0.15 Mcg/mL 1 Month After the Relevant Catch-up Dose

End point title	Percentage of Subjects Achieving Serotype-specific Pneumococcal IgG Antibody Level ≥0.15 Mcg/mL 1 Month After the Relevant Catch-up Dose[48]

End point description:

Percentage of subjects in 13vPnC Groups 4 and 5 achieving predefined antibody threshold ≥0.15 mcg/mL along with the corresponding 95% CI for the 7 common pneumococcal serotypes (serotypes 4, 6B, 9V, 14, 18C, 19F, and 23F) and 6 additional pneumococcal serotypes specific to 13vPnC (serotypes 1, 3, 5, 6A, 7F, and 19A) are presented. Exact 2-sided CI based upon the observed proportion of subjects. Evaluable immunogenicity population.

End point type	Other pre-specified

End point timeframe:

28 to 56 days after vaccination 2 for Group 4, and after the single vaccination in Group 5

Notes:

[48] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: The endpoint was planned to be assessed for subjects who received catch-up dose.

End point values	13vPnC Group 4 (2 Catch-Up Doses)	13vPnC Group 5 (1 Catch-Up Dose)		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	4[49]	9[50]		
Units: Percentage of Subjects				
number (confidence interval 95%)				
Common serotypes - serotype 4	100 (39.8 to 100)	100 (66.4 to 100)		
Common serotypes - serotype 6B	100 (39.8 to 100)	100 (66.4 to 100)		

Common serotypes - serotype 9V	100 (39.8 to 100)	100 (66.4 to 100)		
Common serotypes - serotype 14	100 (39.8 to 100)	100 (66.4 to 100)		
Common serotype - serotype 18C	100 (39.8 to 100)	100 (66.4 to 100)		
Common serotype - serotype 19F	100 (39.8 to 100)	100 (66.4 to 100)		
Common serotype - serotype 23F	100 (39.8 to 100)	100 (66.4 to 100)		
Additional serotype - serotype 1	100 (39.8 to 100)	100 (66.4 to 100)		
Additional serotype - serotype 3	100 (39.8 to 100)	100 (66.4 to 100)		
Additional serotype - serotype 5	100 (39.8 to 100)	100 (66.4 to 100)		
Additional serotype - serotype 6A	100 (39.8 to 100)	100 (66.4 to 100)		
Additional serotype - serotype 7F	100 (39.8 to 100)	100 (66.4 to 100)		
Additional serotype - serotype 19A	100 (39.8 to 100)	100 (66.4 to 100)		

Notes:

[49] - N=number of subjects with a determinate IgG antibody concentration to the given serotype.

[50] - N=number of subjects with a determinate IgG antibody concentration to the given serotype.

Statistical analyses

No statistical analyses for this end point

Other pre-specified: Geometric Mean Concentration (GMC) for Serotype-Specific Pneumococcal IgG Antibodies 1 Month After the Infant Series

End point title	Geometric Mean Concentration (GMC) for Serotype-Specific Pneumococcal IgG Antibodies 1 Month After the Infant Series[51]

End point description:

Antibody GMCs (mcg/mL) for 7 common pneumococcal serotypes (serotypes 4, 6B, 9V, 14, 18C, 19F, and 23F) and 6 additional pneumococcal serotypes specific to 13vPnC (Serotypes 1, 3, 5, 6A, 7F, and 19A) are presented. GMC (13vPnC) with 2-sided 95% CIs were evaluated. CIs are back transformations of confidence levels based on Student t distribution for mean logarithm of concentrations. GMCs were calculated using all subjects with available data for specified blood draw. Evaluable immunogenicity population.

End point type	Other pre-specified

End point timeframe:

28 to 56 days after vaccination 3 for Group 1, after vaccination 2 for Group 2, and after vaccination 1 for Group 3

Notes:

[51] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: The endpoint was planned to be assessed for subjects who received infant dose and toddler dose.

End point values	13vPnC Group 1 (3 Doses Infant Series and 1 Toddler Dose)	13vPnC Group 2 (2 Doses Infant Series and 1 Toddler Dose)	13vPnC Group 3 (1 Dose Infant Series and 1 Toddler Dose)	
Subject group type	Reporting group	Reporting group	Reporting group	
Number of subjects analysed	11[52]	6[53]	3[54]	
Units: Microgram/milliliter (mcg/mL)				
geometric mean (confidence interval 95%)				
Common serotypes - serotype 4	2.64 (1.55 to 4.49)	2.11 (0.93 to 4.8)	4.27 (0.49 to 37.05)	
Common serotypes - serotype 6B	2.74 (1.23 to 6.12)	2.59 (0.55 to 12.26)	28.6 (0.12 to 7043.76)	
Common serotypes - 9V	1.38 (0.97 to 1.98)	1.04 (0.46 to 2.38)	1.6 (0.31 to 8.37)	
Common serotypes - serotype 14	4.78 (1.71 to 13.33)	4.08 (1.68 to 9.88)	10.73 (0.06 to 1894.14)	
Common serotypes - serotype 18C	1.83 (1.04 to 3.21)	1.66 (0.92 to 2.98)	2.99 (0.4 to 22.62)	
Common serotypes - serotype 19F	3.55 (2.1 to 6.01)	1.61 (0.65 to 4.01)	2.37 (1.43 to 3.93)	
Common serotypes - serotype 23F	1.54 (0.79 to 3.01)	1.03 (0.35 to 3.04)	3.37 (1.18 to 9.66)	
Additional serotype - serotype 1	3.33 (1.68 to 6.62)	3.02 (1.06 to 8.6)	2.87 (0.3 to 27.24)	
Additional serotype - serotype 3	0.79 (0.51 to 1.22)	1.61 (0.62 to 4.16)	5.07 (1.53 to 16.79)	
Additional serotype - serotype 5	2.89 (1.57 to 5.31)	1.93 (1.14 to 3.26)	1.75 (0.25 to 12.25)	
Additional serotype - serotype 6A	2.19 (0.93 to 5.16)	2.48 (0.91 to 6.78)	5.41 (0.01 to 2495.5)	
Additional serotype - serotype 7F	3.34 (1.99 to 5.61)	3.18 (1.24 to 8.12)	6.21 (1.84 to 20.92)	
Additional serotype - serotype 19A	1.87 (1.11 to 3.17)	1.35 (0.56 to 3.25)	0.97 (0.37 to 2.53)	

Notes:

[52] - N=number of subjects with a determinate IgG antibody concentration to the given serotype.

[53] - N=number of subjects with a determinate IgG antibody concentration to the given serotype.

[54] - N=number of subjects with a determinate IgG antibody concentration to the given serotype.

Statistical analyses

No statistical analyses for this end point

Other pre-specified: GMC for Serotype-Specific Pneumococcal IgG Antibodies 1 Month After the Toddler Dose

End point title	GMC for Serotype-Specific Pneumococcal IgG Antibodies 1 Month After the Toddler Dose[55]

End point description:

Antibody GMCs (mcg/mL) for 7 common pneumococcal serotypes (serotypes 4, 6B, 9V, 14, 18C, 19F, and 23F) and 6 additional pneumococcal serotypes specific to 13vPnC (Serotypes 1, 3, 5, 6A, 7F, and 19A). GMC (13vPnC) with 2-sided 95% CIs were evaluated. CIs are back transformations of confidence levels based on Student t distribution for mean logarithm of concentrations. GMCs were calculated using all subjects with available data for specified blood draw. Evaluable immunogenicity population. Here "99999" in the 95% CI signifies not available (NA). For reporting group 13vPnC Group 1, in serotype common 6B NA = 3.929*10^11 and in additional serotype 6A NA = 6.518*10^11. For reporting group 13vPnC Group 3, NA signifies that n=2 for this serotype and the value for both subject was 3.96, thus there was no variability so 95% CI could not be determined.

End point type	Other pre-specified

End point timeframe:

28 to 56 days after vaccination 4 for Group 1, after vaccination 3 for Group 2, after vaccination 2 for Group 3

Notes:

[55] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: The endpoint was planned to be assessed for subjects who received infant dose and toddler dose.

End point values	13vPnC Group 1 (3 Doses Infant Series and 1 Toddler Dose)	13vPnC Group 2 (2 Doses Infant Series and 1 Toddler Dose)	13vPnC Group 3 (1 Dose Infant Series and 1 Toddler Dose)	
Subject group type	Reporting group	Reporting group	Reporting group	
Number of subjects analysed	2[56]	6[57]	2[58]	
Units: mcg/mL				
geometric mean (confidence interval 95%)				
Common serotypes - serotype 4	7.16 (0 to 199911.7)	2.55 (1.44 to 4.53)	1.76 (0 to 4004.84)	
Common serotypes - serotype 6B	13.64 (0 to 99999)	12.01 (4.23 to 34.08)	11.32 (0.01 to 11544.32)	
Common serotypes - serotype 9V	2.17 (0.01 to 373.79)	2.2 (1.3 to 3.73)	1.3 (0 to 2387.89)	
Common serotypes - serotype 14	12.18 (0.06 to 2411.14)	11.42 (5.9 to 22.12)	8.54 (0.19 to 377.53)	
Common serotype - serotype 18C	2.35 (0 to 1681.49)	2.22 (1.22 to 4.03)	2.42 (0.08 to 72.55)	
Common serotype - serotype 19F	11 (0 to 11762036)	2.84 (1.03 to 7.79)	3.65 (0 to 130178.5)	
Common serotype - serotype 23F	3.79 (0 to 66245120)	2.35 (0.88 to 6.27)	3.96 (-99999 to 99999)	
Additional serotype - serotype 1	2.15 (0 to 9338.28)	4.19 (1.46 to 12.02)	16.7 (0 to 5685164)	
Additional serotype - serotype 3	0.73 (0 to 276.18)	1.32 (0.63 to 2.78)	1.63 (1.19 to 2.23)	
Additional serotype - serotype 5	4.69 (0 to 36146.55)	3.56 (1.96 to 6.46)	5.85 (0.02 to 1606.11)	
Additional serotype - serotype 6A	17.14 (0 to 99999)	7.79 (3 to 20.21)	3.24 (0.01 to 1555.15)	
Additional serotype - serotype 7F	7.45 (0.47 to 117.4)	5.55 (2.59 to 11.87)	4.5 (0.05 to 440.58)	
Additional serotype - serotype 19A	10.53 (2.63 to 42.07)	3.21 (1.23 to 8.39)	8.86 (0 to 8014211)	

Notes:

[56] - N=number of subjects with a determinate IgG antibody concentration to the given serotype.

[57] - N=number of subjects with a determinate IgG antibody concentration to the given serotype.

[58] - N=number of subjects with a determinate IgG antibody concentration to the given serotype.

Statistical analyses

No statistical analyses for this end point

Other pre-specified: GMC for Serotype-specific Pneumococcal IgG Antibodies 1 Month After the Relevant Catch-up Dose

End point title	GMC for Serotype-specific Pneumococcal IgG Antibodies 1 Month After the Relevant Catch-up Dose[59]

End point description:

Antibody GMCs (mcg/mL) for 7 common pneumococcal serotypes (serotypes 4, 6B, 9V, 14, 18C, 19F, and 23F) and 6 additional pneumococcal serotypes specific to 13vPnC (Serotypes 1, 3, 5, 6A, 7F, and 19A) are presented. GMC (13vPnC) with 2-sided 95% CIs were evaluated. CIs are back transformations of confidence levels based on Student t distribution for mean logarithm of concentrations. GMCs were calculated using all subjects with available data for specified blood draw. Evaluable immunogenicity population.

End point type	Other pre-specified

End point timeframe:

28 to 56 days after vaccination 2 for Group 4, and after the single vaccination in Group 5.

Notes:

[59] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: The endpoint was planned to be assessed for subjects who received catch-up dose.

End point values	13vPnC Group 4 (2 Catch-Up Doses)	13vPnC Group 5 (1 Catch-Up Dose)		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	4[60]	9[61]		
Units: mcg/mL				
geometric mean (confidence interval 95%)				
Common serotypes - serotype 4	4.16 (1.16 to 14.91)	4.92 (1.94 to 12.5)		
Common serotypes - serotype 6B	16.49 (4.31 to 63.06)	22.94 (10.84 to 48.52)		
Common serotypes - serotype 9V	3.22 (1.2 to 8.64)	5.16 (2.34 to 11.37)		
Common serotypes - serotype 14	9.02 (2.18 to 37.29)	16.99 (6.08 to 47.48)		
Common serotype - serotype 18C	4.22 (2.22 to 8.05)	5.08 (2.43 to 10.63)		
Common serotype - serotype 19F	3.39 (0.78 to 14.78)	6.45 (4.23 to 9.83)		
Common serotype - serotype 23F	4.94 (1.82 to 13.41)	5.41 (2.3 to 12.74)		
Additional serotype - serotype 1	9.37 (2.89 to 30.34)	1.25 (0.64 to 2.46)		
Additional serotype - serotype 3	1.65 (1.13 to 2.4)	1.91 (1.06 to 3.46)		
Additional serotype - serotype 5	4.51 (2.77 to 7.34)	2.31 (1.41 to 3.78)		
Additional serotype - serotype 6A	7.42 (1.77 to 31.06)	4.46 (2.09 to 9.48)		
Additional serotype - serotype 7F	8.81 (3.32 to 23.39)	3.14 (1.93 to 5.08)		
Additional serotype - serotype 19A	4.4 (2.96 to 6.54)	5.61 (2.69 to 11.7)		

Notes:

[60] - N=number of subjects with a determinate antibody concentration to the specified serotype.

[61] - N=number of subjects with a determinate antibody concentration to the specified serotype.

Statistical analyses

No statistical analyses for this end point

Other pre-specified: Percentage of Subjects Achieving Opsonophagocytic Assay (OPA) Titers ≥Lower Limit of Quantitation (LLOQ) Measured 1 Month After the Infant Series

End point title	Percentage of Subjects Achieving Opsonophagocytic Assay (OPA) Titers ≥Lower Limit of Quantitation (LLOQ) Measured 1 Month After the Infant Series[62]

End point description:

Percentage of subjects in 13vPnC Groups 1, 2, and 3 achieving OPA with 95% CI for serotypes 4, 6B, 9V, 14, 18C, 19F, 23F, 1, 3, 5, 6A, 7F, and 19A. Exact 2-sided CI based upon the observed proportion of subjects. The LLOQ in titers for each serotype was: Pn001, 18; Pn003, 12; Pn004, 21; Pn005, 29; Pn06A, 37; Pn06B, 43, Pn7F, 210; Pn09V, 345; Pn014, 35; Pn18C, 31; Pn19A, 18; Pn19F, 48; and Pn23F, 13. Limit of detection (LOD) established as lowest titer possible in assay, which was 8. OPA titers below LLOQ set to 0.5*LOD for analysis. Evaluable immunogenicity population.

End point type	Other pre-specified

End point timeframe:

28 to 56 days after vaccination 3 for Group 1, after vaccination 2 for Group 2, and after vaccination 1 for Group 3

Notes:

[62] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: The endpoint was planned to be assessed for subjects who received infant dose and toddler dose.

End point values	13vPnC Group 1 (3 Doses Infant Series and 1 Toddler Dose)	13vPnC Group 2 (2 Doses Infant Series and 1 Toddler Dose)	13vPnC Group 3 (1 Dose Infant Series and 1 Toddler Dose)	
Subject group type	Reporting group	Reporting group	Reporting group	
Number of subjects analysed	11[63]	5[64]	3[65]	
Units: Percentage of Subjects				
number (confidence interval 95%)				
Common serotypes - serotype 4	88.9 (51.8 to 99.7)	75 (19.4 to 99.4)	66.7 (9.4 to 99.2)	
Common serotypes - serotype 6B	100 (66.4 to 100)	100 (39.8 to 100)	100 (15.8 to 100)	
Common serotypes - serotype 9V	30 (6.7 to 65.2)	0 (0 to 70.8)	66.7 (9.4 to 99.2)	
Common serotypes - serotype 14	100 (54.1 to 100)	75 (19.4 to 99.4)	100 (15.8 to 100)	
Common serotype - serotype 18C	100 (59 to 100)	100 (39.8 to 100)	100 (15.8 to 100)	
Common serotype - serotype 19F	88.9 (51.8 to 99.7)	50 (6.8 to 93.2)	0 (0 to 97.5)	
Common serotype - serotype 23F	88.9 (51.8 to 99.7)	66.7 (9.4 to 99.2)	100 (15.8 to 100)	
Additional serotype - serotype 1	60 (26.2 to 87.8)	0 (0 to 60.2)	33.3 (0.8 to 90.6)	
Additional serotype - serotype 3	90.9 (58.7 to 99.8)	100 (47.8 to 100)	100 (29.2 to 100)	
Additional serotype - serotype 5	70 (34.8 to 93.3)	60 (14.7 to 94.7)	33.3 (0.8 to 90.6)	
Additional serotype - serotype 6A	90 (55.5 to 99.7)	100 (47.8 to 100)	100 (29.2 to 100)	
Additional serotype - serotype 7F	100 (69.2 to 100)	100 (47.8 to 100)	100 (15.8 to 100)	
Additional serotype - serotype 19A	80 (44.4 to 97.5)	75 (19.4 to 99.4)	100 (15.8 to 100)	

Notes:

[63] - N=number of subjects with a determinate antibody concentration to the specified serotype.

[64] - N=number of subjects with a determinate antibody concentration to the specified serotype.

[65] - N=number of subjects with a determinate antibody concentration to the specified serotype.

Statistical analyses

No statistical analyses for this end point

Other pre-specified: Percentage of Subjects Achieving OPA Titers ≥LLOQ Measured 1 Month After the Toddler Dose

End point title	Percentage of Subjects Achieving OPA Titers ≥LLOQ Measured 1 Month After the Toddler Dose[66]

End point description:

Percentage of subjects in 13vPnC Groups 1, 2 and 3 achieving OPA with 95% CI for serotypes 4, 6B, 9V, 14, 18C, 19F, 23F, 1, 3, 5, 6A, 7F, and 19A. Exact 2-sided CI based upon the observed proportion of subjects. The LLOQ in titers for each serotype was: Pn001, 18; Pn003, 12; Pn004, 21; Pn005, 29; Pn06A, 37; Pn06B, 43, Pn7F, 210; Pn09V, 345; Pn014, 35; Pn18C, 31; Pn19A, 18; Pn19F, 48; and Pn23F, 13. Limit of detection (LOD) established as lowest titer possible in assay, which was 8. OPA titers below LLOQ set to 0.5*LOD for analysis. Evaluable immunogenicity population.

End point type	Other pre-specified

End point timeframe:

28 to 56 days after vaccination 4 for Group 1, after vaccination 3 for Group 2, and after vaccination 2 for Group 3

Notes:

[66] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: The endpoint was planned to be assessed for subjects who received infant dose and toddler dose.

End point values	13vPnC Group 1 (3 Doses Infant Series and 1 Toddler Dose)	13vPnC Group 2 (2 Doses Infant Series and 1 Toddler Dose)	13vPnC Group 3 (1 Dose Infant Series and 1 Toddler Dose)	
Subject group type	Reporting group	Reporting group	Reporting group	
Number of subjects analysed	2[67]	6[68]	1[69]	
Units: Percentage of Subjects				
number (confidence interval 95%)				
Common serotypes - serotype 4	100 (15.8 to 100)	50 (11.8 to 88.2)	100 (2.5 to 100)	
Common serotypes - serotype 6B	100 (15.8 to 100)	80 (28.4 to 99.5)	100 (2.5 to 100)	
Common serotypes - serotype 9V	100 (15.8 to 100)	83.3 (35.9 to 99.6)	100 (2.5 to 100)	
Common serotypes - serotype 14	100 (15.8 to 100)	100 (29.2 to 100)	100 (2.5 to 100)	
Common serotype - serotype 18C	100 (2.5 to 100)	80 (28.4 to 99.5)	100 (2.5 to 100)	
Common serotype - serotype 19F	100 (15.8 to 100)	60 (14.7 to 94.7)	100 (2.5 to 100)	
Common serotype - serotype 23F	100 (15.8 to 100)	100 (47.8 to 100)	100 (2.5 to 100)	
Additional serotype - serotype 1	100 (15.8 to 100)	83.3 (35.9 to 99.6)	100 (2.5 to 100)	

Additional serotype - serotype 3	100 (15.8 to 100)	100 (54.1 to 100)	100 (2.5 to 100)	
Additional serotype - serotype 5	100 (15.8 to 100)	66.7 (22.3 to 95.7)	100 (2.5 to 100)	
Additional serotype - serotype 6A	100 (15.8 to 100)	100 (54.1 to 100)	100 (2.5 to 100)	
Additional serotype - serotype 7F	100 (15.8 to 100)	100 (54.1 to 100)	100 (2.5 to 100)	
Additional serotype - serotype 19A	100 (15.8 to 100)	100 (47.8 to 100)	100 (2.5 to 100)	

Notes:

[67] - N=number of subjects with a determinate OPA antibody titer to the given serotype.

[68] - N=number of subjects with a determinate OPA antibody titer to the given serotype.

[69] - N=number of subjects with a determinate OPA antibody titer to the given serotype.

Statistical analyses

No statistical analyses for this end point

Other pre-specified: Percentage of Subjects Achieving OPA Titers ≥LLOQ Measured 1 Month After the Relevant Catch-up Dose

End point title	Percentage of Subjects Achieving OPA Titers ≥LLOQ Measured 1 Month After the Relevant Catch-up Dose[70]

End point description:

Percentage of subjects in 13vPnC Groups 4 and 5 achieving OPA with 95% CI for serotypes 4, 6B, 9V, 14, 18C, 19F, 23F, 1, 3, 5, 6A, 7F, and 19A. Exact 2-sided CI based upon the observed proportion of subjects. The LLOQ in titers for each serotype was: Pn001, 18; Pn003, 12; Pn004, 21; Pn005, 29; Pn06A, 37; Pn06B, 43, Pn7F, 210; Pn09V, 345; Pn014, 35; Pn18C, 31; Pn19A, 18; Pn19F, 48; and Pn23F, 13. Limit of detection (LOD) established as lowest titer possible in assay, which was 8. OPA titers below LLOQ set to 0.5*LOD for analysis. Evaluable immunogenicity population.

End point type	Other pre-specified

End point timeframe:

28 to 56 days after vaccination 2 for Group 4, and after the single vaccination in Group 5

Notes:

[70] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: The endpoint was planned to be assessed for subjects who received catch-up dose.

End point values	13vPnC Group 4 (2 Catch-Up Doses)	13vPnC Group 5 (1 Catch-Up Dose)		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	4[71]	8[72]		
Units: Percentage of Subjects				
number (confidence interval 95%)				
Common serotypes - serotype 4	100 (29.2 to 100)	100 (63.1 to 100)		
Common serotypes - serotype 6B	100 (29.2 to 100)	100 (54.1 to 100)		
Common serotypes - serotype 9V	100 (39.8 to 100)	100 (54.1 to 100)		
Common serotypes - serotype 14	100 (15.8 to 100)	100 (54.1 to 100)		
Common serotype - serotype 18C	100 (29.2 to 100)	100 (54.1 to 100)		

Common serotype - serotype 19F	100 (29.2 to 100)	100 (54.1 to 100)		
Common serotype - serotype 23F	100 (29.2 to 100)	100 (54.1 to 100)		
Additional serotype - serotype 1	100 (39.8 to 100)	87.5 (47.3 to 99.7)		
Additional serotype - serotype 3	100 (39.8 to 100)	100 (63.1 to 100)		
Additional serotype - serotype 5	100 (39.8 to 100)	75 (34.9 to 96.8)		
Additional serotype - serotype 6A	100 (39.8 to 100)	87.5 (47.3 to 99.7)		
Additional serotype - serotype 7F	100 (39.8 to 100)	100 (63.1 to 100)		
Additional serotype - serotype 19A	100 (39.8 to 100)	100 (47.8 to 100)		

Notes:

[71] - N=number of subjects with a determinate OPA antibody titer to the given serotype.

[72] - N=number of subjects with a determinate OPA antibody titer to the given serotype.

Statistical analyses

No statistical analyses for this end point

Other pre-specified: Pneumococcal OPA Geometric Mean Titers (GMTs) 1 Month After the Infant Series

End point title	Pneumococcal OPA Geometric Mean Titers (GMTs) 1 Month After the Infant Series[73]

End point description:

Antibody geometric mean titers as measured by OPA assay for 7 common pneumococcal serotypes (serotypes 4, 6B, 9V, 14, 18C, 19F, and 23F) and 6 additional pneumococcal serotypes specific to 13vPnC (Serotypes 1, 3, 5, 6A, 7F, and 19A). GMTs were calculated using all subjects with available data for the specified blood draw. CIs for the GMTs are back transformations of confidence levels based on the Student t distribution for the mean logarithm of the titers. Evaluable immunogenicity population. Here "-99999" and "99999"in the 95% CI signifies NA. For reporting group 13vPnC Group 2 and in common serotype 19F of reporting group 13vPnC Group 3, NA signifies that within group, confidence intervals for serotypes for which subjects all have the same titer value were not computed since variability cannot be estimated. For reporting group 13vPnC Group 3, NA =1.3309*10^8.

End point type	Other pre-specified

End point timeframe:

28 to 56 days after vaccination 3 for Group 1, after vaccination 2 for Group 2, and after vaccination 1 for Group 3

Notes:

[73] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: The endpoint was planned to be assessed for subjects who received infant dose and toddler dose.

End point values	13vPnC Group 1 (3 Doses Infant Series and 1 Toddler Dose)	13vPnC Group 2 (2 Doses Infant Series and 1 Toddler Dose)	13vPnC Group 3 (1 Dose Infant Series and 1 Toddler Dose)	
Subject group type	Reporting group	Reporting group	Reporting group	
Number of subjects analysed	11[74]	5[75]	3[76]	
Units: Titers				
geometric mean (confidence interval				

95%)				
Common serotypes - serotype 4	123 (38.4 to 390.6)	116 (3.1 to 4294.7)	36 (0.3 to 4185.7)	
Common serotypes - serotype 9V	20 (3.1 to 123.1)	4 (-99999 to 99999)	120 (0.1 to 199619.2)	
Common serotype - serotype 19F	151 (49.6 to 457.7)	31 (0.7 to 1348.9)	4 (-99999 to 99999)	
Common serotype - serotype 23F	299 (79.5 to 1124.8)	81 (0.1 to 52488.1)	566 (182.6 to 1755.9)	
Additional serotype - serotype 1	41 (9.2 to 183.5)	4 (-99999 to 99999)	7 (0.5 to 109.4)	
Additional serotype - serotype 3	58 (28.9 to 117.7)	82 (21.7 to 311.6)	206 (66.1 to 639)	
Additional serotype - serotype 5	26 (9.6 to 72.3)	18 (2.9 to 110.8)	9 (0.3 to 234.5)	
Additional serotype - serotype 6A	731 (182.2 to 2933.6)	1069 (337.1 to 3390.4)	984 (179.7 to 5383.5)	
Additional serotype - serotype 7F	1396 (790.5 to 2467)	1177 (692.5 to 2001.6)	3485 (2308.5 to 5261.5)	
Common serotype - serotype 6B	1280 (610.3 to 2683.5)	1450 (370.4 to 5675.6)	1790 (0 to 72147076)	
Common serotype - serotype 14	332 (98 to 1124.2)	71 (3.3 to 1516)	449 (0.1 to 2059598)	
Common serotype - serotype 18C	868 (304.4 to 2475)	452 (67.7 to 3023.6)	906 (0.6 to 1340206)	
Additional serotype - serotype 19A	58 (17.7 to 191.3)	158 (2.5 to 9872.6)	380 (0 to 99999)	

Notes:

[74] - N=number of subjects with a determinate OPA antibody titer to the given serotype.

[75] - N=number of subjects with a determinate OPA antibody titer to the given serotype.

[76] - N=number of subjects with a determinate OPA antibody titer to the given serotype.

Statistical analyses

No statistical analyses for this end point

Other pre-specified: Pneumococcal OPA GMTs 1 Month After the Toddler Dose

End point title	Pneumococcal OPA GMTs 1 Month After the Toddler Dose[77]

End point description:

Antibody geometric mean titers as measured by OPA assay for 7 common pneumococcal serotypes (serotypes 4, 6B, 9V, 14, 18C, 19F, and 23F) and 6 additional pneumococcal serotypes specific to 13vPnC (Serotypes 1, 3, 5, 6A, 7F, and 19A). GMTs were calculated using all subjects with available data for the specified blood draw. CIs for the GMTs are back transformations of confidence levels based on the Student t distribution for the mean logarithm of the titers. Evaluable immunogenicity population. Here "-99999" and "-99999" in 95% CI signifies NA. For reporting group 13vPnC Group 3 and in common serotype 18C of reporting group 13vPnC Group 1, NA signifies that within group, confidence intervals for serotypes with only 1 subject who had a determinate OPA antibody titer were not computed since variability cannot be estimated. For reporting group 13vPnC Group 1, in common serotype 6B, 19F, 23F and additional serotype 6A NA =2.4794*10^8, 2.4862*10^8, 1.208*10^12 and 1.9613*10^9 respectively.

End point type	Other pre-specified

End point timeframe:

28 to 56 days after vaccination 4 for Group 1, after vaccination 3 for Group 2, after vaccination 2 for Group 3

Notes:

[77] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.

Justification: The endpoint was planned to be assessed for subjects who received infant dose and toddler dose.

End point values	13vPnC Group 1 (3 Doses Infant Series and 1 Toddler Dose)	13vPnC Group 2 (2 Doses Infant Series and 1 Toddler Dose)	13vPnC Group 3 (1 Dose Infant Series and 1 Toddler Dose)	
Subject group type	Reporting group	Reporting group	Reporting group	
Number of subjects analysed	2[78]	6[79]	1[80]	
Units: Titers				
geometric mean (confidence interval 95%)				
Common serotypes - serotype 4	599 (422.4 to 850.2)	58 (2.6 to 1317.2)	220 (-99999 to 99999)	
Common serotypes - serotype 9V	428 (68.2 to 2681.9)	460 (35 to 6035.8)	2765 (-99999 to 99999)	
Common serotypes - serotype 14	775 (5.8 to 104290.7)	551 (73.6 to 4120.2)	779 (-99999 to 99999)	
Common serotype - serotype 18C	3311 (-99999 to 99999)	387 (14.8 to 10145.6)	4316 (-99999 to 99999)	
Additional serotype - serotype 1	46 (0.1 to 14754.5)	66 (14.4 to 305.2)	165 (-99999 to 99999)	
Additional serotype - serotype 3	159 (0.2 to 141793.2)	138 (60.9 to 313.7)	402 (-99999 to 99999)	
Additional serotype - serotype 5	87 (0.8 to 9175.7)	30 (5.4 to 164.2)	88 (-99999 to 99999)	
Additional serotype - serotype 7F	4590 (258.5 to 81484.1)	1575 (810.3 to 3061.7)	6105 (-99999 to 99999)	
Additional serotype - serotype 19A	1259 (210.3 to 7542.2)	140 (22.7 to 860.3)	427 (-99999 to 99999)	
Common serotype - serotype 6B	1565 (0 to 99999)	605 (17.1 to 21473.7)	12470 (-99999 to 99999)	
Common serotype - serotype 19F	397 (0 to 99999)	73 (2.5 to 2087.3)	532 (-99999 to 99999)	
Common serotype - serotype 23F	227 (0 to 99999)	313 (30.6 to 3195.2)	1660 (-99999 to 99999)	
Additional serotype - serotype 6A	3207 (0 to 99999)	1688 (480.3 to 5929.3)	6954 (-99999 to 99999)	

Notes:

[78] - N=number of subjects with a determinate OPA antibody titer to the given serotype.

[79] - N=number of subjects with a determinate OPA antibody titer to the given serotype.

[80] - N=number of subjects with a determinate OPA antibody titer to the given serotype.

Statistical analyses

No statistical analyses for this end point

Other pre-specified: Pneumococcal OPA GMTs 1 Month After the Relevant Catch-up Dose

End point title	Pneumococcal OPA GMTs 1 Month After the Relevant Catch-up Dose[81]

End point description:

Antibody geometric mean titers as measured by OPA assay for 13 pneumococcal serotypes (serotypes 1, 3, 4, 5, 6A, 6B, 7F, 9V, 14, 18C, 19A, 19F, and 23F). GMTs were calculated using all subjects with available data for the specified blood draw. CIs for the GMTs are back transformations of confidence levels based on the Student t distribution for the mean logarithm of the titers. Evaluable immunogenicity population.

End point type	Other pre-specified

End point timeframe:

28 to 56 days after vaccination 2 for Group 4, and after the single vaccination in Group 5

Notes:

[81] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.

Justification: The endpoint was planned to be assessed for subjects who received catch-up dose.

End point values	13vPnC Group 4 (2 Catch-Up Doses)	13vPnC Group 5 (1 Catch-Up Dose)		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	4[82]	8[83]		
Units: Titers				
geometric mean (confidence interval 95%)				
Common serotypes - serotype 4	594 (107.1 to 3295.5)	1165 (261.1 to 5197.9)		
Common serotypes - serotype 6B	7236 (3423.7 to 15294)	6795 (795.9 to 58012.5)		
Common serotypes - serotype 9V	2748 (457.9 to 16485.4)	2979 (883.6 to 10040.5)		
Common serotypes - serotype 14	1159 (31.2 to 42979.5)	1547 (347.4 to 6892.2)		
Common serotype - serotype 18C	5910 (1824.6 to 19141.1)	1871 (915.7 to 3824.2)		
Common serotype - serotype 19F	1584 (463.4 to 5417.3)	594 (84.8 to 4164.9)		
Common serotype - serotype 23F	2870 (2572.2 to 3203.4)	1240 (423 to 3632.5)		
Additional serotype - serotype 1	402 (285.8 to 564.7)	62 (18 to 211.6)		
Additional serotype - serotype 3	218 (102 to 466.8)	202 (136 to 299.3)		
Additional serotype - serotype 5	128 (15.9 to 1032)	63 (13.3 to 302.2)		
Additional serotype - serotype 6A	9067 (5187 to 15848.5)	2292 (231.2 to 22724)		
Additional serotype - serotype 7F	7776 (3701.3 to 16335.8)	6086 (2680.9 to 13818.2)		
Additional serotype - serotype 19A	815 (549.9 to 1206.8)	400 (68.4 to 2344.7)		

Notes:

[82] - N=number of subjects with a determinate OPA antibody titer to the given serotype.

[83] - N=number of subjects with a determinate OPA antibody titer to the given serotype.

Statistical analyses

No statistical analyses for this end point

Other pre-specified: Correlation of OPA and IgG Values for Each 13vPnC Serotype 1 Month After the Infant Series

End point title	Correlation of OPA and IgG Values for Each 13vPnC Serotype 1 Month After the Infant Series[84]

End point description:

Correlative analysis output data are only available as figures and were not analyzed or presented statistically. Here "99999" in number signifies NA. Correlative analysis output data are only available as figures and were not analyzed or presented statistically.

End point type	Other pre-specified

End point timeframe:

28 to 56 days after vaccination 3 for Group 1, after vaccination 2 for Group 2, and after vaccination 1

for Group 3

Notes:

[84] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: The endpoint was planned to be assessed for subjects who received infant dose and toddler dose.

End point values	13vPnC Group 1 (3 Doses Infant Series and 1 Toddler Dose)	13vPnC Group 2 (2 Doses Infant Series and 1 Toddler Dose)	13vPnC Group 3 (1 Dose Infant Series and 1 Toddler Dose)	
Subject group type	Reporting group	Reporting group	Reporting group	
Number of subjects analysed	11	5	3	
Units: Subjects				
number (not applicable)	99999	99999	99999	

Statistical analyses

No statistical analyses for this end point

Other pre-specified: Correlation of OPA and IgG Values for Each 13vPnC Serotype 1 Month After the Toddler Dose

End point title	Correlation of OPA and IgG Values for Each 13vPnC Serotype 1 Month After the Toddler Dose[85]

End point description:

Correlative analysis output data are only available as figures and were not analyzed or presented statistically. Here "99999" in number signifies NA. Correlative analysis output data are only available as figures and were not analyzed or presented statistically.

End point type	Other pre-specified

End point timeframe:

28 to 56 days after vaccination 4 for Group 1, after vaccination 3 for Group 2, and after vaccination 2 for Group 3

Notes:

[85] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: The endpoint was planned to be assessed for subjects who received infant dose and toddler dose.

End point values	13vPnC Group 1 (3 Doses Infant Series and 1 Toddler Dose)	13vPnC Group 2 (2 Doses Infant Series and 1 Toddler Dose)	13vPnC Group 3 (1 Dose Infant Series and 1 Toddler Dose)	
Subject group type	Reporting group	Reporting group	Reporting group	
Number of subjects analysed	2	6	1	
Units: Subjects				
number (not applicable)	99999	99999	99999	

Statistical analyses

No statistical analyses for this end point

Other pre-specified: Correlation of OPA and IgG Values for Each 13vPnC Serotype 1 Month After the Relevant Catch-up Dose

End point title	Correlation of OPA and IgG Values for Each 13vPnC Serotype 1 Month After the Relevant Catch-up Dose[86]

End point description:

Correlative analysis output data are only available as figures and were not analyzed or presented statistically. Here "99999" in number signifies NA. Correlative analysis output data are only available as figures and were not analyzed or presented statistically.

End point type	Other pre-specified

End point timeframe:

28 to 56 days after vaccination 2 for Group 4, and after the single vaccination in Group 5

Notes:

[86] - The end point is not reporting statistics for all the arms in the baseline period. It is expected all the baseline period arms will be reported on when providing values for an end point on the baseline period.
Justification: The endpoint was planned to be assessed for subjects who received catch-up dose.

End point values	13vPnC Group 4 (2 Catch-Up Doses)	13vPnC Group 5 (1 Catch-Up Dose)		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	4	8		
Units: Subjects				
number (not applicable)	99999	99999		

Statistical analyses

No statistical analyses for this end point

Adverse events

Adverse events information

Timeframe for reporting adverse events:

Randomization to 6 months after last vaccination

Adverse event reporting additional description:

Same event may appear as both an adverse event (AE) and a serious adverse event (SAE). However, what is presented are distinct events. An event may be categorized as serious in one subject and as non serious in another subject, or one subject may have experienced both events. Version was not captured, here 0.0 is mentioned for dictionary version.

Assessment type	Systematic

Dictionary used

Dictionary name	MedDRA
Dictionary version	0.0

Reporting groups

Reporting group title	Group 1 (Infant Series)

Reporting group description:

Subjects 6 weeks <10 months of age with 0 prior doses of Prevnar received 3 single IM 0.5 mL doses of 13vPnC at least 28 days apart (infant series).

Reporting group title	Group 2 (Infant Series)

Reporting group description:

Subjects <12 months of age with 1 prior dose of Prevnar received 2 single IM 0.5 mL doses of 13vPnC at least 28 days apart (infant series).

Reporting group title	Group 3 (Infant Series)

Reporting group description:

Subjects <12 months of age with 2 prior doses of Prevnar received a single IM 0.5 mL dose of 13vPnC (infant series).

Reporting group title	Group 1 (Toddler Dose)

Reporting group description:

Subjects >12 months of age received a single IM 0.5 mL dose of 13vPnC at least 60 days after last infant dose (toddler dose).

Reporting group title	Group 2 (Toddler Dose)

Reporting group description:

Subjects >12 months of age received a single IM 0.5 mL dose of 13vPnC at least 60 days after last infant dose (toddler dose).

Reporting group title	Group 3 (Toddler Dose)

Reporting group description:

Subjects >12 months of age received a single IM 0.5 mL dose of 13vPnC at least 60 days after last infant dose (toddler dose)

Reporting group title	Group 4 (2 Catch-Up Doses)

Reporting group description:

Subjects >=12 months to <2 years of age received 2 single IM 0.5 mL doses of 13vPnC at least 60 days apart.

Reporting group title	Group 5 (1 Catch-Up Dose)

Reporting group description:

Subjects >=2 years to <5 years of age received a single IM 0.5 mL dose of 13vPnC.

Reporting group title	Group 1 (Infant Series) Follow-up

Reporting group description:

Subjects 6 weeks to <10 months of age with 0 prior doses of Prevnar received 3 single IM 0.5 mL doses of 13vPnC at least 28 days apart were assessed from last 13vPnC Dose blood draw to the 6-month follow-up telephone contact.

Reporting group title	Group 2 (Infant Series) Follow-up

Reporting group description:

Subjects <12 months of age with 1 prior dose of Prevnar received 2 single IM 0.5 mL doses of 13vPnC

at least 28 days apart were assessed from last 13vPnC Dose blood draw to the 6-month follow-up telephone contact.

Reporting group title	Group 3 (Infant Series) Follow-up

Reporting group description:

Subjects <12 months of age with 2 prior doses of Prevnar received a single IM 0.5 mL dose of 13vPnC were assessed from last 13vPnC Dose blood draw to the 6-month follow-up telephone contact.

Reporting group title	Group 4 (2 Catch-Up Doses) Follow-up

Reporting group description:

Subjects >=12 months to <2 years of age received 2 single IM 0.5 mL doses of 13vPnC at least 60 days apart were assessed from last 13vPnC Dose blood draw to the 6-month follow-up telephone contact.

Reporting group title	Group 5 (1 Catch-Up Dose) Follow-up

Reporting group description:

Subjects >=2 years to <5 years of age received a single IM 0.5 mL dose of 13vPnC were assessed from last 13vPnC Dose blood draw to the 6-month follow-up telephone contact.

Reporting group title	Group 1 (After Infant Series)

Reporting group description:

Subjects 6 weeks <10 months of age with 0 prior doses of Prevnar received 3 single IM 0.5 mL doses of 13vPnC at least 28 days apart in infant series.

Reporting group title	Group 2 (After Infant Series)

Reporting group description:

Subjects <12 months of age with 1 prior dose of Prevnar received 2 single IM 0.5 mL doses of 13vPnC at least 28 days apart in infant series.

Reporting group title	Group 3 (After Infant Series)

Reporting group description:

Subjects <12 months of age with 2 prior doses of Prevnar received a single IM 0.5 mL dose of 13vPnC in infant series.

Serious adverse events	Group 1 (Infant Series)	Group 2 (Infant Series)	Group 3 (Infant Series)
Total subjects affected by serious adverse events			
subjects affected / exposed	21 / 151 (13.91%)	2 / 51 (3.92%)	1 / 25 (4.00%)
number of deaths (all causes)	2	0	0
number of deaths resulting from adverse events	0	0	0
Injury, poisoning and procedural complications			
Foreign body			
subjects affected / exposed	0 / 151 (0.00%)	0 / 51 (0.00%)	0 / 25 (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
Congenital, familial and genetic disorders			
Laryngomalacia			
subjects affected / exposed	0 / 151 (0.00%)	0 / 51 (0.00%)	0 / 25 (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			

Convulsion			
subjects affected / exposed	3 / 151 (1.99%)	0 / 51 (0.00%)	0 / 25 (0.00%)
occurrences causally related to treatment / all	0 / 3	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Febrile convulsion			
subjects affected / exposed	1 / 151 (0.66%)	0 / 51 (0.00%)	0 / 25 (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
Blood and lymphatic system disorders			
Anaemia			
subjects affected / exposed	1 / 151 (0.66%)	0 / 51 (0.00%)	0 / 25 (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
Lymphadenitis			
subjects affected / exposed	0 / 151 (0.00%)	0 / 51 (0.00%)	0 / 25 (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			
Pyrexia			
subjects affected / exposed	2 / 151 (1.32%)	0 / 51 (0.00%)	0 / 25 (0.00%)
occurrences causally related to treatment / all	0 / 2	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Sudden infant death syndrome			
subjects affected / exposed	0 / 151 (0.00%)	0 / 51 (0.00%)	0 / 25 (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
Gastrointestinal disorders			
Gastrooesophageal reflux disease			
subjects affected / exposed	0 / 151 (0.00%)	0 / 51 (0.00%)	0 / 25 (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
Respiratory, thoracic and mediastinal disorders			
Pneumonia aspiration			

subjects affected / exposed	2 / 151 (1.32%)	1 / 51 (1.96%)	0 / 25 (0.00%)
occurrences causally related to treatment / all	0 / 2	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Asthma			
subjects affected / exposed	1 / 151 (0.66%)	0 / 51 (0.00%)	0 / 25 (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
Bronchial hyperreactivity			
subjects affected / exposed	0 / 151 (0.00%)	1 / 51 (1.96%)	0 / 25 (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
Respiratory distress			
subjects affected / exposed	1 / 151 (0.66%)	0 / 51 (0.00%)	0 / 25 (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
Respiratory failure			
subjects affected / exposed	1 / 151 (0.66%)	0 / 51 (0.00%)	0 / 25 (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
Apnoea			
subjects affected / exposed	0 / 151 (0.00%)	0 / 51 (0.00%)	1 / 25 (4.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Skin and subcutaneous tissue disorders			
Eczema			
subjects affected / exposed	0 / 151 (0.00%)	0 / 51 (0.00%)	0 / 25 (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			
Breath holding			
subjects affected / exposed	0 / 151 (0.00%)	0 / 51 (0.00%)	0 / 25 (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			

Pneumonia			
subjects affected / exposed	4 / 151 (2.65%)	0 / 51 (0.00%)	1 / 25 (4.00%)
occurrences causally related to treatment / all	0 / 4	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 1	0 / 0	0 / 0
Respiratory syncytial virus bronchiolitis			
subjects affected / exposed	5 / 151 (3.31%)	1 / 51 (1.96%)	0 / 25 (0.00%)
occurrences causally related to treatment / all	0 / 5	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Influenza			
subjects affected / exposed	3 / 151 (1.99%)	1 / 51 (1.96%)	1 / 25 (4.00%)
occurrences causally related to treatment / all	0 / 3	0 / 1	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pneumonia respiratory syncytial viral			
subjects affected / exposed	3 / 151 (1.99%)	0 / 51 (0.00%)	0 / 25 (0.00%)
occurrences causally related to treatment / all	0 / 3	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Bronchiolitis			
subjects affected / exposed	2 / 151 (1.32%)	0 / 51 (0.00%)	0 / 25 (0.00%)
occurrences causally related to treatment / all	0 / 2	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Urinary tract infection			
subjects affected / exposed	2 / 151 (1.32%)	0 / 51 (0.00%)	0 / 25 (0.00%)
occurrences causally related to treatment / all	0 / 2	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Cellulitis staphylococcal			
subjects affected / exposed	1 / 151 (0.66%)	0 / 51 (0.00%)	0 / 25 (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
Meningitis haemophilus			
subjects affected / exposed	1 / 151 (0.66%)	0 / 51 (0.00%)	0 / 25 (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
Septic arthritis haemophilus			

subjects affected / exposed	1 / 151 (0.66%)	0 / 51 (0.00%)	0 / 25 (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
Upper respiratory tract infection			
subjects affected / exposed	1 / 151 (0.66%)	0 / 51 (0.00%)	0 / 25 (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
Meningitis viral			
subjects affected / exposed	0 / 151 (0.00%)	0 / 51 (0.00%)	1 / 25 (4.00%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Abscess			
subjects affected / exposed	0 / 151 (0.00%)	0 / 51 (0.00%)	0 / 25 (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
Herpangina			
subjects affected / exposed	0 / 151 (0.00%)	0 / 51 (0.00%)	0 / 25 (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
Croup infectious			
subjects affected / exposed	0 / 151 (0.00%)	0 / 51 (0.00%)	0 / 25 (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
Arthritis bacterial			
subjects affected / exposed	0 / 151 (0.00%)	0 / 51 (0.00%)	0 / 25 (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
Streptococcal bacteraemia			
subjects affected / exposed	0 / 151 (0.00%)	0 / 51 (0.00%)	0 / 25 (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
Gastroenteritis			

subjects affected / exposed	0 / 151 (0.00%)	0 / 51 (0.00%)	0 / 25 (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
Cellulitis			
subjects affected / exposed	0 / 151 (0.00%)	0 / 51 (0.00%)	0 / 25 (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
Eczema infected			
subjects affected / exposed	0 / 151 (0.00%)	0 / 51 (0.00%)	0 / 25 (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
Genital abscess			
subjects affected / exposed	0 / 151 (0.00%)	0 / 51 (0.00%)	0 / 25 (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
Skin infection			
subjects affected / exposed	0 / 151 (0.00%)	0 / 51 (0.00%)	0 / 25 (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
Tracheobronchitis			
subjects affected / exposed	0 / 151 (0.00%)	0 / 51 (0.00%)	0 / 25 (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			
Hyponatraemia			
subjects affected / exposed	1 / 151 (0.66%)	0 / 51 (0.00%)	0 / 25 (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
Dehydration			
subjects affected / exposed	0 / 151 (0.00%)	0 / 51 (0.00%)	0 / 25 (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	Group 1 (Toddler	Group 2 (Toddler	Group 3 (Toddler

	Dose)	Dose)	Dose)
Total subjects affected by serious adverse events			
subjects affected / exposed	0 / 15 (0.00%)	0 / 23 (0.00%)	2 / 16 (12.50%)
number of deaths (all causes)	0	0	0
number of deaths resulting from adverse events	0	0	0
Injury, poisoning and procedural complications			
Foreign body			
subjects affected / exposed	0 / 15 (0.00%)	0 / 23 (0.00%)	0 / 16 (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
Congenital, familial and genetic disorders			
Laryngomalacia			
subjects affected / exposed	0 / 15 (0.00%)	0 / 23 (0.00%)	1 / 16 (6.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Nervous system disorders			
Convulsion			
subjects affected / exposed	0 / 15 (0.00%)	0 / 23 (0.00%)	0 / 16 (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
Febrile convulsion			
subjects affected / exposed	0 / 15 (0.00%)	0 / 23 (0.00%)	0 / 16 (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
Blood and lymphatic system disorders			
Anaemia			
subjects affected / exposed	0 / 15 (0.00%)	0 / 23 (0.00%)	0 / 16 (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
Lymphadenitis			
subjects affected / exposed	0 / 15 (0.00%)	0 / 23 (0.00%)	0 / 16 (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			

Pyrexia			
subjects affected / exposed	0 / 15 (0.00%)	0 / 23 (0.00%)	0 / 16 (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
Sudden infant death syndrome			
subjects affected / exposed	0 / 15 (0.00%)	0 / 23 (0.00%)	0 / 16 (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
Gastrointestinal disorders			
Gastrooesophageal reflux disease			
subjects affected / exposed	0 / 15 (0.00%)	0 / 23 (0.00%)	1 / 16 (6.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Respiratory, thoracic and mediastinal disorders			
Pneumonia aspiration			
subjects affected / exposed	0 / 15 (0.00%)	0 / 23 (0.00%)	0 / 16 (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
Asthma			
subjects affected / exposed	0 / 15 (0.00%)	0 / 23 (0.00%)	0 / 16 (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
Bronchial hyperreactivity			
subjects affected / exposed	0 / 15 (0.00%)	0 / 23 (0.00%)	0 / 16 (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
Respiratory distress			
subjects affected / exposed	0 / 15 (0.00%)	0 / 23 (0.00%)	1 / 16 (6.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Respiratory failure			
subjects affected / exposed	0 / 15 (0.00%)	0 / 23 (0.00%)	0 / 16 (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

Apnoea			
subjects affected / exposed	0 / 15 (0.00%)	0 / 23 (0.00%)	0 / 16 (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
Skin and subcutaneous tissue disorders			
Eczema			
subjects affected / exposed	0 / 15 (0.00%)	0 / 23 (0.00%)	0 / 16 (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			
Breath holding			
subjects affected / exposed	0 / 15 (0.00%)	0 / 23 (0.00%)	1 / 16 (6.25%)
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			
Pneumonia			
subjects affected / exposed	0 / 15 (0.00%)	0 / 23 (0.00%)	0 / 16 (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
Respiratory syncytial virus bronchiolitis			
subjects affected / exposed	0 / 15 (0.00%)	0 / 23 (0.00%)	0 / 16 (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
Influenza			
subjects affected / exposed	0 / 15 (0.00%)	0 / 23 (0.00%)	0 / 16 (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
Pneumonia respiratory syncytial viral			
subjects affected / exposed	0 / 15 (0.00%)	0 / 23 (0.00%)	0 / 16 (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
Bronchiolitis			

subjects affected / exposed	0 / 15 (0.00%)	0 / 23 (0.00%)	0 / 16 (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 / 15 (0.00%)	0 / 23 (0.00%)	0 / 16 (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
Cellulitis staphylococcal			
subjects affected / exposed	0 / 15 (0.00%)	0 / 23 (0.00%)	0 / 16 (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
Meningitis haemophilus			
subjects affected / exposed	0 / 15 (0.00%)	0 / 23 (0.00%)	0 / 16 (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
Septic arthritis haemophilus			
subjects affected / exposed	0 / 15 (0.00%)	0 / 23 (0.00%)	0 / 16 (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
Upper respiratory tract infection			
subjects affected / exposed	0 / 15 (0.00%)	0 / 23 (0.00%)	0 / 16 (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
Meningitis viral			
subjects affected / exposed	0 / 15 (0.00%)	0 / 23 (0.00%)	0 / 16 (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
Abscess			
subjects affected / exposed	0 / 15 (0.00%)	0 / 23 (0.00%)	0 / 16 (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
Herpangina			

subjects affected / exposed	0 / 15 (0.00%)	0 / 23 (0.00%)	1 / 16 (6.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Croup infectious			
subjects affected / exposed	0 / 15 (0.00%)	0 / 23 (0.00%)	0 / 16 (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
Arthritis bacterial			
subjects affected / exposed	0 / 15 (0.00%)	0 / 23 (0.00%)	0 / 16 (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
Streptococcal bacteraemia			
subjects affected / exposed	0 / 15 (0.00%)	0 / 23 (0.00%)	0 / 16 (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
Gastroenteritis			
subjects affected / exposed	0 / 15 (0.00%)	0 / 23 (0.00%)	0 / 16 (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
Cellulitis			
subjects affected / exposed	0 / 15 (0.00%)	0 / 23 (0.00%)	0 / 16 (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
Eczema infected			
subjects affected / exposed	0 / 15 (0.00%)	0 / 23 (0.00%)	0 / 16 (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
Genital abscess			
subjects affected / exposed	0 / 15 (0.00%)	0 / 23 (0.00%)	0 / 16 (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
Skin infection			

subjects affected / exposed	0 / 15 (0.00%)	0 / 23 (0.00%)	0 / 16 (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
Tracheobronchitis			
subjects affected / exposed	0 / 15 (0.00%)	0 / 23 (0.00%)	0 / 16 (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			
Hyponatraemia			
subjects affected / exposed	0 / 15 (0.00%)	0 / 23 (0.00%)	0 / 16 (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
Dehydration			
subjects affected / exposed	0 / 15 (0.00%)	0 / 23 (0.00%)	1 / 16 (6.25%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0

Serious adverse events	Group 4 (2 Catch-Up Doses)	Group 5 (1 Catch-Up Dose)	Group 1 (Infant Series) Follow-up
Total subjects affected by serious adverse events			
subjects affected / exposed	3 / 66 (4.55%)	0 / 79 (0.00%)	6 / 151 (3.97%)
number of deaths (all causes)	0	0	1
number of deaths resulting from adverse events	0	0	0
Injury, poisoning and procedural complications			
Foreign body			
subjects affected / exposed	0 / 66 (0.00%)	0 / 79 (0.00%)	1 / 151 (0.66%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Congenital, familial and genetic disorders			
Laryngomalacia			
subjects affected / exposed	0 / 66 (0.00%)	0 / 79 (0.00%)	0 / 151 (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			
Convulsion			

subjects affected / exposed	0 / 66 (0.00%)	0 / 79 (0.00%)	0 / 151 (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
Febrile convulsion			
subjects affected / exposed	0 / 66 (0.00%)	0 / 79 (0.00%)	0 / 151 (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
Blood and lymphatic system disorders			
Anaemia			
subjects affected / exposed	0 / 66 (0.00%)	0 / 79 (0.00%)	0 / 151 (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
Lymphadenitis			
subjects affected / exposed	1 / 66 (1.52%)	0 / 79 (0.00%)	0 / 151 (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
General disorders and administration site conditions			
Pyrexia			
subjects affected / exposed	0 / 66 (0.00%)	0 / 79 (0.00%)	0 / 151 (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
Sudden infant death syndrome			
subjects affected / exposed	0 / 66 (0.00%)	0 / 79 (0.00%)	1 / 151 (0.66%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 1
Gastrointestinal disorders			
Gastrooesophageal reflux disease			
subjects affected / exposed	0 / 66 (0.00%)	0 / 79 (0.00%)	0 / 151 (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
Respiratory, thoracic and mediastinal disorders			
Pneumonia aspiration			

subjects affected / exposed	0 / 66 (0.00%)	0 / 79 (0.00%)	0 / 151 (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
Asthma			
subjects affected / exposed	0 / 66 (0.00%)	0 / 79 (0.00%)	0 / 151 (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
Bronchial hyperreactivity			
subjects affected / exposed	0 / 66 (0.00%)	0 / 79 (0.00%)	0 / 151 (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
Respiratory distress			
subjects affected / exposed	0 / 66 (0.00%)	0 / 79 (0.00%)	0 / 151 (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
Respiratory failure			
subjects affected / exposed	0 / 66 (0.00%)	0 / 79 (0.00%)	0 / 151 (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
Apnoea			
subjects affected / exposed	0 / 66 (0.00%)	0 / 79 (0.00%)	0 / 151 (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
Skin and subcutaneous tissue disorders			
Eczema			
subjects affected / exposed	0 / 66 (0.00%)	0 / 79 (0.00%)	0 / 151 (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			
Breath holding			
subjects affected / exposed	0 / 66 (0.00%)	0 / 79 (0.00%)	0 / 151 (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			

Pneumonia			
subjects affected / exposed	1 / 66 (1.52%)	0 / 79 (0.00%)	2 / 151 (1.32%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 2
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Respiratory syncytial virus bronchiolitis			
subjects affected / exposed	0 / 66 (0.00%)	0 / 79 (0.00%)	0 / 151 (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
Influenza			
subjects affected / exposed	1 / 66 (1.52%)	0 / 79 (0.00%)	1 / 151 (0.66%)
occurrences causally related to treatment / all	0 / 1	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Pneumonia respiratory syncytial viral			
subjects affected / exposed	0 / 66 (0.00%)	0 / 79 (0.00%)	0 / 151 (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
Bronchiolitis			
subjects affected / exposed	0 / 66 (0.00%)	0 / 79 (0.00%)	0 / 151 (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 / 66 (0.00%)	0 / 79 (0.00%)	0 / 151 (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
Cellulitis staphylococcal			
subjects affected / exposed	0 / 66 (0.00%)	0 / 79 (0.00%)	0 / 151 (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
Meningitis haemophilus			
subjects affected / exposed	0 / 66 (0.00%)	0 / 79 (0.00%)	0 / 151 (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
Septic arthritis haemophilus			

subjects affected / exposed	0 / 66 (0.00%)	0 / 79 (0.00%)	0 / 151 (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
Upper respiratory tract infection			
subjects affected / exposed	0 / 66 (0.00%)	0 / 79 (0.00%)	0 / 151 (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
Meningitis viral			
subjects affected / exposed	0 / 66 (0.00%)	0 / 79 (0.00%)	0 / 151 (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
Abscess			
subjects affected / exposed	1 / 66 (1.52%)	0 / 79 (0.00%)	0 / 151 (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
Herpangina			
subjects affected / exposed	0 / 66 (0.00%)	0 / 79 (0.00%)	0 / 151 (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
Croup infectious			
subjects affected / exposed	0 / 66 (0.00%)	0 / 79 (0.00%)	0 / 151 (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
Arthritis bacterial			
subjects affected / exposed	0 / 66 (0.00%)	0 / 79 (0.00%)	1 / 151 (0.66%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Streptococcal bacteraemia			
subjects affected / exposed	0 / 66 (0.00%)	0 / 79 (0.00%)	1 / 151 (0.66%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Gastroenteritis			

subjects affected / exposed	0 / 66 (0.00%)	0 / 79 (0.00%)	0 / 151 (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
Cellulitis			
subjects affected / exposed	0 / 66 (0.00%)	0 / 79 (0.00%)	0 / 151 (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
Eczema infected			
subjects affected / exposed	0 / 66 (0.00%)	0 / 79 (0.00%)	0 / 151 (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
Genital abscess			
subjects affected / exposed	0 / 66 (0.00%)	0 / 79 (0.00%)	0 / 151 (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
Skin infection			
subjects affected / exposed	0 / 66 (0.00%)	0 / 79 (0.00%)	0 / 151 (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
Tracheobronchitis			
subjects affected / exposed	0 / 66 (0.00%)	0 / 79 (0.00%)	0 / 151 (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			
Hyponatraemia			
subjects affected / exposed	0 / 66 (0.00%)	0 / 79 (0.00%)	0 / 151 (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
Dehydration			
subjects affected / exposed	0 / 66 (0.00%)	0 / 79 (0.00%)	0 / 151 (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	Group 2 (Infant	Group 3 (Infant	Group 4 (2 Catch-Up

	Series) Follow-up	Series) Follow-up	Doses) Follow-up
Total subjects affected by serious adverse events			
subjects affected / exposed	6 / 51 (11.76%)	1 / 25 (4.00%)	1 / 66 (1.52%)
number of deaths (all causes)	0	0	0
number of deaths resulting from adverse events	0	0	0
Injury, poisoning and procedural complications			
Foreign body			
subjects affected / exposed	1 / 51 (1.96%)	0 / 25 (0.00%)	0 / 66 (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
Congenital, familial and genetic disorders			
Laryngomalacia			
subjects affected / exposed	0 / 51 (0.00%)	0 / 25 (0.00%)	0 / 66 (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			
Convulsion			
subjects affected / exposed	0 / 51 (0.00%)	0 / 25 (0.00%)	0 / 66 (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
Febrile convulsion			
subjects affected / exposed	0 / 51 (0.00%)	0 / 25 (0.00%)	0 / 66 (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
Blood and lymphatic system disorders			
Anaemia			
subjects affected / exposed	1 / 51 (1.96%)	0 / 25 (0.00%)	0 / 66 (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
Lymphadenitis			
subjects affected / exposed	0 / 51 (0.00%)	0 / 25 (0.00%)	0 / 66 (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			

Pyrexia			
subjects affected / exposed	0 / 51 (0.00%)	0 / 25 (0.00%)	0 / 66 (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
Sudden infant death syndrome			
subjects affected / exposed	0 / 51 (0.00%)	0 / 25 (0.00%)	0 / 66 (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
Gastrointestinal disorders			
Gastrooesophageal reflux disease			
subjects affected / exposed	0 / 51 (0.00%)	0 / 25 (0.00%)	0 / 66 (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
Respiratory, thoracic and mediastinal disorders			
Pneumonia aspiration			
subjects affected / exposed	0 / 51 (0.00%)	0 / 25 (0.00%)	0 / 66 (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
Asthma			
subjects affected / exposed	0 / 51 (0.00%)	0 / 25 (0.00%)	0 / 66 (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
Bronchial hyperreactivity			
subjects affected / exposed	0 / 51 (0.00%)	0 / 25 (0.00%)	0 / 66 (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
Respiratory distress			
subjects affected / exposed	0 / 51 (0.00%)	0 / 25 (0.00%)	0 / 66 (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
Respiratory failure			
subjects affected / exposed	0 / 51 (0.00%)	0 / 25 (0.00%)	0 / 66 (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

Apnoea			
subjects affected / exposed	0 / 51 (0.00%)	0 / 25 (0.00%)	0 / 66 (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
Skin and subcutaneous tissue disorders			
Eczema			
subjects affected / exposed	0 / 51 (0.00%)	0 / 25 (0.00%)	0 / 66 (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			
Breath holding			
subjects affected / exposed	0 / 51 (0.00%)	0 / 25 (0.00%)	0 / 66 (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			
Pneumonia			
subjects affected / exposed	3 / 51 (5.88%)	0 / 25 (0.00%)	0 / 66 (0.00%)
occurrences causally related to treatment / all	0 / 3	0 / 0	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Respiratory syncytial virus bronchiolitis			
subjects affected / exposed	0 / 51 (0.00%)	0 / 25 (0.00%)	0 / 66 (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
Influenza			
subjects affected / exposed	0 / 51 (0.00%)	0 / 25 (0.00%)	0 / 66 (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
Pneumonia respiratory syncytial viral			
subjects affected / exposed	1 / 51 (1.96%)	0 / 25 (0.00%)	0 / 66 (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
Bronchiolitis			

subjects affected / exposed	0 / 51 (0.00%)	0 / 25 (0.00%)	0 / 66 (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 / 51 (0.00%)	0 / 25 (0.00%)	0 / 66 (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
Cellulitis staphylococcal			
subjects affected / exposed	0 / 51 (0.00%)	0 / 25 (0.00%)	0 / 66 (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
Meningitis haemophilus			
subjects affected / exposed	0 / 51 (0.00%)	0 / 25 (0.00%)	0 / 66 (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
Septic arthritis haemophilus			
subjects affected / exposed	0 / 51 (0.00%)	0 / 25 (0.00%)	0 / 66 (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
Upper respiratory tract infection			
subjects affected / exposed	0 / 51 (0.00%)	0 / 25 (0.00%)	0 / 66 (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
Meningitis viral			
subjects affected / exposed	0 / 51 (0.00%)	0 / 25 (0.00%)	0 / 66 (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
Abscess			
subjects affected / exposed	0 / 51 (0.00%)	0 / 25 (0.00%)	1 / 66 (1.52%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Herpangina			

subjects affected / exposed	0 / 51 (0.00%)	0 / 25 (0.00%)	0 / 66 (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
Croup infectious			
subjects affected / exposed	1 / 51 (1.96%)	1 / 25 (4.00%)	0 / 66 (0.00%)
occurrences causally related to treatment / all	0 / 1	0 / 1	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Arthritis bacterial			
subjects affected / exposed	0 / 51 (0.00%)	0 / 25 (0.00%)	0 / 66 (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
Streptococcal bacteraemia			
subjects affected / exposed	0 / 51 (0.00%)	0 / 25 (0.00%)	0 / 66 (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
Gastroenteritis			
subjects affected / exposed	0 / 51 (0.00%)	0 / 25 (0.00%)	0 / 66 (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
Cellulitis			
subjects affected / exposed	0 / 51 (0.00%)	0 / 25 (0.00%)	0 / 66 (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
Eczema infected			
subjects affected / exposed	0 / 51 (0.00%)	0 / 25 (0.00%)	0 / 66 (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
Genital abscess			
subjects affected / exposed	0 / 51 (0.00%)	0 / 25 (0.00%)	0 / 66 (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
Skin infection			

subjects affected / exposed	0 / 51 (0.00%)	0 / 25 (0.00%)	0 / 66 (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
Tracheobronchitis			
subjects affected / exposed	0 / 51 (0.00%)	0 / 25 (0.00%)	0 / 66 (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			
Hyponatraemia			
subjects affected / exposed	0 / 51 (0.00%)	0 / 25 (0.00%)	0 / 66 (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
Dehydration			
subjects affected / exposed	0 / 51 (0.00%)	1 / 25 (4.00%)	0 / 66 (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

Serious adverse events	Group 5 (1 Catch-Up Dose) Follow-up	Group 1 (After Infant Series)	Group 2 (After Infant Series)
Total subjects affected by serious adverse events			
subjects affected / exposed	1 / 79 (1.27%)	4 / 151 (2.65%)	2 / 51 (3.92%)
number of deaths (all causes)	0	0	0
number of deaths resulting from adverse events	0	0	0
Injury, poisoning and procedural complications			
Foreign body			
subjects affected / exposed	0 / 79 (0.00%)	0 / 151 (0.00%)	0 / 51 (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
Congenital, familial and genetic disorders			
Laryngomalacia			
subjects affected / exposed	0 / 79 (0.00%)	0 / 151 (0.00%)	0 / 51 (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			
Convulsion			

subjects affected / exposed	0 / 79 (0.00%)	0 / 151 (0.00%)	0 / 51 (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
Febrile convulsion			
subjects affected / exposed	0 / 79 (0.00%)	0 / 151 (0.00%)	0 / 51 (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
Blood and lymphatic system disorders			
Anaemia			
subjects affected / exposed	0 / 79 (0.00%)	0 / 151 (0.00%)	0 / 51 (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
Lymphadenitis			
subjects affected / exposed	0 / 79 (0.00%)	0 / 151 (0.00%)	0 / 51 (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			
Pyrexia			
subjects affected / exposed	0 / 79 (0.00%)	0 / 151 (0.00%)	0 / 51 (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
Sudden infant death syndrome			
subjects affected / exposed	0 / 79 (0.00%)	0 / 151 (0.00%)	0 / 51 (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
Gastrointestinal disorders			
Gastrooesophageal reflux disease			
subjects affected / exposed	0 / 79 (0.00%)	0 / 151 (0.00%)	0 / 51 (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
Respiratory, thoracic and mediastinal disorders			
Pneumonia aspiration			

subjects affected / exposed	0 / 79 (0.00%)	0 / 151 (0.00%)	0 / 51 (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
Asthma			
subjects affected / exposed	0 / 79 (0.00%)	0 / 151 (0.00%)	0 / 51 (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
Bronchial hyperreactivity			
subjects affected / exposed	0 / 79 (0.00%)	0 / 151 (0.00%)	0 / 51 (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
Respiratory distress			
subjects affected / exposed	0 / 79 (0.00%)	0 / 151 (0.00%)	0 / 51 (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
Respiratory failure			
subjects affected / exposed	0 / 79 (0.00%)	0 / 151 (0.00%)	0 / 51 (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
Apnoea			
subjects affected / exposed	0 / 79 (0.00%)	0 / 151 (0.00%)	0 / 51 (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
Skin and subcutaneous tissue disorders			
Eczema			
subjects affected / exposed	0 / 79 (0.00%)	0 / 151 (0.00%)	0 / 51 (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			
Breath holding			
subjects affected / exposed	0 / 79 (0.00%)	0 / 151 (0.00%)	0 / 51 (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			

Pneumonia			
subjects affected / exposed	0 / 79 (0.00%)	2 / 151 (1.32%)	1 / 51 (1.96%)
occurrences causally related to treatment / all	0 / 0	0 / 2	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Respiratory syncytial virus bronchiolitis			
subjects affected / exposed	0 / 79 (0.00%)	2 / 151 (1.32%)	0 / 51 (0.00%)
occurrences causally related to treatment / all	0 / 0	0 / 2	0 / 0
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Influenza			
subjects affected / exposed	0 / 79 (0.00%)	0 / 151 (0.00%)	0 / 51 (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
Pneumonia respiratory syncytial viral			
subjects affected / exposed	0 / 79 (0.00%)	0 / 151 (0.00%)	0 / 51 (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
Bronchiolitis			
subjects affected / exposed	0 / 79 (0.00%)	1 / 151 (0.66%)	0 / 51 (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	0 / 79 (0.00%)	0 / 151 (0.00%)	0 / 51 (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
Cellulitis staphylococcal			
subjects affected / exposed	0 / 79 (0.00%)	0 / 151 (0.00%)	0 / 51 (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
Meningitis haemophilus			
subjects affected / exposed	0 / 79 (0.00%)	0 / 151 (0.00%)	0 / 51 (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
Septic arthritis haemophilus			

subjects affected / exposed	0 / 79 (0.00%)	0 / 151 (0.00%)	0 / 51 (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
Upper respiratory tract infection			
subjects affected / exposed	0 / 79 (0.00%)	0 / 151 (0.00%)	0 / 51 (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
Meningitis viral			
subjects affected / exposed	0 / 79 (0.00%)	0 / 151 (0.00%)	0 / 51 (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
Abscess			
subjects affected / exposed	0 / 79 (0.00%)	0 / 151 (0.00%)	0 / 51 (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
Herpangina			
subjects affected / exposed	0 / 79 (0.00%)	0 / 151 (0.00%)	0 / 51 (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
Croup infectious			
subjects affected / exposed	0 / 79 (0.00%)	0 / 151 (0.00%)	0 / 51 (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
Arthritis bacterial			
subjects affected / exposed	0 / 79 (0.00%)	0 / 151 (0.00%)	0 / 51 (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
Streptococcal bacteraemia			
subjects affected / exposed	0 / 79 (0.00%)	0 / 151 (0.00%)	0 / 51 (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
Gastroenteritis			

subjects affected / exposed	1 / 79 (1.27%)	0 / 151 (0.00%)	0 / 51 (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
Cellulitis			
subjects affected / exposed	0 / 79 (0.00%)	0 / 151 (0.00%)	0 / 51 (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
Eczema infected			
subjects affected / exposed	0 / 79 (0.00%)	0 / 151 (0.00%)	0 / 51 (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
Genital abscess			
subjects affected / exposed	0 / 79 (0.00%)	0 / 151 (0.00%)	1 / 51 (1.96%)
occurrences causally related to treatment / all	0 / 0	0 / 0	0 / 1
deaths causally related to treatment / all	0 / 0	0 / 0	0 / 0
Skin infection			
subjects affected / exposed	0 / 79 (0.00%)	0 / 151 (0.00%)	0 / 51 (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
Tracheobronchitis			
subjects affected / exposed	0 / 79 (0.00%)	0 / 151 (0.00%)	0 / 51 (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			
Hyponatraemia			
subjects affected / exposed	0 / 79 (0.00%)	0 / 151 (0.00%)	0 / 51 (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
Dehydration			
subjects affected / exposed	1 / 79 (1.27%)	0 / 151 (0.00%)	0 / 51 (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	Group 3 (After		

	Infant Series)		
Total subjects affected by serious adverse events			
subjects affected / exposed	4 / 25 (16.00%)		
number of deaths (all causes)	0		
number of deaths resulting from adverse events	0		
Injury, poisoning and procedural complications			
Foreign body			
subjects affected / exposed	0 / 25 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Congenital, familial and genetic disorders			
Laryngomalacia			
subjects affected / exposed	1 / 25 (4.00%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Nervous system disorders			
Convulsion			
subjects affected / exposed	1 / 25 (4.00%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Febrile convulsion			
subjects affected / exposed	0 / 25 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Blood and lymphatic system disorders			
Anaemia			
subjects affected / exposed	0 / 25 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Lymphadenitis			
subjects affected / exposed	0 / 25 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
General disorders and administration site conditions			

Pyrexia				
subjects affected / exposed	0 / 25 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Sudden infant death syndrome				
subjects affected / exposed	0 / 25 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Gastrointestinal disorders				
Gastrooesophageal reflux disease				
subjects affected / exposed	0 / 25 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Respiratory, thoracic and mediastinal disorders				
Pneumonia aspiration				
subjects affected / exposed	1 / 25 (4.00%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Asthma				
subjects affected / exposed	0 / 25 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Bronchial hyperreactivity				
subjects affected / exposed	1 / 25 (4.00%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Respiratory distress				
subjects affected / exposed	1 / 25 (4.00%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Respiratory failure				
subjects affected / exposed	0 / 25 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			

Apnoea				
subjects affected / exposed	1 / 25 (4.00%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Skin and subcutaneous tissue disorders				
Eczema				
subjects affected / exposed	1 / 25 (4.00%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Psychiatric disorders				
Breath holding				
subjects affected / exposed	0 / 25 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Infections and infestations				
Pneumonia				
subjects affected / exposed	1 / 25 (4.00%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Respiratory syncytial virus bronchiolitis				
subjects affected / exposed	1 / 25 (4.00%)			
occurrences causally related to treatment / all	0 / 1			
deaths causally related to treatment / all	0 / 0			
Influenza				
subjects affected / exposed	0 / 25 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Pneumonia respiratory syncytial viral				
subjects affected / exposed	0 / 25 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Bronchiolitis				

subjects affected / exposed	0 / 25 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Urinary tract infection				
subjects affected / exposed	0 / 25 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Cellulitis staphylococcal				
subjects affected / exposed	0 / 25 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Meningitis haemophilus				
subjects affected / exposed	0 / 25 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Septic arthritis haemophilus				
subjects affected / exposed	0 / 25 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Upper respiratory tract infection				
subjects affected / exposed	0 / 25 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Meningitis viral				
subjects affected / exposed	0 / 25 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Abscess				
subjects affected / exposed	0 / 25 (0.00%)			
occurrences causally related to treatment / all	0 / 0			
deaths causally related to treatment / all	0 / 0			
Herpangina				

subjects affected / exposed	0 / 25 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Croup infectious			
subjects affected / exposed	0 / 25 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Arthritis bacterial			
subjects affected / exposed	0 / 25 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Streptococcal bacteraemia			
subjects affected / exposed	0 / 25 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Gastroenteritis			
subjects affected / exposed	0 / 25 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Cellulitis			
subjects affected / exposed	1 / 25 (4.00%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Eczema infected			
subjects affected / exposed	1 / 25 (4.00%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Genital abscess			
subjects affected / exposed	0 / 25 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Skin infection			

subjects affected / exposed	1 / 25 (4.00%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Tracheobronchitis			
subjects affected / exposed	1 / 25 (4.00%)		
occurrences causally related to treatment / all	0 / 1		
deaths causally related to treatment / all	0 / 0		
Metabolism and nutrition disorders			
Hyponatraemia			
subjects affected / exposed	0 / 25 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		
Dehydration			
subjects affected / exposed	0 / 25 (0.00%)		
occurrences causally related to treatment / all	0 / 0		
deaths causally related to treatment / all	0 / 0		

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

Non-serious adverse events	Group 1 (Infant Series)	Group 2 (Infant Series)	Group 3 (Infant Series)
Total subjects affected by non-serious adverse events			
subjects affected / exposed	11 / 151 (7.28%)	4 / 51 (7.84%)	1 / 25 (4.00%)
Nervous system disorders			
Convulsion			
subjects affected / exposed	1 / 151 (0.66%)	0 / 51 (0.00%)	0 / 25 (0.00%)
occurrences (all)	2	0	0
Crying			
subjects affected / exposed	1 / 151 (0.66%)	0 / 51 (0.00%)	0 / 25 (0.00%)
occurrences (all)	1	0	0
General disorders and administration site conditions			
Pyrexia			
subjects affected / exposed	9 / 151 (5.96%)	4 / 51 (7.84%)	1 / 25 (4.00%)
occurrences (all)	10	4	1
Injection site erythema			

subjects affected / exposed	1 / 151 (0.66%)	0 / 51 (0.00%)	0 / 25 (0.00%)
occurrences (all)	1	0	0
Injection site swelling			
subjects affected / exposed	1 / 151 (0.66%)	0 / 51 (0.00%)	0 / 25 (0.00%)
occurrences (all)	1	0	0
Skin and subcutaneous tissue disorders			
Rash maculo-papular			
subjects affected / exposed	0 / 151 (0.00%)	0 / 51 (0.00%)	0 / 25 (0.00%)
occurrences (all)	0	0	0

Non-serious adverse events	Group 1 (Toddler Dose)	Group 2 (Toddler Dose)	Group 3 (Toddler Dose)
Total subjects affected by non-serious adverse events			
subjects affected / exposed	0 / 15 (0.00%)	0 / 23 (0.00%)	0 / 16 (0.00%)
Nervous system disorders			
Convulsion			
subjects affected / exposed	0 / 15 (0.00%)	0 / 23 (0.00%)	0 / 16 (0.00%)
occurrences (all)	0	0	0
Crying			
subjects affected / exposed	0 / 15 (0.00%)	0 / 23 (0.00%)	0 / 16 (0.00%)
occurrences (all)	0	0	0
General disorders and administration site conditions			
Pyrexia			
subjects affected / exposed	0 / 15 (0.00%)	0 / 23 (0.00%)	0 / 16 (0.00%)
occurrences (all)	0	0	0
Injection site erythema			
subjects affected / exposed	0 / 15 (0.00%)	0 / 23 (0.00%)	0 / 16 (0.00%)
occurrences (all)	0	0	0
Injection site swelling			
subjects affected / exposed	0 / 15 (0.00%)	0 / 23 (0.00%)	0 / 16 (0.00%)
occurrences (all)	0	0	0
Skin and subcutaneous tissue disorders			
Rash maculo-papular			
subjects affected / exposed	0 / 15 (0.00%)	0 / 23 (0.00%)	0 / 16 (0.00%)
occurrences (all)	0	0	0

Non-serious adverse events	Group 4 (2 Catch-Up Doses)	Group 5 (1 Catch-Up Dose)	Group 1 (Infant Series) Follow-up
Total subjects affected by non-serious adverse events			

subjects affected / exposed	1 / 66 (1.52%)	0 / 79 (0.00%)	0 / 151 (0.00%)
Nervous system disorders			
Convulsion			
subjects affected / exposed	0 / 66 (0.00%)	0 / 79 (0.00%)	0 / 151 (0.00%)
occurrences (all)	0	0	0
Crying			
subjects affected / exposed	0 / 66 (0.00%)	0 / 79 (0.00%)	0 / 151 (0.00%)
occurrences (all)	0	0	0
General disorders and administration site conditions			
Pyrexia			
subjects affected / exposed	0 / 66 (0.00%)	0 / 79 (0.00%)	0 / 151 (0.00%)
occurrences (all)	0	0	0
Injection site erythema			
subjects affected / exposed	0 / 66 (0.00%)	0 / 79 (0.00%)	0 / 151 (0.00%)
occurrences (all)	0	0	0
Injection site swelling			
subjects affected / exposed	0 / 66 (0.00%)	0 / 79 (0.00%)	0 / 151 (0.00%)
occurrences (all)	0	0	0
Skin and subcutaneous tissue disorders			
Rash maculo-papular			
subjects affected / exposed	1 / 66 (1.52%)	0 / 79 (0.00%)	0 / 151 (0.00%)
occurrences (all)	1	0	0

Non-serious adverse events	Group 2 (Infant Series) Follow-up	Group 3 (Infant Series) Follow-up	Group 4 (2 Catch-Up Doses) Follow-up
Total subjects affected by non-serious adverse events			
subjects affected / exposed	0 / 51 (0.00%)	0 / 25 (0.00%)	0 / 66 (0.00%)
Nervous system disorders			
Convulsion			
subjects affected / exposed	0 / 51 (0.00%)	0 / 25 (0.00%)	0 / 66 (0.00%)
occurrences (all)	0	0	0
Crying			
subjects affected / exposed	0 / 51 (0.00%)	0 / 25 (0.00%)	0 / 66 (0.00%)
occurrences (all)	0	0	0
General disorders and administration site conditions			
Pyrexia			
subjects affected / exposed	0 / 51 (0.00%)	0 / 25 (0.00%)	0 / 66 (0.00%)
occurrences (all)	0	0	0

Injection site erythema			
subjects affected / exposed	0 / 51 (0.00%)	0 / 25 (0.00%)	0 / 66 (0.00%)
occurrences (all)	0	0	0
Injection site swelling			
subjects affected / exposed	0 / 51 (0.00%)	0 / 25 (0.00%)	0 / 66 (0.00%)
occurrences (all)	0	0	0
Skin and subcutaneous tissue disorders			
Rash maculo-papular			
subjects affected / exposed	0 / 51 (0.00%)	0 / 25 (0.00%)	0 / 66 (0.00%)
occurrences (all)	0	0	0

Non-serious adverse events	Group 5 (1 Catch-Up Dose) Follow-up	Group 1 (After Infant Series)	Group 2 (After Infant Series)
Total subjects affected by non-serious adverse events			
subjects affected / exposed	0 / 79 (0.00%)	0 / 151 (0.00%)	0 / 51 (0.00%)
Nervous system disorders			
Convulsion			
subjects affected / exposed	0 / 79 (0.00%)	0 / 151 (0.00%)	0 / 51 (0.00%)
occurrences (all)	0	0	0
Crying			
subjects affected / exposed	0 / 79 (0.00%)	0 / 151 (0.00%)	0 / 51 (0.00%)
occurrences (all)	0	0	0
General disorders and administration site conditions			
Pyrexia			
subjects affected / exposed	0 / 79 (0.00%)	0 / 151 (0.00%)	0 / 51 (0.00%)
occurrences (all)	0	0	0
Injection site erythema			
subjects affected / exposed	0 / 79 (0.00%)	0 / 151 (0.00%)	0 / 51 (0.00%)
occurrences (all)	0	0	0
Injection site swelling			
subjects affected / exposed	0 / 79 (0.00%)	0 / 151 (0.00%)	0 / 51 (0.00%)
occurrences (all)	0	0	0
Skin and subcutaneous tissue disorders			
Rash maculo-papular			
subjects affected / exposed	0 / 79 (0.00%)	0 / 151 (0.00%)	0 / 51 (0.00%)
occurrences (all)	0	0	0

Non-serious adverse events	Group 3 (After Infant Series)		

Total subjects affected by non-serious adverse events subjects affected / exposed	0 / 25 (0.00%)		
Nervous system disorders Convulsion subjects affected / exposed occurrences (all) Crying subjects affected / exposed occurrences (all)	 0 / 25 (0.00%) 0 0 / 25 (0.00%) 0		
General disorders and administration site conditions Pyrexia subjects affected / exposed occurrences (all) Injection site erythema subjects affected / exposed occurrences (all) Injection site swelling subjects affected / exposed occurrences (all)	 0 / 25 (0.00%) 0 0 / 25 (0.00%) 0 0 / 25 (0.00%) 0		
Skin and subcutaneous tissue disorders Rash maculo-papular subjects affected / exposed occurrences (all)	 0 / 25 (0.00%) 0		

More information

Substantial protocol amendments (globally)

Were there any global substantial amendments to the protocol? Yes

Date	Amendment
29 August 2008	Follow up for safety assessment for 6 months after last vaccination was added; a local data safety monitoring board (DSMB) in accordance with the local Alaska Area IRB requirements was added and details on the ongoing safety review by the DSMB were to be provided in a separate charter.
15 October 2008	Timing of nasopharyngeal (NP) colonization swab collection was corrected to reflect annual timing. Study flow charts were updated for clarity regarding vaccine dosing intervals.

Notes:

Interruptions (globally)

Were there any global interruptions to the trial? No

Limitations and caveats

None reported