Clinical trial results:
Long-Term Safety Follow-Up of Thai Children Who Were Included in an Efficacy Study of a Tetravalent Dengue Vaccine

Summary

EudraCT number	2014-001737-88
Trial protocol	Outside EU/EEA
Global end of trial date	

Results information

Result version number	v1
This version publication date	19 May 2016
First version publication date	19 May 2016

Trial information

Trial identification

Sponsor protocol code	CYD57

Additional study identifiers

ISRCTN number	-
ClinicalTrials.gov id (NCT number)	NCT01983553
WHO universal trial number (UTN)	U1111-1127-7380

Notes:

Sponsors

Sponsor organisation name	Sanofi Pasteur SA
Sponsor organisation address	2, avenue Pont Pasteur, Lyon Cedex 07, France, F-69367
Public contact	Director, Clinical Development, Sanofi Pasteur Thailand, Sanofi Pasteur SA, 66 2264 8813, Danaya.chansinghakul@sanofipasteur.com
Scientific contact	Director, Clinical Development, Sanofi Pasteur Thailand, Sanofi Pasteur SA, 66 2264 8813, Danaya.chansinghakul@sanofipasteur.com

Notes:

Paediatric regulatory details

Is trial part of an agreed paediatric investigation plan (PIP)	Yes
EMA paediatric investigation plan number(s)	EMEA-001201-PIP01-11
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	Interim
Date of interim/final analysis	05 February 2015
Is this the analysis of the primary completion data?	Yes
Primary completion date	29 September 2014
Global end of trial reached?	No

Notes:

General information about the trial

Main objective of the trial:

The main objective of this study is to evaluate the continuous long-term follow-up of hospitalized dengue and safety of subjects who received CYD dengue vaccine in study CYD23 (EudraCT No. 2014-001710-25) including retrospective data collection (i.e., including the blood samples collected from March 05th, 2012 to September 9th, 2013) and prospective data collection from CYD57 study start (September 10th, 2013) until the end of the study, and the following activities will be performed.
1) To describe the incidence of virologically-confirmed hospitalized dengue cases.
2) To characterize hospitalized dengue cases.
3) To evaluate the occurrence of related and fatal SAEs.

Protection of trial subjects:

Subjects were vaccinated in a previous study, CYD23. No vaccination was administered as part of this study.

Background therapy:

In study CYD23, subjects age 4 to 11 were randomized to receive 3 injections of either CYD dengue vaccine or control, 1 each at 0, 6, and 12 months. Those that received at least one injection were eligible to be included in this follow-up study.

Evidence for comparator:

Not applicable

Actual start date of recruitment	10 September 2013
Long term follow-up planned	Yes
Long term follow-up rationale	Safety, Efficacy
Long term follow-up duration	4 Years
Independent data monitoring committee (IDMC) involvement?	Yes

Notes:

Population of trial subjects

Subjects enrolled per country

Country: Number of subjects enrolled	Thailand: 3203
Worldwide total number of subjects	3203
EEA total number of subjects	0

Notes:

Subjects enrolled per age group

In utero	0
Preterm newborn - gestational age < 37 wk	0
Newborns (0-27 days)	0

Infants and toddlers (28 days-23 months)	0
Children (2-11 years)	3203
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:

Study subjects were enrolled in the original study CYD23 from 10 September 2013 to 29 November 2013 at 1 clinic center in Thailand.

Pre-assignment

Screening details:

Of the 3997 subjects who were treated in the original study, CYD23 a total of 3203 subjects were included in the 4-year safety follow-up CYD57 study.

Period 1

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

Blinding implementation details:

Subjects and site staff remained blinded to the treatment allocation in CYD23 until the end of the study (CYD57) to avoid any bias in the safety follow-up (detection of dengue cases) until the second year of the follow-up study. In the event of an emergency (i.e., serious adverse event), the code of CYD57 could be broken by the Investigator as explained in the code-breaking procedures described in the Operating Guidelines.

Arms

Are arms mutually exclusive?	Yes
Arm title	CYD Dengue Vaccine Group

Arm description:

Subjects received 3 injections of CYD dengue vaccine, 1 injection each at 0, 6, and 12 months in study CYD23.

Arm type	Experimental
Investigational medicinal product name	CYD Dengue vaccine
Investigational medicinal product code	
Other name	
Pharmaceutical forms	Powder and solvent for suspension for injection
Routes of administration	Subcutaneous use

Dosage and administration details:

0.5 mL, subcutaneous, 1 injection each at 0, 6, and 12 months in the initial CYD23 study

Arm title	Control Group

Arm description:

Subjects received either Rabies vaccine (Verorab®) or placebo control as a first injection on Day 0, placebo for second and third injections at 6 and 12 months, respectively, in the original CYD23 study.

Arm type	Active comparator
Investigational medicinal product name	Rabies vaccine (Verorab®)
Investigational medicinal product code	
Other name	
Pharmaceutical forms	Powder and solvent for suspension for injection
Routes of administration	Subcutaneous use

Dosage and administration details:

0.5 mL, subcutaneous, 1 injection of either Verorab or placebo as a first injection on Day 0, placebo for second and third injections at 6 and 12 months, respectively, in the original CYD23 study

Investigational medicinal product name	Placebo (sodium chloride)
Investigational medicinal product code	
Other name	
Pharmaceutical forms	Suspension for injection
Routes of administration	Subcutaneous use

Dosage and administration details:

0.5 mL, subcutaneous, 1 injection of either Verorab or placebo as a first injection on Day 0, placebo for second and third injections at 6 and 12 months, respectively, in the original CYD23 study

Number of subjects in period 1	CYD Dengue Vaccine Group	Control Group
Started	2131	1072
Completed	2131	1072

Baseline characteristics

Reporting groups	
Reporting group title	CYD Dengue Vaccine Group
Reporting group description:	
Subjects received 3 injections of CYD dengue vaccine, 1 injection each at 0, 6, and 12 months in study CYD23.	
Reporting group title	Control Group
Reporting group description:	
Subjects received either Rabies vaccine (Verorab®) or placebo control as a first injection on Day 0, placebo for second and third injections at 6 and 12 months, respectively, in the original CYD23 study.	

Reporting group values	CYD Dengue Vaccine Group	Control Group	Total
Number of subjects	2131	1072	3203
Age categorical Units: Subjects			
In utero	0	0	0
Preterm newborn infants (gestational age < 37 wks)	0	0	0
Newborns (0-27 days)	0	0	0
Infants and toddlers (28 days-23 months)	0	0	0
Children (2-11 years)	2131	1072	3203
Adolescents (12-17 years)	0	0	0
Adults (18-64 years)	0	0	0
From 65-84 years	0	0	0
85 years and over	0	0	0
Age continuous Units: years			
arithmetic mean	8.11	8.15	
standard deviation	± 2.02	± 2.06	-
Gender categorical Units: Subjects			
Female	1118	579	1697
Male	1013	493	1506

End points

End points reporting groups

Reporting group title	CYD Dengue Vaccine Group
Reporting group description:	
Subjects received 3 injections of CYD dengue vaccine, 1 injection each at 0, 6, and 12 months in study CYD23.	
Reporting group title	Control Group
Reporting group description:	
Subjects received either Rabies vaccine (Verorab®) or placebo control as a first injection on Day 0, placebo for second and third injections at 6 and 12 months, respectively, in the original CYD23 study.	

Primary: Annual Incidence of Hospitalized Virologically-Confirmed Dengue Due to Any and Each Serotype Collected During the Year 1 and Year 2 Passive Surveillance Following Vaccination with Either CYD Dengue Vaccine or a Placebo in a Previous Study

End point title	Annual Incidence of Hospitalized Virologically-Confirmed Dengue Due to Any and Each Serotype Collected During the Year 1 and Year 2 Passive Surveillance Following Vaccination with Either CYD Dengue Vaccine or a Placebo in a Previous Study
End point description:	
Annual incidence rate was defined as the cases among the number of subjects present at the beginning of each period (M) multiplied by 100 converted in annual rate (Annual incidence rate = Cases among M*100).	
End point type	Primary
End point timeframe:	
Year 1 and Year 2 post-vaccination	

End point values	CYD Dengue Vaccine Group	Control Group		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	2131	1072		
Units: Annual incidence rate				
number (confidence interval 95%)				
Year 1: Any of the 4 Serotypes	1.1 (0.7 to 1.7)	1.1 (0.6 to 2)		
Year 1: Serotype 1	0.3 (0.1 to 0.6)	0.5 (0.2 to 1.2)		
Year 1: Serotype 2	0.9 (0.5 to 1.4)	0.4 (0.1 to 1)		
Year 1: Serotype 3	0.1 (0 to 0.3)	0.1 (0 to 0.6)		
Year 1: Serotype 4	0 (0 to 0.2)	0 (0 to 0.4)		
Year 1: Unserotyped	0 (0 to 0.2)	0.2 (0 to 0.7)		
Year 2: Any of the 4 Serotypes	0.8 (0.4 to 1.2)	1.6 (0.9 to 2.5)		
Year 2: Serotype 1	0.2 (0.1 to 0.5)	0.3 (0.1 to 0.8)		
Year 2: Serotype 2	0.2 (0.1 to 0.5)	0.6 (0.2 to 1.2)		
Year 2: Serotype 3	0.3 (0.1 to 0.6)	0.3 (0.1 to 0.8)		
Year 2: Serotype 4	0.1 (0 to 0.3)	0.4 (0.1 to 1)		
Year 2: Unserotyped	0 (0 to 0.2)	0.1 (0 to 0.5)		

Statistical analyses

Statistical analysis title	Relative risk; Year 1; Any of the 4 Serotypes

Statistical analysis description:

Relative risk analysis of the annual incidence rate at Year 1 Any of the 4 Serotypes between the CYD Dengue vaccine group and the Control group.

Comparison groups	CYD Dengue Vaccine Group v Control Group
Number of subjects included in analysis	3203
Analysis specification	Pre-specified
Analysis type	other[1]
Parameter estimate	Relative Risk
Point estimate	1.006
Confidence interval	
level	95 %
sides	2-sided
lower limit	0.47
upper limit	2.3

Notes:

[1] - The 95% confidence interval of the relative risk is calculated using the Exact method described by Breslow and Day.

Statistical analysis title	Relative risk; Year 1; Serotype 1

Statistical analysis description:

Relative risk analysis of the annual incidence rate at Year 1 Serotype 1 between the CYD Dengue vaccine group and the Control group.

Comparison groups	CYD Dengue Vaccine Group v Control Group
Number of subjects included in analysis	3203
Analysis specification	Pre-specified
Analysis type	other[2]
Parameter estimate	Relative Risk
Point estimate	0.503
Confidence interval	
level	95 %
sides	2-sided
lower limit	0.12
upper limit	2.19

Notes:

[2] - The 95% confidence interval of the relative risk is calculated using the Exact method described by Breslow and Day.

Statistical analysis title	Relative risk; Year 1; Serotype 2

Statistical analysis description:

Relative risk analysis of the annual incidence rate at Year 1 Serotype 2 between the CYD Dengue vaccine group and the Control group.

Comparison groups	CYD Dengue Vaccine Group v Control Group

Number of subjects included in analysis	3203
Analysis specification	Pre-specified
Analysis type	other
Parameter estimate	Relative Risk
Point estimate	2.138
Confidence interval	
level	95 %
sides	2-sided
lower limit	0.7
upper limit	8.73

Statistical analysis title	Relative risk; Year 1; Serotype 3

Statistical analysis description:

Relative risk analysis of the annual incidence rate at Year 1 Serotype 3 between the CYD Dengue vaccine group and the Control group.

Comparison groups	CYD Dengue Vaccine Group v Control Group
Number of subjects included in analysis	3203
Analysis specification	Pre-specified
Analysis type	other[3]
Parameter estimate	Relative Risk
Point estimate	0.503
Confidence interval	
level	95 %
sides	2-sided
lower limit	0.01
upper limit	39.49

Notes:

[3] - The 95% confidence interval of the relative risk is calculated using the Exact method described by Breslow and Day.

Statistical analysis title	Relative risk; Year 1; Serotype 4

Statistical analysis description:

Relative risk analysis of the annual incidence rate at Year 1 Serotype 4 between the CYD Dengue vaccine group and the Control group.

Comparison groups	CYD Dengue Vaccine Group v Control Group
Number of subjects included in analysis	3203
Analysis specification	Pre-specified
Analysis type	other[4]
Parameter estimate	Relative Risk
Point estimate	0
Confidence interval	
level	95 %
sides	2-sided
lower limit	0
upper limit	0

Notes:

[4] - The 95% confidence interval of the relative risk is calculated using the Exact method described by Breslow and Day.

Statistical analysis title	Relative risk; Year 1; Unserotyped

Statistical analysis description:

Relative risk analysis of the annual incidence rate at Year 1 Unserotyped between the CYD Dengue vaccine group and the Control group.

Comparison groups	CYD Dengue Vaccine Group v Control Group
Number of subjects included in analysis	3203
Analysis specification	Pre-specified
Analysis type	other[5]
Parameter estimate	Relative Risk
Point estimate	0
Confidence interval	
level	95 %
sides	2-sided
lower limit	0
upper limit	2.68

Notes:

[5] - The 95% confidence interval of the relative risk is calculated using the Exact method described by Breslow and Day.

Statistical analysis title	Relative risk; Year 2; Any of the 4 Serotypes

Statistical analysis description:

Relative risk analysis of the annual incidence rate at Year 2 Any of the 4 Serotypes between the CYD Dengue vaccine group and the Control group.

Comparison groups	CYD Dengue Vaccine Group v Control Group
Number of subjects included in analysis	3203
Analysis specification	Pre-specified
Analysis type	other
Parameter estimate	Relative risk of annual incidence rate
Point estimate	0.473
Confidence interval	
level	95 %
sides	2-sided
lower limit	0.22
upper limit	1

Statistical analysis title	Relative risk; Year 2; Serotype 1

Statistical analysis description:

Relative risk analysis of the annual incidence rate at Year 2 Serotype 1 between the CYD Dengue vaccine group and the Control group.

Comparison groups	CYD Dengue Vaccine Group v Control Group
Number of subjects included in analysis	3203
Analysis specification	Pre-specified
Analysis type	other[6]
Parameter estimate	Relative Risk
Point estimate	0.671
Confidence interval	
level	95 %
sides	2-sided
lower limit	0.11
upper limit	4.58

Notes:

[6] - The 95% confidence interval of the relative risk is calculated using the Exact method described by Breslow and Day.

Statistical analysis title	Relative risk; Year 2; Serotype 2

Statistical analysis description:

Relative risk analysis of the annual incidence rate at Year 2 Serotype 2 between the CYD Dengue vaccine group and the Control group.

Comparison groups	CYD Dengue Vaccine Group v Control Group
Number of subjects included in analysis	3203
Analysis specification	Pre-specified
Analysis type	other[7]
Parameter estimate	Relative Risk
Point estimate	0.335
Confidence interval	
level	95 %
sides	2-sided
lower limit	0.07
upper limit	1.41

Notes:

[7] - The 95% confidence interval of the relative risk is calculated using the Exact method described by Breslow and Day.

Statistical analysis title	Relative risk; Year 2; Serotype 3

Statistical analysis description:

Relative risk analysis of the annual incidence rate at Year 1 Serotype 3 between the CYD Dengue vaccine group and the Control group.

Comparison groups	CYD Dengue Vaccine Group v Control Group
Number of subjects included in analysis	3203
Analysis specification	Pre-specified
Analysis type	other[8]
Parameter estimate	Relative Risk
Point estimate	1.006
Confidence interval	
level	95 %
sides	2-sided
lower limit	0.21
upper limit	6.22

Notes:

[8] - The 95% confidence interval of the relative risk is calculated using the Exact method described by Breslow and Day.

Statistical analysis title	Relative risk; Year 2; Serotype 4

Statistical analysis description:

Relative risk analysis of the annual incidence rate at Year 1 Serotype 4 between the CYD Dengue vaccine group and the Control group.

Comparison groups	CYD Dengue Vaccine Group v Control Group
Number of subjects included in analysis	3203
Analysis specification	Pre-specified
Analysis type	other[9]
Parameter estimate	Relative Risk
Point estimate	0.252

Confidence interval	
level	95 %
sides	2-sided
lower limit	0.02
upper limit	1.76

Notes:

[9] - The 95% confidence interval of the relative risk is calculated using the Exact method described by Breslow and Day.

Statistical analysis title	Relative risk; Year 2; Unserotyped

Statistical analysis description:

Relative risk analysis of the annual incidence rate at Year 1 Unserotyped between the CYD Dengue vaccine group and the Control group.

Comparison groups	CYD Dengue Vaccine Group v Control Group
Number of subjects included in analysis	3203
Analysis specification	Pre-specified
Analysis type	other[10]
Parameter estimate	Relative Risk
Point estimate	0
Confidence interval	
level	95 %
sides	2-sided
lower limit	0
upper limit	19.62

Notes:

[10] - The 95% confidence interval of the relative risk is calculated using the Exact method described by Breslow and Day.

Primary: Number of Cases and Episodes of Hospitalized Virologically-Confirmed Dengue Due to Any and Each Serotype Collected During Year I and Year 2 Passive Surveillance Following Vaccination with Either CYD Dengue Vaccine or a Placebo in a Previous Study

End point title	Number of Cases and Episodes of Hospitalized Virologically-Confirmed Dengue Due to Any and Each Serotype Collected During Year I and Year 2 Passive Surveillance Following Vaccination with Either CYD Dengue Vaccine or a Placebo in a Previous Study[11]

End point description:

Cases were defined as the number of subjects with at least one hospitalized virologically-confirmed dengue episode. Episodes were defined as the number of hospitalized virologically-confirmed dengue episodes.

End point type	Primary

End point timeframe:

Year 1 and Year 2 post-vaccination

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: Descriptive analyses were performed based on the study groups and the study vaccine administered for this outcome.

End point values	CYD Dengue Vaccine Group	Control Group		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	2131	1072		
Units: Number of cases and episodes of dengue				
number (not applicable)				
Year 1 Cases; Any of the 4 Serotypes	22	11		
Year 1 Episodes; Any of the 4 Serotypes	22	11		
Year 1 Cases; Serotype 1	5	5		
Year 1 Episodes; Serotype 1	5	5		
Year 1 Cases; Serotype 2	17	4		
Year 1 Episodes; Serotype 2	17	4		
Year 1 Cases; Serotype 3	1	1		
Year 1 Episodes; Serotype 3	1	1		
Year 1 Cases; Serotype 4	0	0		
Year 1 Episodes; Serotype 4	0	0		
Year 1 Cases; Unserotyped	0	2		
Year 1 Episodes; Unserotyped	0	2		
Year 2 Cases; Any of the 4 Serotypes	16	17		
Year 2 Episodes; Any of the 4 Serotypes	16	17		
Year 2 Cases; Serotype 1	4	3		
Year 2 Episodes; Serotype 1	4	3		
Year 2 Cases; Serotype 2	4	6		
Year 2 Episodes; Serotype 2	4	6		
Year 2 Cases; Serotype 3	6	3		
Year 2 Episodes; Serotype 3	6	3		
Year 2 Cases; Serotype 4	2	4		
Year 2 Episodes; Serotype 4	2	4		
Year 2 Cases; Unserotyped	0	1		
Year 2 Episodes; Unserotyped	0	1		

Statistical analyses

No statistical analyses for this end point

Primary: Annual Incidence of Hospitalized Virologically-Confirmed Dengue Due to Any and Each Serotype Collected By Age Group During the Year I and Year 2 Passive Surveillance Following Vaccination with Either CYD Dengue Vaccine or a Placebo in a Previous Study

End point title	Annual Incidence of Hospitalized Virologically-Confirmed Dengue Due to Any and Each Serotype Collected By Age Group During the Year I and Year 2 Passive Surveillance Following Vaccination with Either CYD Dengue Vaccine or a Placebo in a Previous Study

End point description:

Annual incidence rate was defined as the cases among the number of subjects present at the beginning of each period (M) multiplied by 100 converted in annual rate (Annual incidence rate = Cases among M*100).

End point type	Primary

End point timeframe:

Year 1 and Year 2 post-vaccination

End point values	CYD Dengue Vaccine Group	Control Group		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	2131	1072		
Units: Annual incidence rate				
number (confidence interval 95%)				
Year 1, 4 to 5 Years: Any of the 4 Serotypes	1.4 (0.5 to 3.2)	0.6 (0 to 3.1)		
Year 1, 6 to 11 Years: Any of the 4 Serotypes	1.1 (0.6 to 1.7)	1.2 (0.6 to 2.3)		
Year 1, 4 to 5 Years: Serotype 1	0.6 (0.1 to 2)	0 (0 to 2.1)		
Year 1, 6 to 11 Years: Serotype 1	0.2 (0 to 0.5)	0.6 (0.2 to 1.4)		
Year 1, 4 to 5 Years: Serotype 2	1.1 (0.3 to 2.8)	0 (0 to 2.1)		
Year 1, 6 to 11 Years: Serotype 2	0.8 (0.4 to 1.4)	0.5 (0.1 to 1.3)		
Year 1, 4 to 5 Years: Serotype 3	0 (0 to 1)	0 (0 to 2.1)		
Year 1, 6 to 11 Years: Serotype 3	0.1 (0 to 0.3)	0.1 (0 to 0.7)		
Year 1, 4 to 5 Years: Serotype 4	0 (0 to 1)	0 (0 to 2.1)		
Year 1, 6 to 11 Years: Serotype 4	0 (0 to 0.2)	0 (0 to 0.5)		
Year 1, 4 to 5 Years: Unserotyped	0 (0 to 1)	0.6 (0 to 3.1)		
Year 1, 6 to 11 Years: Unserotyped	0 (0 to 0.2)	0.1 (0 to 0.7)		
Year 2, 4 to 5 Years: Any of the 4 Serotypes	1.3 (0.4 to 2.9)	1.6 (0.3 to 4.5)		
Year 2, 6 to 11 Years: Any of the 4 Serotypes	0.6 (0.3 to 1.1)	1.6 (0.9 to 2.7)		
Year 2, 4 to 5 Years: Serotype 1	1 (0.3 to 2.6)	0 (0 to 1.9)		
Year 2, 6 to 11 Years: Serotype 1	0 (0 to 0.2)	0.3 (0.1 to 1)		
Year 2, 4 to 5 Years: Serotype 2	0 (0 to 0.9)	0.5 (0 to 2.9)		
Year 2, 6 to 11 Years: Serotype 2	0.2 (0.1 to 0.6)	0.6 (0.2 to 1.3)		
Year 2, 4 to 5 Years: Serotype 3	0.3 (0 to 1.4)	0.5 (0 to 2.9)		
Year 2, 6 to 11 Years: Serotype 3	0.3 (0.1 to 0.7)	0.2 (0 to 0.8)		
Year 2, 4 to 5 Years: Serotype 4	0 (0 to 0.9)	0 (0 to 1.9)		
Year 2, 6 to 11 Years: Serotype 4	0.1 (0 to 0.4)	0.5 (0.1 to 1.2)		
Year 2, 4 to 5 Years: Unserotyped	0 (0 to 0.9)	0.5 (0 to 2.9)		
Year 2, 6 to 11 Years: Unserotyped	0 (0 to 0.2)	0 (0 to 0.4)		

Statistical analyses

Statistical analysis title	Relative Risk; Yr 1; 4 to 5 years; Any Serotype

Statistical analysis description:

Relative risk (RR) analysis of the annual incidence rate at Year 1 Any of the 4 Serotypes (4 to 5 years) between the CYD Dengue vaccine group and the Control group.

Comparison groups	CYD Dengue Vaccine Group v Control Group

Number of subjects included in analysis	3203
Analysis specification	Pre-specified
Analysis type	other[12]
Parameter estimate	Relative Risk
Point estimate	2.443
Confidence interval	
level	95 %
sides	2-sided
lower limit	0.27
upper limit	115.54

Notes:

[12] - The 95% confidence interval of the relative risk is calculated using the Exact method described by Breslow and Day.

Statistical analysis title	Relative Risk; Year 1; 6 to 11 years; Any Serotype

Statistical analysis description:

Relative risk analysis of the annual incidence rate at Year 1 Any of the 4 Serotypes (6 to 11 years) between the CYD Dengue vaccine group and the Control group.

Comparison groups	CYD Dengue Vaccine Group v Control Group
Number of subjects included in analysis	3203
Analysis specification	Pre-specified
Analysis type	other[13]
Parameter estimate	Relative Risk
Point estimate	0.861
Confidence interval	
level	95 %
sides	2-sided
lower limit	0.37
upper limit	2.1

Notes:

[13] - The 95% confidence interval of the relative risk is calculated using the Exact method described by Breslow and Day.

Statistical analysis title	Relative Risk; Year 1; 4 to 5 years; Serotype 1

Statistical analysis description:

Relative risk (RR) analysis of the annual incidence rate at Year 1 Serotype 1 (4 to 5 years) between the CYD Dengue vaccine group and the Control group.

Comparison groups	CYD Dengue Vaccine Group v Control Group
Number of subjects included in analysis	3203
Analysis specification	Pre-specified
Analysis type	other[14]
Parameter estimate	Relative Risk
Point estimate	0
Confidence interval	
level	95 %
sides	2-sided
lower limit	0
upper limit	0

Notes:

[14] - The 95% confidence interval of the relative risk is calculated using the Exact method described by Breslow and Day.

Statistical analysis title	Relative Risk; Year 1; 6 to 11 years; Serotype 1

Statistical analysis description:

Relative risk (RR) analysis of the annual incidence rate at Year 1 Serotype 1 (6 to 11 years) between the CYD Dengue vaccine group and the Control group.

Comparison groups	CYD Dengue Vaccine Group v Control Group
Number of subjects included in analysis	3203
Analysis specification	Pre-specified
Analysis type	other[15]
Parameter estimate	Relative Risk
Point estimate	0.304
Confidence interval	
level	95 %
sides	2-sided
lower limit	0.05
upper limit	1.56

Notes:

[15] - The 95% confidence interval of the relative risk is calculated using the Exact method described by Breslow and Day.

Statistical analysis title	Relative Risk; Year 1; 4 to 5 years; Serotype 2

Statistical analysis description:

Relative risk (RR) analysis of the annual incidence rate at Year 1 Serotype 2 (4 to 5 years) between the CYD Dengue vaccine group and the Control group.

Comparison groups	CYD Dengue Vaccine Group v Control Group
Number of subjects included in analysis	3203
Analysis specification	Pre-specified
Analysis type	other[16]
Parameter estimate	Relative Risk
Point estimate	0
Confidence interval	
level	95 %
sides	2-sided
lower limit	0
upper limit	0

Notes:

[16] - The 95% confidence interval of the relative risk is calculated using the Exact method described by Breslow and Day.

Statistical analysis title	Relative Risk; Year 1; 6 to 11 years; Serotype 2

Statistical analysis description:

Relative risk analysis of the annual incidence rate at Year 1 Serotype 2 (6 to 11 years) between the CYD Dengue vaccine group and the Control group.

Comparison groups	CYD Dengue Vaccine Group v Control Group
Number of subjects included in analysis	3203
Analysis specification	Pre-specified
Analysis type	other[17]
Parameter estimate	Relative Risk
Point estimate	1.646
Confidence interval	
level	95 %
sides	2-sided
lower limit	0.51
upper limit	6.93

Notes:

[17] - The 95% confidence interval of the relative risk is calculated using the Exact method described by Breslow and Day.

Statistical analysis title	Relative Risk; Year 1; 4 to 5 years; Serotype 3

Statistical analysis description:

Relative risk analysis of the annual incidence rate at Year 1 Serotype 3 (4 to 5 year) between the CYD Dengue vaccine group and the Control group.

Comparison groups	CYD Dengue Vaccine Group v Control Group
Number of subjects included in analysis	3203
Analysis specification	Pre-specified
Analysis type	other[18]
Parameter estimate	Relative Risk
Point estimate	0
Confidence interval	
level	95 %
sides	2-sided
lower limit	0
upper limit	0

Notes:

[18] - The 95% confidence interval of the relative risk is calculated using the Exact method described by Breslow and Day.

Statistical analysis title	Relative Risk; Year 1; 6 to 11 years; Serotype 3

Statistical analysis description:

Relative risk (RR) analysis of the annual incidence rate at Year 1 Serotype 3 (6 to 11 years) between the CYD Dengue vaccine group and the Control group.

Comparison groups	CYD Dengue Vaccine Group v Control Group
Number of subjects included in analysis	3203
Analysis specification	Pre-specified
Analysis type	other[19]
Parameter estimate	Relative Risk
Point estimate	0.506
Confidence interval	
level	95 %
sides	2-sided
lower limit	0.01
upper limit	39.75

Notes:

[19] - The 95% confidence interval of the relative risk is calculated using the Exact method described by Breslow and Day.

Statistical analysis title	Relative Risk; Year 1; 4 to 5 years; Serotype 4

Statistical analysis description:

Relative risk analysis of the annual incidence rate at Year 1 Serotype 4 (4 to 5 years) between the CYD Dengue vaccine group and the Control group.

Comparison groups	CYD Dengue Vaccine Group v Control Group
Number of subjects included in analysis	3203
Analysis specification	Pre-specified
Analysis type	other[20]
Parameter estimate	Relative Risk
Point estimate	0

Confidence interval	
level	95 %
sides	2-sided
lower limit	0
upper limit	0

Notes:

[20] - The 95% confidence interval of the relative risk is calculated using the Exact method described by Breslow and Day.

Statistical analysis title	Relative Risk; Year 1; 6 to 11 years; Serotype 4

Statistical analysis description:

Relative risk analysis of the annual incidence rate at Year 1 Serotype 4 (6 to 11 years) between the CYD Dengue vaccine group and the Control group.

Comparison groups	CYD Dengue Vaccine Group v Control Group
Number of subjects included in analysis	3203
Analysis specification	Pre-specified
Analysis type	other[21]
Parameter estimate	Relative Risk
Point estimate	0
Confidence interval	
level	95 %
sides	2-sided
lower limit	0
upper limit	0

Notes:

[21] - The 95% confidence interval of the relative risk is calculated using the Exact method described by Breslow and Day.

Statistical analysis title	Relative Risk; Year 1; 4 to 5 years; Unserotyped

Statistical analysis description:

Relative risk analysis of the annual incidence rate at Year 1 Unserotyped (4 to 5 years) between the CYD Dengue vaccine group and the Control group.

Comparison groups	CYD Dengue Vaccine Group v Control Group
Number of subjects included in analysis	3203
Analysis specification	Pre-specified
Analysis type	other[22]
Parameter estimate	Relative Risk
Point estimate	0
Confidence interval	
level	95 %
sides	2-sided
lower limit	0
upper limit	19.05

Notes:

[22] - The 95% confidence interval of the relative risk is calculated using the Exact method described by Breslow and Day.

Statistical analysis title	Relative Risk; Year 1; 6 to 11 years; Unserotyped

Statistical analysis description:

Relative risk (RR) analysis of the annual incidence rate at Year 1 Unserotyped (6 to 11 years) between the CYD Dengue vaccine group and the Control group.

Comparison groups	CYD Dengue Vaccine Group v Control Group

Number of subjects included in analysis	3203
Analysis specification	Pre-specified
Analysis type	other[23]
Parameter estimate	Relative Risk
Point estimate	0
Confidence interval	
level	95 %
sides	2-sided
lower limit	0
upper limit	19.75

Notes:

[23] - The 95% confidence interval of the relative risk is calculated using the Exact method described by Breslow and Day.

Statistical analysis title	Relative Risk; Year 2; 4 to 5 years; Any Serotype

Statistical analysis description:

Relative risk (RR) analysis of the annual incidence rate at Year 2 Any of the 4 Serotypes (4 to 5 years) between the CYD Dengue vaccine group and the Control group.

Comparison groups	CYD Dengue Vaccine Group v Control Group
Number of subjects included in analysis	3203
Analysis specification	Pre-specified
Analysis type	other[24]
Parameter estimate	Relative Risk
Point estimate	0.814
Confidence interval	
level	95 %
sides	2-sided
lower limit	0.16
upper limit	5.24

Notes:

[24] - The 95% confidence interval of the relative risk is calculated using the Exact method described by Breslow and Day.

Statistical analysis title	Relative Risk; Year 2; 6 to 11 years; Any Serotype

Statistical analysis description:

Relative risk (RR) analysis of the annual incidence rate at Year 2 Any of the 4 Serotypes (6 to 11 years) between the CYD Dengue vaccine group and the Control group.

Comparison groups	CYD Dengue Vaccine Group v Control Group
Number of subjects included in analysis	3203
Analysis specification	Pre-specified
Analysis type	other[25]
Parameter estimate	Relative Risk
Point estimate	0.398
Confidence interval	
level	95 %
sides	2-sided
lower limit	0.16
upper limit	0.94

Notes:

[25] - The 95% confidence interval of the relative risk is calculated using the Exact method described by Breslow and Day.

Statistical analysis title	Relative Risk; Year 2; 4 to 5 years; Serotype 1

Statistical analysis description:

Relative risk (RR) analysis of the annual incidence rate at Year 2 Serotype 1 (4 to 5 years) between the CYD Dengue vaccine group and the Control group.

Comparison groups	CYD Dengue Vaccine Group v Control Group
Number of subjects included in analysis	3203
Analysis specification	Pre-specified
Analysis type	other[26]
Parameter estimate	Relative Risk
Point estimate	0
Confidence interval	
level	95 %
sides	2-sided
lower limit	0
upper limit	0

Notes:

[26] - The 95% confidence interval of the relative risk is calculated using the Exact method described by Breslow and Day.

Statistical analysis title	Relative Risk; Year 2; 6 to 11 years; Serotype 1

Statistical analysis description:

Relative risk analysis of the annual incidence rate at Year 2 Serotype 1 (6 to 11 years) between the CYD Dengue vaccine group and the Control group.

Comparison groups	CYD Dengue Vaccine Group v Control Group
Number of subjects included in analysis	3203
Analysis specification	Pre-specified
Analysis type	other[27]
Parameter estimate	Relative Risk
Point estimate	0
Confidence interval	
level	95 %
sides	2-sided
lower limit	0
upper limit	1.23

Notes:

[27] - The 95% confidence interval of the relative risk is calculated using the Exact method described by Breslow and Day.

Statistical analysis title	Relative Risk; Year 2; 4 to 5 years; Serotype 2

Statistical analysis description:

Relative risk analysis of the annual incidence rate at Year 2 Serotype 2 (4 to 5 years) between the CYD Dengue vaccine group and the Control group.

Comparison groups	CYD Dengue Vaccine Group v Control Group
Number of subjects included in analysis	3203
Analysis specification	Pre-specified
Analysis type	other[28]
Parameter estimate	Relative Risk
Point estimate	0
Confidence interval	
level	95 %
sides	2-sided
lower limit	0
upper limit	19.05

Notes:

[28] - The 95% confidence interval of the relative risk is calculated using the Exact method described by Breslow and Day.

Statistical analysis title	Relative Risk; Year 2; 6 to 11 years; Serotype 2

Statistical analysis description:

Relative risk (RR) analysis of the annual incidence rate at Year 2 Serotype 2 (6 to 11 years) between the CYD Dengue vaccine group and the Control group.

Comparison groups	CYD Dengue Vaccine Group v Control Group
Number of subjects included in analysis	3203
Analysis specification	Pre-specified
Analysis type	other[29]
Parameter estimate	Relative Risk
Point estimate	0.405
Confidence interval	
level	95 %
sides	2-sided
lower limit	0.08
upper limit	1.88

Notes:

[29] - The 95% confidence interval of the relative risk is calculated using the Exact method described by Breslow and Day.

Statistical analysis title	Relative Risk; Year 2; 4 to 5 years; Serotype 3

Statistical analysis description:

Relative risk (RR) analysis of the annual incidence rate at Year 2 Serotype 3 (4 to 5 years) between the CYD Dengue vaccine group and the Control group.

Comparison groups	CYD Dengue Vaccine Group v Control Group
Number of subjects included in analysis	3203
Analysis specification	Pre-specified
Analysis type	other[30]
Parameter estimate	Relative Risk
Point estimate	0.489
Confidence interval	
level	95 %
sides	2-sided
lower limit	0.01
upper limit	38.35

Notes:

[30] - The 95% confidence interval of the relative risk is calculated using the Exact method described by Breslow and Day.

Statistical analysis title	Relative Risk; Year 2; 6 to 11 years; Serotype 3

Statistical analysis description:

Relative risk analysis of the annual incidence rate at Year 2 Serotype 3 (6 to 11 years) between the CYD Dengue vaccine group and the Control group.

Comparison groups	Control Group v CYD Dengue Vaccine Group
Number of subjects included in analysis	3203
Analysis specification	Pre-specified
Analysis type	other[31]
Parameter estimate	Relative Risk
Point estimate	1.266

Confidence interval	
level	95 %
sides	2-sided
lower limit	0.21
upper limit	13.29

Notes:

[31] - The 95% confidence interval of the relative risk is calculated using the Exact method described by Breslow and Day.

Statistical analysis title	Relative Risk; Year 2; 4 to 5 years; Serotype 4

Statistical analysis description:

Relative risk (RR) analysis of the annual incidence rate at Year 2 Serotype 4 (4 to 5 years) between the CYD Dengue vaccine group and the Control group.

Comparison groups	CYD Dengue Vaccine Group v Control Group
Number of subjects included in analysis	3203
Analysis specification	Pre-specified
Analysis type	other[32]
Parameter estimate	Relative Risk
Point estimate	0
Confidence interval	
level	95 %
sides	2-sided
lower limit	0
upper limit	0

Notes:

[32] - The 95% confidence interval of the relative risk is calculated using the Exact method described by Breslow and Day.

Statistical analysis title	Relative Risk; Year 2; 6 to 11 years; Serotype 4

Statistical analysis description:

Relative risk analysis of the annual incidence rate at Year 2 Serotype 4 (6 to 11 years) between the CYD Dengue vaccine group and the Control group.

Comparison groups	CYD Dengue Vaccine Group v Control Group
Number of subjects included in analysis	3203
Analysis specification	Pre-specified
Analysis type	other[33]
Parameter estimate	Relative Risk
Point estimate	0.253
Confidence interval	
level	95 %
sides	2-sided
lower limit	0.02
upper limit	1.77

Notes:

[33] - The 95% confidence interval of the relative risk is calculated using the Exact method described by Breslow and Day.

Statistical analysis title	Relative Risk; Year 2; 4 to 5 years; Unserotyped

Statistical analysis description:

Relative risk (RR) analysis of the annual incidence rate at Year 2 Unserotyped (4 to 5 years) between the CYD Dengue vaccine group and the Control group.

Comparison groups	CYD Dengue Vaccine Group v Control Group

Number of subjects included in analysis	3203
Analysis specification	Pre-specified
Analysis type	other[34]
Parameter estimate	Relative Risk
Point estimate	0
Confidence interval	
level	95 %
sides	2-sided
lower limit	0
upper limit	19.05

Notes:

[34] - The 95% confidence interval of the relative risk is calculated using the Exact method described by Breslow and Day.

Statistical analysis title	Relative Risk; Year 2; 6 to 11 years; Unserotyped

Statistical analysis description:

Relative risk analysis of the annual incidence rate at Year 2 Unserotyped (6 to 11 years) between the CYD Dengue vaccine group and the Control group.

Comparison groups	CYD Dengue Vaccine Group v Control Group
Number of subjects included in analysis	3203
Analysis specification	Pre-specified
Analysis type	other[35]
Parameter estimate	Relative Risk
Point estimate	0
Confidence interval	
level	95 %
sides	2-sided
lower limit	0
upper limit	0

Notes:

[35] - The 95% confidence interval of the relative risk is calculated using the Exact method described by Breslow and Day.

Primary: Number of Cases and Episodes of Hospitalized Virologically-Confirmed Dengue Due to Any and Each Serotype Collected By Age Group During Year I and Year 2 Passive Surveillance After Vaccination with Either CYD Dengue Vaccine or a Placebo in a Previous Study

End point title	Number of Cases and Episodes of Hospitalized Virologically-Confirmed Dengue Due to Any and Each Serotype Collected By Age Group During Year I and Year 2 Passive Surveillance After Vaccination with Either CYD Dengue Vaccine or a Placebo in a Previous Study[36]

End point description:

Cases were defined as the number of subjects with at least one hospitalized virologically-confirmed dengue episode. Episodes were defined as the number of hospitalized virologically-confirmed dengue episodes.

End point type	Primary

End point timeframe:

Year 1 and Year 2 post-vaccination

Notes:

[36] - 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: Descriptive analyses were performed based on the study groups and the study vaccine administered for this outcome.

End point values	CYD Dengue Vaccine Group	Control Group		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	2131	1072		
Units: Number of cases and episodes of dengue				
number (not applicable)				
Year 1, 4 to 5 years cases: Any of the 4 Serotypes	5	1		
Year 1, 4 to 5 yrs episodes:Any of the 4 Serotypes	5	1		
Year 1, 6 to 11 yrs cases: Any of the 4 Serotypes	17	10		
Yr 1, 6 to 11 yrs episodes: Any of the 4 Serotypes	17	10		
Year 1, 4 to 5 years cases: Serotype 1	2	0		
Year 1, 4 to 5 years episodes: Serotype 1	2	0		
Year 1, 6 to 11 years cases: Serotype 1	3	5		
Year 1, 6 to 11 years episodes: Serotype 1	3	5		
Year 1, 4 to 5 years cases: Serotype 2	4	0		
Year 1, 4 to 5 years episodes: Serotype 2	4	0		
Year 1, 6 to 11 years cases: Serotype 2	13	4		
Year 1, 6 to 11 years episodes: Serotype 2	13	4		
Year 1, 4 to 5 years cases: Serotype 3	0	0		
Year 1, 4 to 5 years episodes: Serotype 3	0	0		
Year 1, 6 to 11 years cases: Serotype 3	1	1		
Year 1, 6 to 11 years episodes: Serotype 3	1	1		
Year 1, 4 to 5 years cases: Serotype 4	0	0		
Year 1, 4 to 5 years episodes: Serotype 4	0	0		
Year 1, 6 to 11 years cases: Serotype 4	0	0		
Year 1, 6 to 11 years episodes: Serotype 4	0	0		
Year 1, 4 to 5 years cases: Unserotyped	0	1		
Year 1, 4 to 5 years episodes: Unserotyped	0	1		
Year 1, 6 to 11 years cases: Unserotyped	0	1		
Year 1, 6 to 11 years epsiodes: Unserotyped	0	1		
Year 2, 4 to 5 years cases: Any of the 4 Serotypes	5	3		
Year 2, 4 to 5 yrs episodes:Any of the 4 Serotypes	5	3		
Year 2, 6 to 11 yrs cases: Any of the 4 Serotypes	11	14		
Yr 2, 6 to 11 yrs episodes: Any of the 4 Serotypes	11	14		
Year 2, 4 to 5 years cases: Serotype 1	4	0		
Year 2, 4 to 5 years episodes: Serotype 1	4	0		
Year 2, 6 to 11 years cases: Serotype 1	0	3		

Year 2, 6 to 11 years episodes: Serotype 1	0	3		
Year 2, 4 to 5 years cases: Serotype 2	0	1		
Year 2, 4 to 5 years episodes: Serotype 2	0	1		
Year 2, 6 to 11 years cases: Serotype 2	4	5		
Year 2, 6 to 11 years episodes: Serotype 2	4	5		
Year 2, 4 to 5 years cases: Serotype 3	1	1		
Year 2, 4 to 5 years episodes: Serotype 3	1	1		
Year 2, 6 to 11 years cases: Serotype 3	5	2		
Year 2, 6 to 11 years episodes: Serotype 3	5	2		
Year 2, 4 to 5 years cases: Serotype 4	0	0		
Year 2, 4 to 5 years episodes: Serotype 4	0	0		
Year 2, 6 to 11 years cases: Serotype 4	2	4		
Year 2, 6 to 11 years episodes: Serotype 4	2	4		
Year 2, 4 to 5 years cases: Unserotyped	0	1		
Year 2, 4 to 5 years episodes: Unserotyped	0	1		
Year 2, 6 to 11 years cases: Unserotyped	0	0		
Year 2, 6 to 11 years episodes: Unserotyped	0	0		

Statistical analyses

No statistical analyses for this end point

Primary: Annual Incidence of Clinically Severe Hospitalized Virologically-Confirmed Dengue Due to Any and Each Serotype Collected During the Year I and Year 2 Passive Surveillance After Vaccination with Either CYD Dengue Vaccine or a Placebo in a Previous Study

End point title	Annual Incidence of Clinically Severe Hospitalized Virologically-Confirmed Dengue Due to Any and Each Serotype Collected During the Year I and Year 2 Passive Surveillance After Vaccination with Either CYD Dengue Vaccine or a Placebo in a Previous Study[37]

End point description:

Annual incidence rate was defined as the cases among the number of subjects present at the beginning of each period (M) multiplied by 100 converted in annual rate (Annual incidence rate = Cases among M*100).

End point type	Primary

End point timeframe:

Year 1 and Year 2 post-vaccination

Notes:

[37] - 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: Descriptive analyses were performed based on the study groups and the study vaccine administered for this outcome.

End point values	CYD Dengue Vaccine Group	Control Group		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	2131	1072		
Units: Annual incidence rate				
number (confidence interval 95%)				
Year 1: Any of the 4 Serotypes	0.2 (0.1 to 0.5)	0 (0 to 0.4)		
Year 1: Serotype 1	0.2 (0.1 to 0.5)	0 (0 to 0.4)		
Year 1: Serotype 2	0.1 (0 to 0.3)	0 (0 to 0.4)		
Year 1: Serotype 3	0 (0 to 0.2)	0 (0 to 0.4)		
Year 1: Serotype 4	0 (0 to 0.2)	0 (0 to 0.4)		
Year 1: Unserotyped	0 (0 to 0.2)	0 (0 to 0.4)		
Year 2: Any of the 4 Serotypes	0.1 (0 to 0.3)	0.2 (0 to 0.7)		
Year 2: Serotype 1	0.1 (0 to 0.3)	0 (0 to 0.3)		
Year 2: Serotype 2	0 (0 to 0.2)	0.1 (0 to 0.5)		
Year 2: Serotype 3	0 (0 to 0.2)	0 (0 to 0.3)		
Year 2: Serotype 4	0 (0 to 0.2)	0.1 (0 to 0.5)		
Year 2: Unserotyped	0 (0 to 0.2)	0 (0 to 0.3)		

Statistical analyses

No statistical analyses for this end point

Primary: Number of Cases and Episodes of Clinically Severe Hospitalized Virologically-Confirmed Dengue Due to Any and Each Serotype Collected During the Year I and Year 2 Passive Surveillance After Vaccination with CYD Dengue Vaccine or Placebo in a Previous Study

End point title	Number of Cases and Episodes of Clinically Severe Hospitalized Virologically-Confirmed Dengue Due to Any and Each Serotype Collected During the Year I and Year 2 Passive Surveillance After Vaccination with CYD Dengue Vaccine or Placebo in a Previous Study[38]

End point description:

Cases were defined as the number of subjects with at least one hospitalized virologically-confirmed dengue episode. Episodes were defined as the number of hospitalized virologically-confirmed dengue episodes.

End point type	Primary

End point timeframe:

Year 1 and Year 2 post-vaccination

Notes:

[38] - 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: Descriptive analyses were performed based on the study groups and the study vaccine administered for this outcome.

End point values	CYD Dengue Vaccine Group	Control Group		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	2131	1072		
Units: Number of cases and episodes				
number (not applicable)				
Year 1 Cases: Any of the 4 Serotypes	4	0		
Year 1 Episodes: Any of the 4 Serotypes	4	0		

Year 1 Cases: Serotype 1	4	0		
Year 1 Episodes: Serotype 1	4	0		
Year 1 Cases: Serotype 2	1	0		
Year 1 Episodes: Serotype 2	1	0		
Year 1 Cases: Serotype 3	0	0		
Year 1 Episodes: Serotype 3	0	0		
Year 1 Cases: Serotype 4	0	0		
Year 1 Episodes: Serotype 4	0	0		
Year 1 Cases: Unserotyped	0	0		
Year 1 Episodes: Unserotyped	0	0		
Year 2 Cases: Any of the 4 Serotypes	1	2		
Year 2 Episodes: Any of the 4 Serotypes	1	2		
Year 2 Cases: Serotype 1	1	0		
Year 2 Episodes: Serotype 1	1	0		
Year 2 Cases: Serotype 2	0	1		
Year 2 Episodes: Serotype 2	0	1		
Year 2 Cases: Serotype 3	0	0		
Year 2 Episodes: Serotype 3	0	0		
Year 2 Cases: Serotype 4	0	1		
Year 2 Episodes: Serotype 4	0	1		
Year 2 Cases: Unserotyped	0	0		
Year 2 Episodes: Unserotyped	0	0		

Statistical analyses

No statistical analyses for this end point

Primary: Number of Subjects with Non-Serotype Specific Dengue Viremia Among Hospitalized Virologically-Confirmed Dengue Cases Occurring During the Year I and Year 2 Passive Surveillance After Vaccination with CYD Dengue Vaccine or Placebo in a Previous Study

End point title	Number of Subjects with Non-Serotype Specific Dengue Viremia Among Hospitalized Virologically-Confirmed Dengue Cases Occurring During the Year I and Year 2 Passive Surveillance After Vaccination with CYD Dengue Vaccine or Placebo in a Previous Study[39]

End point description:

Quantified viremia was defined as ≥ lower limit of quantitation (log10 pfu/mL).

End point type	Primary

End point timeframe:

Year 1, Year 2, and entire period post-vaccination at each of the following time points: Anytime, 0-3 days, 4-7 days, 0-7 days, After 7 days.

Notes:

[39] - 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: Descriptive analyses were performed based on the study groups and the study vaccine administered for this outcome.

End point values	CYD Dengue Vaccine Group	Control Group		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	2131	1072		
Units: Number of subjects				
number (not applicable)				
Year 1, Cases - Anytime	19	9		
Year 1, Mean Viremia - Anytime	3.98	3.93		
Year 1, Cases - 0-3 days	7	3		
Year 1, Mean Viremia - 0-3 days	4.34	4.84		
Year 1, Cases - 4-7 days	12	5		
Year 1, Mean Viremia - 4-7 days	3.77	3.42		
Year 1, Cases - 0-7 days	19	8		
Year 1, Mean Viremia - 0-7 days	3.98	3.95		
Year 1, Cases - After 7 days	0	1		
Year 1, Mean Viremia - After 7 days	0	3.72		
Year 2, Cases - Anytime	15	16		
Year 2, Mean Viremia - Anytime	3.75	3.36		
Year 2, Cases - 0-3 days	6	5		
Year 2, Mean Viremia - 0-3 days	4.64	3.85		
Year 2, Cases - 4-7 days	9	10		
Year 2, Mean Viremia - 4-7 days	3.16	2.9		
Year 2, Cases - 0-7 days	15	15		
Year 2, Mean Viremia - 0-7 days	3.75	3.22		
Year 2, Cases - After 7 days	0	1		
Year 2, Mean Viremia - After 7 days	0	5.58		
Entire period, Cases - Anytime	34	25		
Entire period, Mean Viremia - Anytime	3.88	3.57		
Entire period, Cases - 0-3 days	13	8		
Entire period, Mean Viremia - 0-3 days	4.48	4.22		
Entire period, Cases - 4-7 days	21	15		
Entire period, Mean Viremia - 4-7 days	3.51	3.07		
Entire period, Cases - 0-7 days	34	23		
Entire period, Mean Viremia - 0-7 days	3.88	3.47		
Entire period, Cases - After 7 days	0	2		
Entire period, Mean Viremia - After 7 days	0	4.65		

Statistical analyses

No statistical analyses for this end point

Primary: Annual Incidence of Hospitalized Virologically-Confirmed Dengue Cases Due to Any and Each Serotype and Meeting WHO 1997 Criteria Collected During Year I and Year 2 Passive Surveillance After Previous Vaccination with Either CYD Dengue Vaccine or Placebo

End point title	Annual Incidence of Hospitalized Virologically-Confirmed Dengue Cases Due to Any and Each Serotype and Meeting WHO 1997 Criteria Collected During Year I and Year 2 Passive Surveillance After Previous Vaccination with Either CYD Dengue Vaccine or Placebo

End point description:

Annual incidence rate was defined as the cases among the number of subjects present at the beginning of each period (M) multiplied by 100 converted in annual rate (Annual incidence rate = Cases among M*100).

End point type	Primary

End point timeframe:

Year 1 and Year 2 post-vaccination

End point values	CYD Dengue Vaccine Group	Control Group		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	2131	1072		
Units: Annual incidence rate				
number (confidence interval 95%)				
Year 1: Any of the 4 Serotypes Any grade	0.2 (0.1 to 0.5)	0 (0 to 0.4)		
Year 1: Any of the 4 Serotypes Grade I	0.1 (0 to 0.3)	0 (0 to 0.4)		
Year 1: Any of the 4 Serotypes Grade II	0.1 (0 to 0.3)	0 (0 to 0.4)		
Year 1: Any of the 4 Serotypes Grade III	0.1 (0 to 0.4)	0 (0 to 0.4)		
Year 1: Any of the 4 Serotypes Grade IV	0 (0 to 0.2)	0 (0 to 0.4)		
Year 1: Serotype 1 Any grade	0.2 (0.1 to 0.5)	0 (0 to 0.4)		
Year 1: Serotype 1 Grade I	0.1 (0 to 0.3)	0 (0 to 0.4)		
Year 1: Serotype 1 Grade II	0.1 (0 to 0.3)	0 (0 to 0.4)		
Year 1: Serotype 1 Grade III	0.1 (0 to 0.4)	0 (0 to 0.4)		
Year 1: Serotype 1 Grade IV	0 (0 to 0.2)	0 (0 to 0.4)		
Year 1: Serotype 2 Any grade	0.1 (0 to 0.3)	0 (0 to 0.4)		
Year 1: Serotype 2 Grade I	0.1 (0 to 0.3)	0 (0 to 0.4)		
Year 1: Serotype 2 Grade II	0 (0 to 0.2)	0 (0 to 0.4)		
Year 1: Serotype 2 Grade III	0 (0 to 0.2)	0 (0 to 0.4)		
Year 1: Serotype 2 Grade IV	0 (0 to 0.2)	0 (0 to 0.4)		
Year 1: Serotype 3 Any grade	0 (0 to 0.2)	0 (0 to 0.4)		
Year 1: Serotype 3 Grade I	0 (0 to 0.2)	0 (0 to 0.4)		
Year 1: Serotype 3 Grade II	0 (0 to 0.2)	0 (0 to 0.4)		
Year 1: Serotype 3 Grade III	0 (0 to 0.2)	0 (0 to 0.4)		
Year 1: Serotype 3 Grade IV	0 (0 to 0.2)	0 (0 to 0.4)		
Year 1: Serotype 4 Any grade	0 (0 to 0.2)	0 (0 to 0.4)		
Year 1: Serotype 4 Grade I	0 (0 to 0.2)	0 (0 to 0.4)		
Year 1: Serotype 4 Grade II	0 (0 to 0.2)	0 (0 to 0.4)		
Year 1: Serotype 4 Grade III	0 (0 to 0.2)	0 (0 to 0.4)		
Year 1: Serotype 4 Grade IV	0 (0 to 0.2)	0 (0 to 0.4)		
Year 1: Unserotyped Any grade	0 (0 to 0.2)	0 (0 to 0.4)		
Year 1: Unserotyped Grade I	0 (0 to 0.2)	0 (0 to 0.4)		
Year 1: Unserotyped Grade II	0 (0 to 0.2)	0 (0 to 0.4)		
Year 1: Unserotyped Grade III	0 (0 to 0.2)	0 (0 to 0.4)		
Year 1: Unserotyped Grade IV	0 (0 to 0.2)	0 (0 to 0.4)		
Year 2: Any of the 4 Serotypes Any grade	0.1 (0 to 0.3)	0.2 (0 to 0.7)		
Year 2: Any of the 4 Serotypes Grade I	0 (0 to 0.2)	0.2 (0 to 0.7)		
Year 2: Any of the 4 Serotypes Grade II	0.1 (0 to 0.3)	0 (0 to 0.3)		

Year 2: Any of the 4 Serotypes Grade III	0 (0 to 0.2)	0 (0 to 0.3)		
Year 2: Any of the 4 Serotypes Grade IV	0 (0 to 0.2)	0 (0 to 0.3)		
Year 2: Serotype 1 Any grade	0.1 (0 to 0.3)	0 (0 to 0.3)		
Year 2: Serotype 1 Grade I	0 (0 to 0.2)	0 (0 to 0.3)		
Year 2: Serotype 1 Grade II	0.1 (0 to 0.3)	0 (0 to 0.3)		
Year 2: Serotype 1 Grade III	0 (0 to 0.2)	0 (0 to 0.3)		
Year 2: Serotype 1 Grade IV	0 (0 to 0.2)	0 (0 to 0.3)		
Year 2: Serotype 2 Any grade	0 (0 to 0.2)	0.1 (0 to 0.5)		
Year 2: Serotype 2 Grade I	0 (0 to 0.2)	0.1 (0 to 0.5)		
Year 2: Serotype 2 Grade II	0 (0 to 0.2)	0 (0 to 0.3)		
Year 2: Serotype 2 Grade III	0 (0 to 0.2)	0 (0 to 0.3)		
Year 2: Serotype 2 Grade IV	0 (0 to 0.2)	0 (0 to 0.3)		
Year 2: Serotype 3 Any grade	0 (0 to 0.2)	0 (0 to 0.3)		
Year 2: Serotype 3 Grade I	0 (0 to 0.2)	0 (0 to 0.3)		
Year 2: Serotype 3 Grade II	0 (0 to 0.2)	0 (0 to 0.3)		
Year 2: Serotype 3 Grade III	0 (0 to 0.2)	0 (0 to 0.3)		
Year 2: Serotype 3 Grade IV	0 (0 to 0.2)	0 (0 to 0.3)		
Year 2: Serotype 4 Any grade	0 (0 to 0.2)	0.1 (0 to 0.5)		
Year 2: Serotype 4 Grade I	0 (0 to 0.2)	0.1 (0 to 0.5)		
Year 2: Serotype 4 Grade II	0 (0 to 0.2)	0 (0 to 0.3)		
Year 2: Serotype 4 Grade III	0 (0 to 0.2)	0 (0 to 0.3)		
Year 2: Serotype 4 Grade IV	0 (0 to 0.2)	0 (0 to 0.3)		
Year 2: Unserotyped Any grade	0 (0 to 0.2)	0 (0 to 0.3)		
Year 2: Unserotyped Grade I	0 (0 to 0.2)	0 (0 to 0.3)		
Year 2: Unserotyped Grade II	0 (0 to 0.2)	0 (0 to 0.3)		
Year 2: Unserotyped Grade III	0 (0 to 0.2)	0 (0 to 0.3)		
Year 2: Unserotyped Grade IV	0 (0 to 0.2)	0 (0 to 0.3)		

Statistical analyses

Statistical analysis title	Relative Risk; Year 2; Any serotype; Any grade

Statistical analysis description:

Relative risk (RR) analysis of the annual incidence rate at Year 2 Any of the 4 Serotypes Any grade between the CYD Dengue vaccine group and the Control group.

The RR was not calculated for Year 1 and is reported for only those serotypes and grades calculated in Year 2.

Comparison groups	CYD Dengue Vaccine Group v Control Group
Number of subjects included in analysis	3203
Analysis specification	Pre-specified
Analysis type	other[40]
Parameter estimate	Relative Risk
Point estimate	0.252
Confidence interval	
level	95 %
sides	2-sided
lower limit	0
upper limit	4.83

Notes:

[40] - The 95% confidence interval of the relative risk is calculated using the Exact method described by Breslow and Day.

Statistical analysis title	Relative Risk; Year 2; Any serotype; Grade I

Statistical analysis description:

Relative risk (RR) analysis of the annual incidence rate at Year 2 Any of the 4 Serotypes Grade I between the CYD Dengue vaccine group and the Control group.

The RR was not calculated for Year 1 and is reported for only those serotypes and grades calculated in Year 2.

Comparison groups	CYD Dengue Vaccine Group v Control Group
Number of subjects included in analysis	3203
Analysis specification	Pre-specified
Analysis type	other[41]
Parameter estimate	Relative Risk
Point estimate	0
Confidence interval	
level	95 %
sides	2-sided
lower limit	0
upper limit	2.68

Notes:

[41] - The 95% confidence interval of the relative risk is calculated using the Exact method described by Breslow and Day.

Statistical analysis title	Relative Risk; Year 2; Serotype 2; Any grade

Statistical analysis description:

Relative risk (RR) analysis of the annual incidence rate at Year 2 Serotype 2 Any grade between the CYD Dengue vaccine group and the Control group.

The RR was not calculated for Year 1 and is reported for only those serotypes and grades calculated in Year 2.

Comparison groups	CYD Dengue Vaccine Group v Control Group
Number of subjects included in analysis	3203
Analysis specification	Pre-specified
Analysis type	other[42]
Parameter estimate	Relative Risk
Point estimate	0
Confidence interval	
level	95 %
sides	2-sided
lower limit	0
upper limit	19.62

Notes:

[42] - The 95% confidence interval of the relative risk is calculated using the Exact method described by Breslow and Day.

Statistical analysis title	Relative Risk; Year 2; Serotype 2; Grade I

Statistical analysis description:

Relative risk (RR) analysis of the annual incidence rate at Year 2 Serotype 2 Grade I between the CYD Dengue vaccine group and the Control group.

The RR was not calculated for Year 1 and is reported for only those serotypes and grades calculated in Year 2.

Comparison groups	Control Group v CYD Dengue Vaccine Group

Number of subjects included in analysis	3203
Analysis specification	Pre-specified
Analysis type	other[43]
Parameter estimate	Relative Risk
Point estimate	0
Confidence interval	
level	95 %
sides	2-sided
lower limit	0
upper limit	19.62

Notes:

[43] - The 95% confidence interval of the relative risk is calculated using the Exact method described by Breslow and Day.

Statistical analysis title	Relative Risk; Year 2; Serotype 4; Any grade

Statistical analysis description:

Relative risk (RR) analysis of the annual incidence rate at Year 2 Serotype 4 Any grade between the CYD Dengue vaccine group and the Control group.

The RR was not calculated for Year 1 and is reported for only those serotypes and grades calculated in Year 2.

Comparison groups	CYD Dengue Vaccine Group v Control Group
Number of subjects included in analysis	3203
Analysis specification	Pre-specified
Analysis type	other[44]
Parameter estimate	Relative Risk
Point estimate	0
Confidence interval	
level	95 %
sides	2-sided
lower limit	0
upper limit	19.62

Notes:

[44] - The 95% confidence interval of the relative risk is calculated using the Exact method described by Breslow and Day.

Statistical analysis title	Relative Risk; Year 2; Serotype 4; Grade I

Statistical analysis description:

Relative risk (RR) analysis of the annual incidence rate at Year 2 Serotype 4 Grade I between the CYD Dengue vaccine group and the Control group.

The RR was not calculated for Year 1 and is reported for only those serotypes and grades calculated in Year 2.

Comparison groups	CYD Dengue Vaccine Group v Control Group
Number of subjects included in analysis	3203
Analysis specification	Pre-specified
Analysis type	other[45]
Parameter estimate	Relative Risk
Point estimate	0
Confidence interval	
level	95 %
sides	2-sided
lower limit	0
upper limit	19.62

Notes:

[45] - The 95% confidence interval of the relative risk is calculated using the Exact method described by Breslow and Day.

Primary: Number of Cases and Episodes of Hospitalized Virologically-Confirmed Dengue Cases Due to Any and Each Serotype and Meeting WHO 1997 Criteria Collected In Year I and Year 2 Passive Surveillance After Previous Vaccination with CYD Dengue Vaccine or Placebo

End point title	Number of Cases and Episodes of Hospitalized Virologically-Confirmed Dengue Cases Due to Any and Each Serotype and Meeting WHO 1997 Criteria Collected In Year I and Year 2 Passive Surveillance After Previous Vaccination with CYD Dengue Vaccine or Placebo[46]

End point description:

Cases were defined as the number of subjects with at least one hospitalized virologically-confirmed dengue episode meeting WHO criteria. Episodes are defined as the number of hospitalized virologically-confirmed dengue episodes meeting WHO criteria. Grade I: Fever accompanied by non-specific constitutional symptoms; the only hemorrhagic manifestation is a positive tourniquet test. Grade II: Spontaneous bleeding in addition to the manifestations of Grade I patients, usually in the form of skin and/or other hemorrhages. Grade III: Circulatory failure manifested by rapid and weak pulse, narrowing of pulse pressure (20 mmHg or less) or hypotension, with the presence of cold clammy skin and restlessness. Grade IV: Profound shock with undetectable blood pressure and pulse.

End point type	Primary

End point timeframe:

Year 1 and Year 2 post-vaccination

Notes:

[46] - 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: Descriptive analyses were performed based on the study groups and the study vaccine administered for this outcome.

End point values	CYD Dengue Vaccine Group	Control Group		
Subject group type	Reporting group	Reporting group		
Number of subjects analysed	2131	1072		
Units: Number of cases and episodes				
number (not applicable)				
Year 1 Cases: Any of the 4 Serotypes Any grade	4	0		
Year 1 Episodes: Any of the 4 Serotypes Any grade	4	0		
Year 1 Cases: Any of the 4 Serotypes Grade I	1	0		
Year 1 Episodes: Any of the 4 Serotypes Grade I	1	0		
Year 1 Cases: Any of the 4 Serotypes Grade II	1	0		
Year 1 Episodes: Any of the 4 Serotypes Grade II	1	0		
Year 1 Cases: Any of the 4 Serotypes Grade III	2	0		
Year 1 Episodes: Any of the 4 Serotypes Grade III	2	0		
Year 1 Cases: Any of the 4 Serotypes Grade IV	0	0		
Year 1 Episodes: Any of the 4 Serotypes Grade IV	0	0		
Year 1 Cases: Serotype 1 Any grade	4	0		
Year 1 Episodes: Serotype 1 Any grade	4	0		
Year 1 Cases: Serotype 1 Grade I	1	0		

Year 1 Episodes: Serotype 1 Grade I	1	0		
Year 1 Cases: Serotype 1 Grade II	1	0		
Year 1 Episodes: Serotype 1 Grade II	1	0		
Year 1 Cases: Serotype 1 Grade III	2	0		
Year 1 Episodes: Serotype 1 Grade III	2	0		
Year 1 Cases: Serotype 1 Grade IV	0	0		
Year 1 Episodes: Serotype 1 Grade IV	0	0		
Year 1 Cases: Serotype 2 Any grade	1	0		
Year 1 Episodes: Serotype 2 Any grade	1	0		
Year 1 Cases: Serotype 2 Grade I	1	0		
Year 1 Episodes: Serotype 2 Grade I	1	0		
Year 1 Cases: Serotype 2 Grade II	0	0		
Year 1 Episodes: Serotype 2 Grade II	0	0		
Year 1 Cases: Serotype 2 Grade III	0	0		
Year 1 Episodes: Serotype 2 Grade III	0	0		
Year 1 Cases: Serotype 2 Grade IV	0	0		
Year 1 Episodes: Serotype 2 Grade IV	0	0		
Year 1 Cases: Serotype 3 Any grade	0	0		
Year 1 Episodes: Serotype 3 Any grade	0	0		
Year 1 Cases: Serotype 3 Grade I	0	0		
Year 1 Episodes: Serotype 3 Grade I	0	0		
Year 1 Cases: Serotype 3 Grade II	0	0		
Year 1 Episodes: Serotype 3 Grade II	0	0		
Year 1 Cases: Serotype 3 Grade III	0	0		
Year 1 Episodes: Serotype 3 Grade III	0	0		
Year 1 Cases: Serotype 3 Grade IV	0	0		
Year 1 Episodes: Serotype 3 Grade IV	0	0		
Year 1 Cases: Serotype 4 Any grade	0	0		
Year 1 Episodes: Serotype 4 Any grade	0	0		
Year 1 Cases: Serotype 4 Grade I	0	0		
Year 1 Episodes: Serotype 4 Grade I	0	0		
Year 1 Cases: Serotype 4 Grade II	0	0		
Year 1 Episodes: Serotype 4 Grade II	0	0		
Year 1 Cases: Serotype 4 Grade III	0	0		
Year 1 Episodes: Serotype 4 Grade III	0	0		
Year 1 Cases: Serotype 4 Grade IV	0	0		
Year 1 Episodes: Serotype 4 Grade IV	0	0		
Year 1 Cases: Unserotyped Any grade	0	0		
Year 1 Episodes: Unserotyped Any grade	0	0		
Year 1 Cases: Unserotyped Grade I	0	0		
Year 1 Episodes: Unserotyped Grade I	0	0		
Year 1 Cases: Unserotyped Grade II	0	0		
Year 1 Episodes: Unserotyped Grade II	0	0		
Year 1 Cases: Unserotyped Grade III	0	0		
Year 1 Episodes: Unserotyped Grade III	0	0		
Year 1 Cases: Unserotyped Grade IV	0	0		
Year 1 Episodes: Unserotyped Grade IV	0	0		
Year 2 Cases: Any of the 4 Serotypes Any grade	1	2		
Year 2 Episodes: Any of the 4 Serotypes Any grade	1	2		

Year 2 Cases: Any of the 4 Serotypes Grade I	0	2		
Year 2 Episodes: Any of the 4 Serotypes Grade I	0	2		
Year 2 Cases: Any of the 4 Serotypes Grade II	1	0		
Year 2 Episodes: Any of the 4 Serotypes Grade II	1	0		
Year 2 Cases: Any of the 4 Serotypes Grade III	0	0		
Year 2 Episodes: Any of the 4 Serotypes Grade III	0	0		
Year 2 Cases: Any of the 4 Serotypes Grade IV	0	0		
Year 2 Episodes: Any of the 4 Serotypes Grade IV	0	0		
Year 2 Cases: Serotype 1 Any grade	1	0		
Year 2 Episodes: Serotype 1 Any grade	1	0		
Year 2 Cases: Serotype 1 Grade I	0	0		
Year 2 Episodes: Serotype 1 Grade I	0	0		
Year 2 Cases: Serotype 1 Grade II	1	0		
Year 2 Episodes: Serotype 1 Grade II	1	0		
Year 2 Cases: Serotype 1 Grade III	0	0		
Year 2 Episodes: Serotype 1 Grade III	0	0		
Year 2 Cases: Serotype 1 Grade IV	0	0		
Year 2 Episodes: Serotype 1 Grade IV	0	0		
Year 2 Cases: Serotype 2 Any grade	0	1		
Year 2 Episodes: Serotype 2 Any grade	0	1		
Year 2 Cases: Serotype 2 Grade I	0	1		
Year 2 Episodes: Serotype 2 Grade I	0	1		
Year 2 Cases: Serotype 2 Grade II	0	0		
Year 2 Episodes: Serotype 2 Grade II	0	0		
Year 2 Cases: Serotype 2 Grade III	0	0		
Year 2 Episodes: Serotype 2 Grade III	0	0		
Year 2 Cases: Serotype 2 Grade IV	0	0		
Year 2 Episodes: Serotype 2 Grade IV	0	0		
Year 2 Cases: Serotype 3 Any grade	0	0		
Year 2 Episodes: Serotype 3 Any grade	0	0		
Year 2 Cases: Serotype 3 Grade I	0	0		
Year 2 Episodes: Serotype 3 Grade I	0	0		
Year 2 Cases: Serotype 3 Grade II	0	0		
Year 2 Episodes: Serotype 3 Grade II	0	0		
Year 2 Cases: Serotype 3 Grade III	0	0		
Year 2 Episodes: Serotype 3 Grade III	0	0		
Year 2 Cases: Serotype 3 Grade IV	0	0		
Year 2 Episodes: Serotype 3 Grade IV	0	0		
Year 2 Cases: Serotype 4 Any grade	0	1		
Year 2 Episodes: Serotype 4 Any grade	0	1		
Year 2 Cases: Serotype 4 Grade I	0	1		
Year 2 Episodes: Serotype 4 Grade I	0	1		
Year 2 Cases: Serotype 4 Grade II	0	0		
Year 2 Episodes: Serotype 4 Grade II	0	0		
Year 2 Cases: Serotype 4 Grade III	0	0		
Year 2 Episodes: Serotype 4 Grade III	0	0		

Year 2 Cases: Serotype 4 Grade IV	0	0		
Year 2 Episodes: Serotype 4 Grade IV	0	0		
Year 2 Cases: Unserotyped Any grade	0	0		
Year 2 Episodes: Unserotyped Any grade	0	0		
Year 2 Cases: Unserotyped Grade I	0	0		
Year 2 Episodes: Unserotyped Grade I	0	0		
Year 2 Cases: Unserotyped Grade II	0	0		
Year 2 Episodes: Unserotyped Grade II	0	0		
Year 2 Cases: Unserotyped Grade III	0	0		
Year 2 Episodes: Unserotyped Grade III	0	0		
Year 2 Cases: Unserotyped Grade IV	0	0		
Year 2 Episodes: Unserotyped Grade IV	0	0		

Statistical analyses

No statistical analyses for this end point

Adverse events

Adverse events information[1]

Timeframe for reporting adverse events:

This study involved a passive surveillance of hospitalized dengue cases in subjects that participated in a previous study. No vaccine were administered in the study and adverse event data were also not collected.

Assessment type	Non-systematic

Dictionary used

Dictionary name	MedDRA
Dictionary version	10

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

Notes:

[1] - There are no non-serious adverse events recorded for these results. It is expected that there will be at least one non-serious adverse event reported.

Justification: This was a long-term immunogenicity follow-up study of subjects who participated in the primary study. No vaccines were administered in the follow-up study and adverse event data were also not collected.

More information

Substantial protocol amendments (globally)

Were there any global substantial amendments to the protocol? Yes

Date	Amendment
08 January 2014	Additional clarifications were added to the study objectives.
17 February 2014	Study objectives were further updated and statistical analyses were revised.

Notes:

Interruptions (globally)

Were there any global interruptions to the trial? No

Limitations and caveats

None reported

Online references

http://www.ncbi.nlm.nih.gov/pubmed/22975340