

EMA/710066/2017

European Medicines Agency decision

P/0359/2017

of 1 December 2017

on the agreement of a paediatric investigation plan and on the granting of a deferral and on the granting of a waiver for *Neisseria meningitidis* serogroup A polysaccharide conjugated to tetanus toxoid / *N. meningitidis* serogroup C polysaccharide conjugated to tetanus toxoid / *N. meningitidis* serogroup Y polysaccharide conjugated to tetanus toxoid / *N. meningitidis* serogroup W polysaccharide conjugated to tetanus toxoid (MenACYW) (EMEA-001930-PIP01-16) in accordance with Regulation (EC) No 1901/2006 of the European Parliament and of the Council

Disclaimer

This decision does not constitute entitlement to the rewards and incentives referred to in Title V of Regulation (EC) No 1901/2006.

Only the English text is authentic.

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The European Medicines Agency,

Having regard to the Treaty on the Functioning of the European Union,

Having regard to Regulation (EC) No 1901/2006 of the European Parliament and of the Council of 12 December 2006 on medicinal products for paediatric use and amending Regulation (EEC) No. 1768/92, Directive 2001/20/EC, Directive 2001/83/EC and Regulation (EC) No 726/2004¹,

Having regard to Regulation (EC) No 726/2004 of the European Parliament and of the Council of 31 March 2004 laying down Community procedures for the authorisation and supervision of medicinal products for human and veterinary use and establishing a European Medicines Agency²,

Having regard to the application submitted by Sanofi Pasteur Inc. on 25 January 2016 under Article 16(1) of Regulation (EC) No 1901/2006 also requesting a deferral under Article 20 of said Regulation and a waiver under Article 13 of said Regulation,

Having regard to the opinion of the Paediatric Committee of the European Medicines Agency, issued on 13 October 2017, in accordance with Article 17 of Regulation (EC) No 1901/2006 and Article 21 of said Regulation and Article 13 of said Regulation,

Having regard to Article 25 of Regulation (EC) No 1901/2006,

Whereas:

- (1) The Paediatric Committee of the European Medicines Agency has given an opinion on the agreement of a paediatric investigation plan and on the granting of a deferral and on the granting of a waiver.
- (2) It is therefore appropriate to adopt a decision agreeing a paediatric investigation plan.
- (3) It is therefore appropriate to adopt a decision granting a deferral.
- (4) It is therefore appropriate to adopt a decision granting a waiver.

¹ OJ L 378, 27.12.2006, p.1.

² OJ L 136, 30.4.2004, p. 1.

Has adopted this decision:

Article 1

A paediatric investigation plan for *Neisseria meningitidis* serogroup A polysaccharide conjugated to tetanus toxoid / *N. meningitidis* serogroup C polysaccharide conjugated to tetanus toxoid / *N. meningitidis* serogroup Y polysaccharide conjugated to tetanus toxoid / *N. meningitidis* serogroup W polysaccharide conjugated to tetanus toxoid (MenACYW), solution for injection, intramuscular use, the details of which are set out in the opinion of the Paediatric Committee of the European Medicines Agency annexed hereto, together with its appendices, is hereby agreed.

Article 2

A deferral for *Neisseria meningitidis* serogroup A polysaccharide conjugated to tetanus toxoid / *N. meningitidis* serogroup C polysaccharide conjugated to tetanus toxoid / *N. meningitidis* serogroup Y polysaccharide conjugated to tetanus toxoid / *N. meningitidis* serogroup W polysaccharide conjugated to tetanus toxoid (MenACYW), solution for injection, intramuscular use, the details of which are set out in the opinion of the Paediatric Committee of the European Medicines Agency annexed hereto, together with its appendices, is hereby granted.

Article 3

A waiver for *Neisseria meningitidis* serogroup A polysaccharide conjugated to tetanus toxoid / *N. meningitidis* serogroup C polysaccharide conjugated to tetanus toxoid / *N. meningitidis* serogroup Y polysaccharide conjugated to tetanus toxoid / *N. meningitidis* serogroup W polysaccharide conjugated to tetanus toxoid (MenACYW), solution for injection, intramuscular use, the details of which are set out in the opinion of the Paediatric Committee of the European Medicines Agency annexed hereto, together with its appendices, is hereby granted.

Article 4

This decision is addressed to Sanofi Pasteur Inc., Discovery Drive, 18370 – Swiftwater, United States.

EMA/PDCO/279193/2016
London, 13 October 2017

Opinion of the Paediatric Committee on the agreement of a Paediatric Investigation Plan and a deferral and a waiver

EMA-001930-PIP01-16

Scope of the application

Active substance(s):

Neisseria meningitidis serogroup A polysaccharide conjugated to tetanus toxoid / *N. meningitidis* serogroup C polysaccharide conjugated to tetanus toxoid / *N. meningitidis* serogroup Y polysaccharide conjugated to tetanus toxoid / *N. meningitidis* serogroup W polysaccharide conjugated to tetanus toxoid (MenACYW)

Condition(s):

Prevention of meningococcal disease

Pharmaceutical form(s):

Solution for injection

Route(s) of administration:

Intramuscular use

Name / corporate name of the PIP applicant:

Sanofi Pasteur Inc.

Basis for opinion

Pursuant to Article 16(1) of Regulation (EC) No 1901/2006 as amended, Sanofi Pasteur Inc. submitted for agreement to the European Medicines Agency on 25 January 2016 an application for a paediatric investigation plan for the above mentioned medicinal product and a deferral under Article 20 of said Regulation and a waiver under Article 13 of said Regulation.

The procedure started on 1 March 2016.

Supplementary information was provided by the applicant on 24 July 2017. The applicant proposed modifications to the paediatric investigation plan.

Opinion

1. The Paediatric Committee, having assessed the proposed paediatric investigation plan in accordance with Article 17 of Regulation (EC) No 1901/2006 as amended, recommends as set out in the appended summary report:

- to agree the paediatric investigation plan in accordance with Article 17(1) of said Regulation;
- to grant a deferral in accordance with Article 21 of said Regulation;
- to grant a waiver for one or more subsets of the paediatric population in accordance with Article 13 of said Regulation and concluded in accordance with Article 11(1)(a) of said Regulation, on the grounds that the specific medicinal product is likely to be ineffective or unsafe in part or all of the paediatric population.

The Norwegian Paediatric Committee member agrees with the above-mentioned recommendation of the Paediatric Committee.

2. The measures and timelines of the agreed paediatric investigation plan and the subset(s) of the paediatric population and condition(s) covered by the waiver are set out in the Annex I.

This opinion is forwarded to the applicant and the Executive Director of the European Medicines Agency, together with its annex and appendix.

Annex I

The subset(s) of the paediatric population and condition(s) covered by the waiver and the measures and timelines of the agreed paediatric investigation plan (PIP)

1. Waiver

1.1. Condition:

Prevention of invasive meningococcal disease caused by *N. meningitidis* serogroups A, C, Y, and W-135

The waiver applies to:

- the paediatric population from birth to less than 6 weeks of age;
- solution for injection, intramuscular use;
- on the grounds that the specific medicinal product is likely to be ineffective.

2. Paediatric investigation plan

2.1. Condition

Prevention of invasive meningococcal disease

2.1.1. Indication(s) targeted by the PIP

Active immunization to prevent invasive meningococcal disease caused by *N. meningitidis* serogroups A, C, Y, and W-135

2.1.2. Subset(s) of the paediatric population concerned by the paediatric development

From 6 weeks to less than 18 years of age

2.1.3. Pharmaceutical form(s)

Solution for injection

2.1.4. Measures

Area	Number of measures	Description
Quality-related studies	0	Not applicable.
Non-clinical studies	0	Not applicable.

Clinical studies	15	Study 1 An open-label, randomized, parallel-group, controlled, multi-centre study to evaluate the immunogenicity and safety profile of a single dose of MenACYW conjugate vaccine when given alone compared to that of the licensed Meningococcal (Groups A, C, Y and W-135) Diphtheria Conjugate Vaccine (Menveo), and when MenACYW conjugate vaccine is given concomitantly with diphtheria, pertussis, and tetanus (Tdap) vaccine and human papilloma virus (HPV) vaccine in paediatric subjects from 10 to less than 17 years of age; (study MET50). Study 2 A randomized, active-controlled, open-label study to evaluate the immunogenicity and safety profile of a single dose of MenACYW conjugate vaccine when given alone compared to that of a licensed meningococcal group A, C, W-135 and Y conjugate vaccine (Nimenrix) in 12 to 23 months old paediatric subjects; (study MET54). Study 3 A modified double-blind, randomized, parallel-group, active-controlled, multi-centre trial to compare the non-inferiority of a single dose of MenACYW conjugate vaccine to a single dose of Nimenrix in paediatric subjects from 12 to less than 24 months old; (study MET51). Study 4 Open-label, randomized, parallel-group, active-controlled, multi-centre study to describe the immunogenicity and safety of a single dose of MenACYW conjugate vaccine when administered alone and when administered concomitantly with other paediatric vaccine(s) in paediatric subjects from 12 to less than 24 months old; (study MET57). Study 5 A double-blind, randomized, parallel-group, active-controlled, trial to evaluate the immunogenicity and describe the safety of MenACYW conjugate vaccine compared to Meningococcal (Groups A, C, Y and W-135) Diphtheria Conjugate Vaccine (Menveo) in paediatric subjects 2 to 9 years of age in the United States; (study MET35). Study 6 Open-label, randomized, parallel-group, active-controlled, multi-centre study to compare the immunogenicity and safety of MenACYW conjugate vaccine when administered concomitantly with Men B vaccine in the 2nd year of life to healthy paediatric subjects in Europe; (study MET52).
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		Study 7 Open-label, randomized, parallel-group, active-controlled, multi-centre study to compare the immunogenicity and safety of MenACYW conjugate vaccine to a licensed quadrivalent meningococcal polysaccharide groups A, C, W-135, and Y conjugate vaccine (Nimenrix) when administered concomitantly with routine paediatric vaccines to healthy paediatric subjects from 2 to 3 months old in EU; (study MET58). Study 8 Open-label, randomized, parallel-group, active-controlled, multi-centre study to compare the immunogenicity and safety of MenACYW conjugate vaccine and Meningococcal (Groups A, C, Y and W-135) Diphtheria Conjugate Vaccine (Menveo) when administered concomitantly with routine paediatric vaccines to paediatric subjects 2 months old in the United States; (study MET42). Study 9 Double-blind, randomized, parallel-group, active-controlled, study to describe the safety of MenACYW conjugate vaccine when administered concomitantly with routine paediatric vaccines given to healthy infants and toddlers in the United States; (study MET41). Study 10 Open-label, randomized, parallel-group, active-controlled, multi-centre study to describe the immunogenicity and safety of a 3-dose immunization schedule of MenACYW conjugate vaccine or a 4-dose immunization schedule of a licensed quadrivalent meningococcal conjugate vaccine (Menveo) when administered concomitantly with routine paediatric vaccines in healthy infants and toddlers in the Philippines, the Russian Federation, and Mexico; (study MET33). Study 11 Open-label, randomized, parallel-group, active-controlled, multi-centre study to compare the immunogenicity and safety of MenACYW conjugate vaccine and Menveo or Meningococcal (Groups A, C, Y and W-135) Polysaccharide Diphtheria Toxoid Conjugate Vaccine (Menactra) when administered concomitantly with routine paediatric vaccines to paediatric subjects 6 to 19 months old, in the United States; (study MET61). Study 12 Modified double-blind, randomized, parallel-group, active-controlled, multi-centre study to evaluate immune lot consistency of MenACYW conjugate vaccine, evaluate the immune non-inferiority versus Meningococcal (Groups A, C, Y and W-135) Polysaccharide Diphtheria Toxoid Conjugate Vaccine (Menactra), and describe the safety and additional immunogenicity of study vaccines in paediatric patients 10 to less than 18 years old; (study MET43).
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		Study 13 Double-blind, randomized, parallel-group, active-controlled, multi-centre trial to compare the immunogenicity and describe the safety of a booster dose of MenACYW conjugate vaccine compared to Meningococcal (Groups A, C, Y and W-135) Polysaccharide Diphtheria Toxoid Conjugate Vaccine (Menactra) in paediatric subjects from 15 to less than 18 years old; (study MET56). Study 14 Open-label, multi-centre study to describe the immune persistence of the priming dose and describe the immunogenicity and safety of a booster dose of MenACYW conjugate vaccine in children in Finland who had been vaccinated 3 years earlier as toddlers with either MenACYW conjugate vaccine or quadrivalent meningococcal polysaccharide groups A, C, W-135, and Y conjugate vaccine (Nimenrix) as part of the MET54 study (Study 2) in paediatric subjects 4 to 5 years old (MET62). Study 15 Open-label, multi-centre study to describe the immune persistence of the priming dose and describe the immunogenicity and safety of a booster dose of MenACYW conjugate vaccine, when co-administered along with MenB vaccines in adolescents (and adults) in USA who had been vaccinated 5 years earlier as adolescents with either MenACYW conjugate vaccine or Meningococcal (Groups A, C, Y and W-135) Diphtheria Conjugate Vaccine (Menveo) as part of the MET50 study (Study 1); (study MET59).
Extrapolation, modelling and simulation studies	0	Not applicable.
Other studies	0	Not applicable.
Other measures	0	Not applicable.

3. Follow-up, completion and deferral of PIP

Concerns on potential long term safety/efficacy issues in relation to paediatric use:	No
Date of completion of the paediatric investigation plan:	By July 2021
Deferral for one or more measures contained in the paediatric investigation plan:	Yes