EMA/823989/2022

European Medicines Agency decision

P/0443/2022

of 28 October 2022

on the acceptance of a modification of an agreed paediatric investigation plan for marstacimab (EMEA-002285-PIP02-19-M02) 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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on the acceptance of a modification of an agreed paediatric investigation plan for marstacimab (EMEA-002285-PIP02-19-M02) in accordance with Regulation (EC) No 1901/2006 of the European Parliament and of the Council

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 726/2004[1],

Having regard to Regulation (EC) No 726/2004 of the European Parliament and of the Council of 31 March 2004 laying down Union procedures for the authorisation and supervision of medicinal products for human use and establishing a European Medicines Agency[2],

Having regard to the European Medicines Agency's decision P/0292/2020 issued on 12 August 2020, and the decision P/0516/2021 issued on 3 December 2021,

Having regard to the application submitted by Pfizer Europe MAA EEIG on 30 May 2022 under Article 22 of Regulation (EC) No 1901/2006 proposing changes to the agreed paediatric investigation plan with a deferral and a waiver,

Having regard to the opinion of the Paediatric Committee of the European Medicines Agency, issued on 9 September 2022, in accordance with Article 22 of Regulation (EC) No 1901/2006,

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 acceptance of changes to the agreed paediatric investigation plan.

(2) It is therefore appropriate to adopt a decision on the acceptance of changes to the agreed paediatric investigation plan.

[1] OJ L 378, 27.12.2006, p.1, as amended.
[2] OJ L 136, 30.4.2004, p. 1, as amended.

Has adopted this decision:

Article 1

Changes to the agreed paediatric investigation plan for marstacimab, solution for injection, subcutaneous use, are hereby accepted in the scope set out in the opinion of the Paediatric Committee of the European Medicines Agency annexed hereto, together with its appendices.

Article 2

This decision is addressed to Pfizer Europe MAA EEIG, 17 Boulevard de la Plaine, 1050 – Brussels, Belgium.

EMA/PDCO/566891/2022
Amsterdam, 9 September 2022

Opinion of the Paediatric Committee on the acceptance of a modification of an agreed Paediatric Investigation Plan

EMEA-002285-PIP02-19-M02

Scope of the application

Active substance(s):

Marstacimab

Condition(s):

Treatment of congenital haemophilia A

Treatment of congenital haemophilia B

Pharmaceutical form(s):

Solution for injection

Route(s) of administration:

Subcutaneous use

Name/corporate name of the PIP applicant:

Pfizer Europe MAA EEIG

Basis for opinion

Pursuant to Article 22 of Regulation (EC) No 1901/2006 as amended, Pfizer Europe MAA EEIG submitted to the European Medicines Agency on 30 May 2022 an application for modification of the agreed paediatric investigation plan with a deferral and a waiver as set out in the European Medicines Agency's decision P/0292/2020 issued on 12 August 2020 and the decision P/0516/2021 issued on 3 December 2021.

The application for modification proposed changes to the agreed paediatric investigation plan.

The procedure started on 11 July 2022.

Scope of the modification

Some measures and timelines of the Paediatric Investigation Plan have been modified.

Opinion

1. The Paediatric Committee, having assessed the application in accordance with Article 22 of Regulation (EC) No 1901/2006 as amended, recommends as set out in the appended summary report:

   - to agree to changes to the paediatric investigation plan in the scope set out in the Annex I of this opinion.

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

2. The measures and timelines of the 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:

Treatment of congenital haemophilia A

The waiver applies to:

- the paediatric population from birth to less than 1 year of age;
- solution for injection, subcutaneous use;
- on the grounds that the specific medicinal product is likely to be unsafe.

1.2. Condition:

Treatment of congenital haemophilia B

The waiver applies to:

- the paediatric population from birth to less than 1 year of age;
- solution for injection, subcutaneous use;
- on the grounds that the specific medicinal product is likely to be unsafe.

2. Paediatric investigation plan

2.1. Condition:

Treatment of congenital haemophilia A

2.1.1. Indication(s) targeted by the PIP

Prophylaxis of bleeding episodes in patients with congenital haemophilia A (factor VIII deficiency), with and without inhibitors

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

From 1 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	2	**Study 1 (B7841005)** Non-randomised, open-label, intra-patient controlled, one-way, cross-over, single-arm study to evaluate the safety and efficacy of marstacimab prophylaxis versus previous standard of care on-demand and prophylaxis treatment in severe congenital haemophilia A patients and moderately severe to severe congenital haemophilia B patients with and without inhibitors from 12 to less than 18 years of age (and adults). **Study 2 (B7841008)** Non-randomised, open-label, intra-patient controlled, one-way, cross-over, single-arm study with age staggered enrolment, to evaluate the safety and efficacy of marstacimab prophylaxis versus previous standard of care on-demand and prophylaxis treatment in severe congenital haemophilia A and B patients with and without inhibitors from 1 to less than 18 years of age.
Extrapolation, modelling and simulation studies	3	**Study 3 (Modelling study 1)** Population PK Modelling and Simulation study to evaluate a suitable dose of marstacimab for adolescents from 12 to less than 18 years of age. **Study 4 (Modelling study 2)** Population PK Modelling and Simulation study to evaluate a suitable dose of marstacimab for children from 1 to less than 18 years of age. **Study 5 (Modelling study 3)** Population PK/PD exposure-response modelling and simulation study to characterize the PK/PD relationship in paediatric haemophilia patients from 1 to less than 18 years of age (and adults).
Other studies	0	Not applicable
Other measures	0	Not applicable

2.2. Condition:

Treatment of congenital haemophilia B

2.2.1. Indication(s) targeted by the PIP

Prophylaxis of bleeding episodes in patients with congenital haemophilia B (factor IX deficiency), with and without inhibitors

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

From 1 to less than 18 years of age

2.2.3. Pharmaceutical form(s)

Solution for injection

2.2.4. Measures

Area	Number of measures	Description
Quality-related studies	0	Not applicable
Non-clinical studies	0	Not applicable
Clinical studies	2	**Study 1** *Same as for condition Treatment of congenital haemophilia A, see above.* **Study 2** *Same as for condition Treatment of congenital haemophilia A, see above.*
Extrapolation, modelling and simulation studies	3	**Study 3** *Same as for condition Treatment of congenital haemophilia A, see above* **Study 4** *Same as for condition Treatment of congenital haemophilia A, see above.* **Study 5** *Same as for condition Treatment of congenital haemophilia A, see above*
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:	Yes.
Date of completion of the paediatric investigation plan:	By November 2028.
Deferral for one or more measures contained in the paediatric investigation plan:	Yes.