EMA/170273/2023

European Medicines Agency decision

P/0156/2023

of 12 May 2023

on the acceptance of a modification of an agreed paediatric investigation plan for valoctocogene roxaparvovec (Roctavian), (EMEA-002427-PIP01-18-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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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/0218/2019 issued on 17 June 2019 and the decision P/0148/2021 issued on 16 April 2021,

Having regard to the application submitted by BioMarin International Limited on 12 December 2022 under Article 22 of Regulation (EC) No 1901/2006 proposing changes to the agreed paediatric investigation plan with a deferral,

Having regard to the opinion of the Paediatric Committee of the European Medicines Agency, issued on 31 March 2023, 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 and to the deferral.

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

[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 valoctocogene roxaparvovec (Roctavian), solution for infusion, intravenous use, including changes to the deferral, 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 BioMarin International Limited, Shanbally, Ringaskiddy, County Cork, P43 R298 - Shanbally, Ireland.

EMA/PDCO/4987/2023
Amsterdam, 31 March 2023

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

EMEA-002427-PIP01-18-M02

Scope of the application

Active substance(s):

Valoctocogene roxaparvovec

Invented name and authorisation status:

See Annex II

Condition(s):

Treatment of haemophilia A

Pharmaceutical form(s):

Solution for infusion

Route(s) of administration:

Intravenous use

Name/corporate name of the PIP applicant:

BioMarin International Limited

Basis for opinion

Pursuant to Article 22 of Regulation (EC) No 1901/2006 as amended, BioMarin International Limited submitted to the European Medicines Agency on 12 December 2022 an application for modification of the agreed paediatric investigation plan with a deferral as set out in the European Medicines Agency's decision P/0218/2019 issued on 17 June 2019 and the decision P/0148/2021 issued on 16 April 2021.

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

The procedure started on 30 January 2023.

Scope of the modification

Some 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 and to the deferral 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 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 annexes 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

Not applicable

2. Paediatric investigation plan

2.1. Condition

Treatment of haemophilia A

2.1.1. Indication(s) targeted by the PIP

Treatment of patients with haemophilia A

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

From birth to less than 18 years of age

2.1.3. Pharmaceutical form(s)

Solution for infusion

2.1.4. Measures

Area	Description
Quality-related studies	Not applicable
Non-clinical studies	**Study 1** Exploratory toxicity and efficacy study comparing FVIII expression levels between neonatal mice and sexually mature mice **Study 2** Definitive juvenile toxicity study to assess the effect of BMN 270 in mice and to evaluate toxicity during post-natal development at varying ages **Study 3** Dose range-finding juvenile toxicity study to assess the effects of BMN 270 in mice at the specific ages identified in study 2 and to support dose-selection
Clinical studies	**Study 4** Open-label, single-arm, single dose trial to evaluate safety and efficacy of BMN 270 in children from 12 years to less than 18 years of age with haemophilia A

	Study 5 Open-label, single-arm, single dose trial to evaluate safety and efficacy of BMN 270 in children from 6 years to less than 12 years of age with haemophilia A **Study 6** Open-label, single-arm, single dose trial to evaluate safety and efficacy of BMN 270 in children from birth to less than 6 years of age with haemophilia A
Extrapolation, modelling and simulation studies	Not applicable
Other studies	Not applicable
Other measures	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 December 2039
Deferral for one or more measures contained in the paediatric investigation plan:	Yes

Annex II

Information about the authorised medicinal product

Information provided by the applicant:

Condition(s) and authorised indication(s)

1. Treatment of haemophilia A

Authorised indication(s):

- Roctavian is indicated for the treatment of severe haemophilia A (congenital factor VIII deficiency) in adult patients without a history of factor VIII inhibitors and without detectable antibodies to adeno-associated virus serotype 5 (AAV5).
  - Invented name(s): Roctavian
  - Authorised pharmaceutical form(s): Solution for infusion
  - Authorised route(s) of administration: Intravenous use
  - Authorised via centralised procedure