EMA/480344/2019

European Medicines Agency decision

P/0348/2019

of 10 September 2019

on the acceptance of a modification of an agreed paediatric investigation plan for 2-hydroxypropyl-ß-cyclodextrin (HP-ß-CD) (EMEA-001866-PIP01-15-M04) 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 Community procedures for the authorisation and supervision of medicinal products for human and veterinary use and establishing a European Medicines Agency[2],

Having regard to the European Medicines Agency's decision P/0030/2017 issued on 31 January 2017 and the decision P/0224/2018 issued on 17 July 2018 and the decision P/0027/2019 issued on 29 January 2019,

Having regard to the application submitted by Mallinckrodt Pharmaceuticals Ireland Ltd on 19 April 2019 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 26 July 2019, 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.
[2] OJ L 136, 30.4.2004, p. 1.

Has adopted this decision:

Article 1

Changes to the agreed paediatric investigation plan for 2-hydroxypropyl-ß-cyclodextrin (HP-ß-CD), solution for injection, intrathecal 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 Mallinckrodt Pharmaceuticals Ireland Ltd, College Business & Technology, Park, Cruiserath Road, D15 TX2V - Blanchardstown, Ireland.

EMA/PDCO/263826/2019
Amsterdam, 26 July 2019

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

EMEA-001866-PIP01-15-M04

Active substance(s):

2-hydroxypropyl-ß-cyclodextrin (HP-ß-CD)

Condition(s):

Treatment of Niemann-Pick disease, type C

Pharmaceutical form(s):

Solution for injection

Route(s) of administration:

Intrathecal use

Name/corporate name of the PIP applicant:

Mallinckrodt Pharmaceuticals Ireland Ltd

Basis for opinion

Pursuant to Article 22 of Regulation (EC) No 1901/2006 as amended, Mallinckrodt Pharmaceuticals Ireland Ltd submitted to the European Medicines Agency on 19 April 2019 an application for modification of the agreed paediatric investigation plan with a deferral as set out in the European Medicines Agency's decision P/0030/2017 issued on 31 January 2017 and the decision P/0224/2018 issued on 17 July 2018 and the decision P/0027/2019 issued on 29 January 2019.

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

The procedure started on 28 May 2019.

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 and to the deferral in the scope set out in the Annex I of this opinion.

   The Norwegian Paediatric Committee member 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 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

Not applicable

2. Paediatric investigation plan

2.1. Condition

Treatment of Niemann-Pick disease, type C

2.1.1. Indication(s) targeted by the PIP

Treatment of progressive neurological manifestations in children and adolescent patients with Niemann-Pick disease, type C

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 injection

2.1.4. Measures

Area	Number of measures	Description
Quality-related studies	0	Not applicable
Non-clinical studies	0	Not applicable
Clinical studies	4	**Study 1** Study deleted during procedure EMEA-001866-PIP01-15-M03 **Study 2** Open-label, single-centre, multiple-dose, dose escalation study in children and adolescents from 2 to less than 18 years of age with NPC (and adults up to 25 years of age), to assess safety, tolerability and pharmacokinetics of 2-hydroxypropyl-ß-cyclodextrin (HP-ß-CD), (Study 13-CH-0001). **Study 3** Randomized, double-blind, sham-controlled, 3-part, efficacy and safety trial of HP-ß-CD, in patients with NPC disease from 4 to less than 18 years of age (and adults up to 21 years of age), (Study VTS301).

		Study 4 Study deleted during procedure EMEA-001866-PIP01-15-M02 **Study 5** Study deleted during procedure EMEA-001866-PIP01-15-M02 **Study 6** Open-label, safety, efficacy and pharmacokinetic study of HP-ß-CD in children from birth to less than 4 years of age with NPC disease, (Study VTS-270-303). **Study 7** Open-label, safety and pharmacokinetic trial of HP-ß-CD, administered by the lumbar intrathecal route (IT) via an implanted delivery device, in children from birth to less than 4 years of age with NPC disease **Study 8** Study deleted during procedure EMEA-001866-PIP01-15-M02
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:	Yes
Date of completion of the paediatric investigation plan:	By April 2022
Deferral for one or more measures contained in the paediatric investigation plan:	Yes