EMA/247011/2017

European Medicines Agency decision

P/0115/2017

of 21 April 2017

on the acceptance of a modification of an agreed paediatric investigation plan for doravirine (EMEA-001676-PIP01-14-M01) 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 doravirine (EMEA-001676-PIP01-14-M01) 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 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/0139/2016 issued on 20 May 2016,

Having regard to the application submitted by Merck Sharp & Dohme (Europe), Inc. on 3 January 2017 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 24 March 2017, 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.
[2] OJ L 136, 30.4.2004, p. 1.

Has adopted this decision:

Article 1

Changes to the agreed paediatric investigation plan for doravirine, tablet, granules, oral 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 Merck Sharp & Dohme (Europe), Inc., Clos du Lynx, 5, B-1200 Brussels, Belgium.

EMA/PDCO/44239/2017
London, 24 March 2017

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

EMEA-001676-PIP01-14-M01

Scope of the application

Active substance(s):

Doravirine

Condition(s):

Treatment of human immunodeficiency virus type 1 (HIV-1) infection

Pharmaceutical form(s):

Tablet

Granules

Route(s) of administration:

Oral use

Name/corporate name of the PIP applicant:

Merck Sharp & Dohme (Europe), Inc.

Basis for opinion

Pursuant to Article 22 of Regulation (EC) No 1901/2006 as amended, Merck Sharp & Dohme (Europe), Inc. submitted to the European Medicines Agency on 3 January 2017 an application for modification of the agreed paediatric investigation plan with a deferral as set out in the European Medicines Agency's decision P/0139/2016 issued on 20 May 2016.

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

The procedure started on 24 January 2017.

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 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 human immunodeficiency virus type 1 (HIV-1) infection

2.1.1. Indication(s) targeted by the PIP

Antiretroviral therapy, in combination with other antiretroviral agents, for the treatment of HIV-1 infection in children aged from birth to 18 years

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)

Tablet

Granules

2.1.4. Measures

Area	Number of measures	Description
Quality-related studies	1	**Study 1** Development of an age-appropriate oral solid dosage form.
Non-clinical studies	2	**Study 2** Dose range-finding juvenile toxicity study **Study 3** Definitive juvenile toxicity study
Clinical studies	6	**Study 4** Open-label two period trial to evaluate pharmacokinetic, safety, and activity, of doravirine and doravirine fixed dose combination with lamivudine and tenofovir disoproxil fumarate in adolescents from 12 to less than 18 years of age weighing at least 35 kg with HIV-1 infection.

		Study 5 Open-label two period trial to evaluate pharmacokinetic, safety and activity, of doravirine and doravirine fixed dose combination with lamivudine and tenofovir disoproxil fumarate in children and adolescents from 2 years of age and less than the minimum weight threshold for administration of the adult dose as identified in study 4 with HIV-1 infection. **Study 6** Open-label two period trial to evaluate pharmacokinetic, safety and activity, of doravirine in paediatric patients from 1 month to less than 2 years of age with HIV-1 infection. **Study 7** Open-label two period trial to evaluate pharmacokinetic, safety and activity of doravirine in children from birth to six weeks of age born to HIV infected mothers. **Study 8** Relative bioavailability study with doravirine oral granules in healthy adults. **Study 9** Relative bioavailability study with doravirine age appropriate granules in healthy adults.
Extrapolation, modelling and simulation studies	1	**Study 10** Pharmacokinetic modelling and simulation study to evaluate appropriate dose of doravirine in children from birth to less than 18 years of age with HIV infection
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 2024
Deferral for one or more measures contained in the paediatric investigation plan:	Yes