EMA/106209/2017

European Medicines Agency decision

P/0074/2017

of 17 March 2017

on the agreement of a paediatric investigation plan and on the granting of a deferral and on the granting of a waiver for lamivudine / dolutegravir (EMEA-001940-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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An agency of the European Union

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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 application submitted by ViiV Healthcare UK Limited on 16 March 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 27 January 2017, in accordance with Article 18 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 lamivudine / dolutegravir, film-coated tablets, age appropriate formulation, oral 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 lamivudine / dolutegravir, film-coated tablets, age appropriate formulation, oral 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 lamivudine / dolutegravir, film-coated tablets, age appropriate formulation, oral 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 ViiV Healthcare UK Limited, 980 Grest West Road, TW8 9GS - Brentford, United Kingdom.

EMA/PDCO/756674/2016
London, 27 January 2017

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

EMEA-001940-PIP01-16

Scope of the application

Active substance(s):

Lamivudine / dolutegravir

Condition(s):

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

Pharmaceutical form(s):

Film-coated tablets

Age appropriate formulation

Route(s) of administration:

Oral use

Name/corporate name of the PIP applicant:

ViiV Healthcare UK Limited

Basis for opinion

Pursuant to Article 16(1) of Regulation (EC) No 1901/2006 as amended, ViiV Healthcare UK Limited submitted for agreement to the European Medicines Agency on 16 March 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 26 April 2016.

Supplementary information was provided by the applicant on 3 November 2016. 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 18 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)(c) of said Regulation, on the grounds that the specific medicinal product does not represent a significant therapeutic benefit over existing treatments for paediatric patients.

   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:

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

The waiver applies to:

- the paediatric population from birth to less than 2 years of age;
- film-coated tablet, age appropriate formulation, oral use;
- on the grounds that the specific medicinal product does not represent a significant therapeutic benefit over existing treatments.

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

Adolescents (between 12 and 18 years old);

- Lamivudine (3TC) / dolutegravir (DTG) fix-dose-combination (FDC) is to be indicated for use as a 2-drug complete regimen for the treatment of ART-naïve adolescents above 12 to less than 18 years of age with plasma HIV-1 RNA less than 500,000 c/mL, or virologically suppressed adolescents, and without known viral drug resistance to either agent.

Children (above 2 to less than 12 years old);

- DTG/3TC FDC is to be indicated for use as a 2-drug complete regimen for the treatment of virologically suppressed children above 2 to less than 12 years of age without known viral drug resistance to either agent.

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

From 2 to less than 18 years of age.

2.1.3. Pharmaceutical form(s)

Film-coated tablet

Age appropriate formulation

2.1.4. Measures

Area	Number of measures	Description
Quality-related studies	1	**Study 1** Paediatric formulation development
Non-clinical studies	0	Not applicable.
Clinical studies	2	**Study 2** Single dose, crossover, relative bioavailability study of DTG plus 3TC paediatric formulation as compared to approved DTG and 3TC formulations. **Study 3** Open-label, randomised, active controlled, non-inferiority study to evaluate the maintenance of Efficacy and Safety of dual therapy, DTG plus 3TC, in anti-retroviral therapy (ART)-experienced HIV-1-Infected Children, from 2 to less than 12 years of age who are virologically suppressed on their current anti-retroviral (ARV) regimen.
Extrapolation, modelling and simulation studies	1	**Study 4** Extrapolation of efficacy from adult to adolescents 12 to less than 18 years old weighing at least 40 kg.
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 December 2023
Deferral for one or more measures contained in the paediatric investigation plan:	Yes