

EMA/758229/2016

European Medicines Agency decision

P/0319/2016

of 2 December 2016

on the refusal of a modification of an agreed paediatric investigation plan for tapentadol (hydrochloride) (Palexia and associated names, Yantil and associated names, Tapentadol and associated names), (EMA-000325-PIP01-08-M06) in accordance with Regulation (EC) No 1901/2006 of the European Parliament and of the Council

Only the English text is authentic.

European Medicines Agency decision

P/0319/2016

of 2 December 2016

on the refusal of a modification of an agreed paediatric investigation plan for tapentadol (hydrochloride) (Palexia and associated names, Yantil and associated names, Tapentadol and associated names), (EMA-000325-PIP01-08-M06) 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 726/2004¹,

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 Agency²,

Having regard to the European Medicines Agency's decision P/48/2009 issued on 24 March 2009, the decision P/0280/2013 issued on 8 November 2013, the decision P/0095/2014 issued on 7 April 2014 and the decision P/0042/2016 issued on 26 February 2016,

Having regard to the application submitted by Grünenthal GmbH on 22 July 2016 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 14 October 2016, 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 refusal of changes to the agreed paediatric investigation plan and to the deferral.
- (2) It is therefore appropriate to adopt a decision on the refusal 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 tapentadol (hydrochloride) (Palexia and associated names, Yantil and associated names, Tapentadol and associated names), prolonged-release tablet, prolonged-release granules, oral use, including changes to the deferral 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, are hereby refused.

Article 2

This decision is addressed to Grünenthal GmbH, Zieglerstrasse 6, 52078 - Aachen, Germany.



EUROPEAN MEDICINES AGENCY
SCIENCE MEDICINES HEALTH

EMA/PDCO/528233/2016

London, 14 October 2016

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

EMA-000325-PIP01-08-M06

Scope of the application

Active substance(s):

Tapentadol (hydrochloride)

Invented name:

Palexia and associated names

Yantil and associated names

Tapentadol and associated names

Condition(s):

Treatment of chronic pain

Authorised indication(s):

See Annex II

Pharmaceutical form(s):

Prolonged-release tablet

Prolonged-release granules

Route(s) of administration:

Oral use

Name / corporate name of the PIP applicant:

Grünenthal GmbH

Information about the authorised medicinal product:

See Annex II



Basis for opinion

Pursuant to Article 22 of Regulation (EC) No 1901/2006 as amended, Grünenthal GmbH submitted to the European Medicines Agency on 22 July 2016 an application for modification of the agreed paediatric investigation plan with a deferral as set out in the European Medicines Agency's decision P/48/2009 issued on 24 March 2009, the decision P/0280/2013 issued on 8 November 2013, the decision P/0095/2014 issued on 7 April 2014 and the decision P/0042/2016 issued on 26 February 2016.

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

The procedure started on 16 August 2016.

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 refuse the changes proposed by the applicant regarding the paediatric investigation plan and the deferral.

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 remain unchanged and are set out in the Annex I.
3. The scientific conclusions and the grounds for refusal are set out in the summary report appended to this opinion.

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 chronic pain

2.1.1. Indication(s) targeted by the PIP

Treatment of chronic pain

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)

Prolonged-release tablet

Prolonged-release granules

2.1.4. Measures

Area	Number of studies	Description
Quality - related studies	1	Study 1 Development of prolonged release granules for oral use
Non-clinical studies	1	Study 2 Juvenile rat toxicity study
Clinical studies	3	Study 3 Randomised, open-label, cross-over relative bioavailability trial to compare a tapentadol multiparticulate (or alternative) formulation and the prolonged release tablets in healthy subjects. Study 4 A randomised, open-label, active-controlled, multi-centre study to evaluate the safety and the efficacy of tapentadol PR in patients from 6 to less than 18 years old suffering from pain requiring prolonged-release (PR) opioid. Study 5 Randomised open-label, active-controlled, multi-centre study to evaluate the safety and the efficacy of tapentadol PR in patients between 2 and 5 years old suffering from chronic malignant or non-malignant pain requiring opioid treatment.

Extrapolation, modelling and simulation studies	1	Study 6 Modelling and simulation study
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 August 2017
Deferral for one or more measures contained in the paediatric investigation plan:	Yes

Annex II

Information about the authorised medicinal product

Condition(s) and authorised indication(s):

1. Treatment of acute pain

Authorised indication(s):

- Relief of moderate to severe acute pain in adults, which can be adequately managed only with opioid analgesics.

2. Treatment of chronic pain

Authorised indication(s):

- Management of severe chronic pain in adults, which can be adequately managed only with opioid analgesics.

Authorised pharmaceutical form(s):

Film-coated tablet

Prolonged-release tablet

Oral solution

Authorised route(s) of administration:

Oral use