EMA/326956/2016

European Medicines Agency decision

P/0134/2016

of 20 May 2016

on the acceptance of a modification of an agreed paediatric investigation plan for cabozantinib (Cometriq) (EMEA-001143-PIP01-11-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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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/0128/2012 issued on 04 July 2012,

Having regard to the application submitted by Exelixis Inc on 1 January 2016 under Article 22 of Regulation (EC) No 1901/2006 proposing changes to the agreed paediatric investigation plan with a deferral and a waiver,

Having regard to the opinion of the Paediatric Committee of the European Medicines Agency, issued on 1 April 2016, in accordance with Article 22 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 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 cabozantinib (Cometriq), age-appropriate oral solid dosage form, capsule, hard, tablet, oral 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 Exelixis Inc, 210 East Grand Avenue, 94080-0511 South San Francisco – USA.

Done at London, 20 May 2016

For the European Medicines Agency
Zaïde Frias
Head of Division
Human Medicines Research and Development Support
(Signature on file)

EMA/PDCO/23916/2016
London, 1 April 2016

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

EMEA-001143-PIP01-11-M01

Scope of the application

Active substance(s):

Cabozantinib

Invented name:

Cometriq

Condition(s):

Treatment of malignant solid tumours

Authorised indication(s):

See Annex II

Pharmaceutical form(s):

Age-appropriate oral solid dosage form

Capsule, hard

Tablet

Route(s) of administration:

Oral use

Name/corporate name of the PIP applicant:

Exelixis Inc

Information about the authorised medicinal product:

See Annex II

Basis for opinion

Pursuant to Article 22 of Regulation (EC) No 1901/2006 as amended, Exelixis Inc submitted to the European Medicines Agency on 1 January 2016 an application for modification of the agreed paediatric investigation plan with a deferral and a waiver as set out in the European Medicines Agency's decision P/0128/2012 issued on 04 July 2012.

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

The procedure started on 2 February 2016.

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 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 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 malignant solid tumours

2.1.1. Indication(s) targeted by the PIP

- Treatment of refractory malignant solid tumours that are associated with MET, VEGFR, and/or RET pathway activation as a result of mutation, overexpression or amplification.
- Treatment of advanced or metastatic medullary thyroid cancer.

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)

Capsule, hard

Tablet

Age-appropriate formulation

2.1.4. Studies

Area	Number of studies	Description
Quality	1	Study 1: Development of an age-appropriate formulation.
Non-clinical	2	Study 2: Juvenile toxicity and toxicokinetic study. Study 3: Comprehensive paediatric non-clinical efficacy testing program.
Clinical	3	Study 4: Open-label trial to evaluate toxicity, tolerability, pharmacokinetics and pharmacodynamics of cabozantinib in children with refractory or relapsed malignant solid tumours. Study 5: Trial to evaluate relative bioavailability (in adults).

		Study 7: Open-label trial to evaluate the safety and activity of cabozantinib in children (and young adults) with a relapsed or refractory solid malignant tumour. Study 6: Randomised, double-blind, controlled, parallel-group safety and efficacy clinical trial of cabozantinib in patients aged from birth to less than 18 years with a malignant solid tumour(s) determined based on results of studies 3 and 4.

3. Follow-up, completion and deferral of PIP

Concerns on potential long term safety issues in relation to paediatric use:	Yes
Date of completion of the paediatric investigation plan:	By January 2023
Deferral for one or more studies contained in the paediatric investigation plan:	Yes

Annex II

Information about the authorised medicinal product

Condition(s) and authorised indication(s):

1. Treatment medullary thyroid carcinoma

Authorised indication(s):

- Treatment of adult patients with progressive, unresectable locally advanced or metastatic medullary thyroid carcinoma.

Authorised pharmaceutical form(s):

Capsule, hard

Authorised route(s) of administration:

Oral use