MHRA
10 South Colonnade
Canary Wharf
London
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United Kingdom

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Decision Cover Letter

Decision of the licensing authority to:

agree a paediatric investigation plan and grant a deferral and grant a waiver

MHRA-100312-PIP01-21

Scope of the Application

Active Substance(s)

Ritlectinib

Condition(s)

Treatment of alopecia areata

Pharmaceutical Form(s)

Capsule, hard; Age-appropriate oral dosage form

Route(s) of Administration

Oral use

Name / Corporate name of the PIP applicant

Pfizer Limited

Basis for the Decision

Pursuant to the Human Medicines Regulations 2012, Pfizer Limited submitted to the licensing authority on 26/01/2022 12:31 GMT an application for a Paediatric Investigation Plan

The procedure started on 24/05/2022 16:02 BST

1. The licensing authority, having assessed the application in accordance with the Human Medicines Regulations 2012, decides, as set out in the appended summary report:

to agree a paediatric investigation plan and grant a deferral and grant a waiver

2. The measures and timelines of the paediatric investigation plan are set out in the Annex I.

This decision is forwarded to the applicant, together with its annex and appendix.

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Final Decision Letter

MHRA-100312-PIP01-21

Of 16/06/2022 16:02 BST

On the adopted decision for Ritlectinib (MHRA-100312-PIP01-21) in accordance with the Human Medicines Regulations 2012.

The licensing authority, in accordance with the Human Medicines Regulations 2012, has adopted this decision:

Agreement on a paediatric investigation plan

This decision applies to a Paediatric Investigation Plan for Ritlectinib, Capsule, hard; Age-appropriate oral dosage form , Oral use .

This decision is addressed to Pfizer Limited, Ramsgate Road, Sandwich, United Kingdom, CT13 9NJ

ANNEX I

1. Waiver

1.1 Condition:

Treatment of alopecia areata The waiver applies / applied to: Paediatric Subset(s): The paediatric population from birth to less than 6 years of age Pharmaceutical form(s): Capsule, hard; Age-appropriate oral dosage form Route(s) of administration: Oral use Reason for granting waiver: on the grounds that clinical studies with the specific medicinal product cannot be expected to be of significant therapeutic benefit to or fulfil a therapeutic need of the specified paediatric subset(s).

2. Paediatric Investigation Plan:

2.1 Condition(s):

Treatment of alopecia areata (including alopecia universalis and alopecia totalis)

2.2 Indication(s) targeted by the PIP:

Treatment of severe alopecia areata

2.3 Subset(s) of the paediatric population concerned by the paediatric development:

The paediatric population from 6 years to less than 18 years of age

2.4 Pharmaceutical Form(s):

Capsule, hard; Age-appropriate oral dosage form

2.5 Studies:

Study Type	Number of Studies	Study Description
Quality Measures	1	Study 1 Development of age-appropriate oral formulation suitable for children less than 12 years of age.
Non-Clinical Studies	1	Study 2 Definitive juvenile toxicity study in rats with a 2 month recovery period.
Clinical Studies	3	Study 3 (B7981031) Open label, non-randomised, multiple once daily dose, PK/PD study in children 6 years to less than 12 years of age with severe alopecia areata. Study 4 (B7981027) Randomised, double-blind, 24-week, placebo-controlled study to evaluate the safety and efficacy of ritlecitinib in children 6 years to less than 12 years of age with severe alopecia areata. Study 5 (B7981028) Long-term, extension study to evaluate the long-term safety and long-term efficacy of ritlecitinib in children 6 years to less than 12 years of age with severe alopecia areata (AA).
Extrapolation, Modeling & Simulation Studies	4	Study 6 Population PK analysis to characterise the PK of ritlecitinib in adult and adolescent AA participants and for dose-prediction in children 6 years to less than 12 years of age. Study 7 Population PK analysis to characterise the PK of ritlecitinib in adult and paediatric AA subjects and to evaluate overall dosing recommendation in the paediatric AA population. Study 8 Longitudinal exposure-response analysis of absolute SALT (severity of alopecia tool) score to characterise the

		temporal relationship of exposure-response of ritlecitinib on scalp hair growth in AA subjects. Study 9 Extrapolation study to support the extrapolation of efficacy, safety and clinical PK data of ritlecitinib to adolescents with severe AA.
Other Studies	0	Not applicable
Other Measures	0	Not applicable

3. Follow-up, completion and deferral of a PIP:

Concerns on potential long term safety and efficacy issues in relation to paediatric use:	Yes
Date of completion of the paediatric investigation plan:	30/11/2029
Deferral of one or more studies contained in the paediatric investigation plan:	Yes