MHRA
10 South Colonnade
Canary Wharf
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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-100297-PIP01-21

Scope of the Application

Active Substance(s)

Human, recombinant, non-fucosylated IgG1k monoclonal antibody targeting OX-40 receptor on activated T cells (AMG 451)

Condition(s)

Treatment of atopic dermatitis

Pharmaceutical Form(s)

Solution for injection

Route(s) of Administration

Subcutaneous use

Name / Corporate name of the PIP applicant

Amgen Ltd.

Basis for the Decision

Pursuant to the Human Medicines Regulations 2012, Amgen Ltd. submitted to the licensing authority on 20/12/2021 09:20 GMT an application for a Paediatric Investigation Plan

The procedure started on 31/08/2022 08:03 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-100297-PIP01-21

Of 09/11/2022 08:03 GMT

On the adopted decision for Human, recombinant, non-fucosylated IgG1k monoclonal antibody targeting OX-40 receptor on activated T cells (AMG 451) (MHRA-100297-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 Human, recombinant, non-fucosylated IgG1k monoclonal antibody targeting OX-40 receptor on activated T cells (AMG 451), Solution for injection , Subcutaneous use .

This decision is addressed to Amgen Ltd., 216 Cambridge Science Park, Milton Road, Cambridge, United Kingdom, CB4 0WA

ANNEX I

1. Waiver

1.1 Condition:

Treatment of atopic dermatitis The waiver applies / applied to: Paediatric Subset(s): The paediatric population from birth to less than 6 months of age Pharmaceutical form(s): Solution for injection Route(s) of administration: Subcutaneous use Reason for granting waiver: on the grounds that the specific medicinal product is likely to be unsafe.

2. Paediatric Investigation Plan:

2.1 Condition(s):

Treatment of atopic dermatitis

2.2 Indication(s) targeted by the PIP:

Treatment of moderate to severe atopic dermatitis (AD) with or without topical corticosteroids

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

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

2.4 Pharmaceutical Form(s):

Solution for injection

2.5 Studies:

Study Type	Number of Studies	Study Description
Quality Measures	1	Study 1 Compatibility study of the solution for injection to ensure that the dosage preparation procedure and presentation is age appropriate.
Non-Clinical Studies	1	Study 2 (SBL303-238) Enhanced pre- and postnatal development reproductive toxicity study.
Clinical Studies	3	Study 3 (20210145) A randomised, double-blind, placebo-controlled, parallel group two part study to investigate the efficacy and safety of human, recombinant, non-fucosylated IgG1k monoclonal antibody targeting OX-40 receptor on activated T cells (referred to as AMG 451) monotherapy in adolescents 12 years to less than 18 years of age (and adults) with moderate to severe atopic dermatitis. Study 4 (20210261) A randomised, double-blind, placebo-controlled study to investigate the safety and efficacy of AMG 451 in combination with topical corticosteroids (TCS) in subjects aged 6 years to less than 12 years with moderate to severe atopic dermatitis Study 5 (20210262) A two part open label dose finding (Part A) and randomised, double blind, placebo controlled study (Part B) to investigate pharmacokinetics

		(PK), pharmacodynamics (PD), safety and efficacy, of AMG 451 in combination with topical corticosteroids (TCS) in children aged 6 months to less than 6 years, with moderate to severe atopic dermatitis.
Extrapolation, Modeling & Simulation Studies	1	Study 6 Modelling and simulation study to evaluate the use of the product in the treatment of moderate to severe atopic dermatitis in children from 6 months to less than 18 years of age with moderate to severe atopic dermatitis.
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:	31/08/2035
Deferral of one or more studies contained in the paediatric investigation plan:	Yes