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:

accept change(s) to the agreed paediatric investigation plan (MHRA-100283-PIP01-21-M02) and to the deferral

MHRA-100283-PIP01-21-M03

Scope of the Application

Active Substance(s)

USTEKINUMAB

Condition(s)

Treatment of Crohn's Disease

Pharmaceutical Form(s)

Concentrate for solution for infusion Solution for injection

Route(s) of Administration

SUBCUTANEOUS USE INTRAVENOUS USE

Name / Corporate name of the PIP applicant

Janssen-Cilag Ltd

Basis for the Decision

Pursuant to the Human Medicines Regulations 2012, Janssen-Cilag Ltd submitted to the licensing authority on 21/12/2023 12:56 GMT an application for a Modification

The procedure started on 12/02/2024 12:21 GMT

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 accept change(s) to the agreed paediatric investigation plan and to the deferral

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-100283-PIP01-21-M03

Of 04/04/2024 16:49 BST

On the adopted decision for USTEKINUMAB (MHRA-100283-PIP01-21-M03) 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 modification of a paediatric investigation plan (including modification of a waiver or deferral included in that paediatric investigation plan)

This decision applies to a Modification for USTEKINUMAB, Concentrate for solution for infusion Solution for injection , SUBCUTANEOUS USE INTRAVENOUS USE .

This decision is addressed to Janssen-Cilag Ltd, 50-100 Holmers Farm Way, High Wycombe, UNITED KINGDOM, HP12 4EG

ANNEX I

1. Waiver

1.1 Condition:

Treatment of Crohn's disease The waiver applies / applied to: Paediatric Subset(s): The paediatric population from birth to less than 2 years of age Pharmaceutical form(s): Concentrate for solution for infusion Solution for injection Route(s) of administration: SUBCUTANEOUS USE INTRAVENOUS USE Reason for granting waiver: 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(s):

Treatment of Crohn's disease

2.2 Indication(s) targeted by the PIP:

Treatment of Crohn's disease

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

The paediatric population from 2 years to less than 18 years of age

2.4 Pharmaceutical Form(s):

Concentrate for solution for infusion Solution for injection

2.5 Studies:

Study Type	Number of Studies	Study Description
Quality Measures	0	Study 1 Deleted during procedure EMEA-000311-PIP04-13-M03.
Non-Clinical Studies	0	Not applicable.
Clinical Studies	3	Study 2 (CNTO1275CRD1001) Randomised, double-blind, pharmacokinetic (PK) study of intravenous (IV) ustekinumab induction followed by subcutaneous (SC) ustekinumab maintenance in children and adolescents 2 to less than 18 years (6 to less than 18 years in the EU) with moderately to severely active Crohn's disease (CD) who have had an inadequate response and/or intolerance to conventional therapies. Study 3 (CNTO1257CRD3004) Open-label single administration of IV induction dose of ustekinumab followed by a randomised, double-blind, 2-arm study of two different SC ustekinumab maintenance dose regimens to assess the pharmacokinetics, safety and clinical response in children and adolescents 2 to less than 18 years with moderately to severely active Crohn’s disease who have had an inadequate response and/or intolerance to biologic therapy and/or conventional therapies.

		Study 7 (added during procedure MHRA-100283-PIP01-21-M02) Retrospective, single-arm, non-interventional, observational, real world evidence study to evaluate the effectiveness of ustekinumab by estimating clinical remission at week 52 since initiation of ustekinumab in children and adolescents from 2 years to less than 18 years of age with moderately to severely active Crohn's disease.
Extrapolation, Modeling & Simulation Studies	3	Study 4 (added during procedure EMEA-000311-PIP04-13-M01) Population PK model. Study 5 (added during procedure EMEA-000311-PIP04-13-M01) Exposure-response model. Study 6 (added during procedure EMEA-000311-PIP04-13-M01) Analysis of internal and literature data to support the assumptions of similarity of disease, treatment effects, and exposure-response relationship between paediatric and adult subjects with Crohn's disease (CD).
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:	31/01/2026
Deferral of one or more studies contained in the paediatric investigation plan:	Yes