MHRA
10 South Colonnade
Canary Wharf
London
E14 4PU
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-100130-PIP01-21-M02) and to the deferral

MHRA-100130-PIP01-21-M03

Scope of the Application

Active Substance(s)

BUPIVACAINE

Condition(s)

Postsurgical analgesia

Pharmaceutical Form(s)

Prolonged release dispersion for injection

Route(s) of Administration

INFILTRATION USE PERINEURAL USE

Name / Corporate name of the PIP applicant

Pacira Limited

Basis for the Decision

Pursuant to the Human Medicines Regulations 2012, Pacira Limited submitted to the licensing authority on 25/11/2024 09:52 GMT an application for a Modification

The procedure started on 13/01/2025 13:52 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-100130-PIP01-21-M03

Of 01/08/2025 08:38 BST

On the adopted decision for BUPIVACAINE (MHRA-100130-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 BUPIVACAINE, Prolonged release dispersion for injection , INFILTRATION USE; PERINEURAL USE .

This decision is addressed to Pacira Limited, C/O Hiller Hopkins LLP, Radius House, First Floor, 51 Clarendon Road, Watford, UNITED KINGDOM, WD17 1HP

ANNEX I

1. Waiver

1.1 Condition:

Not Applicable

2. Paediatric Investigation Plan:

2.1 Condition(s):

Postsurgical analgesia

2.2 Indication(s) targeted by the PIP:

Local or regional analgesia when administered into the surgical site or as a nerve block in children from birth to less than 18 years of age.

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

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

2.4 Pharmaceutical Form(s):

Prolonged release dispersion for injection

2.5 Studies:

Study Type	Number of Studies	Study Description
Quality Measures	0	Not applicable.
Non-Clinical Studies	0	Not applicable.
Clinical Studies	3	Study 1 (402-C-319) A multicentre study to evaluate the pharmacokinetics and safety of liposomal bupivacaine (hereinafter: Exparel) for postsurgical analgesia in paediatric subjects aged 6 to less than 17 years of age. Study 2 (402-C-WI2) Multicentre, double-blind, randomised, bupivacaine-controlled study to evaluate the pharmacokinetics and safety of Exparel for postsurgical analgesia in subjects from birth to less than 6 years undergoing cardiac surgery. Study 3 (402-C-NB1) Deleted during procedure MHRA-100130-PIP01-21-M03 Study 4 (402-C-NB2) Deleted during procedure MHRA-100130-PIP01-21-M03 Study 5 (402-C-WI3) (Added during procedure MHRA-100130-PIP01-21-M02) Multicentre, double blind, randomised, bupivacaine-controlled study to evaluate the safety and efficacy of Exparel for postoperative analgesia in paediatric subjects from birth to less than 6 years undergoing cardiac surgery.
Extrapolation, Modeling & Simulation Studies	0	Not applicable.
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:	No
Date of completion of the paediatric investigation plan:	31/03/2031
Deferral of one or more studies contained in the paediatric investigation plan:	Yes