16 November 2016
EMA/PDCO/102084/2009
Inspections, Human Medicines Pharmacovigilance and Committees Division

PDCO monthly report of opinions on paediatric investigation plans and other activities

8-11 November 2016

Opinions on paediatric investigation plans

The Paediatric Committee (PDCO) adopted opinions agreeing paediatric investigation plans (PIPs) for the following medicines:

- Octenidine dihydrochloride, EMEA-001384-PIP01-12, from Schülke & Mayr GmbH, for skin disinfection;
- Dexamethasone / Complex of povidone and iodine, EMEA-001936-PIP01-16, from Shire Pharmaceuticals Ireland Ltd, for the treatment of infectious conjunctivitis;
- Eculizumab, EMEA-000876-PIP03-14, from Alexion Europe SAS, for the treatment of neuromyelitis optica spectrum disorders;
- Galcanezumab, EMEA-001860-PIP04-16, from Eli Lilly and Company Limited, for the prevention of cluster headache;
- synthetic surfactant protein B analogue / synthetic surfactant protein C analogue / 1-palmitoyl-2-oleoyl-sn-glycero-3-phosphoglycerol sodium salt / dipalmitoylphosphatidylcholine, EMEA-001780-PIP01-15, from Chiesi Farmaceutici SpA, for the treatment of respiratory distress syndrome (RDS);
- Pimavanserin, EMEA-001688-PIP03-16, from ACADIA Pharmaceuticals Inc., for the treatment of schizophrenia and other psychotic disorders;
- Betrixaban, EMEA-001834-PIP02-16, from Portola Pharma UK Limited, for the prevention of venous thromboembolism;
- Peramivir, EMEA-001856-PIP02-16, from BioCryst UK Ltd., for the treatment of influenza.

A PIP sets out a programme for the development of a medicine in the paediatric population. The PIP aims to generate the necessary quality, safety and efficacy data through studies to support the authorisation of the medicine for use in children of all ages. These data have to be submitted to the European Medicines Agency, or national competent authorities, as part of an application for a marketing authorisation for a new medicine, or for one covered by a patent. In some cases, a PIP may include a waiver of the studies in one or more paediatric subsets, or a deferral.

30 Churchill Place ● Canary Wharf ● London E14 5EU ● United Kingdom
Telephone +44 (0)20 3660 6000 **Facsimile** +44 (0)20 3660 5510
Send a question via our website www.ema.europa.eu/contact

Opinions on product-specific waivers

The PDCO adopted positive opinions for product-specific waivers, recommending that the obligation to submit data obtained through clinical studies with children be waived in all subsets of the paediatric population, for the following medicines:

- Ibuprofen, EMEA-002017-PIP01-16, from Strides Shasun Limited, for the treatment of febrile disorders and treatment of pain;
- Netarsudil, EMEA-002037-PIP01-16, from Aerie Pharmaceuticals Ireland, Ltd., for the treatment of glaucoma;
- DNA plasmid encoding HPV type 18 consensus E6 and E7 proteins (pGX3002) / DNA plasmid encoding HPV type 16 consensus E6 and E7 proteins (pGX3001), EMEA-002022-PIP01-16, from Inovio Pharmaceuticals Inc., for the treatment of squamous intraepithelial lesions of the cervix caused by HPV types 16 and 18;
- Botulinum toxin, Type A, EMEA-002038-PIP01-16, from Evolus Inc., for the treatment of glabellar lines.

The PDCO adopted 1 opinion on the **refusal** of a request for waiver for:

- palonosetron / fosnetupitant, EMEA-001198-PIP02-16, from Helsinn Birex Pharmaceuticals Limited, for the prevention of chemotherapy-induced nausea and vomiting.

Waivers can be issued if there is evidence that the medicine concerned is likely to be ineffective or unsafe in the paediatric population, or that the disease or condition targeted occurs only in adult populations, or that the medicine, or the performance of trials, does not represent a significant therapeutic benefit over existing treatments for paediatric patients.

Opinions on modifications to an agreed PIP

The PDCO also adopts, every month, opinions on modifications to an agreed PIP, which can be requested by the applicant when the plan is no longer appropriate or when there are difficulties that render the plan unworkable. The PDCO adopted positive opinions, agreeing change(s), for the following products:

- Estetrol / Drospirenone, EMEA-001332-PIP01-12-M02, from Estetra SPRL, for the prevention of pregnancy;
- sacubitril / valsartan, EMEA-000316-PIP02-11-M03, from Novartis Europharm Ltd., for the treatment of heart failure;
- Liquid extract ethanolic 30 per cent (w/w) of Allium cepa L. (fresh bulb) and Citrus limon (L.) Burm. f. (fresh fruit), Paullinia cupana Kunth, Theobroma cacao L., EMEA-001835-PIP01-15-M01, from Legacy Healthcare, for the treatment of alopecia;
- Talimogene laherparepvec, EMEA-001251-PIP01-11-M02, from Amgen Europe B.V., for the treatment of solid malignant non-CNS tumours;
- Allantoin, EMEA-001590-PIP01-13-M03, from Scioderm, Inc., for the treatment of epidermolysis bullosa;
- Finerenone, EMEA-001623-PIP01-14-M01, from Bayer Pharma AG, for the treatment of chronic kidney disease;

- Ceftaroline fosamil, EMEA-000769-PIP01-09-M06, from AstraZeneca AB, for the treatment of community-acquired pneumonia and treatment of complicated skin and soft tissue infections;
- Metreleptin, EMEA-001701-PIP01-14-M01, from Aegerion Pharmaceuticals Ltd, for the treatment of lipodystrophy;
- Deferiprone, EMEA-001126-PIP01-10-M02, from Consorzio per Valutazioni Biologiche e Farmacologiche, for the treatment of chronic iron overload.

The PDCO adopted 1 opinion on the **refusal** of modifications to an agreed PIP for the following application:

- Rubidium Rb-82 Chloride, EMEA-000882-PIP03-11-M02, from Jubilant DraxImage Inc., for the visualization of myocardial perfusion for diagnostic purposes;

Opinion on compliance check

The PDCO adopted a positive opinion on (full) compliance check for:

- Nilotinib, EMEA-C-000290-PIP01-08-M04, from Novartis Europharm Ltd., for the treatment of chronic myeloid leukaemia.

A compliance check is performed to verify that all the measures agreed in a PIP and reflected in the Agency's decision have been conducted in accordance with the decision, including the agreed timelines. Full compliance with all studies/measured contained in the PIP is one of several prerequisites for obtaining the rewards and incentives provided for in Articles 36 to 38 of the Paediatric Regulation.

Before the submission of a request for a compliance check, applicants are encouraged to consult the Agency's Procedural advice for validation of a new marketing authorisation application or extension/variation application and compliance check with an agreed PIP.

Withdrawals

The PDCO noted that 2 applications were withdrawn during the late stages of the evaluation (30 days or less before completion of the procedure).

Interaction with Enpr-EMA

In the framework of the planned exchange of information and interaction between PDCO and Enpr-EMA, some national clinical research networks (from France, Finland, Norway and United Kingdom) were invited to PDCO to present their activities and their views on how to maximize synergies in promoting paediatric research.

The PDCO congratulated the representatives of the networks for their excellent work and discussed options to improve a mutual interchange of information. Other members of the Enpr-EMA will be invited to PDCO throughout 2017.

Committee interactions with the Formulations Working Group

On 11 November 2016 the PDCO and FWG Members discussed ways to improve the interaction between the two groups and highlighted relevant formulation-specific topics on which the groups should concentrate their attention.

Strategic Review and Learning Meeting

On 19-21 October 2016, the PDCO together with the Committee for Medicinal Products for Human Use (CHMP) held a joint Strategic Review and Learning Meeting (SRLM) organised by the Belgian Federal Agency for Medicines and Health Products (FAMHP), under the auspices of the Slovak Presidency of the Council of the European Union. The PDCO and CHMP discussed proposals for improving inter-committee interactions, extrapolation case studies, work plans for 2017 and other topics relevant for the optimisation of their work.

Other matters

The next meeting of the PDCO will be held on 13-16 December 2016.

– END –

Notes:

1. As of 26 January 2009, pharmaceutical companies that submit an application for a marketing authorisation for a medicinal product, or those that submit an application for an extension of indication, a new route of administration, or a new pharmaceutical form of a medicinal product already authorised in the European Union, have to provide either the results of studies in children conducted in accordance with an approved PIP, or an Agency's decision on a waiver or on a deferral.

2. PDCO opinions on PIPs and waivers are transformed into Agency's decisions within the timeframe laid down by the Paediatric Regulation (Regulation (EC) No 1901/2006, as amended). The decisions can be found on the Agency's website at: http://www.ema.europa.eu/ema/index.jsp?curl=pages/medicines/landing/pip_search.jsp&murl=menus/medicines/medicines.jsp&mid=WC0b01ac058001d129

3. More information about the PDCO and the Paediatric Regulation is available in the Regulatory section of the Agency's website: http://www.ema.europa.eu/ema/index.jsp?curl=pages/regulation/general/general_content_000023.jsp&murl=menus/regulations/regulations.jsp&mid=WC0b01ac05800240cd

4. This meeting report, together with other information on the work of the Agency's, can be found on the Agency's website: http://www.ema.europa.eu

Enquiries to: AskEMA
(http://www.ema.europa.eu/ema/index.jsp?curl=pages/about_us/landing/ask_ema_landing_page.jsp&mid=)