13 November 2020
EMA/PDCO/615662/2020
Human Medicines Division

Paediatric Committee (PDCO)

Minutes for the meeting on 10-13 November 2020

Chair: Koenraad Norga – Vice-Chair: Sabine Scherer

10 November 2020, 14:00- 19:00, Virtual meeting

11 November 2020, 08:30- 19:00, Virtual meeting

12 November 2020, 08:30- 19:00, Virtual meeting

13 November 2020, 08:30- 13:00, Virtual meeting

Disclaimers

Some of the information contained in this set of minutes is considered commercially confidential or sensitive and therefore not disclosed. With regard to intended therapeutic indications or procedure scopes listed against products, it must be noted that these may not reflect the full wording proposed by applicants and may also vary during the course of the review. Additional details on some of these procedures will be published in the PDCO Committee meeting reports (after the PDCO Opinion is adopted), and on the Opinions and decisions on paediatric investigation plans webpage (after the EMA Decision is issued).

Of note, this set of minutes is a working document primarily designed for PDCO members and the work the Committee undertakes.

Further information with relevant explanatory notes can be found at the end of this document.

Note on access to documents

Some documents mentioned in the minutes cannot be released at present following a request for access to documents within the framework of Regulation (EC) No 1049/2001 as they are subject to on-going procedures for which a final decision has not yet been adopted. They will become public when adopted or considered public according to the principles stated in the Agency policy on access to documents (EMA/127362/2006).

Official address Domenico Scarlattilaan 6 ● 1083 HS Amsterdam ● The Netherlands
Address for visits and deliveries Refer to www.ema.europa.eu/how-to-find-us
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1. Introductions

1.1. Welcome and declarations of interest of members, alternates and experts

The Chairperson opened the meeting by welcoming all participants. Due to the current coronavirus (COVID-19) outbreak, and the associated EMA Business Continuity Plan (BCP), the meeting was held remotely.

In accordance with the Agency's policy on handling of declarations of interests of scientific Committees' members and experts, based on the declarations of interest submitted by the Committee members, alternates and experts and based on the topics in the agenda of the current meeting, the Committee Secretariat announced the restricted involvement of some meeting participants in upcoming discussions as included in the pre-meeting list of participants and restrictions.

Participants in this meeting were asked to declare any changes, omissions or errors to their declared interests and/or additional restrictions concerning the matters for discussion. No new or additional interests or restrictions were declared.

Discussions, deliberations and voting took place in full respect of the restricted involvement of Committee members and experts in line with the relevant provisions of the Rules of Procedure and as included in the list of participants. All decisions taken at this meeting were made in the presence of a quorum of members (i.e. 17 or more members were present in the room). All decisions, recommendations and advice were agreed by consensus, unless otherwise specified. Due to restricted involvement, the Chair Koenraad Norga was replaced by the Vice-Chair Sabine Scherer for the discussion on agenda topic 3.3.32.

1.2. Adoption of agenda

PDCO agenda for 10-13 November 2020

The agenda of the PDCO meeting 10-13 November 2020 was adopted.

1.3. Adoption of the minutes

PDCO minutes for 13-16 October 2020

The minutes of the PDCO meeting 13-16 October 2020 were adopted and will be published on the EMA website.

2. Opinions

Disclosure of some information related to this section cannot be released at the present time as it is deemed to contain commercially confidential information.

2.1. Opinions on Products

2.1.1. Venglustat - Orphan - EMEA-001716-PIP05-20

Genzyme Europe B.V.; Polycystic kidney, autosomal dominant / For long term treatment to slow the progression of cysts development in paediatric patients from 12 years to <18 years old with autosomal dominant polycystic kidney disease

Day 120 opinion

Endocrinology-Gynaecology-Fertility-Metabolism / Uro-nephrology

Summary of committee discussion:

The PDCO's view expressed at D90 was endorsed and the PDCO recommends granting a paediatric investigation plan for venglustat for the paediatric population from 12 years of age to less than 18 years of age for the condition treatment of autosomal dominant polycystic kidney disease with a deferral and a waiver for the age subset based 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).

2.1.2. Etranacogene dezaparvovec - Orphan - EMEA-002722-PIP01-19

uniQure biopharma B.V.; Treatment of Haemophilia B

Day 120 opinion

Haematology-Hemostaseology

Summary of committee discussion:

The PDCO discussed outstanding issues.

In conclusion, a positive opinion was made at Day 120 for a PIP for etranacogene dezaparvovec in the condition, treatment of haemophilia B, as well as for a deferral.

2.1.3. Etrasimod L-arginine - EMEA-002713-PIP01-19

Arena Pharmaceuticals, Inc.; Treatment of ulcerative colitis

Day 120 opinion

Gastroenterology-Hepatology

Summary of committee discussion:

The PDCO discussed this procedure on D120. The applicant's responses to the D90 issues were in general considered acceptable. A positive opinion for the proposed product was adopted with a deferral for the treatment of a subset of children with ulcerative colitis. A waiver on the grounds lack of significant benefit was adopted for a subset of children.

2.1.4. Olinciguat - EMEA-002759-PIP01-19

Cyclerion Therapeutics Inc.; Treatment of Sickle Cell Disease (SCD)

Day 120 opinion

Haematology-Hemostaseology

Summary of committee discussion:

The Committee acknowledged the clarification received from the applicant after D90.
In conclusion, the PDCO adopted a positive opinion for olinciguat for the treatment of sickle cell disease with a deferral and a waiver for a subset of children on the ground of the disease not occurring in this age group, and in another subset of children on the ground of lack of safety and efficacy.

2.1.5. Fully human IgG1 RB-1 YTE anti-RSV F monoclonal antibody (MK-1654) - EMEA-002755-PIP01-19

Merck Sharp & Dohme (Europe), Inc.; Prevention of lower respiratory tract infection caused by respiratory syncytial virus

Day 120 opinion

Infectious Diseases

Summary of committee discussion:

Based on the final discussion during the November PDCO plenary meeting taking all information into account which have been provided following the PDCO discussion in October 2020, the PDCO came to an agreement on a PIP opinion with the applicant referring to a fully human monoclonal antibody for prophylaxis of respiratory syncytial virus (RSV) infection.

2.1.6. (R)-2-(1-(6-Amino-5-chloropyrimidine-4- carboxamido)ethyl)-N-(5-chloro-4-(trifluoromethyl)pyridin- 2-yl)thiazole-5-carboxamide - EMEA-002763-PIP01-20

DOT Therapeutics-1 Inc; Paediatric low grade glioma / Relapsed or refractory paediatric low grade glioma in adolescents and children 6 months of age and older

Day 120 opinion

Oncology

Summary of committee discussion:

The PDCO took note of the additional information received, which could be followed by the committee.
To conclude, the PDCO agreed to a PIP for (R)-2-(1-(6-Amino-5-chloropyrimidine-4-carboxamido)ethyl)-N-(5-chloro-4-(trifluoromethyl)pyridin- 2-yl)thiazole-5-carboxamide for the condition of treatment of paediatric low grade glioma, including a waiver for a subset of patients based on the ground of lack of significant therapeutic benefit and a deferral on own motions which applies to the pivotal study in patients with newly diagnosed low grade glioma only.

2.1.7. Bisoprolol (fumarate) / ramipril - EMEA-002860-PIP01-20

Egis Pharmaceuticals PLC; Essential hypertension

Day 60 opinion

Cardiovascular Diseases

Summary of committee discussion:

Based on the assessment of this application and further discussions at the Paediatric Committee, the PDCO agrees with the applicant's request for a waiver. The PDCO recommends granting a waiver for bisoprolol fumarate / ramipril for all subsets of the paediatric population (0 to less than 18 years of age) in the condition "treatment of hypertension" based on lack of significant therapeutic benefit over existing treatments. The PDCO emphasises that the granting of a waiver for the condition mentioned above should not prevent the applicant from considering a development in the paediatric population in indications where there is a paediatric need.

2.1.8. Dapagliflozin - EMEA-000694-PIP06-20

AstraZeneca AB; Prevention of hospitalisation for heart failure and cardiovascular death in adults who have had a myocardial infarction

Day 60 opinion

Cardiovascular Diseases

Summary of committee discussion:

Based on the assessment of this application and in line with the day 30 PDCO discussion, the PDCO adopted a positive Opinion on a full waiver in children from birth to less than 18 years of age, on the grounds of likely lack of safety.
PDCO agreed that treatment of ischaemic heart disease would be the most suitable condition wording, considering also that 'ischaemic heart disease' is an incurable disorder, and any treatment, in fact, aims at prevention of worsening.

2.1.9. Ziltivekimab - EMEA-002840-PIP01-20

Novo Nordisk A/S; Prevention of cardiovascular events in patients with atherosclerosis

Day 60 opinion

Cardiovascular Diseases

Summary of committee discussion:

Based on the assessment of this application and further discussions at the Paediatric Committee, the PDCO agrees with the applicant's request for a waiver. The PDCO recommends granting a waiver for ziltivekimab for all subsets of the paediatric population (0 to 18 years of age) in the condition of prevention of cardiovascular events in patients with atherosclerosis.
The PDCO emphasises that the granting of a waiver for the condition mentioned above should not prevent the applicant from considering a development in the paediatric population in indications where there is a paediatric need. In principle according to the

Paediatric Regulation, incentives for the development for use in the paediatric population are available even if a waiver has been granted in another condition.

2.1.10. Efgartigimod alfa - EMEA-002597-PIP03-20

argenx BV; Treatment of pemphigus

Day 60 opinion

Dermatology

Summary of committee discussion:

Based on the assessment of this application and further discussions at the Paediatric Committee, the PDCO agrees with the applicant's request for a waiver. The PDCO recommends granting a waiver for efgartigimod alfa for all subsets of the paediatric population (from birth to 18 years of age) in the condition of "treatment of pemphigus", on the grounds that the specific medicinal product is likely to be unsafe in a subset of children and on the grounds that the specific medicinal product does not represent a significant therapeutic benefit as clinical studies are not feasible in a subset of children.
The PDCO emphasises that the granting of a waiver for the condition mentioned above should not prevent the applicant from considering a development in the paediatric population in indications where there is a paediatric need. In principle according to the Paediatric Regulation, incentives for the development for use in the paediatric population are available even if a waiver has been granted in another condition.

2.1.11. Obinutuzumab - Orphan - EMEA-001207-PIP03-20

Roche Registration GmbH; Treatment of membranous glomerulonephritis also known as membranous nephropathy (MN)

Day 60 opinion

Immunology-Rheumatology-Transplantation

Summary of committee discussion:

The PDCO in November 2020 reiterated the points already agreed at D30. The PDCO highlighted that there is a paediatric need in adolescent patients with primary membranous nephropathy (pMN). Therefore, a request for a waiver for all subset of the paediatric population cannot be granted and the applicant shall present a proposal to address this in line with the comments provided in the Summary Report. Based on the pharmacological rationale and subject to further discussion, this does not need to be necessarily restricted only to patients with pMN.
Therefore, the PDCO adopted a negative Opinion on the request of a waiver for all subset of the paediatric population for treatment of glomerulonephritis and nephrotic syndrome.

2.1.12. N-[(1R)-1-(1H-indol-3-ylmethyl)pentyl]-2-(4-methylpiperazin-1-yl)thiazole-5-carboxamide - EMEA-002866-PIP01-20

UCB Pharma S.A.; Treatment of idiopathic Parkinson's disease (PD)

Day 60 opinion

Neurology

Summary of committee discussion:

Based on the assessment of this application and further discussions at the Paediatric Committee including contributions of external expert(s), the PDCO agrees with the applicant's request for a waiver. The PDCO recommends granting a waiver for N-[(1R)-1-(1H-indol-3-ylmethyl)pentyl]-2-(4-methylpiperazin-1-yl)thiazole-5-carboxamide for all subsets of the paediatric population (from birth to less than 18 years of age) in the condition of treatment of Parkinson's disease (PD) on the grounds, that the medicinal product is likely to be ineffective in all of the paediatric population.
The PDCO emphasises that the granting of a waiver for the condition mentioned above should not prevent the applicant from considering a development in the paediatric population in indications where there is a paediatric need. In principle according to the Paediatric Regulation, incentives for the development for use in the paediatric population are available even if a waiver has been granted in another condition.

2.1.13. Tauroursodeoxycholic acid / (Sodium) phenylbutyrate - Orphan - EMEA-002876-PIP01-20

Drug Development and Regulation SL; Treatment of Amyotrophic Lateral Sclerosis

Day 60 opinion

Neurology

Summary of committee discussion:

Based on the assessment of this application and further discussions at the Paediatric Committee, the PDCO agrees with the applicant's request for a waiver. The PDCO recommends granting a waiver for tauroursodeoxycholic acid / Sodium phenylbutyrate for all subsets of the paediatric population (0 to 18 years of age) in the condition of treatment of amyotrophic lateral sclerosis.
The committee agreed on a waiver due to lack of significant therapeutic benefit because clinical studies were deemed to not be feasible.
The PDCO emphasises that the granting of a waiver for the condition mentioned above should not prevent the applicant from considering a development in the paediatric population in indications where there is a paediatric need.

2.1.14. Delolimogene mupadenorepvec - Orphan - EMEA-002864-PIP01-20

Lokon Pharma AB; Colorectal cancer / Ovarian cancer / Pancreatic cancer / Malignant melanoma / Biliary cancer

Day 60 opinion

Oncology

Summary of committee discussion:

The PDCO discussed this procedure at Day 60 during the November 2020 plenary meeting and confirmed all the conclusions reached at Day 30.
The PDCO adopted positive Opinion for a waiver for delolimogene mupadenorepvec for all subsets of the paediatric population (from birth to less than 18 years of age) in the

condition of treatment of pancreatic cancer on the grounds that pancreatic cancer does not occur in the paediatric population.
The PDCO emphasises that the granting of a waiver for the condition mentioned above should not prevent the applicant from considering a development in the paediatric population in indications where there is a paediatric need. In principle according to the Paediatric Regulation, incentives for the development for use in the paediatric population are available even if a waiver has been granted in another condition.

2.1.15. Tipifarnib - EMEA-002871-PIP01-20

Kura Oncology, Inc.; Treatment of HRAS mutant head and neck squamous cell carcinoma ('Malignant Neoplasms of Head, Face, and Neck')

Day 60 opinion

Oncology

Summary of committee discussion:

The PDCO re-discussed this product in line with the outcome conclusion made at the D30 discussion.
Based on the assessment of this application and further discussions at the Paediatric Committee, the PDCO agrees with the applicant's request for a waiver. The PDCO recommends granting a waiver for tipifarnib for all subsets of the paediatric population (0 to 18 years of age) in the condition of 'treatment of head and neck epithelial malignant neoplasms', because the disease does not occur in children.
The PDCO emphasises that the granting of a waiver for the condition mentioned above should not prevent the applicant from considering a development in the paediatric population in indications where there is a paediatric need. In principle according to the Paediatric Regulation, incentives for the development for use in the paediatric population are available even if a waiver has been granted in another condition.

2.1.16. Sulbactam / durlobactam - EMEA-002807-PIP01-20

Entasis Therapeutics Inc.; Treatment of infections due to organisms of the Acinetobacter baumannii-calcoaceticus complex / Treatment of infections due to Acinetobacter baumannii-calcoaceticus complex in patients with limited treatment options

Day 120 opinion

Infectious Diseases

Summary of committee discussion:

Based on the assessment of this application and in line with previous discussions at the Paediatric Committee, the PDCO agrees on a PIP for all subsets of the paediatric population (0 to 18 years of age) in the revised condition, treatment of infections due to organisms of the Acinetobacter baumannii-calcoaceticus complex (ABC), as well as a deferral.

2.1.17. (7S,13R) -11-fluoro-7,13-dimethyl-6,7,13,14-tetrahydro-1,15-ethenopyrazolo[4,3-f][1,4,8,10]benzoxatriaza-cyclotridecin-4(5H)-one - EMEA-002635-PIP01-19

Treatment of advanced or metastatic malignancies harbouring ALK, ROS1, or NTRK1-3

alterations

Day 120 opinion

Oncology

Summary of committee discussion:

Withdrawal request received on 02/11/2020.

2.1.18. Tabelecleucel - Orphan - EMEA-002025-PIP04-19

Atara Biotherapeutics, Inc.; Treatment of Epstein-Barr virus associated post-transplant lymphoproliferative disorder / Treatment of allogeneic haematopoietic cell transplant patients with Epstein-Barr virus associated post-transplant lymphoproliferative disease who have received one prior therapy/ Treatment of solid organ transplant patients with Epstein-Barr virus associated post-transplant lymphoproliferative disease who have received one prior therapy

Day 120 opinion

Oncology

Summary of committee discussion:

The PDCO re-discussed this procedure at Day 120 during the November 2020 plenary meeting.

The PDCO noted that all issues were resolved and adopted a positive opinion at D120 for tabelecleucel on this paediatric investigation plan with a deferral in the condition of treatment of Epstein-Barr virus associated post-transplant lymphoproliferative disorder.

2.1.19. Neisseria meningitidis serogroup B fHbp subfamily B / Neisseria meningitidis serogroup B fHbp subfamily A / Neisseria meningitidis group Y polysaccharide 1conjugated to tetanus toxoid carrier protein / Neisseria meningitidis group W-135 polysaccharide 1conjugated to tetanus toxoid carrier protein / Neisseria meningitidis group C polysaccharide 1conjugated to tetanus toxoid carrier protein / Neisseria meningitidis group A polysaccharide 1conjugated to tetanus toxoid carrier protein - EMEA-002814-PIP01-20

Invasive disease caused by Neisseria meningitidis group A, B, C, W and Y from 2 months of age

Day 120 opinion

Vaccines

Post-meeting note: This procedure was withdrawn on the 13/11/ 2020.

2.1.20. Efgartigimod alfa - EMEA-002597-PIP03-20

argenx BV; Treatment of pemphigus

Day 60 opinion

Dermatology

Summary of committee discussion:

Based on the assessment of this application and further discussions at the Paediatric Committee, the PDCO agrees with the applicant's request for a waiver. The PDCO recommends granting a waiver for efgartigimod alfa for all subsets of the paediatric population (from birth to 18 years of age) in the condition of "treatment of pemphigus", on the grounds that the specific medicinal product is likely to be unsafe in a subset of children and on the grounds that the specific medicinal product does not represent a significant therapeutic benefit as clinical studies are not feasible in a subset of children .

The PDCO emphasises that the granting of a waiver for the condition mentioned above should not prevent the applicant from considering a development in the paediatric population in indications where there is a paediatric need. In principle according to the Paediatric Regulation, incentives for the development for use in the paediatric population are available even if a waiver has been granted in another condition.

2.2. Opinions on Compliance Check

The following compliance checks have been put up for discussion and the members of the PDCO have been invited to comment on issues of possible non-compliance.

2.2.1. Human normal immunoglobulin for subcutaneous administration - EMEA-C-001853-PIP01-15-M02

Grifols Therapeutics LLC; Treatment of primary immunodeficiency

Day 60 opinion

Immunology-Rheumatology-Transplantation

Summary of committee discussion:

The PDCO adopted on 13 November 2020 an opinion confirming the compliance of all studies in the agreed paediatric investigation plan as set out in the latest Agency's Decision P/0264/2019 of 23 July 2019.

2.2.2. Isavuconazonium (sulfate) - EMEA-C1-001301-PIP02-12-M03

Basilea Pharmaceutica International Ltd.; Treatment of mucormycosis

Day 30 letter

Infectious Diseases

Summary of committee discussion:

The applicant requested to check compliance of 2 studies.
Both studies are considered to be compliant.
The compliance procedure was concluded by the EMA.

2.3. Opinions on Modification of an Agreed Paediatric Investigation Plan

2.3.1. Captopril - EMEA-001544-PIP01-13-M02

Proveca Pharma Limited; Heart failure / Treatment of heart failure in children from birth to 18 years

Day 60 opinion

Cardiovascular Diseases

Summary of committee discussion:

The applicant's response to the PDCO's Day 30 comments were noted. The outstanding issues were deemed resolved.

Based on the review of the rationale submitted by the application for modifying the agreed paediatric investigation plan, the PDCO considered that the proposed changes could be accepted.

The PDCO therefore adopted a favourable Opinion on the modification of the agreed PIP as set in the Agency's latest decision (P/0206/2014 of 8 August 2014).

The new PDCO Opinion on the modified agreed PIP supersedes the previous PDCO Opinion.

2.3.2. Fluciclovine (18F) - Orphan - EMEA-001644-PIP02-14-M02

Blue Earth Diagnostics Ireland Ltd; Diagnosis of amino acid metabolism in solid malignant tumours / Diagnosis of primary and recurrent brain tumours

Day 60 opinion

Diagnostic / Oncology

Summary of committee discussion:

The PDCO discussed the information provided by the Applicant after Day 30 regarding the paediatric clinical development .
The Committee is still of the view that fluciclovine (18F) could be of diagnostic benefit to children with glioma.
The PDCO therefore adopted a negative Opinion on the modification of the agreed PIP as set in the Agency's latest decision (P/0285/2019 of 16 August 2019).

2.3.3. Cotadutide - EMEA-002287-PIP01-17-M01

AstraZeneca AB; Treatment of Type 2 Diabetes Mellitus

Day 60 opinion

Endocrinology-Gynaecology-Fertility-Metabolism

Summary of committee discussion:

In line with the day 30 discussion, the PDCO adopted a positive Opinion on the modification of the agreed PIP as set in the Agency's latest decision (P/0235/2018 of 15/08/2018).
The new PDCO Opinion on the modified agreed PIP supersedes the previous PDCO Opinion.

2.3.4. Testosterone - EMEA-001529-PIP02-14-M03

Acerus Biopharma Inc.; Treatment of Male hypogonadism

Day 60 opinion

Endocrinology-Gynaecology-Fertility-Metabolism

Summary of committee discussion:

The PDCO discussed this modification procedure on D60.
Based on the review of the rationale submitted by the application for modifying the agreed paediatric investigation plan, the PDCO considered that the proposed changes could be accepted.
The PDCO therefore adopted a favourable Opinion on the modification of the agreed PIP as set in the Agency's latest decision (P/0236/2019 issued on 16 July 2019).
The new PDCO Opinion on the modified agreed PIP supersedes the previous PDCO Opinion.

2.3.5. Naloxegol (as naloxegol oxalate) - EMEA-001146-PIP01-11-M06

Kyowa Kirin Pharmaceutical Development Limited; Treatment of opioid-induced constipation

Day 60 opinion

Gastroenterology-Hepatology

Summary of committee discussion:

The PDCO's view expressed at Day 30 was re-discussed and endorsed.
Based on the review of the rationale submitted by the application for modifying the agreed paediatric investigation plan, the PDCO considered that the proposed changes could be accepted.
The PDCO therefore adopted a favourable Opinion on the modification of the agreed PIP as set in the Agency's latest decision (P/0381/2019 of 04/12/2019).
The new PDCO Opinion on the modified agreed PIP supersedes the previous PDCO Opinion.

2.3.6. Avatrombopag - EMEA-001136-PIP02-19-M01

Dova Pharmaceuticals Ireland Limited; Chemotherapy-induced thrombocytopenia / Treatment of chemotherapy-induced thrombocytopenia (CIT) in adult patients receiving myelosuppressive chemotherapy for solid tumours

Day 60 opinion

Haematology-Hemostaseology

Summary of committee discussion:

The PDCO discussed the proposed modification considering the update provided by the applicant on the adult development.
In conclusion, the PDCO considered that at this stage the changes of study 1, study 2, study 3 and study 4 only could be agreed.
The PDCO therefore adopted a favourable Opinion on the modification of the agreed PIP as set in the Agency's latest decision (P/0096/2020 of 18/03/2020).
The new PDCO Opinion on the modified agreed PIP supersedes the previous PDCO Opinion.

2.3.7. Eltrombopag - EMEA-000170-PIP03-13-M04

Novartis Europharm Limited; Bone Marrow Depression and Hypoplastic Anaemia / Treatment of cytopenias in paediatric patients with severe aplastic anaemia who are not receiving hematopoietic stem cell transplant

Day 60 opinion

Haematology-Hemostaseology

Summary of committee discussion:

The PDCO discussed the Request for Modification for eltrombopag for treatment of aplastic anaemia during its plenary on 13 November 2020.
The PDCO therefore adopted a favourable Opinion on the modification of the agreed PIP as set in the Agency's latest decision (P/0280/2017 of 04/10/2017).
The new PDCO Opinion on the modified agreed PIP supersedes the previous PDCO Opinion.

2.3.8. Aztreonam - Orphan - EMEA-000827-PIP01-09-M05

Gilead Sciences International Ltd.; Cystic Fibrosis with pulmonary manifestations / Treatment of initial Pseudomonas aeruginosa pulmonary infection/colonisation in patients with Cystic Fibrosis / Treatment of chronic Pseudomonas aeruginosa pulmonary infection/ colonisation in patients with Cystic Fibrosis

Day 60 opinion

Infectious Diseases

Summary of committee discussion:

Based on the review of the rationale submitted by the application for modifying the agreed paediatric investigation plan, the PDCO considered that the proposed changes could be accepted.
The PDCO therefore adopted a positive Opinion on the modification of the agreed PIP as set in the Agency's latest decision (P/0064/2016 of 18 March 2016).
The new PDCO Opinion on the modified agreed PIP supersedes the previous PDCO Opinion.

2.3.9. Cobicistat / atazanavir sulphate - EMEA-001465-PIP01-13-M03

Bristol-Myers Squibb Pharma EEIG; Treatment of HIV-1 infection / combination with other ARV medicinal products for the treatment of HIV-1 infected adults and children from 3 years of age without known mutations associated with resistance to atazanavir.

Day 60 opinion

Infectious Diseases

Summary of committee discussion:

The PDCO discussed the Applicant responses after Day 30 in line with other cobicistat PIPs and considered that all issues are resolved now.
The PDCO therefore adopted a favourable Opinion on the modification of the agreed PIP as set in the Agency's latest decision (P/0009/2018 of 30 January 2018).
The new PDCO Opinion on the modified agreed PIP supersedes the previous PDCO Opinion.

2.3.10. Eculizumab - Orphan - EMEA-000876-PIP03-14-M05

Alexion Europe SAS; Neuromyelitis optica spectrum disorders / Treatment of relapsing neuromyelitis optica spectrum disorders (NMOSD) in the paediatric population

Day 60 opinion

Neurology

Summary of committee discussion:

Based on the review of the rationale submitted by the application for modifying the agreed paediatric investigation plan, the PDCO considered that the proposed changes could be accepted.
The PDCO therefore adopted a favourable Opinion on the modification of the agreed PIP as set in the Agency's latest decision (P/0075/2020 of 21/03/2020).
The new PDCO Opinion on the modified agreed PIP supersedes the previous PDCO Opinion.

2.3.11. Ocrelizumab - EMEA-000310-PIP03-10-M04

Roche Registration GmbH; Treatment of Multiple Sclerosis / Treatment of relapsing remitting multiple sclerosis

Day 60 opinion

Neurology

Summary of committee discussion:

The PDCO re-discussed this procedure at Day 60 during the November 2020 plenary meeting.
The Committee confirmed all the conclusions reached at Day 30 and, in addition, it considered the information the applicant provided between Day 30 and Day 60.
Based on the review of the documentation submitted by the applicant for modifying the agreed paediatric investigation plan, the PDCO considered that the proposed changes could be acceptable. The main changes relate to study design and change of active control.
Therefore, the PDCO adopted a positive opinion on the modification of the agreed PIP as set in the Agency's latest decision (P/0028/2019 of 29 January 2019).
The new PDCO opinion on the modified agreed PIP supersedes the previous PDCO opinion.

2.3.12. Avelumab - EMEA-001849-PIP02-15-M03

Merck Healthcare GmbH; Treatment of all conditions included in the category of malignant neoplasms (except central nervous system tumours, haematopoietic and lymphoid tissue neoplams) / Treatment of malignant neoplasms of lymphoid tissue / Treatment of malignant neoplasms of the central nervous system / Treatment of paediatric patients from 2 years to less than 18 years old with a relapsed or refractory solid tumour or with a solid tumour as part of the first line treatment

Day 60 opinion

Oncology

Summary of committee discussion:

The PDCO re-discussed the proposed modification considering the clarifications provided by the applicant after D30
Based on the review of the rationale submitted by the application for modifying the agreed paediatric investigation plan, the PDCO considered that the proposed changes could be accepted.
The PDCO therefore adopted a favourable Opinion on the modification of the agreed PIP as set in the Agency's latest decision (P/0242/2018 of 15 August 2018).
The new PDCO Opinion on the modified agreed PIP supersedes the previous PDCO Opinion.

2.3.13. Brigatinib - EMEA-002296-PIP01-17-M02

Takeda Pharm A/S; Inflammatory Myofibroblastic Tumors (IMT) / Non-small cell lung cancer (NSCLC) / Anaplastic large cell lymphoma (ALCL) / Treatment of anaplastic lymphoma kinase (ALK) positive advanced non-small cell lung cancer (NSCLC) / Treatment of paediatric patients ≥1 years of age with ALK+ unresectable or recurrent IMT / Treatment in combination with standard chemotherapy in paediatric patients ≥1 years of age with newly diagnosed ALK+ ALCL at high risk for recurrence

Day 60 opinion

Oncology

Summary of committee discussion:

The PDCO re-discussed the proposed modification considering the clarifications provided by the applicant after D30.
The PDCO therefore adopted a favourable Opinion on the modification of the agreed PIP as set in the Agency's latest decision (P/0430/2019 of 5 December 2019).

2.3.14. Burosumab - Orphan - EMEA-001659-PIP01-15-M05

Kyowa Kirin Holdings B.V.; X-linked Hypophosphataemia / Treatment of X-linked Hypophosphataemia

Day 60 opinion

Other

Summary of committee discussion:

Based on the review of the rationale submitted by the application for modifying the agreed paediatric investigation plan, the PDCO considered that the proposed changes could be accepted.
The PDCO therefore adopted a favourable Opinion on the modification of the agreed PIP as set in the Agency's latest decision (P/0093/2020 of 18/03/2020).
The new PDCO Opinion on the modified agreed PIP supersedes the previous PDCO Opinion.

2.3.15. Ivacaftor / lumacaftor - EMEA-001582-PIP01-13-M10

Vertex Pharmaceuticals (Europe) Ltd; Cystic Fibrosis / Treatment of Cystic Fibrosis

Day 60 opinion

Other

Summary of committee discussion:

Based on the committee's day 30 discussion, the PDCO discussed the applicant's additional clarifications.
Therefore, based on the review of the rationale submitted by the application for modifying the agreed paediatric investigation plan, the PDCO considered that the proposed changes could be accepted.
The PDCO therefore adopted a favourable Opinion on the modification of the agreed PIP as set in the Agency's latest decision (P/0431/2019 of 6 December 2019).
The new PDCO Opinion on the modified agreed PIP supersedes the previous PDCO Opinion.

2.3.16. Levofloxacin hemihydrate - EMEA-001211-PIP01-11-M02

Chiesi Farmaceutici S.p.A.; Cystic fibrosis

Day 60 opinion

Pneumology - Allergology

Summary of committee discussion:

Based on the review of the rationale submitted by the application for modifying the agreed paediatric investigation plan, the PDCO considered that the proposed changes could be accepted.
The PDCO therefore adopted a favourable Opinion on the modification of the agreed PIP as set in the Agency's latest decision (P/0187/2013 of 8 August 2013) and to grant a full product specific waiver on the grounds of lack of significant therapeutic benefit over existing treatments.
The new PDCO Opinion on the modified agreed PIP supersedes the previous PDCO Opinion.

2.3.17. Calcifediol - EMEA-002093-PIP02-17-M01

Vifor Fresenius Medical Care Renal Pharma France; Endocrine, nutritional and metabolic diseases, and immunity disorders

Day 60 opinion

Uro-nephrology

Summary of committee discussion:

The PDCO's view expressed at Day 30 was re-discussed and endorsed.
Based on the review of the rationale submitted by the application for modifying the agreed paediatric investigation plan, the PDCO considered that the proposed changes could be accepted.
The PDCO therefore adopted a favourable Opinion on the modification of the agreed PIP as set in the Agency's latest decision (P/0339/2018 of 08/11/2018).
The new PDCO Opinion on the modified agreed PIP supersedes the previous PDCO Opinion.

2.3.18. Cholera vaccine, recombinant, live, oral (strain CVD 103-HgR) - EMEA-001490-PIP01-13-M02

Emergent Netherlands B.V.; Cholera disease caused by Vibrio cholerae serogroup O1 / Prevention of disease caused by Vibrio cholerae serogroup O1

Day 60 opinion

Vaccines

Summary of committee discussion:

The PDCO discussed the Applicant feedback on the draft Opinion and considered that the proposed modifications could be accepted.
The PDCO adopted a favourable Opinion on the modification of the agreed PIP as set in the Agency's latest decision (P/0381/2018 of 7 December 2018).
The new PDCO Opinion on the modified agreed PIP supersedes the previous PDCO Opinion.

2.3.19. Landiolol (hydrochloride) - EMEA-001150-PIP02-13-M03

AOP Orphan Pharmaceuticals AG; Treatment of supraventricular arrythmias

Day 60 opinion

Cardiovascular Diseases

Post-meeting note: Withdrawal request received on 11/11/2020

2.3.20. Ladarixin - EMEA-002642-PIP01-19-M02

Treatment of type 1 diabetes / Treatment of new-onset type 1 diabetes mellitus

Day 60 opinion

Endocrinology-Gynaecology-Fertility-Metabolism

Post- meeting note: Withdrawal request received on 02/11/2020

2.4. Opinions on Re-examinations

No items

2.5. Opinions on Review of Granted Waivers

No items

2.6. Finalisation and adoption of opinions

2.7. Partial Compliance Checks completed by EMA

The following partial compliance check has concluded positively without PDCO discussion. The Committee has been informed in writing.

2.7.1. Bictegravir / Tenofovir alafenamide / emtricitabine - EMEA-C2-001766-PIP01-15-M02

Gilead Sciences International Ltd; Treatment of human immunodeficiency virus (HIV-1) infection

Day 30 discussion

Infectious Diseases

2.7.2. Burosumab- Recombinant Human IgG1 monoclonal antibody targeting fibroblast growth factor 23 (FGF23) - EMEA-C4-001659-PIP01-15-M04

Kyowa Kirin Holdings B.V; Treatment of X-linked hypophosphatemia

Day 30 letter

Other

2.7.3. Maribavir - EMEA-C1-000353-PIP02-16-M01

Shire Pharmaceuticals Ireland Ltd; Treatment of cytomegalovirus (CMV) infection

Day 30 discussion

Infectious Diseases

2.7.4. Isavuconazonium (sulfate) - EMEA-C1-001301-PIP02-12-M03

Basilea Pharmaceutica International Ltd.; Treatment of mucormycosis

Day 30 discussion

Infectious Diseases

2.7.5. Bumetanide - EMEA-C1-001303-PIP01-12-M02

Les Laboratoires Servier; Treatment of autistic spectrum disorder

Day 30 discussion

Neurology

3. Discussion of applications

Disclosure of some information related to this section cannot be released at the present time as it is deemed to contain commercially confidential information.

3.1. Discussions on Products D90-D60-D30

3.1.1. EMEA-002757-PIP01-19

Treatment of ulcerative colitis

Day 90 discussion

Gastroenterology-Hepatology

3.1.2. Obinutuzumab - Orphan - EMEA-001207-PIP02-19

Roche Registration GmbH; Systemic Lupus Erythemathosus

Day 90 discussion

Immunology-Rheumatology-Transplantation

3.1.3. Lenacapavir - EMEA-002740-PIP01-19

Treatment of human immunodeficiency virus (HIV-1) infection

Day 90 discussion

Infectious Diseases

3.1.4. Retinol (Vitamin A) - Orphan - EMEA-002790-PIP01-20

Orphanix GmbH; Prevention of Bronchopulmonary Dysplasia (BPD)

Day 90 discussion

Neonatology - Paediatric Intensive Care

3.1.5. Arimoclomol citrate - Orphan - EMEA-001748-PIP03-19

Orphazyme A/S; Treatment of Amyotrophic Lateral Sclerosis

Day 90 discussion

Neurology

3.1.6. Autologous tumor-infiltrating lymphocytes - EMEA-002776-PIP01-20

Treatment of all conditions included in the category of malignant neoplasms (except haematopoietic and lymphoid tissue neoplasms)

Day 90 discussion

Oncology

3.1.7. Carfilzomib - Orphan - EMEA-001806-PIP04-19

Amgen Europe BV; Treatment of acute lymphoblastic leukemia / Treatment of paediatric patients aged 1 month or older and young adult patients up to 21 years of age with bone marrow relapse of T-cell ALL treated with at least one prior therapy or B-cell ALL treated with prior targeted immune therapy, with or without extramedullary disease

Day 90 discussion

Oncology / Haematology-Hemostaseology

3.1.8. KH176: (S)-6-hydroxy-2,5,7,8-tetramethyl-N-((R)-piperidin-3-yl)chroman-2-carboxamide hydrochloride - Orphan - EMEA-002113-PIP01-16

Khondrion BV; Treatment of mitochondrial respiratory chain/oxidative phosphorylation defects

Day 90 discussion

Other

3.1.9. Dexmedetomidine - EMEA-002758-PIP01-19

Treatment of acute Agitation in Bipolar Disorder / Treatment of acute Agitation in Schizophrenia

Day 90 discussion

Psychiatry

3.1.10. Esketamine - EMEA-002772-PIP01-20

Bipolar depression / Major depressive disorder / Treatment-resistant bipolar depression / Treatment-resistant depression in the course of major depressive disorder

Day 90 discussion

Psychiatry

3.1.11. Allopurinol / verinurad - EMEA-002754-PIP01-19

Chronic Kidney Disease / Treatment of Chronic Kidney Disease in children and adolescents (6 to <18 years old) with hyperuricaemia and albuminuria

Day 90 discussion

Uro-nephrology

3.1.12. Sparsentan - Orphan - EMEA-001984-PIP02-20

Retrophin Europe Limited; Treatment of Focal segmental glomerular sclerosis (FSGS)

Day 90 discussion

Uro-nephrology

3.1.13. Sparsentan - EMEA-001984-PIP03-20

Treatment of IgA Nephropathy (IgAN)

Day 90 discussion

Uro-nephrology

3.1.14. EMEA-002870-PIP01-20

Induction of general anesthesia in adults and children

Day 60 discussion

Anaesthesiology

3.1.15. Allogeneic skin-derived ABCB5-positive mesenchymal stem cells - Orphan - EMEA-002875-PIP01-20

RHEACELL GmbH & Co. KG; Treatment of epidermolysis bullosa / Treatment of recessive dystrophic epidermolysis bullosa and junctional epidermolysis bullosa

Day 60 discussion

Dermatology

3.1.16. Infigratinib - EMEA-002594-PIP02-20

Achondroplasia

Day 60 discussion

Endocrinology-Gynaecology-Fertility-Metabolism

3.1.17. Calmangfodipir - EMEA-002865-PIP01-20

Treatment of paracetamol overdose

Day 60 discussion

Gastroenterology-Hepatology

3.1.18. Lanifibranor - EMEA-002872-PIP01-20

Treatment of non-alcoholic steatohepatitis (NASH)

Day 60 discussion

Gastroenterology-Hepatology

3.1.19. Odevixibat - Orphan - EMEA-002054-PIP03-20

Albireo AB; Treatment of Alagille syndrome

Day 60 discussion

Gastroenterology-Hepatology

3.1.20. Vedolizumab - EMEA-000645-PIP04-20

Treatment of pouchitis

Day 60 discussion

Gastroenterology-Hepatology

3.1.21. EMEA-002863-PIP01-20

Paroxysmal Nocturnal Haemoglobinuria

Day 60 discussion

Haematology-Hemostaseology

3.1.22. EMEA-002861-PIP02-20

Prevention of Coronavirus disease 2019 (COVID-19) Day 60 discussion

Infectious Diseases

3.1.23. Autologous CD4+ and CD8+ T cells transduced with a lentiviral vector containing an affinity-enhanced T-cell receptor targeting the MAGE-A4 antigen - Orphan - EMEA-002867-PIP01-20

Adaptimmune Ltd; Soft Tissue Sarcoma / Myxoid / Round Cell Liposarcoma / Synovial Sarcoma

Day 60 discussion

Oncology

3.1.24. Magrolimab - Orphan - EMEA-002819-PIP01-20

Gilead Sciences International Ltd; Treatment of acute myeloid leukaemia, Treatment of myelodysplastic syndromes (including juvenile myelomonocytic leukaemia)

Day 60 discussion

Oncology

3.1.25. Pralsetinib - EMEA-002575-PIP02-20

Treatment of all conditions included in the category of malignant neoplasms (except

haematopoietic and lymphoid tissue neoplasms) / Treatment of pediatric patients from 12 years to less than 18 years of age with RET-altered solid tumors who require systemic therapy and have no satisfactory alternative treatment options

Day 60 discussion

Oncology

3.1.26. Docosahexaenoic Acid - Orphan - EMEA-002808-PIP01-20

Natac Pharma S.L.; Retinitis Pigmentosa

Day 60 discussion

Ophthalmology

3.1.27. (2S,4S)-2-(4-Carboxyphenyl)-4-ethoxy-1-[(5-methoxy-7-methyl-1H-indol-4-yl)methyl]piperidin-1-ium chloride—water (1/1) - Orphan - EMEA-002705-PIP03-20

Novartis Europharm Limited; Paroxysmal Nocturnal Haemoglobinuria

Day 60 discussion

Other / Haematology-Hemostaseology

3.1.28. Autologous selected renal cells - EMEA-002844-PIP01-20

Treatment of chronic kidney disease

Day 60 discussion

Uro-nephrology

3.1.29. EMEA-002873-PIP01-20

Active immunisation for the prevention of disease caused by chikungunya virus

Day 60 discussion

Vaccines

3.1.30. Rosuvastatin / acetylsalicylic acid - EMEA-002891-PIP01-20

prevention of cardiovascular events

Day 30 discussion

Cardiovascular Diseases

3.1.31. Heparin - EMEA-002885-PIP01-20

Prevention and treatment of thromboembolic events

Day 30 discussion

Cardiovascular Diseases / Haematology-Hemostaseology

3.1.32. Human, recombinant, non-fucosylated IgG1k monoclonal antibody targeting OX-40 receptor on activated T cells - EMEA-002886-PIP01-20

Atopic dermatitis / Treatment of moderate to severe atopic dermatitis

Day 30 discussion

Dermatology

3.1.33. Semaglutide - EMEA-001441-PIP05-20

Treatment of non-alcoholic fatty liver disease (NAFLD)

Day 30 discussion

Gastroenterology-Hepatology

3.1.34. Fenebrutinib - EMEA-002349-PIP03-20

Treatment of multiple sclerosis / Treatment of Relapsing multiple sclerosis in patients 10 years of age to less than 18 years of age

Day 30 discussion

Immunology-Rheumatology-Transplantation

3.1.35. Trimodulin - EMEA-002883-PIP01-20

Treatment of bacterial infections / Treatment of community-acquired sepsis / Treatment of neonatal sepsis / Treatment of severe community-acquired pneumonia / Treatment of community-acquired septic shock / Treatment of community-acquired severe sepsis

Day 30 discussion

Neonatology - Paediatric Intensive Care / Infectious Diseases / Pneumology - Allergology

3.1.36. Zidebactam / cefepime - EMEA-002892-PIP01-20

Treatment of complicated Urinary Tract Infections (cUTI)

Day 30 discussion

Neonatology - Paediatric Intensive Care / Infectious Diseases / Uro-nephrology

3.1.37. Edasalonexent [N-(2-((4Z,7Z,10Z,13Z,16Z,19Z)-docosa-4,7,10,13,16,19-hexaenamido)ethyl)-2-hydroxybenzamide] - Orphan - EMEA-001960-PIP03-20

Catabasis Pharmaceuticals Inc.; Duchenne Muscular Dystrophy / Treatment of Duchenne Muscular Dystrophy

Day 30 discussion

Neurology

3.1.38. Ravulizumab - EMEA-001943-PIP03-20

Acetylcholine receptor-antibody positive generalized myasthenia gravis / Treatment of acetylcholine receptor-antibody positive generalized myasthenia gravis

Day 30 discussion

Neurology

3.1.39. Reldesemtiv - Orphan - EMEA-002868-PIP01-20

Cytokinetics, Inc.; Amyotrophic lateral sclerosis (ALS)

Day 30 discussion

Neurology

3.1.40. EMEA-002884-PIP01-20

Progressive supranuclear palsy

Day 30 discussion

Neurology

3.1.41. EMEA-002745-PIP02-20

Treatment of lymphoplasmacytic lymphoma

Day 30 discussion

Oncology

3.1.42. Allogeneic anti-CD19 CAR T cells produced using CRISPR/Cas9 to disrupt the T cell receptor alpha constant (TRAC) and β2-microglobulin (B2M) genomic loci and a recombinant adeno-associated viral vector to deliver donor template for insertion of the anti-CD19 CAR expression cassette into the TRAC locus - EMEA-002881-PIP01-20

B cell lymphoblastic leukemia/lymphoma / Mature B cell neoplasms / Treatment of relapsed/refractory B cell ALL / Treatment of relapsed/refractory B cell NHL

Day 30 discussion

Oncology

3.1.43. Autologous peripheral blood T cells CD4- and CD8-selected and CD3- and CD28-activated transduced with retroviral vector expressing anti-CD19 CD28/CD3-zeta chimeric antigen receptor and cultured - Orphan - EMEA-001862-PIP03-20

Kite Pharma EU B.V.; Treatment of mature B-cell neoplasms / Treatment of paediatric and

adolescent subjects with relapsed or refractory B-cell non-Hodgkin lymphoma (NHL)

Day 30 discussion

Oncology

3.1.44. Catumaxomab - EMEA-002879-PIP01-20

Treatment of malignant ascites

Day 30 discussion

Oncology

3.1.45. Lorlatinib - EMEA-002669-PIP02-20

Treatment of lung malignant neoplasms

Day 30 discussion

Oncology

3.1.46. Ublituximab - Orphan - EMEA-002889-PIP01-20

CambPharma Solutions (CY) Ltd; Treatment of chronic lymphocytic leukemia (CLL)

Day 30 discussion

Oncology

3.1.47. Umbralisib - EMEA-002890-PIP01-20

Treatment chronic lymphocytic leukemia (CLL)

Day 30 discussion

Oncology

3.1.48. Ribitol - EMEA-002887-PIP01-20

Treatment of Limb-Girdle Muscular Dystrophy / Treatment of paediatric patients aged 5 years and older with a genetically confirmed diagnosis of Limb-Girdle Muscular Dystrophy type 2i (LGMD2I)/ LGMD R9-fukutin related

Day 30 discussion

Other

3.1.49. Paracetamol / nefopam - EMEA-002877-PIP01-20

Treatment of acute pain / Symptomatic short-term treatment of moderate to severe somatic acute pain

Day 30 discussion

Pain

3.1.50. Eliapixant - EMEA-002882-PIP01-20

Treatment of Refractory and/or unexplained chronic cough (RUCC)

Day 30 discussion

Pneumology - Allergology

3.1.51. Alpha1-Proteinase Inhibitor (Human) - EMEA-002888-PIP01-20

Treatment of emphysema secondary to alpha 1-proteinase inhibitor deficiency

Day 30 discussion

Pneumology - Allergology / Haematology-Hemostaseology

3.1.52. Acetylcysteine / Ibuprofen - EMEA-002561-PIP02-20

Treatment of upper respiratory tract infections

Day 30 discussion

Pneumology - Allergology / Oto-rhino-laryngology

3.1.53. mRNA that encodes for the pre-fusion stabilized Spike glycoprotein of 2019-novel Coronavirus - EMEA-002893-PIP01-20

Prevention of COVID-19 / Active immunisation against SARS-CoV-2

Day 30 discussion

Vaccines

3.2. Discussions on Compliance Check

The following compliance checks have been put up for discussion and the members of the PDCO have been invited to comment on issues of possible non-compliance.

3.2.1. Riociguat - EMEA-C-000718-PIP01-09-M06

Bayer AG; Treatment of pulmonary hypertension

Day 30 discussion

Cardiovascular Diseases

3.2.2. Idursulfase - EMEA-C-000294-PIP02-12-M01

Shire Human Genetic Therapies AB; Treatment of mucopolysaccharidosis II (Hunter syndrome)

Day 30 discussion

Endocrinology-Gynaecology-Fertility-Metabolism

3.2.3. Elbasvir / grazoprevir - EMEA-C-001604-PIP01-13-M03

Merck Sharp & Dohme B.V.; Treatment of chronic hepatitis C

Day 30 discussion

Infectious Diseases

3.3. Discussions on Modification of an Agreed Paediatric Investigation Plan

3.3.1. Remimazolam (as besylate) - EMEA-001880-PIP02-19-M01

PAION Deutschland GmbH; Sedation / General anesthesia / Sedation of mechanically ventilated patients / Procedural sedation

Day 30 discussion

Anaesthesiology

3.3.2. Denosumab - EMEA-000145-PIP02-12-M03

Amgen Europe B.V.; Treatment of Osteoporosis / Treatment of osteogenesis imperfecta / Treatment of glucocorticoid induced osteoporosis

Day 30 discussion

Endocrinology-Gynaecology-Fertility-Metabolism

3.3.3. Darvadstrocel - Orphan - EMEA-001561-PIP01-13-M02

Takeda Pharma A/S; Treatment of perianal fistula

Day 30 discussion

Gastroenterology-Hepatology

3.3.4. Obeticholic Acid - Orphan - EMEA-001304-PIP02-13-M05

Intercept Pharma International Ltd.; Treatment of Primary Biliary Cholangitis (PBC) / Treatment of Biliary Atresia

Day 30 discussion

Gastroenterology-Hepatology

3.3.5. Tofacitinib - EMEA-000576-PIP03-12-M05

Pfizer Europe MA EEIG; Treatment of ulcerative colitis

Day 30 discussion

Gastroenterology-Hepatology

3.3.6. Betibeglogene autotemcel - Orphan - EMEA-001665-PIP01-14-M04

bluebird bio (Netherlands) B.V.; Treatment of β-thalassaemia / Treatment of beta-thalassaemia major and severe intermedia

Day 30 discussion

Haematology-Hemostaseology

3.3.7. Luspatercept - Orphan - EMEA-001521-PIP01-13-M05

Celgene Europe B.V.; Treatment of beta-thalassaemia / Treatment of anaemia in patients with beta-thalassemia intermedia and major

Day 30 discussion

Haematology-Hemostaseology

3.3.8. Risankizumab - EMEA-001776-PIP02-17-M01

AbbVie Ltd; Chronic idiopathic arthritis

Day 30 discussion

Immunology-Rheumatology-Transplantation

3.3.9. Tofacitinib citrate - EMEA-000576-PIP01-09-M13

Pfizer Europe MA EEIG; Treatment of chronic idiopathic arthritis (including rheumatoid arthritis, psoriatic arthritis, ankylosing spondylitis and juvenile idiopathic arthritis) / Juvenile idiopathic arthritis

Day 30 discussion

Immunology-Rheumatology-Transplantation

3.3.10. 3-({5-chloro-1-[3-(methylsulfonyl)propyl]-1Hindol-2-yl}methyl)-1-(2,2,2-trifluoroethyl)-1,3-dihydro-2H-imidazo[4,5-C]pyridin-2-one) (JNJ-53718678) - EMEA-001838-PIP01-15-M03

Janssen-Cilag International NV; Treatment of respiratory tract disease caused by human respiratory syncytial virus (RSV)

Day 30 discussion

Infectious Diseases

3.3.11. Aztreonam / Avibactam - EMEA-002283-PIP01-17-M01

Pfizer Europe MA EEIG; Treatment of infections caused by aerobic gram-negative bacteria / Treatment of infections caused by aerobic gram-negative bacteria in patients with limited therapeutic options

Day 30 discussion

Infectious Diseases

3.3.12. Baloxavir marboxil - EMEA-002440-PIP01-18-M01

Roche Registration GmbH; Prevention of Influenza, Treatment of Influenza / Treatment of influenza type A/B in otherwise healthy and high risk patients / Prevention (post-exposure prophylaxis) of influenza type A/B / Reduction of transmission of influenza type A/B

Day 30 discussion

Infectious Diseases

3.3.13. Ceftazidime / avibactam - EMEA-001313-PIP01-12-M10

Pfizer Europe MA EEIG; Treatment of bacterial infections / Treatment of complicated urinary tract infections / Treatment of complicated intraabdominal infections / Treatment of pneumonia / Treatment of infections due to aerobic Gram-negative organisms

Day 30 discussion

Infectious Diseases

3.3.14. Cobicistat - EMEA-000969-PIP01-10-M05

Gilead Sciences International Ltd.; Treatment of human immunodeficiency virus type-1 (HIV-1) infection. / Treatment of human immunodeficiency virus type-1 (HIV-1) infection - pharmacoenhancer for use in combination with antiretroviral agents

Day 30 discussion

Infectious Diseases

3.3.15. Cobicistat / darunavir - EMEA-001280-PIP01-12-M03

Janssen-Cilag International NV; Treatment of HIV-1 Infection / Treatment of HIV-1 infection in paediatric patients from 3 to less than 18 years

Day 30 discussion

Infectious Diseases

3.3.16. Tenofovir Alafenamide / Emtricitabine - EMEA-001577-PIP02-14-M04

Gilead Sciences International Ltd.; Treatment of human immunodeficiency virus (HIV-1) infection

Day 30 discussion

Infectious Diseases

3.3.17. Tenofovir alafenamide / Emtricitabine / Bictegravir - EMEA-001766-PIP01-15-M03

Gilead Sciences International Ltd.; Treatment of Human immunodeficiency virus [HIV] disease resulting in other conditions / treatment of adults and paediatrics aged less than 2 years weighing more than 4 kg infected with human immunodeficiency virus-1 (HIV-1) without any known mutations associated with resistance to the individual components

Day 30 discussion

Infectious Diseases

3.3.18. Cannabidiol - Orphan - EMEA-001964-PIP01-16-M03

GW Pharma (International) B.V.; Lennox Gastaut Syndrome / Tuberous Sclerosis Complex / Infantile Spasms / Dravet Syndrome / Treatment of Seizures

Day 30 discussion

Neurology

3.3.19. Lasmiditan - EMEA-002166-PIP01-17-M05

Eli Lilly and Company Limited; Migraine with and without aura

Day 30 discussion

Neurology

3.3.20. Ofatumumab - EMEA-002397-PIP01-18-M01

Novartis Europharm Limited; Treatment of Multiple Sclerosis / Treatment of relapsing remitting multiple sclerosis

Day 30 discussion

Neurology

3.3.21. Pitolisant - Orphan - EMEA-001176-PIP01-11-M06

BIOPROJET PHARMA; Narcolesy / Narcolepsy with or without cataplexy

Day 30 discussion

Neurology

3.3.22. Siponimod hemifumarate - EMEA-000716-PIP01-09-M03

Novartis Europharm Ltd; Treatment of Multiple Sclerosis / Treatment of children/adolescent patients (10-18 years old) with relapsing forms of multiple sclerosis

Day 30 discussion

Neurology

3.3.23. Brentuximab vedotin - Orphan - EMEA-000980-PIP01-10-M07

Takeda Pharma A/S; Treatment of Hodgkin lymphoma / Treatment of paediatric patients with newly diagnosed relapse or refractory Hodgkin lymphoma (from 5 years of age)

Day 30 discussion

Oncology

3.3.24. Crizotinib - EMEA-001493-PIP03-18-M01

Pfizer Europe MA EEIG; Inflammatory myofibroblastic tumour (IMT) / Anaplastic large cell lymphoma (ALCL) / Treatment of paediatric patients that are able to swallow capsules (age range: 6 years to less than 18 years of age) with relapsed/refractory systemic ALK-positive ALCL / Treatment of paediatric patients that are able to swallow capsules (age range: 6 years to less than 18 years of age) with unresectable or relapsed/refractory ALK-positive IMT / Treatment of paediatric patients with relapsed/refractory systemic ALK-positive ALCL / Treatment of paediatric patients with unresectable or relapsed/refractory ALK-positive IMT

Day 30 discussion

Oncology

3.3.25. Eribulin - EMEA-001261-PIP01-11-M06

Eisai GmbH; Soft Tissue Sarcoma

Day 30 discussion

Oncology

3.3.26. Fosdenopterin - Orphan - EMEA-001491-PIP01-13-M01

Origin Biosciences, Inc.; Treatment of molybdenum cofactor deficiency type A

Day 30 discussion

Other

3.3.27. Human thrombin (component 2) / Human fibrinogen (component 1) - EMEA-001598-PIP01-13-M03

Instituto Grifols, S.A.; Treatment of haemorrhage resulting from a surgical procedure / Supportive treatment in surgery where standard surgical techniques are insufficient for improvement of haemostasis, and as a suture support in vascular surgery

Day 30 discussion

Other

3.3.28. Human Thrombin / Human Fibrinogen - EMEA-001149-PIP01-11-M06

Omrix Biopharmaceuticals N.V.; Treatment of haemorrhage resulting from a surgical procedure / Treatment of cerebrospinal fluid leakage resulting from a surgical procedure / for supportive treatment in surgery where standard surgical techniques are insufficient, for improvement of haemostasis / for suture line sealing in dura mater closure / for supportive treatment in surgery where standard surgical techniques are insufficient / for improvement of haemostasis

Day 30 discussion

Other

3.3.29. In vitro expanded autologous human articular chondrocytes - EMEA-001823-PIP01-15-M02

TETEC Tissue Engineering Technologies AG; Treatment of cartilage disorders

Day 30 discussion

Other

3.3.30. In vitro expanded autologous human articular chondrocytes - EMEA-002217-PIP01-17-M01

TETEC Tissue Engineering Technologies AG; Treatment of cartilage disorders

Day 30 discussion

Other

3.3.31. Vortioxetine - EMEA-000455-PIP02-10-M07

H. Lundbeck A/S; Major Depressive Disorder

Day 30 discussion

Psychiatry

3.3.32. Daprodustat - EMEA-001452-PIP01-13-M03

GlaxoSmithKline Trading Services Limited; Treatment of anaemia associated with chronic kidney disease

Day 30 discussion

Uro-nephrology / Haematology-Hemostaseology

3.3.33. Split influenza virus, inactivated containing antigens equivalent to the B-like strain (Yamagata lineage) / Split influenza virus, inactivated containing antigens equivalent to the B-like strain (Victoria lineage) / Split influenza virus, inactivated containing antigens equivalent to the A/H3N2-like strain / Split influenza virus,

inactivated containing antigens equivalent to the A/H1N1-like strain - EMEA-002359-PIP01-18-M02

Sanofi Pasteur; Prevention of influenza infection

Day 30 discussion

Vaccines

4. Nominations

Disclosure of information related to this section cannot be released at the present time as it is deemed to contain commercially confidential information.

4.1. List of submission of applications with start of procedure 1 December 2020 for Nomination of Rapporteur and Peer reviewer

Summary of committee discussion:

The PDCO approved the lists of Rapporteurs and Peer Reviewers.

4.2. Nomination of Rapporteur for requests of confirmation on the applicability of the EMA decision on class waiver

Summary of committee discussion:

The PDCO approved the lists of Rapporteurs and Peer Reviewers.

4.3. Nominations for other activities

Summary of committee discussion:

The PDCO approved the lists of Rapporteurs and Peer Reviewers.

5. Scientific Advice Working Party (SAWP) and Paediatric Committee (PDCO) Interaction

Disclosure of information related to this section cannot be released at the present time as it is deemed to contain commercially confidential information.

5.1. Ongoing Scientific Advice

Information related to this section cannot be released at the present time as it is deemed to contain commercially confidential information.

6. Discussion on the applicability of class waivers

Disclosure of some information related to this section cannot be released at the present time as it is deemed to contain commercially confidential information.

6.1. Discussions on the applicability of class waiver for products

No items

7. Discussion on the inclusion of an indication within a condition in an agreed PIP/waiver

7.1. Discussion on the possibility to include an indication within a condition in an agreed PIP/waiver

No items

8. Annual reports on deferrals

The members of the PDCO took note of the products listed in the Annex B.

9. Organisational, regulatory and methodological matters

9.1. Mandate and organisation of the PDCO

No items

9.2. Coordination with EMA Scientific Committees or CMDh-v

9.2.1. Committee for Medicinal Products for Human Use (CHMP)

Summary of committee discussion:

The PDCO members were informed about the final CHMP Opinions on six medicinal products with recommended paediatric indications adopted in October 2020 by CHMP. These include Fintepla (fenfluramine), Oxlumo (lumasiran), Palforzia (defatted powder of arachis hypogaea l., semen (peanuts) / defatted powder of arachis hypogaea l., semen (peanuts)), Dupixent (dupilumab), Humira (adalimumab), Vimpat (lacosamide).

The list of procedures with paediatric indications to be evaluated by the CHMP, starting in October 2020, was presented to the PDCO members.

9.3. Coordination with EMA Working Parties/Working Groups/Drafting Groups

9.3.1. Non-clinical Working Group: D30 Products identified

PDCO member: Karen van Malderen

Summary of committee discussion:

The Chair of the Non-clinical Working Group (NcWG) identified the products which will require NcWG evaluation and discussion.

9.3.2. Formulation Working Group

PDCO member: Brian Aylward

Summary of committee discussion:

The Chair of the Formulation Working Group (FWG) identified the products which will require FWG evaluation and discussion.

9.4. Cooperation within the EU regulatory network

9.4.1. European Network of Paediatric Research (Enpr) - European Medicines Agency (EMA)

No items

9.5. Cooperation with International Regulators

No items

9.6. Contacts of the PDCO with external parties and interaction with the Interested Parties to the Committee

No items

9.7. PDCO work plan

No items

9.8. Planning and reporting

No items

10. Any other business

10.1.1. COVID -19 update

Summary of committee discussion:

The PDCO was updated on the most relevant aspects of therapeutics and vaccines for COVID-19.

10.1.2. EMA IT tool on marketing status information

Summary of committee discussion:

PDCO was briefed on a new electronic reporting tool of Marketing Status data electronically by Industry for centrally authorized human medicinal products. The tool is another expansion of the IRIS platform and will enable data to be reported in a structured way and will provide an overview of such reported data through a dashboard available to all EU/EEA Member States.

The information on marketing status to be reported is in line with legal requirements (i.e. data at presentation level (EU number) of CAPs for each EU/EEA MS on the date of placing on the market, the date of cessation to market, the type of cessation (permanent/ temporary), the reasons for cessation and where applicable, the expected date of reintroduction in the market).

The new reporting system is intended to be launched in Q1 2021. For CAPs not yet launched in any EU/EEA MS, the MAH should report initial placing on the market and any subsequent updates via IRIS from the launch of the system. For CAPs already marketed in at least one EU/EEA MS, the MAH will require a baseline submission of the current marketing status in all EU/EEA MS for all presentations to be provided within 6 months after launch of the tool. From that moment all future reporting of changes of marketing status will be performed via IRIS.

The management of access to the tool will be done through the IRIS contact points in the Member States.

10.1.3. PDCO Meeting dates 2022-2024

Summary of committee discussion:

The meeting dates for 2022-2024 have been agreed and adopted.

10.1.4. Template for PDCO Opinion - Proposed revision of the extrapolation related section

Summary of committee discussion:

Proposed modifications to the PDCO opinion template were presented.

10.1.5. PDCO Member at the HMPC

Summary of committee discussion:

A reminder was circulated on the open call for an expert in paediatric medicines.

10.1.6. PDCO Eudramail mailbox - update

Summary of committee discussion:

An update was presented on the status of a eudranet mailbox for PDCO.

11. Breakout sessions

11.1.1. Neonatology

Summary of committee discussion:

Neonatal issues related to PIPs were discussed.

Duchenne Muscular Dystrophy (DMD): In accordance with the PDCO Work Plan (section on Development of Therapeutic Areas strategies) a TC was organised. The outcome of the TC was reported to the PDCO.

The Chair thanked all participants and closed the meeting.

12. List of participants

List of participants including any restrictions with respect to involvement of members / alternates / experts following evaluation of declared interests for the 10-13 November 2020 meeting.

Name	Role	Member state or affiliation	Outcome restriction following evaluation of e-DoI	Topics on agenda for which restrictions apply
Koenraad Norga	Chair	Belgium	When chairing the meeting: To be replaced for discussions, final deliberations and voting When not chairing the meeting: No participation in final deliberations and voting	3.3.32. Daprodustat - EMEA-001452-PIP01-13-M03
Karl-Heinz Huemer	Member	Austria	No interests declared	
Johanna Wernsperger	Alternate	Austria	No interests declared	
Marleen Renard	Member	Belgium	No participation in final deliberations and voting on:	2.3.14. Avelumab - EMEA-001849-PIP02-15-M03
Karen Van Malderen	Alternate	Belgium	No interests declared	
Dimitar Roussinov	Member	Bulgaria	No restrictions applicable to this meeting	
Georgios Savva	Member	Cyprus	No interests declared	
Lucie Kravackova	Member	Czech Republic	No interests declared	
Petra Dominikova	Alternate	Czech Republic	No interests declared	
Nanna Borup Johansen	Member	Denmark	No interests declared	
Irja Lutsar	Member	Estonia	No interests declared	
Jana Lass	Alternate	Estonia	No interests declared	
Sylvie Benchetrit	Member	France	No interests declared	
Dominique Ploin	Alternate	France	No interests declared	
Sabine Scherer	Member (Vice-Chair)	Germany	No interests declared	
Yuansheng Sun	Alternate	Germany	No interests declared	
Eleni Katsomiti	Member	Greece	No interests declared	
Anastasia Mountaki	Alternate	Greece	No interests declared	
Ágnes Gyurasics	Member (CHMP member)	Hungary	No interests declared	
Brian Aylward	Member	Ireland	No interests declared	

Name	Role	Member state or affiliation	Outcome restriction following evaluation of e-DoI	Topics on agenda for which restrictions apply
Sara Galluzzo	Member	Italy	No interests declared	
Dina Apele-Freimane	Member	Latvia	No restrictions applicable to this meeting	
Sigita Burokiene	Member	Lithuania	No interests declared	
Carola de Beaufort	Member (CHMP alternate)	Luxembourg	No interests declared	
Herbert Lenicker	Alternate	Malta	No interests declared	
Maaike van Dartel	Alternate	Netherlands	No interests declared	
Siri Wang	Member	Norway	No interests declared	
Anette Solli Karlsen	Alternate	Norway	No interests declared	
Marek Migdal	Member	Poland	No interests declared	
Helena Fonseca	Member	Portugal	No interests declared	
Hugo Tavares	Alternate	Portugal	No interests declared	
Dana Gabriela Marin	Member (CHMP alternate)	Romania	No interests declared	
Peter Sisovsky	Member	Slovakia	No interests declared	
Peter Szitanyi	Alternate	Slovakia	No interests declared	
Stefan Grosek	Member	Slovenia	No interests declared	
Fernando de Andrés Trelles	Member	Spain	No interests declared	
Maria Jesus Fernández Cortizo	Alternate	Spain	No interests declared	
Eva Agurell	Member	Sweden	No interests declared	
Johannes Taminiau	Alternate	Healthcare Professionals' Representative	No interests declared	
Fernando Cabanas	Member	Healthcare Professionals' Representative	No restrictions applicable to this meeting	
Francesca Rocchi	Member	Healthcare Professionals' Representative	No restrictions applicable to this meeting	
Catherine Cornu	Alternate	Healthcare Professionals' Representative	No participation in final deliberations and voting on:	3.3.21. Pitolisant - Orphan - EMEA-001176-PIP01-11-M06
Jaroslav Sterba	Member	Patients' Organisation Representative	No interests declared	
Dimitrios Athanasiou	Member	Patients' Organisation Representative	No interests declared	
Tomasz Grybek	Alternate	Patients' Organisation Representative	No interests declared	
Nora Kriauzaite	Alternate	Patients' Organisation Representative	No participation in discussion, final deliberations and	2.1.17.Obinutuzumab - Orphan - EMEA-001207-PIP03-20

Name	Role	Member state or affiliation	Outcome restriction following evaluation of e-DoI	Topics on agenda for which restrictions apply
			voting on:	2.3.13. Ocrelizumab - EMEA-000310-PIP03-10-M04 3.1.2. Obinutuzumab - Orphan - EMEA-001207-PIP02-19 3.1.24. Pralsetinib - EMEA-002575-PIP02-20 3.1.34. Fenebrutinib - EMEA-002349-PIP03-20 3.3.12. Baloxavir marboxil - EMEA-002440-PIP01-18-M01
María Estela Moreno Martín	Expert	Spain - AGEMED/AEMPS	No interests declared	
Lutz Wiesner	Expert	Germany - BfArM	No interests declared	
Emmely de Vries	Expert	Netherlands - CBG/MEB	No interests declared	
Michal Odermarsky*	Expert	European Union - EMA	No participation in final deliberations and voting on:	3.1.31. Heparin Sodium - EMEA-002885-PIP01-20
Karri Penttila	Expert	Finland - FIMEA	No interests declared	
Sara Vennberg	Expert	Sweden - MPA	No interests declared	
Rune Kjeken*	Expert	Norway - NOMA	No restrictions applicable to this meeting	
Serena Marchetti*	Expert	Netherlands - CBG/MEB	No restrictions applicable to this meeting	
Kristian Wennmalm	Expert	Sweden - MPA	No interests declared	
Andreas Kirisits	Expert	Austria Ages	No interests declared	
Juha Kolehmainen	Expert	Finland - FIMEA	No interests declared	
A representative from the European Commission attended the meeting				
Meeting run with support from relevant EMA staff				

* Experts were only evaluated against the agenda topics or activities they participated in

13. Explanatory notes

The Notes give a brief explanation of relevant agenda items and should be read in conjunction with the agenda.

Paediatric investigation plan (PIP) (section 2.1 Opinion on PIPs and section 3.1 Discussions on PIPs)
A paediatric investigation plan (PIP) is a development plan aimed at ensuring that the necessary data are obtained through studies in children, when it is safe to do so, to support the authorisation of a medicine for children. Pharmaceutical companies submit proposals for PIPs to the European Medicines Agency's Paediatric Committee (PDCO). This Committee is responsible for agreeing or refusing the plan.

Compliance checks (section 2.2 Opinions on Compliance check, section 3.2 Discussions on Compliance check)
A compliance check may be necessary before any application for marketing authorisation (even for an adult indication) can be considered valid, if there was no deferral for at least one of the studies agreed in the PIP, or after the due date of initiation or completion of a study/measure. The same applies to some regulatory applications for authorised products, as described above.

Modification of an Agreed Paediatric Investigation Plan (section 2.3 Opinions on Modification of an agreed PIP, section 3.3 Discussions on Modification of an agreed PIP)
The development plan for a medicine can be modified at a later stage as knowledge increases. Modifications can also be made if the applicant encounters such difficulties with the implementation of a PIP, which render it unworkable or no longer appropriate.
In some cases, studies can be deferred until after the studies in adults have been conducted. This ensures that research in children is done only when it is safe and ethical to do so. Even when studies are deferred, the PIP will include details of the paediatric studies and their timelines.

Class waiver (section 6 Discussion on the applicability of class waiver)
As some diseases do not affect children (for example Parkinson's disease), the development of medicines for these diseases should not be performed in children. In these cases, a PIP is not required and it will be waived**.** For more information on the classes of diseases subject to waivers, see class waivers.

Annual reports on deferrals (section 8)
If the medicinal product is approved in the EU, annual reports on the deferred measures in the PIP must be submitted to the Agency.

More detailed information on the above terms can be found on the EMA website: www.ema.europa.eu/