28 April 2020
EMA/PDCO/173462/2020
Human Medicines Division

Paediatric Committee (PDCO)

Minutes for the meeting on 28-30 April 2020

Disclaimers

Some of the information contained in these 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).

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1. Introductions

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

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 2.1.4, 2.1.5, 2.3.18, 3.1.39, 3.3.1, 3.3.5, 3.3.7.

1.2. Adoption of agenda

PDCO agenda for 28-30 April 2020

The agenda from the PDCO plenary meeting 28th -30th April 2020 was adopted. An additional topic of Covid-19 update was added under section 10.

1.3. Adoption of the minutes

PDCO minutes for 24-27 March 2020

The minutes from the PDCO plenary 24th -27th March 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. Bis-choline tetrathiomolybdate - Orphan - EMEA-002232-PIP02-19

Alexion Europe S.A.S.; Treatment of Wilson disease

Day 120 opinion

Endocrinology-Gynaecology-Fertility-Metabolism

Summary of committee discussion:

The PDCO discussed this application on D120. In general, the applicant's responses to the issues raised on D90 were considered acceptable.
A positive opinion for Bis-choline tetrathiomolybdate for the treatment of Wilson disease in children from 3 years of age to less than 18 years of age was adopted and a waiver was granted in children on the grounds that the specific medicinal product does not represent a significant therapeutic benefit over existing treatments.

2.1.2. Fidanacogene elaparvovec - Orphan - EMEA-002362-PIP02-19

Pfizer Europe MA EEIG; Treatment of congenital factor IX deficiency (haemophilia B)

Day 120 opinion

Haematology-Hemostaseology

Summary of committee discussion

During its plenary on 30 April 2020, the PDCO adopted a positive Opinion on the PIP for fidanacogene elaparvovec for the prophylactic treatment of congenital factor IX deficiency (haemophilia B), in patients without inhibitors.

2.1.3. Gepotidacin - EMEA-002443-PIP01-18

GlaxoSmithKline Trading Services Limited; Treatment of uncomplicated urinary tract infection (uUTI)

Day 120 opinion

Infectious Diseases

Summary of committee discussion

During its plenary on 30th April 2020 the PDCO adopted a favourable Opinion on the PIP application for gepotidacin, an antibacterial agent, for the treatment of uncomplicated lower urinary tract infection (uUTI).

2.1.4. Gepotidacin - EMEA-002443-PIP02-18

GlaxoSmithKline Trading Services Limited; Treatment of uncomplicated urogenital gonorrhoea (GC)

Day 120 opinion

Infectious Diseases

Summary of committee discussion:

During its plenary on 30th April 2020 the PDCO adopted a favourable Opinion on the PIP application for gepotidacin, an antibacterial agent, for the treatment of uncomplicated urogenital gonorrhoea.

2.1.5. Dexamethasone (sodium phosphate) encapsulated in human autologous erythrocytes - Orphan - EMEA-001957-PIP02-19

EryDel S.p.A; Treatment of ataxia telangiectasia (AT)

Day 120 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 paediatric investigation plan for dexamethasone (sodium phosphate) encapsulated in human autologous erythrocytes for the treatment of ataxia telangiectasia. The plan includes a waiver based on lack of safety. Some studies are deferred, and the plan is expected to complete in May 2024.

2.1.6. Allogeneic haptenised and irradiated lysates derived from glioma - Orphan - EMEA-002663-PIP01-19

ERC Belgium; Treatment of glioma

Day 120 opinion

Oncology

Summary of committee discussion:

The PDCO discussed this procedure at Day 120 during the April 2020 plenary meeting.

The PDCO took into consideration the information the applicant provided following the PDCO’s comments at Day 90.

Taking the above into consideration, the PDCO adopted a positive Opinion for a paediatric investigation plan, with a waiver for an age subset and a deferral.

2.1.7. Allogeneic haptenised and irradiated cells derived from glioma - Orphan - EMEA-002662-PIP01-19

ERC Belgium; Treatment of glioma

Day 120 opinion

Oncology

Summary of committee discussion:

The PDCO discussed this procedure at Day 120 during the April 2020 plenary meeting. The PDCO took into consideration the information the applicant provided following the

PDCO's comments at Day 90, and agreed that they were appropriate.
Taking the above into consideration, the PDCO adopted a positive Opinion for a paediatric investigation plan, for a waiver for an age subset and for a deferral.

2.1.8. Autologous haptenised and irradiated lysates derived from glioma - Orphan - EMEA-002664-PIP01-19

ERC Belgium; Treatment of glioma

Day 120 opinion

Oncology

Summary of committee discussion:

The PDCO discussed this procedure at Day 120 during the April 2020 plenary meeting.
The PDCO took into consideration the information the applicant provided following the PDCO's comments at Day 90, and agreed that they were appropriate.
Taking the above into consideration, the PDCO adopted a positive Opinion for a paediatric investigation plan, for a waiver for an age subset and for a deferral.

2.1.9. Autologous haptenised and irradiated cells derived from glioma - Orphan - EMEA-002661-PIP01-19

ERC Belgium; Treatment of glioma

Day 120 opinion

Oncology

Summary of committee discussion:

The PDCO discussed this procedure at Day 120 during the April 2020 plenary meeting.
The PDCO took into consideration the information the applicant provided following the PDCO's comments at Day 90, and agreed that they were appropriate.
Taking the above into consideration, the PDCO adopted a positive Opinion for a paediatric investigation plan, for a waiver for an age subset and for a deferral.

2.1.10. Lenvatinib - EMEA-001119-PIP03-19

Eisai GmbH; Treatment of all conditions included in the category of malignant neoplasms except haematopoietic and lymphoid tissue neoplasms, papillary thyroid cancer, follicular thyroid cancer and osteosarcoma

Day 120 opinion

Oncology

Summary of committee discussion:

The PDCO took note of the clarifications provided by the applicant after the D90 discussion and the comments received by the applicant on the draft opinion.
All pending issues were considered solved.
In conclusion, the PDCO recommended granting a paediatric investigation plan for the paediatric population from 2 years to less than 18 years of age with a waiver for the age

subset based on the on the grounds that the specific medicinal product is likely to be unsafe and a deferral in the condition 'treatment of all conditions included in the category of malignant neoplasms except haematopoietic and lymphoid tissue neoplasms, papillary thyroid cancer, follicular thyroid cancer and osteosarcoma'.

2.1.11. Adeno-associated viral vector serotype 8 containing the human RPGR gene - Orphan - EMEA-002601-PIP01-19

Nightstar Europa Limited; Treatment of X-linked retinitis pigmentosa

Day 120 opinion

Ophthalmology

Summary of committee discussion

Based on the assessment of this application, a positive opinion for Adeno-associated viral vector serotype 8 containing the human RPGR gene for a subset of boys in the condition of Treatment of X-Linked Retinitis Pigmentosa was adopted. A waiver in a subset of boys is based on the grounds of lack of significant benefit and in girls from birth to less than 18 years of age based on the grounds disease not occurring was also adopted.

2.1.12. Atropine - EMEA-002545-PIP01-19

Fondazione Per La Ricerca Farmacologica Gianni Benzi Onlus; Treatment of myopia

Day 120 opinion

Ophthalmology

Summary of committee discussion:

During its plenary on 30 April 2020, the PDCO discussed the applicant's feedback to the PDCO's day 90 minutes for atropine, (eye drops, ocular use) for the treatment of progressive myopia in children and adolescents.
In conclusion, the applicant has satisfactorily addressed all remaining issues and the PIP Opinion has been revised accordingly. Therefore, the PDCO adopted a positive Opinion.

2.1.13. Sodium alginate oligosaccharide - Orphan - EMEA-002321-PIP01-17

AlgiPharma AS; Treatment of cystic fibrosis

Day 120 opinion

Pneumology – Allergology

Summary of committee discussion:

Based on the assessment of this application and further discussions at the Paediatric Committee and considering the applicant's responses following the Day 90 discussion, the PDCO adopted a favourable opinion on the paediatric investigation plan for sodium alginate oligosaccharide for the treatment of cystic fibrosis from 6 to less than 18 years of age. A waiver was requested for a subset of patients on the grounds that the specific medicinal product does not represent a significant therapeutic benefit as the needs are already covered

and was granted. Deferrals for the completion dates of some studies were requested and granted.

2.1.14. Chloroprocaine hydrochloride - EMEA-000639-PIP06-20

Sintetica GmbH; Ocular surface anaesthesia

Day 60 opinion

Anaesthesiology

Summary of committee discussion:

The PDCO confirmed the outcome of the discussion at Day 30. 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 chloroprocaine for all subsets of the paediatric population (0 to less than 18 years of age) in the condition of ocular surface anaesthesia on the grounds that the specific medicinal product does not represent a 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. 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. Ezetimibe / atorvastatin - EMEA-002649-PIP02-20

ELPEN Pharmaceutical Co. Inc.; Prevention of cardiovascular events

Day 60 opinion

Cardiovascular Diseases

Summary of committee discussion:

Based on the assessment of this application, the PDCO agrees with the applicant's request for a waiver. The PDCO recommends granting a waiver for Ezetimibe / Atorvastatin for all subsets of the paediatric population (0 to 18 years of age) in the condition of Prevention of cardiovascular events on the grounds that the specific medicinal product does not represent a significant therapeutic benefit over existing treatments for paediatric patients.
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. Bilastine - EMEA-000347-PIP05-20

FAES FARMA S.A.; Treatment of acute type I hypersensitivity reactions

Day 60 opinion

Dermatology / Pneumology - Allergology / Oto-rhino-laryngology

Summary of committee discussion

Based on the assessment of this application taking into the applicant's additional clarifications, the PDCO did not agree with the applicant's request for a waiver in the condition "treatment of acute type I hypersensitivity reactions". There is also a need for the planned formulation in the paediatric population be considered in severe cases irrespective of the patient age and should, therefore, also be developed for children.

The committee, therefore, adopted a negative opinion.

2.1.17. N-[(2S)-5-{[(1R, 2S)-2-(4-fluorophenyl)cyclopropyl] amino}-1-(4-methylpiperazin-1-yl)-1-oxopentan-2-yl]-4-(1H-1,2,3-triazol-1-yl)benzamide, bis-tosylate salt - Orphan - EMEA-002752-PIP01-19

Imago Biosciences BV; Treatment of myeloproliferative neoplasms

Day 60 opinion

Haematology-Hemostaseology

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 N-[(2S)-5-{[(1R, 2S)-2-(4-fluorophenyl)cyclopropyl] amino}-1-(4-methylpiperazin-1-yl)-1-oxopentan-2-yl]-4-(1H-1,2,3-triazol-1-yl)benzamide, bis-tosylate salt for all subsets of the paediatric population (0 to 18 years of age) in the condition of Treatment of myeloproliferative neoplasms.
Considering the rarity of these disease in the paediatric population, the PDCO agrees as proposed by the applicant to grant a full product specific waiver with the justification that clinical trials in the paediatric population are of no significant therapeutic benefit as clinical trial would not be feasible.

2.1.18. Adeno-associated virus serotype 2 (AAV2) encoding human aromatic L-amino acid decarboxylase (hAADC) - EMEA-002753-PIP01-19

Neurocrine Therapeutics, Ltd.; Treatment of Parkinson's disease

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 on the grounds that the product is expected to be both ineffective and unsafe in the paediatric population. A positive opinion was therefore adopted.
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.

2.1.19. Monalizumab - EMEA-002751-PIP01-19

AstraZeneca AB; Treatment of head and neck epithelial malignant neoplasms

Day 60 opinion

Oncology

Summary of committee discussion:

The PDCO re-discussed this application in line with the D30 discussion outcome. Overall the PDCO agreed to product specific waiver for the condition of treatment of head and neck epithelial malignant neoplasms. the most appropriate waiver ground was concluded to be based on lack of significant therapeutic benefit as studies are not feasible.
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 monalizumab 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.
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.

2.1.20. Ribociclib - EMEA-002765-PIP01-19

Novartis Europharm Limited; Treatment of breast cancer

Day 60 opinion

Oncology

Summary of committee discussion:

The PDCO discussed this procedure at Day 60 during the April 2020 plenary meeting.
The PDCO took into consideration the clarifications the applicant provided between Day 30 and Day 60 and considered them satisfactory.
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 ribociclib for all subsets of the paediatric population (0 to 18 years of age) in the condition of treatment of breast cancer on the grounds that this medicinal product does not represent a significant therapeutic benefit over existing treatments for paediatric patients.
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. The PDCO, also considering the information provided by the applicant on currently available results, identified neuroblastoma and medulloblastoma as an unmet need that could be targeted by this product. 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.

2.1.21. AdAptVEGF-C, [Adenovirus encoding vascular endothelial growth factor C (VEGF-C)] - EMEA-002748-PIP01-20

Herantis Pharma Plc; Treatment of secondary lymphoedema associated with the treatment of breast cancer

Day 60 opinion

Other

Summary of committee discussion:

The PDCO re-discussed this application in line with the D30 conclusions.
Based on the assessment of this application and further discussions at the Paediatric, the PDCO agrees with the applicant's request for a waiver. The PDCO recommends granting a waiver for AdAptVEGF-C, [Adenovirus encoding vascular endothelial growth factor C (VEGF-C)] for all subsets of the paediatric population (0 to 18 years of age) in the condition of treatment of secondary lymphedema associated with the treatment of breast cancer based on the ground that the disease 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.

2.1.22. Betahistine dihydrochloride - EMEA-002652-PIP01-19

Auris Medical Ltd.; Treatment of peripheral vertigo

Day 60 opinion

Oto-rhino-laryngology / Neurology

Summary of committee discussion

Following the D30 discussion, the PDCO in April 2020 agreed with the applicant's request and recommends granting a waiver for betahistine (dihydrochloride) nasal solution for all subsets of the paediatric population (0 to 18 years of age) in the condition of "treatment of peripheral vertigo" nasal spray, solution, intranasal use; on the grounds that the specific medicinal product does not represent a 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.

2.2. 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. Adalimumab - EMEA-C-000366-PIP02-09-M06

AbbVie Limited; Treatment of ulcerative colitis

Day 60 opinion

Gastroenterology-Hepatology

Summary of committee discussion:

The PDCO took note of outcomes of preceding partial compliance check procedures: EMEA-C1-000366-PIP02-09-M01.
The PDCO adopted on 30 April 2020 an opinion confirming the compliance of all studies in the agreed paediatric investigation plan as set out in the EMA's decision P/0174/2019 of 15 May 2019.

2.2.2. Lipegfilgrastim - EMEA-C-001019-PIP01-10-M05

UAB "Sicor Biotech"; Treatment of chemotherapy-induced neutropenia

Day 30 opinion

Oncology

Summary of committee discussion:

PDCO has discussed the procedure and adopted on 30 April 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/0034/2020) of 29 January 2020.

2.2.3. Zoledronic acid - EMEA-C-000057-PIP01-07-M07

Novartis Europharm Limited; Treatment of osteoporosis / Treatment of Paget's disease of the bone

Day 60 opinion

Gastroenterology-Hepatology

Summary of committee discussion:

The PDCO took note of outcomes of preceding partial compliance check procedures:

- EMEA-C1-000057-PIP01-07-M05.

The PDCO accepted the Applicant's responses to the day 30 minutes and adopted on 30 April 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/0028/2020 of 17/01/2020.

2.2.4. (3R,4R)-4-methyl-3-(methyl-7H-pyrrolo[2,3-d]pyrimidin-4-ylamino)-β-oxo-1-piperidinepropanenitrile, 2-hydroxy-1,2,3-propanetricarboxylate (1:1) - EMEA-C4-000576-PIP01-09-M10

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

Day 60 letter

Immunology-Rheumatology-Transplantation

Summary of committee discussion:

Between Day 30 and Day 60 the applicant provided the requested clarifications. The PDCO discussed the completed studies and considered that these are compliant with the latest Agency's Decision P/0134/2019 of 17 April 2019.
The PDCO finalised this partially completed compliance procedure on 30 April 2020.

2.2.5. Inebilizumab - EMEA-C3-001911-PIP01-15-M02

Viela Bio; Treatment of neuromyelitis optica spectrum disorders

Day 60 letter

Action: For adoption

Neurology

Summary of committee discussion:

The PDCO view expressed at Day 30 was re-discussed and endorsed.

Some studies are considered not compliant with the latest Agency's Decision (P/0129/2019 of 17 April 2019) as the proof for completion of a study cannot be accepted and issues were raised concerning another study.

The PDCO finalised this partially completed compliance procedure on 30 April 2020.

2.3. Opinions on Modification of an Agreed Paediatric Investigation Plan

2.3.1. Apixaban - EMEA-000183-PIP01-08-M08

Bristol-Myers Squibb / Pfizer EEIG; Prevention of arterial thromboembolism / Prevention of venous thromboembolism

Day 60 opinion

Cardiovascular 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 favourable Opinion on the modification of the agreed PIP as set in the Agency’s latest decision (P/0154/2018 of 25/05/2018).

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

2.3.2. Apixaban - EMEA-000183-PIP02-12-M03

Bristol-Myers Squibb / Pfizer EEIG; Treatment of venous thromboembolism

Day 60 opinion

Cardiovascular 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 favourable Opinion on the modification of the agreed PIP as set in the Agency's latest decision (P/0153/2018 of 25/05/2018).

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

2.3.3. Ticagrelor - EMEA-000480-PIP01-08-M13

AstraZeneca AB; Prevention of thromboembolic events

Day 60 opinion

Cardiovascular Diseases / Haematology-Hemostaseology

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/0205/2018 of 19 July 2018).
The new PDCO Opinion on the modified agreed PIP supersedes the previous PDCO Opinion.

2.3.4. Ligelizumab - EMEA-001811-PIP02-15-M03

Novartis Europharm Limited; Treatment of chronic spontaneous urticaria

Day 60 opinion

Dermatology

Summary of committee discussion:

The PDCO adopted a favourable Opinion on the modification of the agreed PIP as set in the Agency's latest decision (P/0221/2018 of 17 July 2018).
The new PDCO Opinion on the modified agreed PIP supersedes the previous PDCO Opinion.

2.3.5. Lucerastat - Orphan - EMEA-002095-PIP01-16-M01

Idorsia Pharmaceuticals Deutschland GmbH; Treatment of Fabry disease

Day 60 opinion

Endocrinology-Gynaecology-Fertility-Metabolism

Summary of committee discussion:

Based on the review of the rationale submitted by the application for modifying the agreed

paediatric investigation plan including the additional clarifications submitted after Day 30, 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.
The new PDCO Opinion on the modified agreed PIP supersedes the previous PDCO Opinion.

2.3.6. Darvadstrocel - Orphan - EMEA-001561-PIP01-13-M01

Takeda Pharma A/S; Treatment of perianal fistula

Day 60 opinion

Gastroenterology-Hepatology

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/0253/2014 of 29 September 2014).
The new PDCO Opinion on the modified agreed PIP supersedes the previous PDCO Opinion.

2.3.7. Autologous CD34+ cells transduced with lentiviral vector encoding the human beta-globin gene - Orphan - EMEA-001933-PIP01-16-M01

Orchard Therapeutics (Europe) Ltd; Treatment of beta-thalassemia

Day 60 opinion

Haematology-Hemostaseology

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

2.3.8. Human fibrinogen concentrate - EMEA-001931-PIP01-16-M02

Biotest AG; Treatment of congenital fibrinogen deficiency

Day 60 opinion

Haematology-Hemostaseology

Summary of committee discussion:

Based on the review of the rationale submitted by the application for modifying the agreed paediatric investigation plan, the PDCO thus 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/0285/2018 of 12 September 2018).
The new PDCO Opinion on the modified agreed PIP supersedes the previous PDCO Opinion.

2.3.9. Tofacitinib citrate - EMEA-000576-PIP01-09-M12

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

Day 60 opinion

Immunology-Rheumatology-Transplantation

Summary of committee discussion:

In April 2020, the PDCO discussed the outstanding points after receiving the replies of the applicant to the points discussed at D30.
The PDCO therefore adopted a favourable Opinion on the modification of the agreed PIP as set in the Agency's latest decision (P/0211/2017of 9/8/2017).
The new PDCO Opinion on the modified agreed PIP supersedes the previous PDCO Opinion.

2.3.10. Upadacitinib - EMEA-001741-PIP04-17-M01

AbbVie Ltd; Treatment of atopic dermatitis

Day 60 opinion

Immunology-Rheumatology-Transplantation / Dermatology

Summary of committee discussion

The PDCO discussed the responses submitted by the Applicant and considered that the justifications / clarifications and proposed changes are acceptable.
The PDCO adopted a favourable Opinion on the modification of the agreed PIP as set in the Agency's latest decision (P/0394/2018 of 7 December 2018).
The new PDCO Opinion on the modified agreed PIP supersedes the previous PDCO Opinion.

2.3.11. Tenofovir alafenamide / emtricitabine / cobicistat / elvitegravir - EMEA-001460-PIP01-13-M04

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

Day 60 opinion

Infectious Diseases

Summary of committee discussion:

In April 2020, the PDCO discussed the outstanding points after receiving the replies of the applicant to the points discussed at D30. The PDCO agreed with the previously proposed changes but, in addition, agreed to change the ground of the previously agreed waiver from lack of safety to lack of therapeutic benefit over existing treatment. In fact, all other agreed PIPs for tenofovir alafenamide share this ground and the only exception was the PIP for this formulation based on the fact that when the PIP was initially agreed the expected dose of TAF was not established and could have been potentially higher than in other products. This

is no longer the case after new PK results have become available.
The PDCO therefore adopted a favourable Opinion on the modification of the agreed PIP as set in the Agency's latest decision (P/0211/2017of 9/8/2017).
The new PDCO Opinion on the modified agreed PIP supersedes the previous PDCO Opinion.

2.3.12. Eptinezumab - EMEA-002243-PIP01-17-M01

H. Lundbeck A/S; Prevention of migraine headaches

Day 60 opinion

Neurology

Summary of committee discussion:

In April 2020 the PDCO noted the responses of the applicant to the points raised at D30.
The PDCO therefore adopted a favourable Opinion on the modification of the agreed PIP as set in the Agency's latest decision (P/0314/2019 of 11/9/2019).
The new PDCO Opinion on the modified agreed PIP supersedes the previous PDCO Opinion.

2.3.13. Erenumab - EMEA-001664-PIP02-15-M04

Novartis Europharm Limited; Prevention of migraine headaches

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 change can 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/0107/2019 of 22/3/2019).
The new PDCO Opinion on the modified agreed PIP supersedes the previous PDCO Opinion.

2.3.14. Humanised anti-IL-6 receptor (IL-6R) monoclonal antibody (INN: satralizumab) - Orphan - EMEA-001625-PIP01-14-M05

Roche Registration GmbH; Treatment of neuromyelitis optica

Day 60 opinion

Neurology

Summary of committee discussion:

The PDCO re-discussed this modification request in line with the D30 discussion outcome. The committee noted the additional information received by the applicant. The PDCO therefore adopted a favourable Opinion on the modification of the agreed PIP as set in the Agency's latest decision (P/0220/2019 of 17 June 2019).
The new PDCO Opinion on the modified agreed PIP supersedes the previous PDCO Opinion.

2.3.15. Meningococcal group Y oligosaccharides conjugated to *Corynebacterium diphtheriae* CRM197 protein (MenABCWY) / recombinant *Neisseria meningitis* group B Protein 961c / recombinant *Neisseria meningitis* group B Protein 287- 953 / meningococcal group C oligosaccharides conjugated to *Corynebacterium diphtheriae* CRM197 protein / meningococcal group A oligosaccharides conjugated to *Corynebacterium diphtheriae* CRM197 protein / Outer Membrane Vesicles (OMV) from *Neisseria meningitidis* Strain NZ 98/254 / meningococcal group W-135 oligosaccharides conjugated to *Corynebacterium diphtheriae* CRM197 protein / recombinant *Neisseria meningitis* group B Protein 936-741 - EMEA-001260-PIP01-11-M01

GSK Vaccines s.r.l.; Prevention of meningococcal infection

Day 60 opinion

Vaccines

Summary of committee discussion:

In line with the Day 30 discussion the PDCO adopted a favourable Opinion on the modification of the agreed PIP as set in the Agency's latest decision (P/0304/2012 of 20 December 2012).
The new PDCO Opinion on the modified agreed PIP supersedes the previous PDCO Opinion.

2.3.16. Monovalent, recombinant, replication-incompetent human adenovirus serotype 26-vectored vaccine encoding the pre-fusion conformation-stabilised F protein derived from the RSV A2 strain - EMEA-002172-PIP02-17-M01

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

Day 60 opinion

Vaccines / Infectious Diseases

Summary of committee discussion:

The PDCO took note of the clarifications provided by the applicant after the D30 discussion and the comments received from the applicant on the draft opinion.
The PDCO therefore adopted a favourable Opinion on the modification of the agreed PIP as set in the Agency's latest decision (P/0318/2018 of 12 September 2018).
The new PDCO Opinion on the modified agreed PIP supersedes the previous PDCO Opinion.

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

2.6.1. Cilastatin sodium / relebactam / imipenem monohydrate - EMEA-C2-001809-PIP01-15-M01

Merck Sharp & Dohme (Europe), Inc.; Treatment of Gram-negative bacterial infections

Day 60 letter

Infectious Diseases

Summary of committee discussion

The PDCO considered that the initiation of Study 3 and Study 6 is compliant with the latest Agency's Decision P/0173/2019 of 15 May 2019.

The PDCO finalised this partial compliance check via written procedure on 9 April 2020.

2.7. Partial Compliance Checks completed by EMA

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

2.7.1. Vosoritide - EMEA-C3-002033-PIP01-16-M01

BioMarin International Limited; Treatment of achondroplasia

Day 30 letter

Other

2.7.2. Autologous CD4+ and CD8+ cells expressing a CD19-specific chimeric antigen receptor (JCAR017) - EMEA-C2-001995-PIP01-16-M02

Celgene Europe B.V.; Treatment of mature B-cell neoplasms

Day 30 letter

Oncology

2.7.3. Cannabidiol - EMEA-C2-001964-PIP01-16-M01

GW Pharma (International) B.V.; Dravet syndrome / Lennox-Gastaut syndrome / Tuberous sclerosis complex / Infantile spasms

Day 30 letter

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. Pegzilarginase - Orphan - EMEA-001925-PIP02-19

Aeglea BioTherapeutics, Inc.; Arginase 1 deficiency

Day 90 discussion

Endocrinology-Gynaecology-Fertility-Metabolism

3.1.2. Ravagalimab - EMEA-002617-PIP01-19

Ulcerative colitis

Day 90 discussion

Immunology-Rheumatology-Transplantation

3.1.3. Tecovirimat monohydrate - Orphan - EMEA-001205-PIP02-19

SIGA Technologies, Inc.; Orthopoxvirus disease (smallpox, monkeypox, cowpox, and vaccinia complications)

Day 90 discussion

Infectious Diseases

3.1.4. Chimeric 2'-O-(2-methoxyethyl) modified antisense oligonucleotide - Orphan - EMEA-002546-PIP01-19

Roche Registration GmbH; Huntington's disease

Day 90 discussion

Neurology

3.1.5. Rozanolixizumab - Orphan - EMEA-002681-PIP01-19

UCB Pharma S.A.; Treatment of myasthenia gravis

Day 90 discussion

Neurology

3.1.6. Ondansetron - EMEA-002623-PIP01-19

Treatment of alcohol use disorder (AUD)

Day 90 discussion

Other / Neurology

3.1.7. Autologous CD34+ hematopoietic stem cells with a CRISPR-edited erythroid enhancer region of the BCL11A gene - Orphan - EMEA-002730-PIP02-19

Vertex Pharmaceuticals (Ireland) Limited; Treatment of sickle cell disease

Day 60 discussion

Haematology-Hemostaseology

3.1.8. Bimekizumab - EMEA-002189-PIP03-19

Treatment of chronic idiopathic arthritis (including rheumatoid arthritis, spondyloarthritis, psoriatic arthritis and juvenile idiopathic arthritis (JIA)) / Treatment of JIA (enthesitis-related arthritis (ERA) and juvenile psoriatic arthritis (JPsA)) in patients from ≥2 years to <18 years of age

Day 60 discussion

Immunology-Rheumatology-Transplantation

3.1.9. EMEA-002742-PIP01-19

Treatment of parainfluenza infection

Day 60 discussion

Infectious Diseases

3.1.10. 1-[({4-[(4-fluoro-2-methyl-1H-indol-5-yl)oxy]-6-methoxyquinolin-7-yl}oxy)methyl]cyclopropan-1-amine - Orphan - EMEA-002486-PIP03-20

Advenchen Laboratories, LLC.; Treatment of soft tissue sarcomas / Treatment of alveolar soft part sarcoma / Treatment of synovial sarcoma

Day 60 discussion

Oncology

3.1.11. Atropine - EMEA-002744-PIP01-19

Myopia / Treatment to slow myopia progression

Day 60 discussion

Ophthalmology

3.1.12. Rituximab / $CD3^{+}CD4^{+}CD25^{+}CD127\text{-}FoxP3^{+}$ regulatory T cells - EMEA-002737-PIP01-19

Treatment of type 1 diabetes mellitus (T1DM)

Day 60 discussion

Other / Endocrinology-Gynaecology-Fertility-Metabolism

3.1.13. Dexmedetomidine - EMEA-002758-PIP01-19

Treatment of acute agitation in bipolar disorder / Treatment of acute agitation in schizophrenia

Day 60 discussion

Psychiatry

3.1.14. 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 60 discussion

Uro-nephrology

3.1.15. EMEA-002630-PIP01-19

Chikungunya virus infection

Day 60 discussion

Vaccines / Infectious Diseases

Summary of committee discussion:

The PDCO re-discussed this PIP application and agreed, in line with the outcome at D30, on a request for modification.

3.1.16. EMEA-002735-PIP01-19

Cardioplegia / For induction of cardioplegia in paediatric patients of all ages undergoing cardiac surgery to correct congenital heart malformation in operations requiring CPB support

Day 30 discussion

Cardiovascular Diseases

3.1.17. Ezetimibe / rosuvastatin - EMEA-002202-PIP02-20

Prevention of cardiovascular events

Day 30 discussion

Cardiovascular Diseases

3.1.18. EMEA-002786-PIP01-20

Prevention of cardiovascular events due to atherosclerotic cardiovascular disease in patients with elevated lipoprotein a (Lp(a))

Day 30 discussion

Cardiovascular Diseases

3.1.19. Recombinant human IgG1λ monoclonal Fab antibody - EMEA-002766-PIP01-20

Reversal of antiplatelet effects of ticagrelor in patients with uncontrolled major or life-threatening bleeding or requiring urgent surgery or invasive procedure

Day 30 discussion

Cardiovascular Diseases

3.1.20. Trimetazidine / bisoprolol - EMEA-002768-PIP01-20

Treatment of ischaemic coronary artery disorders

Day 30 discussion

Cardiovascular Diseases

3.1.21. Insulin - EMEA-002761-PIP01-20

Type 2 diabetes mellitus / Type 1 diabetes mellitus

Day 30 discussion

Endocrinology-Gynaecology-Fertility-Metabolism

3.1.22. Metformin / Sitagliptin - EMEA-002732-PIP02-20

Improvement of glycaemic control in patients with type 2 diabetes

Day 30 discussion

Endocrinology-Gynaecology-Fertility-Metabolism

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

Genzyme Europe B.V.; Polycystic kidney, autosomal dominant / 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 30 discussion

Endocrinology-Gynaecology-Fertility-Metabolism / Uro-nephrology

3.1.24. Bispecific antibody binding to clotting factor IX and X - EMEA-002762-PIP02-20

Treatment of haemophilia A / Routine prophylaxis to prevent or reduce frequency of bleeding episodes in patients with haemophilia A (congenital factor VIII deficiency) with or without factor VIII inhibitors / For use in all age groups

Day 30 discussion

Haematology-Hemostaseology

3.1.25. Human fibrinogen - EMEA-002769-PIP01-20

Treatment of congenital fibrinogen deficiency

Day 30 discussion

Haematology-Hemostaseology

3.1.26. Ravagalimab - EMEA-002617-PIP02-19

Sjogrens syndrome

Day 30 discussion

Immunology-Rheumatology-Transplantation

3.1.27. Secukinumab - EMEA-000380-PIP06-19

Lupus nephritis

Day 30 discussion

Immunology-Rheumatology-Transplantation

3.1.28. Sutimlimab - EMEA-002542-PIP02-19

Immune thrombocytopenia purpura

Day 30 discussion

Immunology-Rheumatology-Transplantation / Haematology-Hemostaseology

3.1.29. Fully human IgG1 RB-1 YTE anti-RSV F monoclonal antibody (mAb) - EMEA-002755-PIP01-19

Prevention of lower respiratory tract infection caused by respiratory syncytial virus

Day 30 discussion

Infectious Diseases

3.1.30. Itraconazole - Orphan - EMEA-002787-PIP01-20

Laboratoires SMB S.A.; Prevention of invasive mould disease / Prevention of invasive mould

disease in adolescents with acute leukaemia and neutropaenia

Day 30 discussion

Infectious Diseases / Oncology

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

orphanix GmbH; Prevention of bronchopulmonary dysplasia (BPD)

Day 30 discussion

Neonatology - Paediatric Intensive Care

3.1.32. Acetyl-L-leucine - Orphan - EMEA-002796-PIP01-20

IntraBio Ltd.; Niemann-Pick disease type C

Day 30 discussion

Neurology

3.1.33. Adeno-associated viral vector serotype 9 containing the human mini-dystrophin gene - Orphan - EMEA-002741-PIP01-20

Pfizer Europe MA EEIG; Treatment of Duchenne muscular dystrophy

Day 30 discussion

Neurology

3.1.34. Edaravone - Orphan - EMEA-002785-PIP01-20

Treeway B.V.; Amyotrophic lateral sclerosis

Day 30 discussion

Neurology

3.1.35. EMEA-002763-PIP01-20

Paediatric low grade glioma / Relapsed or refractory paediatric low grade glioma in adolescents and children 1 year of age and older

Day 30 discussion

Oncology

3.1.36. Abiraterone / niraparib - EMEA-002789-PIP01-20

Treatment of prostate malignant neoplasms

Day 30 discussion

Oncology

3.1.37. Autologous tumor-infiltrating lymphocytes: lifileucel (LN-144)/LN-145 - EMEA-002776-PIP01-20

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

Day 30 discussion

Oncology

3.1.38. Humanised antibody targeting the inducible T cell co-stimulatory receptor - EMEA-002781-PIP01-20

Head and neck squamous cell carcinoma / Treatment of head and neck squamous cell carcinoma

Day 30 discussion

Oncology

3.1.39. Neratinib - EMEA-002783-PIP01-20

Malignant neoplasm of breast

Day 30 discussion

Oncology

3.1.40. Rucaparib - EMEA-002760-PIP01-19

Treatment of fallopian tube cancer / Treatment of ovarian cancer / Treatment of prostate malignant neoplasms / Treatment of primary peritoneal cancer

Day 30 discussion

Oncology

3.1.41. 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 30 discussion

Oncology

3.1.42. Adrenaline (epinephrine) - EMEA-002749-PIP01-19

Treatment of allergic reactions / The emergency treatment of allergic reactions including anaphylaxis

Day 30 discussion

Pneumology – Allergology

3.1.43. Codon-optimised human cystic fibrosis transmembrane conductance regulator messenger ribonucleic acid complexed with lipid-based nanoparticles - Orphan - EMEA-002733-PIP01-19

Real Regulatory Limited; Cystic fibrosis / Treatment of cystic fibrosis

Day 30 discussion

Pneumology – Allergology

3.1.44. Timapiprant - EMEA-002788-PIP01-20

Treatment of asthma / Add-on maintenance treatment for severe eosinophilic asthmatic patients, aged 6-17 years, inadequately controlled on ICS plus other controller medication

Day 30 discussion

Pneumology – Allergology

3.1.45. 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 30 discussion

Psychiatry

3.1.46. EMEA-002329-PIP02-20

Treatment of dermatitis and eczema / Treatment of chronic hand eczema, Treatment of atopic dermatitis

Dermatology

3.1.47. Remdesivir - EMEA-002826-PIP01-20

Coronavirus disease 2019 (COVID-19) / Treatment of paediatric patients aged from birth to < 18 years (weighing < 40kg) with coronavirus disease 2019 (COVID-19) / Indicated for the treatment of adults and adolescents 12 years or older ⩾ 40kg with coronavirus disease 2019 (COVID-19)

Immunology-Rheumatology-Transplantation

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. Live, attenuated, chimeric dengue virus, serotype 3 / Live, attenuated, chimeric dengue virus, serotype 4 / Live, attenuated, chimeric dengue virus, serotype 1 / Live, attenuated, chimeric dengue virus, serotype 2 - EMEA-C-001545-PIP01-13-M02

Sanofi Pasteur; Prevention of dengue disease

Day 30 discussion

Vaccines

3.3. Discussions on Modification of an Agreed Paediatric Investigation Plan

3.3.1. Ambrisentan - Orphan - EMEA-000434-PIP01-08-M07

Glaxo Group Limited; Pulmonary arterial hypertension / Treatment of pulmonary arterial hypertension

Day 30 discussion

Cardiovascular Diseases

3.3.2. Alogliptin benzoate (as alogliptin) - EMEA-000496-PIP01-08-M08

Takeda Development Centre Europe Ltd; Type 2 diabetes mellitus

Day 30 discussion

Endocrinology-Gynaecology-Fertility-Metabolism

3.3.3. Drospirenone / estetrol monohydrate - EMEA-001332-PIP01-12-M03

Estetra SPRL; Prevention of pregnancy

Day 30 discussion

Endocrinology-Gynaecology-Fertility-Metabolism

3.3.4. Maralixibat chloride - Orphan - EMEA-001475-PIP03-17-M02

Mirum Pharmaceuticals; Treatment of progressive familial intrahepatic cholestasis

Day 30 discussion

Gastroenterology-Hepatology

3.3.5. Mepolizumab - Orphan - EMEA-000069-PIP01-07-M06

GSK Trading Services Limited; Treatment of hypereosinophilic syndrome (HES)

Day 30 discussion

Haematology-Hemostaseology

3.3.6. RFVIIIFc-VWF-XTEN - Orphan - EMEA-002501-PIP01-18-M01

Bioverativ Therapeutics, Inc., a Sanofi Company; Treatment of haemophilia A / Treatment and prophylaxis of bleeding in patients with haemophilia A (congenital FVIII deficiency)

Day 30 discussion

Haematology-Hemostaseology

3.3.7. Otilimab - EMEA-001882-PIP02-16-M02

GlaxoSmithKline Trading Services Limited; Treatment of chronic idiopathic arthritis (including rheumatoid arthritis, psoriatic arthritis, ankylosing spondylarthritis and juvenile idiopathic arthritis) / Treatment of active systemic juvenile idiopathic arthritis (sJIA) in patients 1 year of age and older / Treatment of active polyarticular juvenile idiopathic arthritis (pJIA) in patients 2 years of age and older

Day 30 discussion

Immunology-Rheumatology-Transplantation

3.3.8. Doravirine - EMEA-001676-PIP01-14-M03

Merck Sharp & Dohme (Europe), Inc.; Treatment of human immunodeficiency virus-1 (HIV-1) infection / Antiretroviral therapy, in combination with other antiretroviral agents, for the treatment of HIV-1 infection in children aged from birth to 18 years

Day 30 discussion

Infectious Diseases

3.3.9. Eravacycline - EMEA-001555-PIP01-13-M04

Tetraphase Pharmaceuticals, Inc.; Complicated intra-abdominal infection

Day 30 discussion

Infectious Diseases

3.3.10. Fostemsavir (tromethamine) - EMEA-001687-PIP01-14-M04

ViiV Healthcare UK Ltd; Treatment of human immunodeficiency virus (HIV-1) infection / Treatment of HIV-1 infection as part of a combination therapy in paediatric patients who have no more than 2 remaining available fully active antiretroviral therapies

Day 30 discussion

Infectious Diseases

3.3.11. Tenofovir disoproxil fumarate / lamivudine / doravirine - EMEA-001695-PIP01-14-M03

Merck Sharp & Dohme (Europe), Inc.; Treatment of human immunodeficiency virus-1 (HIV-1) infection / Antiretroviral combination therapy, for the treatment of HIV-1 infection in adults and children aged 12 to 18 years

Day 30 discussion

Infectious Diseases

3.3.12. Vaborbactam / meropenem - EMEA-001731-PIP01-14-M02

Menarini International Operations Luxembourg S.A.; Treatment of Gram-negative bacterial infections

Day 30 discussion

Infectious Diseases

3.3.13. Alemtuzumab - EMEA-001072-PIP01-10-M04

Genzyme Europe B.V.; Multiple sclerosis / For paediatric patients with relapsing remitting multiple sclerosis (RRMS) with active disease on prior disease modifying treatment (DMT) defined by clinical or imaging features

Day 30 discussion

Neurology

3.3.14. Brivaracetam - Orphan - EMEA-000332-PIP02-17-M01

UCB Pharma S.A.; Treatment of paediatric epilepsy syndromes / Monotherapy in patients 4 to 25 years of age with childhood absence epilepsy (CAE) and juvenile absence epilepsy (JAE)

Day 30 discussion

Neurology

3.3.15. Bempegaldesleukin - EMEA-002492-PIP01-18-M01

Nektar Therapeutics; Treatment of all conditions included in the category of malignant neoplasms (except nervous system, haematopoetic, and lymphoid tissue neoplasms) / Bempegaldesleukin in combination with nivolumab for treatment of patients with unresectable or metastatic melanoma in the age group from 12 years to less than 18 years of age / Bempegaldesleukin in combination with nivolumab for treatment of a relapsed or refractory paediatric malignant solid tumour in paediatric patients less than 18 years of age

Day 30 discussion

Oncology

3.3.16. Bosutinib - EMEA-000727-PIP01-09-M04

Pfizer Europe MA EEIG; Treatment of chronic myeloid leukaemia (CML) / Treatment of chronic, accelerated or blast phase CML with resistance or intolerance to prior tyrosine kinase inhibitor (TKI therapy) / Treatment of newly-diagnosed chronic phase of Philadelphia chromosome positive chronic myeloid leukaemia (Ph+ CML)

Day 30 discussion

Oncology

3.3.17. Brentuximab vedotin - Orphan - EMEA-000980-PIP01-10-M06

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

3.3.18. Olaparib - EMEA-002269-PIP01-17-M01

AstraZeneca AB; Treatment of all conditions included in the category of malignant neoplasms (except haematopoietic, and lymphoid tissue) / Treatment of paediatric patients from 6 months to ≤18 years old with homologous recombination repair (HRR) mutated solid tumours

Day 30 discussion

Oncology

3.3.19. Tapentadol - EMEA-000325-PIP01-08-M10

Grünenthal GmbH; Chronic pain / Treatment of chronic pain

Day 30 discussion

Pain

3.3.20. Vortioxetine - EMEA-000455-PIP02-10-M06

H. Lundbeck A/S; Major depressive disorder

Day 30 discussion

Psychiatry

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 letters of intent received for submission of applications with start of procedure 30 April 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. New Scientific Advice

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

5.2. Ongoing Scientific Advice

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

5.3. Final Scientific Advice (Reports and Scientific Advice letters)

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

Note: The annual reports on deferrals to be noted by the members of the PDCO are flagged in the Annex B.

9. Organisational, regulatory and methodological matters

9.1. Mandate and organisation of the PDCO

9.1.1. Updated meeting dates of PDCO plenaries for 2020-2021

Summary of committee discussion:

The new PDCO plenary meeting dates for 2020 and 2021 have been presented during the February 2020 PDCO meeting. The PDCO has adopted the changes as now published on the EMA website.

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 5 medicinal products with recommended paediatric indications adopted in March 2020 by CHMP. These include Fluad Tetra (influenza vaccine (surface antigen, inactivated), Zolgensma (Onasemnogene abeparvovec), INTELENCE (Etravirine), Kineret (Anakinra) and Ruconest (onestat alfa).

The list of procedures with paediatric indications to be evaluated by the CHMP, starting in March 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 identified the products which will require Non-clinical Working Group evaluation and discussion.

9.3.2. Formulation Working Group

PDCO member: Brian Aylward

Summary of committee discussion:

Relevant products for FWG discussion were identified.

9.3.3. HMPC request for PDCO input

Summary of committee discussion:

The PDCO was asked input from the HMPC in the framework of the finalisation of monograph.

9.4. Cooperation within the EU regulatory network

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 Annex to reflection paper on wording of the indication

PDCO Member: Siri Wang, Agnes Gyurasics

Summary of committee discussion:

The PDCO discussed wording of an indication concerning the paediatric population relating to age and/or body weight.

10.2 COVID-19 update

Summary of committee discussion:

Update on ongoing development for COVID treatments and vaccines.

11. Breakout sessions

11.1.1. Neonatology

Summary of committee discussion:

The Neonatology breakout session discussed a PIP for a condition specific to premature neonates.

12. List of Participants

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 on: -When not chairing the meeting: No participation in final deliberations and voting on:	- 2.1.4. Gepotidacin - EMEA-002443-PIP01-18; -2.1.5. Gepotidacin - EMEA-002443-PIP02-18 -2.3.18. Meningococcal group Y oligosaccharides conjugated to Corynebacterium diphtheriae CRM197 protein (MenABCWY) / recombinant Neisseria meningitis group B Protein 961c / recombinant Neisseria meningitis group B Protein 287- 953 / meningococcal group C oligosaccharides conjugated to Corynebacterium diphtheriae CRM197 protein / meningococcal group A oligosaccharides conjugated to Corynebacterium diphtheriae CRM197 protein / Outer Membrane Vesicles (OMV) from Neisseria meningitidis Strain NZ 98/254 / meningococcal group W-135 oligosaccharides conjugated to Corynebacterium diphtheriae CRM197 protein / recombinant Neisseria meningitis group B Protein 936-741 - EMEA-001260-PIP01-11-M01; -3.3.7. Otilimab - EMEA-001882-PIP02-16-M02 -3.1.39. Humanised antibody targeting the

Name	Role	Member state or affiliation	Outcome restriction following evaluation of e-DoI	Topics on agenda for which restrictions apply
				inducible T cell co-stimulatory receptor - EMEA-002781-PIP01-20 - 3.3.1. Ambrisentan - Orphan - EMEA-000434-PIP01-08-M07 - 3.3.5. Mepolizumab - Orphan - EMEA-000069-PIP01-07-M06
Karl-Heinz Huemer	Member	Austria	No interests declared	
Johanna Wernsperger	Alternate	Austria	No interests declared	
Karen Van Malderen	Alternate	Belgium	No interests declared	
Dimitar Roussinov	Member	Bulgaria	No restrictions applicable to this meeting	N/A
Georgios Savva	Member	Cyprus	No interests declared	
Lucie Kravackova	Member	Czech Republic	No interests declared	
Kirstine Moll Harboe	Member	Denmark	No interests declared	
Irja Lutsar	Member	Estonia	No interests declared	
Jana Lass	Alternate	Estonia	No interests declared	
Ann Marie Totterman	Member	Finland	No interests declared	
Sylvie Benchetrit	Member	France	No interests declared	
Dominique Ploin	Alternate	France	No interests declared	
Sabine Scherer	Member	Germany	No interests declared	
Yuansheng Sun	Alternate	Germany	No interests declared	
Eleni Katsomiti	Member	Greece	No interests declared	
Anastastia Mountaki	Alternate	Greece	No interests declared	
Ágnes Gyurasics	Member (CHMP member)	Hungary	No interests declared	
Brian Aylward	Member	Ireland	No interests declared	
Sara Galluzzo	Member	Italy	No interests declared	
Dina Apele-Freimane	Member	Latvia	No restrictions applicable to this meeting	N/A
Sigita Burokiene	Member	Lithuania	No interests declared	
Carola de Beaufort	Member (CHMP alternate)	Luxembourg	No interests declared	

Name	Role	Member state or affiliation	Outcome restriction following evaluation of e-DoI	Topics on agenda for which restrictions apply
Herbert Lenicker	Alternate	Malta	No interests declared	
Roel Bolt	Member	Netherlands	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	
Ninna Gullberg	Member	Sweden	No interests declared	
Eva Agurell	Alternate	Sweden	No interests declared	
Fernando Cabanas	Member	Healthcare Professionals' Representative	No restrictions applicable to this meeting	N/A
Jorrit Gerritsen	Alternate	Healthcare Professionals' Representative	No interests declared	
Francesca Rocchi	Member	Healthcare Professionals' Representative	No restrictions applicable to this meeting	N/A
Catherine Cornu	Alternate	Healthcare Professionals' Representative	No restrictions applicable to this meeting	N/A
Johannes Taminiau	Member	Healthcare Professionals' Representative	No interests declared	
Doina Plesca	Alternate	Healthcare Professionals' Representative	No interests declared	
Günter Karl-Heinz Auerswald	Member	Patients' Organisation Representative	No restrictions applicable to this meeting	N/A
Paola	Alternate	Patients'	No restrictions	N/A

Name	Role	Member state or affiliation	Outcome restriction following evaluation of e-DoI	Topics on agenda for which restrictions apply
Baiardi		Organisation Representative	applicable to this meeting	
Michal Odermarsky	Member	Patients' Organisation Representative	No restrictions applicable to this meeting	N/A
Dimitrios Athanasiou	Member	Patients' Organisation Representative	No interests declared	
Viviana Giannuzzi	Alternate	Patients' Organisation Representative	No participation in final deliberations and voting on:	- 2.1.14. Atropine - EMEA-002545-PIP01-19 (as informed during last meeting)
Maria Estela Moreno Martin	Expert – via telephone*	Spain	No interests declared	
Silvia Girotto	Expert– via telephone*	Italy	No interests declared	
Janet Koening	Expert– via telephone*	Germany	No interests declared	
Filip Josepsson	Expert– via telephone*	Sweden	No interests declared	
Andrea Laslop	Expert– via telephone*	Austria	No interests declared	
Eeva Leinonen	Expert– via telephone*	Finland	No interests declared	
David Khan	Expert– via telephone*	Sweden	No interests declared	
Kristin Karlsson	Expert– via telephone*	Sweden	No restrictions applicable to this meeting	
Doreen Fagan	Expert– via telephone*	Ireland	No restrictions applicable to this meeting	
Sheena Kennedy	Expert– via telephone*	Ireland	No restrictions applicable to this meeting	
Meeting run with support from relevant EMA staff				

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/