19 June 2023
EMA/CHMP/243980/2023
Human Medicines Division

Committee for medicinal products for human use (CHMP)

Draft agenda for the meeting on 19-22 June 2023

Chair: Harald Enzmann – Vice-Chair: Bruno Sepodes

19 June 2023, 09:00 – 19:30, virtual meeting/room 1C

20 June 2023, 08:30 – 19:30, virtual meeting/room 1C

21 June 2023, 08:30 – 19:30, virtual meeting/room 1C

22 June 2023, 08:30 – 15:00, virtual meeting/room 1C

Disclaimers

Some of the information contained in this agenda 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 CHMP meeting highlights once the procedures are finalised and start of referrals will also be available.

Of note, this agenda is a working document primarily designed for CHMP members and the work the Committee undertakes.

Note on access to documents

Some documents mentioned in the agenda 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. Introduction

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

Pre-meeting list of participants and restrictions in relation to declarations of interests applicable to the items of the agenda for the CHMP plenary session to be held 19-22 June 2023. See June 2023 CHMP minutes (to be published post July 2023 CHMP meeting).

1.2. Adoption of agenda

CHMP agenda for 19-22 June 2023.

1.3. Adoption of the minutes

CHMP minutes for 22-25 May 2023.

Minutes from PReparatory and Organisational Matters (PROM) meeting held on 12 June 2023.

2. Oral Explanations

2.1. Pre-authorisation procedure oral explanations

2.1.1. adagrasib - EMEA/H/C/006013

treatment of patients with advanced non-small cell lung cancer (NSCLC) with KRAS G12C mutation

Scope: Oral explanation

Action: Oral explanation to be held on 20 June 2023 at 16:00

List of Outstanding Issues adopted on 25.05.2023, 23.02.2023. List of Questions adopted on 15.09.2022.

2.1.2. gefapixant - EMEA/H/C/005884

treatment of refractory or unexplained chronic cough

Scope: Possible oral explanation

Action: Oral explanation to be held on 21 June 2023 at 14:00

List of Outstanding Issues adopted on 26.04.2023, 27.01.2022, 16.12.2021, 14.10.2021. List of Questions adopted on 24.06.2021.

2.1.3. gefapixant - EMEA/H/C/005476

treatment of refractory or unexplained chronic cough

Scope: Possible oral explanation

Action: Oral explanation to be held on 21 June 2023 at 14:00

List of Outstanding Issues adopted on 26.04.2023, 27.01.2022, 16.12.2021, 14.10.2021. List of Questions adopted on 24.06.2021.

2.1.4. natalizumab - EMEA/H/C/005752

Therapy for active relapsing remitting multiple sclerosis (RRMS)

Scope: Oral explanation

Action: Oral explanation to be held on 19 June 2023 at 14:00

List of Outstanding Issues adopted on 26.04.2023. List of Questions adopted on 10.11.2022.

2.2. Re-examination procedure oral explanations

2.2.1. Lagevrio - molnupiravir - EMEA/H/C/005789

Merck Sharp & Dohme B.V.; treatment of coronavirus disease 2019 (COVID-19)

Scope: Oral explanation

Action: Oral explanation to be held on 20 June 2023 at 14:00

New active substance (Article 8(3) of Directive No 2001/83/EC)

Opinion adopted on 23.02.2023. List of Outstanding Issues adopted on 22.04.2022. List of Questions adopted on 24.02.2022, 16.12.2021.

See 5.1

2.3. Post-authorisation procedure oral explanations

No items

2.4. Referral procedure oral explanations

No items

3. Initial applications

3.1. Initial applications; Opinions

3.1.1. sodium phenylbutyrate / ursodoxicoltaurine - Orphan - EMEA/H/C/005901

Amylyx Pharmaceuticals EMEA B.V.; treatment of amyotrophic lateral sclerosis (ALS)

Scope: Opinion; third party interventions

Action: For adoption

List of Outstanding Issues adopted on 23.02.2023, 10.11.2022. List of Questions adopted on 23.06.2022.

3.1.2. atogepant monohydrate - EMEA/H/C/005871

Prophylaxis of migraine in adults who have at least 4 migraine days per month.

Scope: Opinion

Action: For adoption

List of Outstanding Issues adopted on 30.03.2023. List of Questions adopted on 10.11.2022.

3.1.3. enalapril maleate - PUMA - EMEA/H/C/005731

treatment of heart failure

Scope: Opinion

Action: For adoption

List of Outstanding Issues adopted on 25.05.2023, 23.02.2023. List of Questions adopted on 21.07.2022.

3.1.4. daprodustat - EMEA/H/C/005746

treatment of anaemia associated with chronic kidney disease (CKD) in adults

Scope: Opinion

Action: For adoption

List of Outstanding Issues adopted on 25.05.2023, 30.03.2023, 10.11.2022. List of Questions adopted on 23.06.2022.

3.2. Initial applications; List of outstanding issues (Day 180; Day 120 for procedures with accelerated assessment timetable)

3.2.1. respiratory syncytial virus vaccines - EMEA/H/C/006027

Accelerated assessment

prevention of respiratory tract disease

Scope: List of outstanding issues

Action: For adoption

List of Questions adopted on 24.04.2023.

3.2.2. dantrolene sodium, hemiheptahydrate - Orphan - EMEA/H/C/006009

Norgine B.V.; treatment of malignant hyperthermia (including suspected cases)

Scope: List of outstanding issues

Action: For adoption

List of Questions adopted on 10.11.2022.

3.2.3. latanoprost - EMEA/H/C/005933

Reduction of elevated intraocular pressure (IOP)

Scope: List of outstanding issues

Action: For adoption

List of Questions adopted on 26.01.2023.

3.2.4. talquetamab - PRIME - Orphan - EMEA/H/C/005864

Accelerated assessment

Janssen-Cilag International N.V.; monotherapy treatment of adult patients with relapsed and refractory multiple myeloma

Scope: List of outstanding issues

Action: For adoption

List of Questions adopted on 24.04.2023.

3.2.5. fezolinetant - EMEA/H/C/005851

treatment of moderate to severe vasomotor symptoms (VMS) associated with menopause

Scope: List of outstanding issues

Action: For adoption

List of Questions adopted on 26.01.2023.

3.2.6. zilucoplan - Orphan - EMEA/H/C/005450

UCB Pharma S.A.; treatment of generalised myasthenia gravis in adults

Scope: List of outstanding issues

Action: For adoption

List of Questions adopted on 26.01.2023.

3.3. Initial applications; List of questions (Day 120; Day 90 for procedures with accelerated assessment timetable)

3.3.1. methylphenidate hydrochloride - PUMA - EMEA/H/C/005975

treatment of Attention Deficit Hyperactivity Disorder (ADHD) in children aged 6 years of age and over

Scope: List of questions

Action: For adoption

3.3.2. buprenorphine - EMEA/H/C/006188

treatment of opioid drug dependence

Scope: List of questions

Action: For adoption

3.3.3. rdESAT-6 / rCFP-10 - EMEA/H/C/006177

Diagnosis of infection with Mycobacterium tuberculosis

Scope: List of questions

Action: For adoption

3.3.4. sugemalimab - EMEA/H/C/006088

treatment of adults with metastatic non-small-cell lung cancer (NSCLC)

Scope: List of questions

Action: For adoption

3.3.5. aprocitentan - EMEA/H/C/006080

treatment of resistant hypertension

Scope: List of questions

Action: For adoption

3.4. Update on on-going initial applications for Centralised procedure

3.4.1. polihexanide - Orphan - EMEA/H/C/005858

SIFI SPA; For the treatment of acanthamoeba keratitis

Scope: Letter by the applicant dated 16.06.2023 requesting an extension to the clock stop to respond to the list of questions adopted in September 2022.

Action: For adoption

List of Questions adopted on 15.09.2022.

3.4.2. aumolertinib - EMEA/H/C/006069

treatment of non-small cell lung cancer

Scope: Letter by the applicant dated 09.06.2023 requesting an extension to the clock stop to respond to the list of questions adopted in March 2023.

Action: For adoption

List of Questions adopted on 30.03.2023.

3.4.3. exagamglogene autotemcel - PRIME - Orphan - ATMP - EMEA/H/C/005763

Vertex Pharmaceuticals (Ireland) Limited; treatment of transfusion-dependent β-thalassemia and sickle cell disease

Scope: Letter by the applicant requesting a change to the clock stop to respond to the list of questions adopted in May 2023.

Action: For information

List of Questions adopted on 17.05.2023.

3.4.4. dabigatran etexilate - EMEA/H/C/006023

Prevention of venous thromboembolic events

Scope: Letter by the applicant dated 14.06.2023 requesting an extension to the clock stop to respond to the list of outstanding issues adopted in March 2023.

Action: For adoption

List of outstanding issues adopted on 30.03.2023. List of Questions adopted on 21.07.2022.

3.4.5. omecamtiv mecarbil - EMEA/H/C/006112

treatment of adult patients with symptomatic chronic heart failure and reduced ejection fraction less than 30%

Scope: Letter by the applicant dated 09.06.2023 requesting an extension to the clock stop to respond to the list of questions adopted in April 2023.

Action: For adoption

List of questions adopted on 26.04.2023.

3.4.6. bevacizumab - EMEA/H/C/005723

Treatment of neovascular (wet) age-related macular degeneration (nAMD).

Scope: Letter by the applicant dated 09.06.2023 requesting an extension to the clock stop to respond to the list of questions adopted in April 2023.

Action: For adoption

List of questions adopted on 26.04.2023.

3.4.7. omaveloxolone - Orphan - EMEA/H/C/006084

Reata Ireland Limited; Treatment of Friedreich's ataxia

Scope: Letter by the applicant dated 06.06.2023 requesting an extension to the clock stop to respond to the list of questions adopted in April 2023.

Action: For adoption

List of questions adopted on 26.04.2023.

3.5. Re-examination of initial application procedures under Article 9(2) of Regulation no 726/2004

3.5.1. Lagevrio - molnupiravir - EMEA/H/C/005789

Merck Sharp & Dohme B.V.; treatment of coronavirus disease 2019 (COVID-19)

Scope: Opinion

Action: For adoption

New active substance (Article 8(3) of Directive No 2001/83/EC)

Opinion adopted on 23.02.2023. List of Outstanding Issues adopted on 22.04.2022. List of Questions adopted on 24.02.2022, 16.12.2021.

See 2.2

3.6. Initial applications in the decision-making phase

No items

3.7. Withdrawals of initial marketing authorisation application

3.7.1. pegfilgrastim - EMEA/H/C/005810

Treatment of neutropenia

Scope: Withdrawal of marketing authorisation application

Action: For information

List of outstanding issues adopted on 13.10.2022. List of Questions adopted on 27.01.2022.

3.7.2. filgrastim - EMEA/H/C/005888

Reduction in the duration of neutropenia and the incidence of febrile neutropenia, indicated for the mobilisation of peripheral blood progenitor cells and persistent neutropenia in patients with advanced HIV infection

Scope: Withdrawal of marketing authorisation application

Action: For information

List of outstanding issues adopted on 10.11.2022. List of Questions adopted on 23.06.2022.

4. Extension of marketing authorisation according to Annex I of Commission Regulation (EC) No 1234/2008

4.1. Extension of marketing authorisation according to Annex I of Commission Regulation (EC) No 1234/2008; Opinion

4.1.1. Adtralza - tralokinumab - EMEA/H/C/005255/X/0007

LEO Pharma A/S

Rapporteur: Jayne Crowe, PRAC Rapporteur: Kimmo Jaakkola

Scope: "Extension application to add a new strength of 300 mg (150 mg/ml) tralokinumab solution for injection in pre-filled pen for subcutaneous administration.
The RMP (version 1.1) is updated accordingly."

Action: For adoption

List of Questions adopted on 30.03.2023.

4.1.2. Comirnaty - COVID-19 mRNA vaccine (nucleoside-modified) - EMEA/H/C/005735/X/0180

BioNTech Manufacturing GmbH

Rapporteur: Filip Josephson, PRAC Rapporteur: Menno van der Elst

Scope: "Extension application to add a new pharmaceutical form of Comirnaty 5/5 µg (tozinameran, famtozinameran) dispersion for injection for active immunisation for children aged 5 to 11 years of age."

Action: For adoption

4.1.3. Erleada - apalutamide - EMEA/H/C/004452/X/0028/G

Janssen-Cilag International N.V.

Rapporteur: Carolina Prieto Fernandez, PRAC Rapporteur: Tiphaine Vaillant

Scope: "Extension application to add a new strength (240 mg) film-coated tablets grouped with the IB variation (C.I.z).
The RMP (version 6.1) has also been submitted.
C.I.z (IB): to align the SmPC/PL for Erleada 60 mg with the SmPC/PL proposed for the registration of the new Erleada film-coated tablet strength, 240 mg.
The PL for Erleada 60 mg is proposed to be updated to ensure consistency.
In addition, few minor revisions are proposed to the SmPC for Erleada 60 mg, to align the SmPC proposed for the 240 mg strength:
- SmPC sections 5.1 and 5.2: Orthographic corrections
- SmPC section 6.5: Further details on the description of the current packaging have been added, this change does not result from a change to the container.
- SmPC section 6.6: The title of the section has been aligned with QRD template."

Action: For adoption

List of Questions adopted on 30.03.2023.

4.1.4. Ofev - nintedanib - EMEA/H/C/003821/X/0052/G

Boehringer Ingelheim International GmbH

Rapporteur: Finbarr Leacy, Co-Rapporteur: Ewa Balkowiec Iskra, PRAC Rapporteur: Nikica Mirošević Skvrce

Scope: "Extension application to add a new strength of 25 mg soft capsule grouped with a type II variation C.I.6.a to add a new indication of treatment of fibrosing Interstitial Lung Diseases (ILDs) in children and adolescents from 6 to 17 years of age, based on results from study 1199 0337 (InPedILD); a randomised, placebo-controlled, double-blind, multicentre, multinational, phase III clinical trial undertaken to evaluate dose-exposure and safety of nintedanib on top of standard of care in children and adolescents (6 to 17 years old) with clinically significant fibrosing ILD. As a consequence, sections 4.1, 4.2, 4.4, 4.5, 4.8, 5.1, 5.2 and 5.3 of the SmPC are updated. The Package Leaflet is updated in accordance. In addition, the MAH took the opportunity to implement minor editorial changes to the list of local representatives in the Package Leaflet. The updated RMP version 12.0 is also submitted."

Action: For adoption

List of Outstanding Issues adopted on 26.04.2023. List of Questions adopted on 26.01.2023.

4.1.5. Tenkasi - oritavancin - EMEA/H/C/003785/X/0036

Menarini International Operations Luxembourg S.A.

Rapporteur: Janet Koenig, PRAC Rapporteur: Adam Przybylkowski

Scope: "Extension application to add a new strength of 1200 mg for powder for concentrate for solution for infusion. The RMP (version 4) is updated in accordance."

Action: For adoption

List of Outstanding Issues adopted on 26.04.2023, 23.02.2023. List of Questions adopted on 21.07.2022.

4.1.6. Xolair - omalizumab - EMEA/H/C/000606/X/0115/G

Novartis Europharm Limited

Rapporteur: Kristina Dunder, PRAC Rapporteur: Mari Thorn

Scope: "Extension application to add a new strength of 300 mg (150 mg/ml) for Xolair solution for injection grouped with quality type II, IB and IAIN variations. The RMP (version 17.0) is updated in accordance."

Action: For adoption

List of Questions adopted on 15.12.2022.

4.2. Extension of marketing authorisation according to Annex I of Commission Regulation (EC) No 1234/2008; Day 180 list of outstanding issues

No items

4.3. Extension of marketing authorisation according to Annex I of Commission Regulation (EC) No 1234/2008; Day 120 List of question

4.3.1. Comirnaty - COVID-19 mRNA vaccine (nucleoside-modified) - EMEA/H/C/005735/X/0176

BioNTech Manufacturing GmbH

Rapporteur: Filip Josephson, PRAC Rapporteur: Menno van der Elst

Scope: "Extension application to add a new strength of 1.5/1.5 µg (tozinameran, famtozinameran) for active immunisation in infants and children between 6 months to 4 years of age."

Action: For adoption

4.3.2. Eylea - aflibercept - EMEA/H/C/002392/X/0084/G

Bayer AG

Rapporteur: Jean-Michel Race, PRAC Rapporteur: Nathalie Gault

Scope: "Extension application to add a new strength of Aflibercept 114.3 mg/ml solution for injection (in a vial), to be indicated in adults for the (1) treatment of neovascular (wet) age-related macular degeneration (nAMD) and (2) visual impairment due to diabetic macular oedema (DME), grouped with a type II variation (B.II.g.2) to introduce a post-approval change management protocol to add a new presentation for Aflibercept solution 114.3 mg/ml in a single-use pre-filled syringe for intravitreal injection."

Action: For adoption

4.3.3. Lumykras - sotorasib - EMEA/H/C/005522/X/0009

Amgen Europe B.V.

Rapporteur: Alexandre Moreau

Scope: "Extension application to add a new strength of 240 mg film-coated tablet."

Action: For adoption

4.3.4. Talzenna - talazoparib - EMEA/H/C/004674/X/0015/G

Pfizer Europe MA EEIG

Rapporteur: Filip Josephson, Co-Rapporteur: Hrefna Gudmundsdottir, PRAC Rapporteur: Inês Ribeiro-Vaz

Scope: "Extension application for Talzenna to introduce a new strength of 0.1 mg hard capsules, grouped with a type II variation (C.I.6.a) in order to extend the indication for Talzenna in combination with enzalutamide for the treatment of adult patients with metastatic castration-resistant prostate cancer (mCRPC), based on final results from study C3441021 (TALAPRO-2) as well as supplemental data from study C3441006 (TALAPRO-1). Study C3441021 (TALAPRO-2) is a randomized, double-blind, placebo-controlled, phase 3 study of talazoparib in combination with enzalutamide in mCRPC, while study C3441006 (TALAPRO-1) is a phase 2, open-label, response rate study of talazoparib in men with DNA repair defects and mCRPC who previously received taxane-based chemotherapy and progressed on at least one novel hormonal agent. As a consequence, sections 1, 2, 3, 4.1, 4.2, 4.5, 4.7, 4.8, 5.1, 5.2, 6.1, 6.5 and 8 of the SmPC are updated. The Package Leaflet and Labelling are updated in accordance. Version 1.1 of the RMP has also been submitted. In addition, the MAH took the opportunity to introduce minor editorial changes to the PI."

Action: For adoption

4.3.5. Uptravi - selexipag - EMEA/H/C/003774/X/0038

Janssen-Cilag International N.V.

Rapporteur: Martina Weise, PRAC Rapporteur: Nathalie Gault

Scope: "Extension application to add a new strength of 100 µg film-coated tablets in HDPE bottle. The RMP (version 10.1) is updated in accordance."

Action: For adoption

4.4. Update on on-going extension application according to Annex I of Commission Regulation (EC) No 1234/2008

No items

4.5. Re-examination procedure of extension of marketing authorisation according to Annex I of Commission Regulation (EC) No 1234/2008

No items

5. Type II variations - variation of therapeutic indication procedure according to Annex I of Commission Regulation (EC) No 1234/2008

5.1. Type II variations - variation of therapeutic indication procedure according to Commission Regulation (EC) No 1234/2008; Opinions or Requests for supplementary information

5.1.1. Abecma - idecabtagene vicleucel - Orphan - ATMP - EMEA/H/C/004662/II/0031

Bristol-Myers Squibb Pharma EEIG

Rapporteur: Rune Kjeken, Co-Rapporteur: Heli Suila, CHMP Coordinators: Ingrid Wang, Johanna Lähteenvuo, PRAC Rapporteur: Ulla Wändel Liminga

Scope: "Extension of indication to include treatment of adult patients with relapsed and refractory multiple myeloma (RRMM) who have received at least two prior therapies, including an immunomodulatory agent, a proteasome inhibitor and an anti-CD-38 antibody and have demonstrated disease progression on the last therapy for Abecma (idecabtagene vicleucel, ide-cel), based on results from study BB2121-MM-003 (MM-003, KarMMa-3). This is a Phase 3, multicentre, randomised, open-label study to compare the efficacy and safety of ide-cel versus standard regimens in subjects with RRMM. As a consequence, sections 2.1, 2.2, 4.1, 4.2, 4.4, 4.5, 4.8, 5.1, 5.2, 6.3, 6.4 and 6.6 of the SmPC are updated. The Package Leaflet and Labelling are updated in accordance. Version 3.0 of the RMP has also been submitted. Furthermore, the PI is brought in line with the Guideline on core SmPC, Labelling and Package Leaflet for advanced therapy medicinal products (ATMPs) containing genetically modified cells.", Request for 1 year of market protection for a new indication (Article 14(11) of Regulation (EC) 726/2004)

Action: For adoption

5.1.2. Adcetris - brentuximab vedotin - Orphan - EMEA/H/C/002455/II/0107

Takeda Pharma A/S

Rapporteur: Johann Lodewijk Hillege

Scope: "Extension of indication to include treatment of adult patients with previously untreated CD30+ advanced (including Stage III) Hodgkin lymphoma (HL), in combination with doxorubicin, vinblastine and dacarbazine (AVD), for Adcetris, based on the second interim analysis of OS data from ECHELON-1 study (C25003); this is a randomized, open-label, phase 3 trial of A+AVD versus ABVD as frontline therapy in patients with advanced classical HL. As a consequence, sections 4.1 and 5 of the SmPC are updated."

Action: For adoption

5.1.3. Brukinsa - zanubrutinib - EMEA/H/C/004978/II/0014

BeiGene Ireland Ltd

Rapporteur: Aaron Sosa Mejia, Co-Rapporteur: Johanna Lähteenvuo, PRAC Rapporteur:

Menno van der Elst

Scope: "Extension of indication to include in combination with obinutuzumab treatment of adult patients with relapsed or refractory follicular lymphoma who have received at least two prior systemic treatments for Brukinsa; based on results from studies BGB-3111-212 and BGB-3111-GA101-001. BGB-3111-212 is an ongoing international, Phase 2, open-label, randomized (2:1), active control study of zanubrutinib plus obinutuzumab (Arm A) versus obinutuzumab monotherapy (Arm B) in patients with R/R FL. The primary efficacy endpoint is overall response rate (ORR); while BGB-3111-GA101-001 is a Phase 1b Study to Assess Safety, Tolerability and Antitumor Activity of the Combination of BGB-3111 with Obinutuzumab in Subjects with B-Cell Lymphoid Malignanciesa. As a consequence, sections 4.1, 4.4, 4.8 and 5.1 of the SmPC are updated. The Package Leaflet is updated in accordance. Version 3.1 of the RMP has also been submitted. In addition, the MAH took the opportunity to implement editorial changes to the SmPC."

Action: For adoption

5.1.4. Bylvay - odevixibat - Orphan - EMEA/H/C/004691/II/0011

Albireo

Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Adam Przybylkowski

Scope: "Extension of indication to include treatment of cholestasis and pruritus in Alagille syndrome (ALGS) in patients from birth and older for Bylvay, based on final results from study A4250-012 and interim results from study A4250-015. Study A4250-012 is a 24-week, randomised, double-blind, placebo-controlled Phase III study conducted in 52 patients with a genetically confirmed diagnosis of ALGS and presence of pruritus and high serum bile acid levels at baseline. Study A4250-015 is an ongoing 72-week open-label extension trial for patients who completed study A4250-012 and evaluates the long-term safety and efficacy of Bylvay in patients with ALGS. As a consequence, sections 4.1, 4.2, 4.8, 5.1 and 5.2 of the SmPC are updated. The Package Leaflet is updated in accordance. Version 2.1 of the RMP has also been submitted." Request for 1 year of market protection for a new indication (Article 14(11) of Regulation (EC) 726/2004)

Action: For adoption

Request for Supplementary Information adopted on 30.03.2023.

5.1.5. Comirnaty - COVID-19 mRNA vaccine (nucleoside-modified) - EMEA/H/C/005735/II/0177/G

BioNTech Manufacturing GmbH

Rapporteur: Filip Josephson, PRAC Rapporteur: Menno van der Elst

Scope: "C.I.6: Extension of indication to include Comirnaty Original/Omicron BA.4-5 (5/5 micrograms)/dose concentrate for dispersion for injection and Comirnaty Original/Omicron BA.4-5 (15/15 micrograms)/dose dispersion for injection for use as primary vaccination course against COVID-19 in children aged 5 to 11 years and in individuals 12 years of age and older, respectively, based on interim results from studies C4591044 and C4591048. Study C4591044 is an interventional, randomized, active-controlled, phase 2/3 study to investigate the safety, tolerability, and immunogenicity of bivalent BNT162b RNA-based vaccine candidates as a booster dose in COVID-19 vaccine–experienced healthy individuals,

while study C4591048 is a phase 1/2/3 master study to investigate the safety, tolerability, and immunogenicity of a bivalent BNT162b2 RNA-based vaccine candidate. As a consequence, sections 4.1, 4.2, 4.8 and 5.1 of the SmPC are updated. The Package Leaflet is updated in accordance. Version 9.1 of the RMP has also been submitted. In addition, the MAH took the opportunity to introduce minor editorial changes to the PI.
A.6: To change the ATC Code of tozinameran, riltozinameran and famtozinameran from J07BX03 to J07BN01."

Action: For adoption

5.1.6. Imjudo - tremelimumab - EMEA/H/C/006016/II/0001

AstraZeneca AB

Rapporteur: Aaron Sosa Mejia

Scope: "Extension of indication to include in combination with durvalumab and platinum-based chemotherapy, the first-line treatment of adults with metastatic non-small cell lung cancer (NSCLC) with no sensitising EGFR mutations or ALK positive mutations for Imjudo, based on the final analysis from the pivotal study D419MC00004, a Randomised, Multi-center, Open-Label, Comparative Global Study to Determine the Efficacy of Durvalumab or Durvalumab and Tremelimumab in Combination with Platinum-Based Chemotherapy for First-Line Treatment in Patients with Metastatic Non Small-Cell Lung Cancer (NSCLC) (POSEIDON). As a consequence, sections 2, 4.1, 4.2, 4.4, 4.5, 4.8, 5.1, 5.2, 5.3 and 6.6 of the SmPC are updated. The Package Leaflet is updated in accordance. In addition, the MAH took the opportunity to include editorial changes."

Action: For adoption

Request for Supplementary Information adopted on 25.05.2023.

5.1.7. Jardiance - empagliflozin - EMEA/H/C/002677/II/0074

Boehringer Ingelheim International GmbH

Rapporteur: Johann Lodewijk Hillege, Co-Rapporteur: Karin Janssen van Doorn, PRAC Rapporteur: Maria del Pilar Rayon

Scope: "Extension of indication to include treatment of chronic kidney disease (CKD) for Jardiance, based on final results from study EMPA-KIDNEY (1245-0137) listed as a category 3 study in the RMP; this is a Phase III, multicentre international randomised parallel group double-blind placebo controlled clinical trial of empagliflozin once daily to assess cardio-renal outcomes in patients with chronic kidney disease. As a consequence, sections 4.1, 4.2, 4.4, 4.8, 5.1 and 5.2 of the SmPC are updated. The Package Leaflet is updated in accordance. Version 19.0 of the RMP has also been submitted. Furthermore, the PI is brought in line with the latest QRD template version 10.3."

Action: For adoption

Request for Supplementary Information adopted on 30.03.2023.

5.1.8. Keytruda - pembrolizumab - EMEA/H/C/003820/II/0134

Merck Sharp & Dohme B.V.

Rapporteur: Armando Genazzani, PRAC Rapporteur: Menno van der Elst

Scope: "Extension of indication to include in combination with platinum-containing chemotherapy as neoadjuvant treatment, and then continued as monotherapy as adjuvant, treatment of resectable stage II, IIIA, or IIIB (T3 4N2) non small cell lung carcinoma in adults for Keytruda based on study KEYNOTE-671, a phase III, randomized, double-blind trial of platinum doublet chemotherapy +/- pembrolizumab as neoadjuvant/adjuvant therapy for participants with resectable stage II, IIIA, and resectable IIIB (T3-4N2) non-small cell lung cancer. As a consequence, sections 4.1, 4.2 and 5.1 of the SmPC are updated. The Package Leaflet is updated in accordance. Version 41.1 of the RMP has also been submitted."

Action: For adoption

5.1.9. Keytruda - pembrolizumab - EMEA/H/C/003820/II/0135

Merck Sharp & Dohme B.V.

Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Menno van der Elst

Scope: "Extension of indication to include in combination with chemotherapy the first-line treatment of locally advanced unresectable or metastatic HER2-negative gastric or gastroesophageal junction adenocarcinoma in adults based on study KEYNOTE-859, a randomized, double-blind phase 3 trial, evaluating Keytruda in combination with chemotherapy compared to placebo in combination with chemotherapy for the first-line treatment of patients with HER2-negative locally advanced unresectable or metastatic gastric or GEJ adenocarcinoma. As a consequence, sections 4.1 and 5.1 of the SmPC are updated. The Package Leaflet and Annex II are updated in accordance. Version 42.1 of the RMP has also been submitted."

Action: For adoption

5.1.10. Lonsurf - trifluridine / tipiracil - EMEA/H/C/003897/II/0026

Les Laboratoires Servier

Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Ulla Wändel Liminga

Scope: "Extension of indication to include treatment of patients with refractory metastatic colorectal cancer, for Lonsurf in combination with bevacizumab based on results from study SUNLIGHT (CL3-95005-007); This is an open-label, randomised, phase III study comparing trifluridine/tipiracil in combination with bevacizumab to trifluridine/tipiracil monotherapy in patients with refractory metastatic colorectal cancer. As a consequence, sections 4.1, 4.2, 4.4, 4.8 and 5.1 of the SmPC. The package leaflet is updated in accordance. The updated RMP version 9.1 has also been submitted. In addition, the MAH took the opportunity to update section 4.6 of the SmPC and the package leaflet accordingly."

Action: For adoption

Request for Supplementary Information adopted on 25.05.2023.

5.1.11. Mircera - methoxy polyethylene glycol-epoetin beta - EMEA/H/C/000739/II/0092

Roche Registration GmbH

Rapporteur: Maria Concepcion Prieto Yerro

Scope: "Extension of indication to include treatment of paediatric patients from 3 months to less than 18 years of age requiring dialysis or not yet on dialysis and switching from another ESA to Mircera, based on final results from study NH19708; this is a single-arm, open-label, Phase II study of Mircera in patients aged 3 months to <18 years with CKD on dialysis or not yet on dialysis to generate PK, efficacy, and safety data for subcutaneous (SC) administration of Mircera. In addition, supportive data from studies NH19707, Modelling & Simulation study (study 3) and MH40258 were included. As a consequence, sections 4.1, 4.2, 4.8, 5.1 and 5.2 of the SmPC are updated. The Package Leaflet is updated in accordance. In addition, the MAH took the opportunity to introduce minor editorial changes to the PI and to update the Instruction for Use in the Package Leaflet."

Action: For adoption

Request for Supplementary Information adopted on 23.02.2023, 15.09.2022.

5.1.12. Mounjaro - tirzepatide - EMEA/H/C/005620/II/0007

Eli Lilly Nederland B.V.

Rapporteur: Martina Weise, Co-Rapporteur: Kristina Dunder, PRAC Rapporteur: Menno van der Elst

Scope: "Extension of indication to include chronic weight management, including weight loss and weight maintenance, for Mounjaro, as an adjunct to a reduced-calorie diet and increased physical activity in adults with an initial body mass index (BMI) of ≥ 30 kg/m2 (obesity), or ≥ 27 kg/m2 to < 30 kg/m2 (overweight) in the presence of at least one weight-related comorbid condition, based on a global, pivotal phase 3 study I8F-MC-GPHK (SURMOUNT-1) and five supportive phase 3 studies (SURPASS-1 to -5) in participants with T2DM and BMI ≥ 27 kg/m2. SURMOUNT-1 is a phase 3, randomized, double-blind, placebo-controlled trial to investigate the efficacy and safety of tirzepatide once weekly in participants without type 2 diabetes who have obesity or are overweight with weight related comorbidities. As a consequence, sections 4.1, 4.8, 5.1 and 5.2 of the SmPC are updated. The Package Leaflet is updated in accordance. Version 2.1 of the RMP has also been submitted. In addition, the MAH took the opportunity to introduce minor editorial changes to the PI.", Request for 1 year of market protection for a new indication (Article 14(11) of Regulation (EC) 726/2004)

Action: For adoption

5.1.13. NexoBrid - concentrate of proteolytic enzymes enriched in bromelain - EMEA/H/C/002246/II/0058

MediWound Germany GmbH

Rapporteur: Janet Koenig, PRAC Rapporteur: Martin Huber

Scope: "Extension of current indication for removal of eschar in adults with deep partial- and full-thickness thermal burns to the paediatric population for NexoBrid based on interim results from study MW2012-01-01 (CIDS study), listed as study MW2012-01-01 is a 3-stage, multi-centre, multi-national, randomised, controlled, open label, 2 arm study aiming to demonstrate the superiority of NexoBrid treatment over SOC treatment in paediatric patients (aged 0 to 18 years) with deep partial thickness (DPT) and full thickness (FT)

thermal burns of 1% to 30% of total body surface area (TBSA).
As a consequence, sections 4.1, 4.2, 4.8, 5.1, 5.2 and 5.3 of the SmPC are updated. The Package Leaflet is updated accordingly. Version 9 of the RMP has also been submitted."

Action: For adoption

Request for Supplementary Information adopted on 15.12.2022.

5.1.14. Pepaxti - melphalan flufenamide - EMEA/H/C/005681/II/0002

Oncopeptides AB

Rapporteur: Johann Lodewijk Hillege, Co-Rapporteur: Elita Poplavska, PRAC Rapporteur: Martin Huber

Scope: "Extension of indication to include treatment of patients with Multiple Myeloma who have received at least two prior lines of therapies for Pepaxti, based on final results from study OP-103 OCEAN; this is a randomized, open-label phase III study in patients with relapsed or refractory multiple myeloma following two to four lines of prior therapies and who were refractory to lenalidomide and the last line of therapy. As a consequence, sections 4.1, 4.4, 4.8 and 5.1 of the SmPC are updated. The Package Leaflet is updated in accordance. Version 1.1 of the RMP has also been submitted. In addition, the MAH took the opportunity to implement editorial changes in the SmPC."

Action: For adoption

Request for Supplementary Information adopted on 30.03.2023.

5.1.15. Praluent - alirocumab - EMEA/H/C/003882/II/0078

Sanofi Winthrop Industrie

Rapporteur: Johann Lodewijk Hillege, Co-Rapporteur: Alar Irs, PRAC Rapporteur: Gabriele Maurer

Scope: "Extension of indication to include treatment of paediatric patients 8 years of age and older with heterozygous familial hypercholesterolemia (HeFH) as an adjunct to diet, alone or in combination with other LDL-C lowering therapies, based on final results from study EFC14643 listed as a category 3 study in the RMP; this is a randomized, double-blind, placebo-controlled study followed by an open-label treatment period to evaluate the efficacy and safety of alirocumab in children and adolescents with heterozygous familial hypercholesterolemia. As a consequence, sections 4.1, 4.2, 4.8, 5.1 and 5.2 of the SmPC are updated. The Package Leaflet is updated in accordance. Version 8.0 of the RMP is also submitted."

Action: For adoption

5.1.16. Prevymis - letermovir - Orphan - EMEA/H/C/004536/II/0033/G

Merck Sharp & Dohme B.V.

Rapporteur: Filip Josephson, Co-Rapporteur: Aaron Sosa Mejia, PRAC Rapporteur: Kirsti Villikka

Scope: "Grouped application consisting of 1) Extension of indication to include prophylaxis

of cytomegalovirus in kidney transplant recipients (KTR) for Prevymis, based on final results from study P002MK8228; this is a Phase III, randomized, double-blind, active comparator-controlled study to evaluate the efficacy and safety of letermovir versus valganciclovir for the prevention of Human Cytomegalovirus (CMV) Disease in adult kidney transplant recipients. As a consequence, sections 4.1, 4.2, 4.4, 4.5, 4.8, 5.1, 5.2 and 5.3 of the SmPC are updated. The Package Leaflet is updated in accordance. Version 4.1 of the RMP has also been submitted. In addition, the MAH took the opportunity to implement editorial changes; 2) Update of section 4.2 of the SmPC in order to update duration of treatment recommendation based on final results from study P040MK8228; this is a Phase III randomized, double-blind, placebo-controlled clinical trial to evaluate the safety and efficacy of letermovir (LET) prophylaxis when extended from 100 days to 200 days post-transplant in cytomegalovirus (CMV) seropositive recipients (R+) of an allogeneic hematopoietic stem cell transplant (HSCT)."

Action: For adoption

5.1.17. Reblozyl - luspatercept - Orphan - EMEA/H/C/004444/II/0021

Bristol-Myers Squibb Pharma EEIG

Rapporteur: Daniela Philadelphy, Co-Rapporteur: Ewa Balkowiec Iskra, PRAC Rapporteur: Jo Robays

Scope: "Extension of indication to include treatment of adult patients with anaemia due to very low, low and intermediate-risk myelodysplastic syndromes (MDS), who may require RBC transfusions for Reblozyl, based on results from study ACE-536-MDS-002 (COMMANDS), an active-controlled, open-label, randomized Phase 3 study comparing the efficacy and safety of luspatercept vs epoetin alfa in adult subjects with anemia due to IPSS-R very low, low or intermediate risk MDS, who are ESA naïve and require RBC transfusions, and studies ACE-536-MDS-001(MEDALIST), ACE-536-MDS-004 , A536-03, A536-05 and ACE-536-LTFU-001; as a consequence, sections 4.1, 4.2, 4.4, 4.8, 5.1 and 5.2 of the SmPC are updated. The Package Leaflet is updated in accordance. Version 2.0 of the RMP has also been submitted."

Action: For adoption

5.1.18. Refixia - nonacog beta pegol - EMEA/H/C/004178/II/0032

Novo Nordisk A/S

Rapporteur: Daniela Philadelphy, Co-Rapporteur: Alexandre Moreau, PRAC Rapporteur: Gabriele Maurer

Scope: "Extension of indication to include treatment and prophylaxis of bleeding in children below 12 years of age with haemophilia B including previously untreated patients for Refixia, based on interim results from studies NN7999-3774 and NN7999-3895. NN7999-3774 is a multicentre, open-label, non-controlled study evaluating the safety, efficacy and pharmacokinetics of nonacog beta pegol in previously treated children with haemophilia B, while NN7999-3895 is a multicentre, open-label, single-arm, non-controlled trial evaluating the safety and efficacy of nonacog beta pegol in previously untreated patients with haemophilia B. As a consequence, sections 4.1, 4.2, 4.4, 4.8, 5.1, 5.2 and 5.3 of the SmPC are updated. The Package Leaflet is updated accordingly. Version 5.0 of the RMP has also

been submitted. In addition, the MAH took the opportunity to introduce minor editorial changes to the PI."

Action: For adoption

Request for Supplementary Information adopted on 30.03.2023.

5.1.19. Ryeqo - relugolix / estradiol / norethisterone acetate - EMEA/H/C/005267/II/0013/G

Gedeon Richter Plc.

Rapporteur: Johann Lodewijk Hillege, Co-Rapporteur: Jean-Michel Race, PRAC Rapporteur: Martin Huber

Scope: "Extension of indication to include treatment of moderate to severe pain associated with endometriosis for Ryeqo in adult women of reproductive age with a history of previous medical or surgical treatment for their endometriosis, based on final results from studies MVT-601-3101 and MVT-601-3102 and final results up to 104 weeks from study MVT-601-3103. Studies 3101 and 3102 are pivotal, phase III, randomised, double-blind, placebo-controlled, safety and efficacy studies to evaluate relugolix with E2 and NETA as a combination therapy for pain associated with endometriosis. Study 3103 is an open-label extension study including patients who completed one of the two pivotal studies and met the eligibility criteria, regardless of their treatment assignment in the pivotal studies. In the extension part all patients received relugolix combination therapy. As a consequence, sections 4.1, 4.2, 4.4, 4.8 and 5.1 of the SmPC were updated. The Package Leaflet is updated in accordance.
Update of section 4.5 of the SmPC to update information regarding Drug-Drug Interaction based on final results of DDI studies MVT-601-54, MVT-601-55 and MVT-601-57. Study MVT-601-54 is a 2-part interventional open-label study to assess the potential effects of erythromycin on the PK of the 3 components of Ryeqo. Study MVT-601-55 is an interventional open label fixed single sequence cross-over study to assess whether a 6-hour dose separation is sufficient to mitigate absorption mediated increased exposure to relugolix and study MVT-601-057 is a 2-part study to assess the potential effect of relugolix on the PK of total dabigatran.
The updated RMP version (2.0) has also been submitted. As part of the application, the MAH also requests an extension of the market protection by one additional year." Request for 1 year of market protection for a new indication (Article 14(11) of Regulation (EC) 726/2004)

Action: For adoption

Request for Supplementary Information adopted on 23.02.2023.

5.1.20. Soliris - Eculizumab - Orphan - EMEA/H/C/000791/II/0126

Alexion Europe SAS

Rapporteur: Carolina Prieto Fernandez, PRAC Rapporteur: Monica Martinez Redondo

Scope: "Extension of indication to include treatment of paediatric patients with refractory generalised myasthenia gravis (gMG) for Soliris, based on interim results from study ECU-MG-303; this is an open-label, multicenter, phase 3 study to evaluate the efficacy, safety, pharmacokinetics and pharmacodynamics of intravenous (IV) eculizumab in paediatric patients aged 6 to less than 18 years with acetylcholine receptor-antibody (AChR-Ab)

positive (+) refractory gMG. As a consequence, sections 4.1, 4.2, 4.4, 4.5, 4.8, 5.1 and 5.2 of the SmPC are updated. The Package Leaflet is updated in accordance. Version 20.0 of the RMP has also been submitted. In addition, the MAH took the opportunity to update section 4.8 of the SmPC in order to update the frequency of the list of adverse drug reactions (ADRs) based on cumulative safety data and to introduce minor editorial changes to the PI."

Action: For adoption

Request for Supplementary Information adopted on 23.02.2023.

5.1.21. Spikevax - COVID-19 mRNA vaccine (nucleoside-modified) - EMEA/H/C/005791/II/0104/G

Moderna Biotech Spain, S.L.

Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Marie Louise Schougaard Christiansen

Scope: "Grouped variation:
- C.I.6.a: Extension of indication to include the use of Spikevax bivalent Original/Omicron BA.4-5 (50 micrograms/50 micrograms)/mL dispersion for injection as a two-dose primary vaccination course in children 6 months through 5 years of age, based on data from study mRNA-1273-P306 (NCT05436834), an Open-Label, Phase 3 Study to Evaluate the Safety and Immunogenicity of the mRNA-1273.214 Vaccine for SARS-CoV-2 Variants of Concern in Participants Aged 6 Months to < 6 Years; as a consequence, sections 2, 4.1, 4.2 and 6.6 of the SmPC are updated. The Package Leaflet and Labelling are updated in accordance. Version 6.7 of the RMP has also been submitted.
- C.I.6.a: Extension of indication to include the use of Spikevax bivalent Original/Omicron BA.4-5 as a single-dose primary vaccination course against COVID-19 in individuals 6 years of age and older, irrespective of vaccination history, based on epidemiology and clinical data from study P306 part 1; as a consequence, sections 2, 4.1, 4.2 and 6.6 of the SmPC are updated. The Package Leaflet is updated in accordance.
In addition, the Marketing authorisation holder (MAH) took the opportunity to make minor editorial changes/corrections throughout the product information."

Action: For adoption

5.1.22. Trodelvy - sacituzumab govitecan - EMEA/H/C/005182/II/0020

Gilead Sciences Ireland UC

Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Menno van der Elst

Scope: "Extension of indication to include treatment of adult patients with unresectable or metastatic hormone receptor (HR)-positive, human epidermal growth factor receptor 2 (HER2)-negative breast cancer who have received endocrine-based therapy and at least two additional systemic therapies in the metastatic setting, based on final results from study IMMU-132-09 (TROPiCS-02); this is an open-label, randomized, multicenter phase 3 study of sacituzumab govitecan (IMMU-132) versus treatment of physician's choice (TPC) in subjects with hormonal receptor-positive (HR+) human epidermal growth factor receptor 2 (HER2) negative metastatic breast cancer (mBC) who have failed at least two prior chemotherapy regimens. As a consequence, sections 4.1, 4.2, 4.4, 4.5, 4.8, 5.1, 5.2 and 6.6 of the SmPC are updated. The Package Leaflet is updated in accordance. Version 2.1 of the RMP has also been submitted. In addition, the MAH took the opportunity to introduce

minor editorial changes to the PI and to update the list of local representatives in the Package Leaflet.", Request for 1 year of market protection for a new indication (Article 14(11) of Regulation (EC) 726/2004)

Action: For adoption

Request for Supplementary Information adopted on 30.03.2023.

5.1.23. Valdoxan - agomelatine - EMEA/H/C/000915/II/0051

Les Laboratoires Servier

Rapporteur: Eva Skovlund, PRAC Rapporteur: Pernille Harg

Scope: "Extension of indication to include new therapeutic indication in adolescents aged 12 to 17 years for the treatment of moderate to severe major depressive episodes, if depression is unresponsive to psychological therapy alone, for Valdoxan, further to the results of the phase 2 (CL2-20098-075) and phase 3 (CL3-20098-076) paediatric clinical studies included in the Paediatric Investigation Plan number EMEA-001181-PIP-11; As a consequence, the sections 4.1, 4.2, 4.4, 4.8, 5.1 and 5.2 of the SmPC are updated.
The Package Leaflet is updated accordingly. The updated RMP version 25.1 has also been submitted."

Action: For adoption

Request for Supplementary Information adopted on 26.01.2023.

5.1.24. Veyvondi - vonicog alfa - EMEA/H/C/004454/II/0030

Baxalta Innovations GmbH

Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Mari Thorn

Scope: "Extension of indication to include "prophylactic treatment to prevent or reduce the frequency of bleeding episodes" for Veyvondi based on final results from study 071301 and interim results from study SHP677-304. Study 071301 is a prospective, phase 3, open-label, international multicenter study on efficacy and safety of prophylaxis with rVWF in severe von Willebrand disease; while study SHP677-304 is a phase 3B, prospective, open-label, uncontrolled, multicenter study on long-term safety and efficacy of rVWF in paediatric and adult subjects with severe von Willebrand disease. As a consequence, sections 4.1, 4.2, 4.4, 5.1, 5.2, 6.2 and 6.6 of the SmPC are updated. The Package Leaflet is updated in accordance. Version 4.0 of the RMP has also been submitted."

Action: For adoption

5.1.25. Zinplava - bezlotoxumab - EMEA/H/C/004136/II/0037

Merck Sharp & Dohme B.V.

Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Adam Przybylkowski

Scope: "Extension of indication to include treatment of the paediatric population (1 to 18 years of age) for Zinplava, based on final results from study MK-6072-001 (MODIFY III) listed as a category 3 study in the RMP; this is a phase 3, randomised, placebo-controlled, parallel-group, multi-site, double-blind trial evaluating the safety, tolerability,

pharmacokinetics (PK) and efficacy of a single infusion of bezlotoxumab in paediatric participants from 1 to <18 years of age receiving antibacterial drug treatment for Clostridioides difficile infection (CDI). As a consequence, sections 4.1, 4.2, 4.8, 5.1 and 5.2 of the SmPC are updated. The Package Leaflet is updated in accordance. Version 2.3 of the RMP has also been submitted. In addition, the marketing authorisation holder (MAH) took the opportunity to introduce minor editorial changes to the PI."

Action: For adoption

5.2. Update on on-going Type II variation; variation of therapeutic indication procedure according to Commission Regulation (EC) No 1234/2008

5.2.1. Epidyolex - cannabidiol - Orphan - EMEA/H/C/004675/II/0020

GW Pharma (International) B.V.

Rapporteur: Thalia Marie Estrup Blicher, Co-Rapporteur: Tomas Radimersky, PRAC Rapporteur: Ana Sofia Diniz Martins

Scope: Final assessment report re-tabled

Action: For information

Opinion adopted on 26.04.2023. Request for Supplementary Information adopted on 30.03.2023, 15.12.2022, 15.09.2022.

5.3. Re-examination of Type II variation; variation of therapeutic indication procedure according to Commission Regulation (EC) No 1234/2008

No items

6. Medical devices

6.1. Ancillary medicinal substances - initial consultation

6.1.1. human albumin solution / gentamicin sulfate - EMEA/H/D/006141

human assisted reproductive techniques including in-vitro fertilisation procedures

Scope: Opinion

Action: For adoption

List of Outstanding Issues adopted on 26.04.2023. List of Questions adopted on 10.11.2022.

6.2. Ancillary medicinal substances – post-consultation update

No items

6.3. Companion diagnostics - initial consultation

6.3.1. in vitro diagnostic medical device - EMEA/H/D/006232

to detect rearrangements involving the ALK gene via fluorescence

Scope: List of questions

Action: For adoption

6.4. Companion diagnostics – follow-up consultation

No items

7. Procedure under Article 83(1) of Regulation (EC) 726/2004 (Compassionate Use)

7.1. Procedure under Article 83(1) of Regulation (EC) 726/2004 (Compassionate Use)

No items

8. Pre-submission issues

8.1. Pre-submission issue

8.1.1. avacincaptad pegol – H0006153

Treatment of adults with geographic atrophy (GA) secondary to age-related macular degeneration (AMD)

Scope: Briefing note and the Rapporteurs' recommendation on the request for accelerated assessment.

Action: For adoption

8.1.2. mRNA-1345 - H0006278

Active immunisation for the prevention of respiratory syncytial virus (RSV)-associated lower respiratory tract disease (LRTD) in individuals 60 years of age and older

Scope: Briefing note and the Rapporteurs' recommendation on the request for accelerated assessment.

Action: For adoption

8.1.3. Troriluzole hydrochloride - Orphan - H0006068

Biohaven Bioscience Ireland Limited; Treatment of hereditary spinocerebellar ataxia

Scope: Briefing note and the Rapporteurs' recommendation on the request for accelerated assessment.

Action: For adoption

8.2. Priority Medicines (PRIME)

Information related to priority medicines cannot be released at present time as these contain commercially confidential information

8.2.1. List of applications received

Action: For information

8.2.2. Recommendation for PRIME eligibility

Action: For adoption

9. Post-authorisation issues

9.1. Post-authorisation issues

9.1.1. Mysimba - naltrexone hydrochloride / bupropion hydrochloride - EMEA/H/C/003687/II/0056

Orexigen Therapeutics Ireland Limited

Scope: Re-examination; List of questions to the SAG

Action: For adoption

Opinion adopted on 30.03.2023. Request for Supplementary Information adopted on 26.01.2023, 15.09.2022, 24.03.2022.

9.1.2. Blenrep - belantamab mafodotin - EMEA/H/C/004935/R/0017, Orphan

GlaxoSmithKline (Ireland) Limited

Rapporteur: Johanna Lähteenvuo, Co-Rapporteur: Carolina Prieto Fernandez, PRAC Rapporteur: Ulla Wändel Liminga

Scope: Renewal of conditional marketing authorisation

Action: For adoption

Request for Supplementary Information adopted on 26.04.2023.

9.1.3. Brintellix - vortioxetine - EMEA/H/C/002717/II/0038

H. Lundbeck A/S

Rapporteur: Karin Janssen van Doorn

Scope: "Update of sections 4.2, 4.4, 4.8, 5.1 and 5.2 of the SmPC in order to include clinically relevant information on the efficacy, safety, tolerability, and PK of vortioxetine in the paediatric population based on final results from studies 12709A, 12712A and 12712B. Study 12709A is an interventional, randomized, double-blind, placebo-controlled, active-reference (fluoxetine), fixed-dose study of vortioxetine in paediatric patients aged 7 to 11 years, with Major Depressive Disorder (MDD) to evaluate efficacy and safety. Whereas studies 12712A and 12712B are 2 open-label, long-term safety and efficacy studies in children and adolescents: one 6-month extension study (study 12712A) to studies 12709A and 12710A, and one 18-month extension study (study 12712B) to study 12712A. The Package Leaflet is updated accordingly."

Action: For adoption

Request for Supplementary Information adopted on 30.03.2023, 15.12.2022.

10. Referral procedures

10.1. Procedure for Centrally Authorised products under Article 20 of Regulation (EC) No 726/2004

No items

10.2. Requests for CHMP Opinion under Article 5(3) of Regulation (EC) No 726/2004

10.2.1. Colistimethate sodium (CMS) – EMEA/H/A-5(3)/1524

Various MAHs

Referral Rapporteur: Martina Weise, Referral Co-Rapporteur: Ewa Balkowiec Iskra

Review of the ratio of polymyxins E1 and E2 in colistin starting material and of the (sulfomethylation) composition profile of CMS finished product.

Scope: List of questions

Action: For adoption

10.3. Procedure under Articles 5(2) and 10 of Regulation (EC) No 726/2004

No items

10.4. Disagreement between Member States on application for medicinal product (potential serious risk to public health) –under Article 29(4) of Directive 2001/83/EC

No items

10.5. Harmonisation - Referral procedure under Article 30 of Directive 2001/83/EC

10.5.1. Havrix and associated names - Inactivated hepatitis A virus - EMEA/H/A-30/1527

MAH: GlaxoSmithKline Biologicals

Referral Rapporteur: TBC, Referral Co-Rapporteur: TBC

Harmonisation exercise for Havrix and associated names. Product Information harmonisation was triggered by the MAH.

Scope: Appointment of referral rapporteurs

Action: For adoption

10.6. Community Interests - Referral under Article 31 of Directive 2001/83/EC

No items

10.7. Re-examination Procedure under Article 32(4) of Directive 2001/83/EC

No items

10.8. Procedure under Article 107(2) of Directive 2001/83/EC

No items

10.9. Disagreement between Member States on Type II variation– Arbitration procedure initiated by MAH under Article 6(13) of Commission Regulation (EC) No 1084/2003

No items

10.10. Procedure under Article 29 of Regulation (EC) 1901/2006

No items

10.11. Referral under Article 13 Disagreement between Member States on Type II variation– Arbitration procedure initiated by Member State under Article 13 (EC) of Commission Regulation No 1234/2008

No items

11. Pharmacovigilance issue

11.1. Early Notification System

June 2023 Early Notification System on envisaged CHMP/CMDh outcome accompanied by communication to the general public.

Action: For information

12. Inspections

12.1. GMP inspections

Information related to GMP inspections will not be published as it undermines the purpose of such inspections

12.2. GCP inspections

Information related to GCP inspections will not be published as it undermines the purpose of such inspections

12.3. Pharmacovigilance inspections

Information related to Pharmacovigilance inspections will not be published as it undermines the purpose of such inspections

12.4. GLP inspections

Information related to GLP inspections will not be published as it undermines the purpose of such inspections

13. Innovation Task Force

13.1. Minutes of Innovation Task Force

No items

13.2. Innovation Task Force briefing meetings

No items

13.3. Requests for CHMP Opinion under Article 57(1)J and (1)P of Regulation (EC) No 726/2004

No items

13.4. Nanomedicines activities

No items

14. Organisational, regulatory and methodological matters

14.1. Mandate and organisation of the CHMP

14.1.1. Timetable for August 2023 Written Procedure

Action: For adoption

14.1.2. CHMP learnings

Collection, discussion and recording of CHMP learnings.

CHMP: Outi Mäki-Ikola

Action: For discussion

14.1.3. CHMP meetings

Alternating face-to-face and virtual meetings.

Action: For discussion

14.2. Coordination with EMA Scientific Committees

14.2.1. Pharmacovigilance Risk Assessment Committee (PRAC)

List of Union Reference Dates and frequency of submission of Periodic Safety Update Reports (EURD list) for June 2023

Action: For adoption

14.2.2. Paediatric Committee (PDCO)

Agenda of the June 2023 PDCO plenary meeting

Action: For information

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

14.3.1. Biologics Working Party (BWP)

Chair: Sean Barry/Francesca Luciani

Reports from BWP June 2023 meeting to CHMP for adoption:

- 13 reports on products in scientific advice and protocol assistance
- 6 reports on products in pre-authorisation procedures
- 2 reports on products in post-authorisation procedures
- 5 reports on products in plasma master file

Action: For adoption

14.3.2. Nomination of Biologics Working Party (BWP) members

Following the call for nominations launched in April, the BWP selection panel has recommended new BWP members.

Action: For endorsement

14.3.3. Call for nominations for BWP Chair

Launch call for nomination for a chair for the Biologics Working Party (BWP).

Members of BWP are invited to indicate their interest in standing for this position of chair for BWP.

Applications are to be submitted before the start of the CHMP meeting on 17 July 2023.

Action: For information

14.3.4. Nomination of Quality Working Party (QWP) members

Following the call for nominations launched in April, the QWP selection panel has recommended new QWP members.

Action: For endorsement

14.3.5. Call for nominations for QWP Chair and Vice-chair

Launch call for nomination for chair and vice-chair for Quality Working Party (QWP).

Members of QWP are invited to indicate their interest in standing for this position of chair and vice-chair for QWP.

Applications are to be submitted before the start of the CHMP meeting on 17 July 2023.

Action: For information

14.3.6. Nomination of Biosimilar Medicinal Products Working Party (BMWP) members

Following the call for nominations launched in April, the BMWP selection panel has recommended new BMWP members.

Action: For endorsement

14.3.7. Call for nominations for BMWP Chair

Launch call for nomination for chairs for Biosimilar Medicinal Products Working Party (BMWP).

Members of BMWP are invited to indicate their interest in standing for this position of chair for BMWP.

Applications are to be submitted before the start of the CHMP meeting on 17 July 2023.

Action: For information

14.3.8. Scientific Advice Working Party (SAWP)

Chair: Paolo Foggi

Report from the SAWP meeting held on 05-08 June 2023. Table of conclusions

Action: For information

Scientific advice letters:

Information related to scientific advice letters cannot be released at present time as these contain commercially confidential information.

14.3.9. Guideline on clinical investigation of medicinal products in the treatment or prevention of diabetes mellitus Rev. 2 – Cardiovascular Working Party (CVSWP)

Final Guideline

Action: For adoption

14.3.10. CHMP Question to Healthcare Professionals Working Party (HCPWP)

CHMP questions to HCPWP on SmPC guidance section 5.1.

Action: For adoption

Following-up on the discussions during the Swedish Strategic Review and Learning Meeting (SRLM) on the guidance document on the information to be included in section 5.1 "Pharmacodynamic properties" of the SmPC, which also includes description of clinical trials, and given that HCP are the primary users of the SmPC, the CHMP has drafted a LoQ to consult the HCPWP on a few specific aspects.

14.4. Cooperation within the EU regulatory network

No items

14.5. Cooperation with International Regulators

No items

14.6. Contacts of the CHMP with external parties and interaction with the Interested Parties to the Committee

No items

14.7. CHMP work plan

No items

14.8. Planning and reporting

No items

14.9. Others

No items

15. Any other business

15.1. AOB topic

15.1.1. Update on COVID-19

Action: For information

15.1.2. Control options for nitrosamines

Action: For information

15.1.3. Gap analysis assessment in support of member state shortage management activities

Action: For adoption

15.1.4. Product Information for upcoming Comirnaty and Spikevax variant vaccine

Product Information for upcoming Comirnaty and Spikevax variant vaccine for preliminary

discussion

Action: For discussion

15.1.5. Preparation of oncology product-related discussions

Action: For discussion

15.1.6. Update on EMA collaborative systems

Update on the use of EMA collaborative systems – IRIS Network Portal, IRIS SharePoint and Teams.

Action: For discussion

Explanatory notes

The notes below give a brief explanation of the main sections and headings in the CHMP agenda and should be read in conjunction with the agenda or the minutes.

Oral explanations *(section 2)*

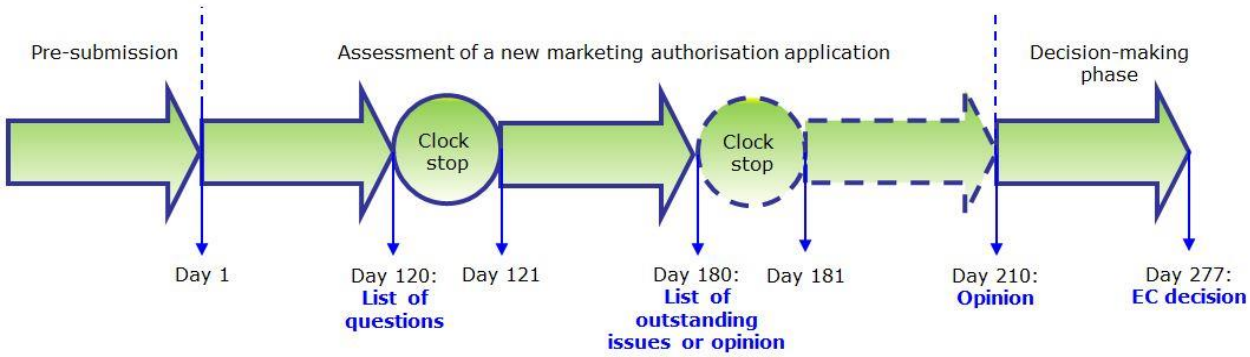
The items listed in this section are those for which marketing authorisation holders (MAHs) or applicants have been invited to the CHMP plenary meeting to address questions raised by the Committee. Oral explanations normally relate to on-going applications (section 3, 4 and 5) or referral procedures (section 10) but can relate to any other issue for which the CHMP would like to discuss with company representatives in person.

Initial applications *(section 3)*

This section lists applications for marketing authorisations of new medicines that are to be discussed by the Committee.

Section 3.1 is for medicinal products nearing the end of the evaluation and for which the CHMP is expected to adopt an **opinion** at this meeting on whether marketing authorisation should be granted. Once adopted, the CHMP opinion will be forwarded to the European Commission for a final legally binding decision valid throughout the EU.

The other items in the section are listed depending on the stage of the evaluation, which is shown graphically below:



The assessment of an application for a new medicine takes up to 210 'active' days. This active evaluation time is interrupted by at least one 'clock-stop' during which time the applicant prepares the answers to questions from the CHMP. The clock stop happens after day 120 and may also happen after day 180, when the CHMP has adopted a list of questions or outstanding issues to be addressed by the company. Related discussions are listed in the agenda under sections 3.2 (**Day 180 List of outstanding issues**) and 3.3 (**Day 120 list of questions**).

CHMP discussions may also occur at any other stage of the evaluation, and these are listed under section 3.4, **update on ongoing new applications for centralised procedures**.

The assessment leads to an opinion from the CHMP by day 210. Following a CHMP opinion the European Commission takes usually 67 days to issue a legally binding decision (i.e. by day 277 of the procedure). CHMP discussions on products that have received a CHMP opinion and are awaiting a decision are listed under section 3.6, **products in the decision making phase**.

Extension of marketing authorisations according to Annex I of Reg. 1234/2008 *(section 4)*

Extensions of marketing authorisations are applications for the change or addition of new strengths, formulations or routes of administration to existing marketing authorisations. Extension applications follow a 210-day evaluation process, similarly to applications for new medicines (see figure above).

Type II variations - Extension of indication procedures *(section 5)*

Type II variations are applications for a change to the marketing authorisation which requires an update of the product information and which is not covered in section 4. Type II variations include applications for a new use of the medicine (extension of indication), for which the assessment takes up to 90 days. For the applications listed in this section, the CHMP may adopt an opinion or request supplementary information from the applicant.

Ancillary medicinal substances in medical devices *(section 6)*

Although the EMA does not regulate medical devices it can be asked by the relevant authorities (the so-called Notified Bodies) that are responsible for regulating these devices to give a scientific opinion on a medicinal substance contained in a medical device.

Re-examination procedures (new applications) under article 9(2) of regulation no 726/2004 *(section 3.5)*

This section lists applications for new marketing authorisation for which the applicant has requested a re-examination of the opinion previously issued by the CHMP.

Re-examination procedures *(section5.3)*

This section lists applications for type II variations (including extension of indication applications) for which the applicant has requested re-examination of the opinion previously issued by the CHMP.

Withdrawal of application *(section 3.7)*

Applicants may decide to withdraw applications at any stage during the assessment and a CHMP opinion will therefore not be issued. Withdrawals are included in the agenda for information or discussion, as necessary.

Procedure under article 83(1) of regulation (EC) 726/2004 (compassionate use) *(section 7)*

Compassionate use is a way of making available to patients with an unmet medical need a promising medicine which has not yet been authorised (licensed) for their condition. Upon request, the CHMP provides recommendations to all EU Member States on how to administer, distribute and use certain medicines for compassionate use.

Pre-submission issues *(section 8)*

In some cases the CHMP may discuss a medicine before a formal application for marketing authorisation is submitted. These cases generally refer to requests for an accelerated assessment for medicines that are of major interest for public health or can be considered a therapeutic innovation. In case of an accelerated assessment the assessment timetable is reduced from 210 to 150 days.

Post-authorisation issues *(section 9)*

This section lists other issues concerning authorised medicines that are not covered elsewhere in the agenda. Issues include supply shortages, quality defects, some annual reassessments or renewals or type II variations to marketing authorisations that would require specific discussion at the plenary.

Referral procedures *(section 10)*

This section lists referrals that are ongoing or due to be started at the plenary meeting. A referral is a procedure used to resolve issues such as concerns over the safety or benefit-risk balance of a medicine or a class of medicines. In a referral, the EMA is requested to conduct a scientific assessment of a particular medicine or class of medicines on behalf of the EU. Further information on such procedures can be found here.

Pharmacovigilance issues *(section 11)*

This section lists issues that have been discussed at the previous meeting of the PRAC, the EMA's committee responsible for evaluating and monitoring safety issues for medicines. Feedback is provided by the PRAC. This section also refers to the early notification system, a system used to notify the European regulatory network on proposed EMA communication on safety of medicines.

Inspections Issues *(section 12)*

This section lists inspections that are undertaken for some medicinal products. Inspections are carried out by regulatory agencies to ensure that marketing authorisation holders comply with their obligations. Inspection can relate to good manufacturing practice (GMP), good clinical practice (GCP), good laboratory practice (GLP) or good pharmacovigilance practice (GVP).

Innovation task force *(section 13)*

The Innovation Task Force (ITF) is a body set up to encourage early dialogue with applicants developing innovative medicines. Minutes from the last ITF meeting as well as any related issue that requires discussion with the CHMP are listed in this section of the agenda. Further information on the ITF can be found here.

Scientific advice working party (SAWP) *(section 14.3.1)*

This section refers to the monthly report from the CHMP's Scientific Advice Working Party (SAWP) on scientific advice given to companies during the development of medicines. Further general information on SAWP can be found here.

Satellite groups / other committees *(section 14.2)*

This section refers to the reports from groups and committees making decisions relating to human medicines: the Coordination Group for Mutual Recognition and Decentralised Procedures – Human (CMDh), the Committee for Orphan Medicinal Products (COMP), the Committee for Herbal Medicinal Products (HMPC), Paediatric Committee (PDCO), the Committee for Advanced Therapies (CAT) and the Pharmamacovigilance Risk Assessment Committee (PRAC).

Invented name issues *(section 14.3)*

This section list issues related to invented names proposed by applicants for new medicines. The CHMP has established the Name Review Group (NRG) to perform reviews of the invented names. The group's main role is to consider whether the proposed names could create a public-health concern or potential safety risk. Further information can be found here.

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

EUROPEAN MEDICINES AGENCY
SCIENCE MEDICINES HEALTH

19 June 2023
EMA/CHMP/243981/2023

Annex to 19-22 June 2023 CHMP Agenda

Pre-submission and post-authorisations issues

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A. PRE-SUBMISSION ISSUES

A.1. ELIGIBILITY REQUESTS

Report on Eligibility to Centralised Procedure for June 2023: **For adoption**

A.2. Appointment of Rapporteur / Co-Rapporteur Full Applications

Final Outcome of Rapporteurship allocation for June 2023: **For adoption**

A.3. PRE-SUBMISSION ISSUES FOR INFORMATION

Information related to pre-submission of initial applications cannot be released at the present time as these contain commercially confidential information.

B. POST-AUTHORISATION PROCEDURES OUTCOMES

B.1. Annual re-assessment outcomes

B.1.1. Annual reassessment for products authorised under exceptional circumstances

Evoltra - clofarabine - EMEA/H/C/000613/S/0078
Sanofi B.V., Rapporteur: Alexandre Moreau, PRAC Rapporteur: Tiphaine Vaillant

Lamzede - velmanase alfa - EMEA/H/C/003922/S/0031, Orphan
Chiesi Farmaceutici S.p.A., Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Jan Neuhauser

SCENESSE - afamelanotide - EMEA/H/C/002548/S/0045, Orphan
Clinuvel Europe Limited, Rapporteur: Janet Koenig, PRAC Rapporteur: Martin Huber
Request for Supplementary Information adopted on 26.04.2023.

B.2. RENEWALS OF MARKETING AUTHORISATIONS OUTCOMES

B.2.1. Renewals of Marketing Authorisations requiring 2nd Renewal

B.2.2. Renewals of Marketing Authorisations for unlimited validity

Dengvaxia - dengue tetravalent vaccine (live, attenuated) - EMEA/H/C/004171/R/0027
Sanofi Pasteur, Rapporteur: Christophe Focke, Co-Rapporteur: Sol Ruiz, PRAC Rapporteur: Sonja Hrabcik

Fortacin - lidocaine / prilocaine - EMEA/H/C/002693/R/0038
Recordati Ireland Ltd, Rapporteur: Maria Concepcion Prieto Yerro, Co-Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Maria del Pilar Rayon

Lenalidomide Accord - lenalidomide - EMEA/H/C/004857/R/0021
Accord Healthcare S.L.U., Generic, Generic of Revlimid, Rapporteur: Ewa Balkowiec Iskra, PRAC Rapporteur: Tiphaine Vaillant
Request for Supplementary Information adopted on 26.04.2023.

Namuscla - mexiletine - EMEA/H/C/004584/R/0014, Orphan
Lupin Europe GmbH, Rapporteur: Bruno Sepodes, Co-Rapporteur: Kristina Dunder, PRAC Rapporteur: Eva Jirsová

Ogivri - trastuzumab - EMEA/H/C/004916/R/0054
Viatris Limited, Rapporteur: Karin Janssen van Doorn, Co-Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Gabriele Maurer

Pelmeg - pegfilgrastim - EMEA/H/C/004700/R/0025
Mundipharma Corporation (Ireland) Limited, Rapporteur: Karin Janssen van Doorn, Co-Rapporteur: Christian Gartner, PRAC Rapporteur: Menno van der Elst

POTELIGEO - mogamulizumab - EMEA/H/C/004232/R/0021, Orphan
Kyowa Kirin Holdings B.V., Rapporteur: Johann Lodewijk Hillege, Co-Rapporteur: Armando Genazzani, PRAC Rapporteur: Marie Louise

Schougaard Christiansen	
Symkevi - tezacaftor / ivacaftor - EMEA/H/C/004682/R/0038, Orphan Vertex Pharmaceuticals (Ireland) Limited, Rapporteur: Johann Lodewijk Hillege, Co-Rapporteur: Beata Maria Jakline Ullrich, PRAC Rapporteur: Rhea Fitzgerald Request for Supplementary Information adopted on 30.03.2023.	
TAKHZYRO - lanadelumab - EMEA/H/C/004806/R/0035, Orphan Takeda Pharmaceuticals International AG Ireland Branch, Rapporteur: Kristina Dunder, Co-Rapporteur: Jean-Michel Race, PRAC Rapporteur: Kirsti Villikka Request for Supplementary Information adopted on 25.05.2023.	
Venclyxto - venetoclax - EMEA/H/C/004106/R/0046 AbbVie Deutschland GmbH & Co. KG, Rapporteur: Filip Josephson, Co-Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Eva Jirsová Request for Supplementary Information adopted on 26.04.2023.	

B.2.3. Renewals of Conditional Marketing Authorisations

Adakveo - crizanlizumab - EMEA/H/C/004874/R/0014, Orphan Novartis Europharm Limited, Rapporteur: Daniela Philadelphy, PRAC Rapporteur: Jo Robays	
Blenrep - belantamab mafodotin - EMEA/H/C/004935/R/0017, Orphan GlaxoSmithKline (Ireland) Limited, Rapporteur: Johanna Lähteenvuo, Co-Rapporteur: Carolina Prieto Fernandez, PRAC Rapporteur: Ulla Wändel Liminga Request for Supplementary Information adopted on 26.04.2023.	See 9.1

B.3. POST-AUTHORISATION PHARMACOVIGILANCE OUTCOMES

Signal detection

PRAC recommendations on signals adopted at the PRAC meeting held on 05-08 June 2023
PRAC:

Signal of cytokine release syndrome

Opdivo (CAP) – nivolumab

(Please note this is part of the signal of Ipilimumab; nivolumab; pembrolizumab – Capillary leak syndrome (Opdivo, Yervoy, Keytruda) and cytokine release syndrome (Opdivo) – EPITT 19880)

Rapporteur: Carolina Prieto Fernandez, Co-Rapporteur: Johann Lodewijk Hillege, PRAC
Rapporteur: Martin Huber
PRAC recommendation on a variation
Action: For adoption

Signal of acnes

Xeljanz (CAP) – tofacitinib

Rapporteur: Armando Genazzani, Co-Rapporteur: Johann Lodewijk Hillege, PRAC
Rapporteur: Liana Gross-Martirosyan
PRAC recommendation on a variation
Action: For adoption

PSUR procedures for which PRAC adopted a recommendation for variation of the terms of the MA at its June 2023 meeting:

EMEA/H/C/PSUSA/00001172/202211
(doxorubicin)
CAPS:
Caelyx pegylated liposomal (EMEA/H/C/000089) (doxorubicin), Baxter Holding B.V., Rapporteur: Petr Vrbata
Celdoxome pegylated liposomal (EMEA/H/C/005330) (doxorubicin), YES Pharmaceutical Development Services GmbH, Rapporteur: Fátima Ventura
Myocet liposomal (EMEA/H/C/000297) (doxorubicin hydrochloride), Teva B.V., Rapporteur: Filip Josephson
Zolsketil pegylated liposomal (EMEA/H/C/005320) (doxorubicin), Accord Healthcare S.L.U., Rapporteur: Carolina Prieto Fernandez
NAPS:
NAPs - EU
PRAC Rapporteur: Eva Jirsová, "12/11/2021 To: 12/11/2022"

EMEA/H/C/PSUSA/00002652/202211
(rituximab)
CAPS:
Blitzima (EMEA/H/C/004723) (rituximab), Celltrion Healthcare Hungary Kft., Rapporteur: Sol Ruiz
MabThera (EMEA/H/C/000165) (rituximab), Roche Registration GmbH, Rapporteur: Aaron Sosa Mejia
Rixathon (EMEA/H/C/003903) (rituximab), Sandoz GmbH, Rapporteur: Jan Mueller-Berghaus
Riximyo (EMEA/H/C/004729) (rituximab), Sandoz GmbH, Rapporteur: Jan Mueller-Berghaus
Ruxience (EMEA/H/C/004696) (rituximab), Pfizer Europe MA EEIG, Rapporteur: Johann Lodewijk Hillege
Truxima (EMEA/H/C/004112) (rituximab), Celltrion Healthcare Hungary Kft., Rapporteur: Sol Ruiz, PRAC Rapporteur: Anette Kirstine Stark, "17/11/2021 To: 17/11/2022"

EMEA/H/C/PSUSA/00002841/202210
(tadalafil)
CAPS:
Adcirca (EMEA/H/C/001021) (tadalafil), Eli Lilly Nederland B.V., Rapporteur: Maria Concepcion Prieto Yerro
Cialis (EMEA/H/C/000436) (tadalafil), Eli Lilly Nederland B.V., Rapporteur: Maria Concepcion Prieto Yerro
Tadalafil Lilly (EMEA/H/C/004666) (tadalafil), Eli Lilly Nederland B.V., Rapporteur: Maria Concepcion Prieto Yerro
NAPS:
NAPs - EU
PRAC Rapporteur: Maria del Pilar Rayon, "14/10/2019 To: 14/10/2022"

EMEA/H/C/PSUSA/00009176/202211
(hydrocortisone (for centrally authorised products for adrenal insufficiency, congenital adrenal hyperplasia, modified-release formulations))
CAPS:
Efmody (EMEA/H/C/005105) (hydrocortisone), Diurnal Europe BV, Rapporteur: Johann Lodewijk Hillege
Plenadren (EMEA/H/C/002185) (hydrocortisone), Takeda Pharmaceuticals International AG Ireland Branch, Rapporteur: Kristina Dunder, PRAC Rapporteur: Mari Thorn, "02/11/2019 To: 02/11/2022"

EMEA/H/C/PSUSA/00010301/202211
(ibrutinib)
CAPS:
Imbruvica (EMEA/H/C/003791) (ibrutinib), Janssen-Cilag International N.V., Rapporteur: Filip Josephson, PRAC Rapporteur: Nikica Mirošević Skvrce, "13/11/2021 To: 12/11/2022"

EMEA/H/C/PSUSA/00010318/202210
(nintedanib (oncology indications))
CAPS:
Vargatef (EMEA/H/C/002569) (nintedanib), Boehringer Ingelheim International GmbH, Rapporteur: Aaron Sosa Mejia, PRAC Rapporteur: Georgia Gkegka, "15/10/2019 To: 15/10/2022"

EMEA/H/C/PSUSA/00010887/202210 (acalabrutinib) CAPS: **Calquence** (EMEA/H/C/005299) (acalabrutinib), AstraZeneca AB, Rapporteur: Filip Josephson, PRAC Rapporteur: Željana Margan Koletić, "31/10/2021 To: 30/10/2022"	
EMEA/H/C/PSUSA/00010917/202211 (selpercatinib) CAPS: **Retsevmo** (EMEA/H/C/005375) (selpercatinib), Eli Lilly Nederland B.V., Rapporteur: Alexandre Moreau, PRAC Rapporteur: Menno van der Elst, "08/05/2022 To: 08/11/2022"	
EMEA/H/C/PSUSA/00010992/202211 (tixagevimab / cilgavimab (Evusheld)) CAPS: **EVUSHELD** (EMEA/H/C/005788) (tixagevimab / cilgavimab), AstraZeneca AB, Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Kimmo Jaakkola, "14/05/2022 To: 13/11/2022"	
EMEA/H/C/PSUSA/00011019/202211 (tirzepatide) CAPS: **Mounjaro** (EMEA/H/C/005620) (tirzepatide), Eli Lilly Nederland B.V., Rapporteur: Martina Weise, PRAC Rapporteur: Menno van der Elst, "13/05/2022 To: 13/11/2022"	

B.4. EPARs / WPARs

in vitro diagnostic medical device - EMEA/H/D/006255 is indicated as an aid in the selection of adult hemophilia A patients for whom valoctocogene roxaparvovec treatment is being considered, Companion Diagnostics (Article 48 (3), (4), (7), (8) of Regulation (EU) 2017/746)	For information only. Comments can be sent to the PL in case necessary.
Briumvi - ublituximab - EMEA/H/C/005914 Propharma Group The Netherlands B.V., treatment of relapsing forms of multiple sclerosis (RMS), New active substance (Article 8(3) of Directive No 2001/83/EC)	For information only. Comments can be sent to the PL in case necessary.
Pylclari - piflufolastat (18f) - EMEA/H/C/005520 Curium Pet France, imaging in patients	For information only. Comments can be sent to the PL in case necessary.

undergoing oncologic diagnostic procedures when increased expression of prostate specific membrane antigen is a diagnostic target, New active substance (Article 8(3) of Directive No 2001/83/EC)	
SOHONOS - palovarotene - EMEA/H/C/004867, Orphan Ipsen Pharma, treatment of fibrodysplasia ossificans progressiva, New active substance (Article 8(3) of Directive No 2001/83/EC)	For information only. Comments can be sent to the PL in case necessary.
Sugammadex Piramal - sugammadex - EMEA/H/C/006083 Piramal Critical Care B.V., Reversal of neuromuscular blockade induced by rocuronium or vecuronium in adults, Generic, Generic of Bridion, Generic application (Article 10(1) of Directive No 2001/83/EC)	For information only. Comments can be sent to the PL in case necessary.
Ztalmy - ganaxolone - EMEA/H/C/005825, Orphan Marinus Pharmaceuticals Emerald Limited, treatment of epileptic seizures associated with cyclindependent kinase-like 5 deficiency disorder (CDD), New active substance (Article 8(3) of Directive No 2001/83/EC)	For information only. Comments can be sent to the PL in case necessary.

B.5. TYPE II VARIATION, WORKSHARING PROCEDURE OUTCOMES

Scopes related to Chemistry, Manufacturing, and Controls cannot be released at the present time as these contain commercially confidential information.

B.5.1. CHMP assessed procedures scope: Pharmaceutical aspects

Advate - octocog alfa - EMEA/H/C/000520/II/0120/G Takeda Manufacturing Austria AG, Rapporteur: Jan Mueller-Berghaus	
Aflunov - prepandemic influenza vaccine (H5N1) (surface antigen, inactivated, adjuvanted) - EMEA/H/C/002094/II/0084/G Seqirus S.r.l, Rapporteur: Maria Grazia Evandri	
Apexxnar - pneumococcal polysaccharide conjugate vaccine (20-valent, adsorbed) - EMEA/H/C/005451/II/0014/G Pfizer Europe MA EEIG, Rapporteur: Daniela Philadelphy Opinion adopted on 01.06.2023.	Positive Opinion adopted by consensus on 01.06.2023.

Azacitidine betapharm - azacitidine - EMEA/H/C/005075/II/0015 betapharm Arzneimittel GmbH, Generic, Generic of Vidaza, Rapporteur: Petr Vrbata	
Azacitidine betapharm - azacitidine - EMEA/H/C/005075/II/0016 betapharm Arzneimittel GmbH, Generic, Generic of Vidaza, Rapporteur: Petr Vrbata	
Brintellix - vortioxetine - EMEA/H/C/002717/II/0033 H. Lundbeck A/S, Rapporteur: Karin Janssen van Doorn Request for Supplementary Information adopted on 01.12.2022, 02.06.2022.	
Cetrotide - cetrorelix - EMEA/H/C/000233/II/0090 Merck Europe B.V., Rapporteur: Martina Weise	
COMIRNATY - COVID-19 mRNA vaccine (nucleoside-modified) - EMEA/H/C/005735/II/0178 BioNTech Manufacturing GmbH, Rapporteur: Filip Josephson Opinion adopted on 01.06.2023.	Positive Opinion adopted by consensus on 01.06.2023.
COMIRNATY - COVID-19 mRNA vaccine (nucleoside-modified) - EMEA/H/C/005735/II/0181 BioNTech Manufacturing GmbH, Rapporteur: Filip Josephson	
COMIRNATY - COVID-19 mRNA vaccine (nucleoside-modified) - EMEA/H/C/005735/II/0182/G BioNTech Manufacturing GmbH, Rapporteur: Filip Josephson	
ECALTA - anidulafungin - EMEA/H/C/000788/II/0052/G Pfizer Europe MA EEIG, Rapporteur: Johann Lodewijk Hillege Opinion adopted on 08.06.2023. Request for Supplementary Information adopted on 16.02.2023.	Positive Opinion adopted by consensus on 08.06.2023.
Epidyolex - cannabidiol - EMEA/H/C/004675/II/0025/G, Orphan Jazz Pharmaceuticals Ireland Limited, Rapporteur: Thalia Marie Estrup Blicher Opinion adopted on 01.06.2023. Request for Supplementary Information adopted	Positive Opinion adopted by consensus on 01.06.2023.

on 23.03.2023.	
Eptifibatide Accord - eptifibatide - EMEA/H/C/004104/II/0016/G Accord Healthcare S.L.U., Generic, Generic of Integrilin, Rapporteur: Jayne Crowe Opinion adopted on 08.06.2023. Request for Supplementary Information adopted on 04.05.2023.	Positive Opinion adopted by consensus on 08.06.2023.
Eylea - aflibercept - EMEA/H/C/002392/II/0086 Bayer AG, Rapporteur: Alexandre Moreau Opinion adopted on 01.06.2023.	Positive Opinion adopted by consensus on 01.06.2023.
Gazyvaro - obinutuzumab - EMEA/H/C/002799/II/0053/G, Orphan Roche Registration GmbH, Rapporteur: Aaron Sosa Mejia	
Idacio - adalimumab - EMEA/H/C/004475/II/0018/G Fresenius Kabi Deutschland GmbH, Rapporteur: Johann Lodewijk Hillege Opinion adopted on 15.06.2023. Request for Supplementary Information adopted on 23.03.2023.	Positive Opinion adopted by consensus on 15.06.2023.
LIVMARLI - maralixibat - EMEA/H/C/005857/II/0002, Orphan Mirum Pharmaceuticals International B.V., Rapporteur: Martina Weise	
Lonsurf - trifluridine / tipiracil - EMEA/H/C/003897/II/0025 Les Laboratoires Servier, Rapporteur: Johann Lodewijk Hillege Request for Supplementary Information adopted on 19.01.2023.	
Ontruzant - trastuzumab - EMEA/H/C/004323/II/0046/G Samsung Bioepis NL B.V., Rapporteur: Karin Janssen van Doorn	
Pluvicto - lutetium (177Lu) vipivotide tetraxetan - EMEA/H/C/005483/II/0003 Novartis Europharm Limited, Rapporteur: Janet Koenig Request for Supplementary Information adopted on 01.06.2023.	Request for supplementary information adopted with a specific timetable.
POTELIGEO - mogamulizumab - EMEA/H/C/004232/II/0020, Orphan Kyowa Kirin Holdings B.V., Rapporteur: Johann	

Lodewijk Hillege Request for Supplementary Information adopted on 14.04.2023.	
Roclanda - latanoprost / netarsudil - EMEA/H/C/005107/II/0011 Santen Oy, Rapporteur: Jayne Crowe Opinion adopted on 01.06.2023. Request for Supplementary Information adopted on 09.02.2023.	Positive Opinion adopted by consensus on 01.06.2023.
Spikevax - COVID-19 mRNA vaccine (nucleoside-modified) - EMEA/H/C/005791/II/0100/G Moderna Biotech Spain, S.L., Rapporteur: Jan Mueller-Berghaus Request for Supplementary Information adopted on 15.06.2023.	Request for supplementary information adopted with a specific timetable.
Supemtek - influenza quadrivalent vaccine (rDNA) - EMEA/H/C/005159/II/0011/G Sanofi Pasteur, Rapporteur: Jan Mueller-Berghaus Request for Supplementary Information adopted on 08.06.2023, 09.02.2023.	Request for supplementary information adopted with a specific timetable.
Surgiflo Haemostatic Matrix Kit - human thrombin - EMEA/H/D/002301/II/0034/G Ferrosan Medical Devices A/S, Rapporteur: Jan Mueller-Berghaus	
TOBI Podhaler - tobramycin - EMEA/H/C/002155/II/0057/G, Orphan Viatris Healthcare Limited, Rapporteur: Johann Lodewijk Hillege	
Toujeo - insulin glargine - EMEA/H/C/000309/II/0121/G Sanofi-Aventis Deutschland GmbH, Duplicate, Duplicate of Lantus, Rapporteur: Johann Lodewijk Hillege	
TRODELVY - sacituzumab govitecan - EMEA/H/C/005182/II/0023/G Gilead Sciences Ireland UC, Rapporteur: Jan Mueller-Berghaus Request for Supplementary Information adopted on 08.06.2023.	Request for supplementary information adopted with a specific timetable.
TRODELVY - sacituzumab govitecan - EMEA/H/C/005182/II/0024/G Gilead Sciences Ireland UC, Rapporteur: Jan Mueller-Berghaus Request for Supplementary Information adopted	Request for supplementary information adopted with a specific timetable.

on 08.06.2023.	
Vaxelis - diphtheria, tetanus, pertussis (acellular, component), hepatitis B (rDNA), poliomyelitis (inact.) and haemophilus type B conjugate vaccine (adsorbed) - EMEA/H/C/003982/II/0119/G MCM Vaccine B.V., Rapporteur: Christophe Focke Opinion adopted on 08.06.2023.	Positive Opinion adopted by consensus on 08.06.2023.
VidPrevtyn Beta - SARS-CoV-2, B.1.351 variant, prefusion Spike delta TM protein, recombinant - EMEA/H/C/005754/II/0003 Sanofi Pasteur, Rapporteur: Jan Mueller-Berghaus Request for Supplementary Information adopted on 01.06.2023.	Request for supplementary information adopted with a specific timetable.
Zinplava - bezlotoxumab - EMEA/H/C/004136/II/0038/G Merck Sharp & Dohme B.V., Rapporteur: Jan Mueller-Berghaus	
WS2479 **Hexacima-** **EMEA/H/C/002702/WS2479/0146** **Hexyon-** **EMEA/H/C/002796/WS2479/0150** Sanofi Pasteur, Lead Rapporteur: Jan Mueller-Berghaus Opinion adopted on 08.06.2023.	Positive Opinion adopted by consensus on 08.06.2023.

B.5.2. CHMP assessed procedures scope: Non-Clinical and Clinical aspects

Adtralza - tralokinumab - EMEA/H/C/005255/II/0008 LEO Pharma A/S, Rapporteur: Jayne Crowe, "To update section 4.8 of the SmPC in order to update safety information based on interim results from the ECZTEND study, listed as a category 3 study in the RMP. This is a phase 3 open-label, single-arm, multi-centre, long-term extension trial to evaluate the safety and efficacy of tralokinumab in subjects with moderate-to-severe atopic dermatitis who participated in previous tralokinumab clinical trials. In addition, the MAH is taking this opportunity to update the list of local representatives in the Package Leaflet." Request for Supplementary Information adopted	Request for supplementary information adopted with a specific timetable.

on 01.06.2023, 09.02.2023.	
Aimovig - erenumab - EMEA/H/C/004447/II/0026/G Novartis Europharm Limited, Rapporteur: Kristina Dunder, "Update of section 5.1 of the SmPC in order to update clinical efficacy and safety information based on final results from studies CAMG334A2301 (LIBERTY) and CAMG334ADE01 (HER-MES). The 'LIBERTY' study is a randomized, double-blind, parallel-group, placebo-controlled phase 3 study to assess the efficacy and tolerability of Aimovig in adult patients with episodic migraine who had previously failed 2- 4 prophylactic migraine treatments, while the 'HER-MES' study is a randomized, double-blind, double-dummy, multicenter, parallel group, phase 4 study to assess tolerability and efficacy of Aimovig against topiramate in adult patients with episodic and chronic migraine." Request for Supplementary Information adopted on 12.05.2023.	
Brintellix - vortioxetine - EMEA/H/C/002717/II/0038 H. Lundbeck A/S, Rapporteur: Karin Janssen van Doorn, "Update of sections 4.2, 4.4, 4.8, 5.1 and 5.2 of the SmPC in order to include clinically relevant information on the efficacy, safety, tolerability, and PK of vortioxetine in the paediatric population based on final results from studies 12709A, 12712A and 12712B. Study 12709A is an interventional, randomized, double-blind, placebo-controlled, active-reference (fluoxetine), fixed-dose study of vortioxetine in paediatric patients aged 7 to 11 years, with Major Depressive Disorder (MDD) to evaluate efficacy and safety. Whereas studies 12712A and 12712B are 2 open-label, long-term safety and efficacy studies in children and adolescents: one 6-month extension study (study 12712A) to studies 12709A and 12710A, and one 18-month extension study (study 12712B) to study 12712A. The Package Leaflet is updated accordingly." Request for Supplementary Information adopted on 30.03.2023, 15.12.2022.	See 9.1
CABOMETYX - cabozantinib - EMEA/H/C/004163/II/0032 Ipsen Pharma, Rapporteur: Ingrid Wang,	Positive Opinion adopted by consensus on 08.06.2023.

"Update of sections 4.4 and 4.8 of the SmPC in order to add a new warning on Vanishing Bile Duct Syndrome (VBDS), to add embolism arterial to the list of adverse drug reactions (ADRs) with frequency Uncommon and to add vanishing bile duct syndrome to the list of adverse drug reactions (ADRs) with frequency Not known based on the cumulative review of the global safety database and literature search. The Package Leaflet is updated accordingly. In addition, the MAH took the opportunity to introduce minor editorial changes to the PI." Opinion adopted on 08.06.2023.	
Dupixent - dupilumab - EMEA/H/C/004390/II/0071 Sanofi Winthrop Industrie, Rapporteur: Jan Mueller-Berghaus, "Update of section 4.8 of the SmPC in order to support the longer-term (5-year) safety of dupilumab in adults with moderate-to-severe Atopic Dermatitis (AD) based on final results from study R668-AD-1225 listed as a specific PASS category 3 study in the RMP. The study R668-AD-1225 was a phase 3, multicenter, open-label extension (OLE) study of dupilumab in adults with moderate-to-severe atopic dermatitis (AD) who had previously participated in dupilumab clinical trials. The main objective of this study is to assess the long-term safety of dupilumab administered in adult patients with AD." Opinion adopted on 15.06.2023.	Positive Opinion adopted by consensus on 15.06.2023.
Dupixent - dupilumab - EMEA/H/C/004390/II/0072 Sanofi Winthrop Industrie, Rapporteur: Jan Mueller-Berghaus, "Update of sections 4.8, 5.1 and 5.2 of the SmPC in order to update the safety and efficacy information relevant to patients with hand and foot Atopic Dermatitis based on the results from study R668-AD-1924. This is a Phase 3, Multicenter, Randomized, Double-blind, Placebo-controlled, Parallel-group Study to Evaluate the Efficacy and Safety of Dupilumab in Adult and Adolescent Patients with Moderate-to-Severe Atopic Hand and Foot Dermatitis."	
Invokana - canagliflozin - EMEA/H/C/002649/II/0062 Janssen-Cilag International N.V., Rapporteur:	

Martina Weise, "Update of section 4.5 of the SmPC in order to add drug-drug interaction information with lithium, based on a safety review. In addition, the MAH took the opportunity to introduce minor editorial changes to the PI."
Request for Supplementary Information adopted on 14.04.2023.

Kanuma - sebelipase alfa - EMEA/H/C/004004/II/0044, Orphan
Alexion Europe SAS, Rapporteur: Karin Janssen van Doorn, "Update of sections 4.2 and 6.6 of the SmPC in order to limit the 1-hour infusion time only to those patients receiving the 1 mg/kg dose and to modify the table for 'recommended infusion volumes' to address the USP endotoxin limit. The Package Leaflet is updated accordingly. In addition, the MAH took the opportunity to introduce minor editorial changes to the PI."

Kisqali - ribociclib - EMEA/H/C/004213/II/0041/G
Novartis Europharm Limited, Rapporteur: Filip Josephson, "Grouped application comprising two type II variations as follows:
- Update of section 5.2 of the SmPC in order to update absorption information based on final results from study CLEE011A2117, a Phase I, single center, two-period, two-treatment, open label, randomized crossover study to investigate the absolute bioavailability of a single oral dose of 600 mg of ribociclib relative to an intravenous (i.v.) infusion of 150 mg ribociclib in healthy subjects.
- Update of sections 4.2 and 4.5 of the SmPC in order to update the recommended dose modification when ribociclib is administered in combination with CYP3A4 inhibitors and update the drug-drug interaction information on substances that may increase ribociclib plasma concentrations based on the updated PBPK modelling.
In addition, the MAH took this opportunity to introduce minor editorial changes to the Package Leaflet."

Koselugo - selumetinib - EMEA/H/C/005244/II/0013, Orphan
AstraZeneca AB, Rapporteur: Alexandre Moreau, "Update of sections 4.2 and 5.2 of the SmPC in

order to update the recommended dosage regimen to remove the fasting state and update pharmacokinetic information, based on the final results from study D1346C00015; this is a phase 1, single-arm, sequential study to evaluate the effect of food on the gastrointestinal tolerability and pharmacokinetics of selumetinib after multiple doses in adolescent children with neurofibromatosis type 1 (NF1) related plexiform neurofibromas (PN). The Package Leaflet is updated accordingly. In addition, the MAH took the opportunity to introduce minor editorial changes to the PI."	
Nexviadyme - avalglucosidase alfa - EMEA/H/C/005501/II/0008 Sanofi B.V., Rapporteur: Christian Gartner, "Update of sections 4.4, 4.8, 5.1 and 5.2 of the SmPC in order to update the list of adverse drug reactions (ADRs) and to update the safety and efficacy information, based on interim results from the open-label extension period of study EFC14028 as well as pooled safety and immunogenicity data. EFC14028 is a phase 3 randomized, multicenter, multinational, double-blinded study comparing the efficacy and safety of repeated biweekly infusions of avalglucosidase alfa and alglucosidase alfa in treatment naïve patients with late-onset Pompe disease. The Package Leaflet is updated accordingly. In addition, the MAH took the opportunity to introduce minor editorial changes to the PI." Request for Supplementary Information adopted on 08.06.2023.	Request for supplementary information adopted with a specific timetable.
NUVAXOVID - covid-19 vaccine (recombinant, adjuvanted) - EMEA/H/C/005808/II/0030 Novavax CZ, a.s., Rapporteur: Johann Lodewijk Hillege, "Submission of 6-month efficacy and safety interim data from the ongoing randomized, observer-blinded, placebo-controlled clinical studies 2019nCoV-501, 2019nCoV-301 and 2019nCoV-302." Request for Supplementary Information adopted on 15.12.2022.	
Orgovyx - relugolix - EMEA/H/C/005353/II/0012 Accord Healthcare S.L.U., Rapporteur: Johann	Positive Opinion adopted by consensus on 01.06.2023.

Lodewijk Hillege, "Update of section 4.5 of the SmPC based on final results from study MVT-601-9039; this is an In vitro Interaction Study of Relugolix with human OATP2B1 Uptake Transporter."
Opinion adopted on 01.06.2023.

Paxlovid - nirmatrelvir / ritonavir - EMEA/H/C/005973/II/0040/G

Pfizer Europe MA EEIG, Rapporteur: Jean-Michel Race, "Grouped application comprising two type II variations as follows:
- Update of section 4.3 of the SmPC in order to add 'Mineralocorticoid receptor antagonists: finerenone' and 'Opioid antagonists: naloxegol' under Medicinal products that are highly dependent on CYP3A for clearance and for which elevated plasma concentrations are associated with serious and/or life-threatening reactions and to add 'primidone' and 'Cystic fibrosis transmembrane conductance regulator potentiators: lumacaftor/ivacaftor' under Medicinal products that are potent CYP3A inducers where significantly reduced nirmatrelvir/ritonavir plasma concentrations may be associated with the potential for loss of virologic response and possible resistance based on the review of the PI for a number of medicines from different drug classes that are metabolised by CYP3A4 or CYP2D6, transported by P-gp, or induce CYP3A4.
- Update of section 4.5 of the SmPC in order to add drug-drug interaction information with Alpha1-adrenoreceptor antagonist, Analgesics, Antiarrhythmics, Anticoagulants, Anticonvulsants, Anti-HIV, Anti-infectives, β2-agonist (long acting), Calcium channel antagonists, Cardiovascular agents and Migraine medicinal products, to add drug-drug interaction information with Cystic fibrosis transmembrane conductance regulator potentiators, Dipeptidyl peptidase 4 (DPP4) inhibitors, Janus kinase (JAK) inhibitors, Mineralocorticoid receptor antagonists, Muscarinic receptor antagonists, Neuropsychiatric agents and Opioid antagonists and to remove cross reference to section 4.4 from information regarding coadministration of Paxlovid with Antidepressants based on the review of the PI for a number of medicines from different drug classes that are metabolised by CYP3A4 or CYP2D6, transported by P-gp, or

induce CYP3A4."	
Reblozyl - luspatercept - EMEA/H/C/004444/II/0016, Orphan Bristol-Myers Squibb Pharma EEIG, Rapporteur: Daniela Philadelphy, "Update of section 5.1 of the SmPC in order to reflect the correct values of late-reported transfusions and modifications of previously reported transfusions based on final results from study ACE-536-B-THAL-001 (BELIEVE), A Phase 3, Double-Blind, Randomized, Placebo-Controlled, Multicenter Study To Determine The Efficacy And Safety Of Luspatercept (Ace-536) Versus Placebo In Adults Who Require Regular Red Blood Cell Transfusions Due To Beta-Thalassemia." Request for Supplementary Information adopted on 16.03.2023.	
RINVOQ - upadacitinib - EMEA/H/C/004760/II/0034 AbbVie Deutschland GmbH & Co. KG, Rapporteur: Kristina Dunder, "Submission of the final report from study M13-542, listed as a category 3 study in the RMP. This is a phase 3, randomized, double-blind study comparing upadacitinib (ABT-494) to placebo on stable conventional synthetic disease-modifying anti rheumatic drugs (csDMARDs) in subjects with moderately to severely active rheumatoid arthritis with inadequate response or intolerance to biologic DMARDs (bDMARDs)." Request for Supplementary Information adopted on 08.06.2023.	Request for supplementary information adopted with a specific timetable.
RINVOQ - upadacitinib - EMEA/H/C/004760/II/0035 AbbVie Deutschland GmbH & Co. KG, Rapporteur: Kristina Dunder, "Submission of the final report from study M13-549 listed as a category 3 study in the RMP. This is a Phase III, Randomized, Double-Blind Study Comparing Upadacitinib (ABT-494) to Placebo in Subjects with Moderately to Severely Active Rheumatoid Arthritis Who Are on a Stable Dose of Conventional Synthetic Disease-Modifying Anti Rheumatic Drugs (csDMARDs) and Have an Inadequate Response to csDMARDs." Request for Supplementary Information adopted on 08.06.2023.	Request for supplementary information adopted with a specific timetable.
Rydapt - midostaurin -	

EMEA/H/C/004095/II/0029, Orphan
Novartis Europharm Limited, Rapporteur: Johann Lodewijk Hillege, "Update of section 4.8 of the SmPC in order to add "Acute febrile neutrophilic dermatosis (Sweet syndrome)" to the list of adverse drug reactions (ADRs) with frequency not known based on pre-clinical data, clinical trial datasets, scientific literature and safety databases. The Package Leaflet is updated accordingly."
Request for Supplementary Information adopted on 30.03.2023.

Scemblix - asciminib - EMEA/H/C/005605/II/0004/G, Orphan
Novartis Europharm Limited, Rapporteur: Janet Koenig, "Grouped application comprising two type II variations as follows:
- Submission of the final reports from studies DMPK-R2200470 (REC). This is an in vitro evaluation of inducibility of OATP1V1, MDR1 and CYP3A4 by asciminib using human hepatocytes.
- Submission of the final report from study DMPK-R2270399 (REC). This is a physiologically based PK modelling and simulations to characterise the effect of cyclodextrins on the exposure of asciminib."

Spinraza - nusinersen - EMEA/H/C/004312/II/0029, Orphan
Biogen Netherlands B.V., Rapporteur: Bruno Sepodes, "Update of section 5.3 of the SmPC in order to update non-clinical information based on final results from study P058-17-02. This is a 24-month carcinogenicity study when administered by subcutaneous injection in mouse."

Synagis - palivizumab - EMEA/H/C/000257/II/0132
AstraZeneca AB, Rapporteur: Thalia Marie Estrup Blicher, "Update of sections 4.2 and 5.1 of the SmPC in order to update safety information based on results from safety data evaluations from multiple sources, including the clinical study W00-350, post-Marketing Clinical Surveillance Programme (REACH), literature searches and the AstraZeneca Global Patient Safety database."
Request for Supplementary Information adopted on 30.03.2023.

TAGRISSO - osimertinib - EMEA/H/C/004124/II/0050

AstraZeneca AB, Rapporteur: Carolina Prieto Fernandez, "Update of sections 4.2, 4.4 and 4.8 of the SmPC in order to modify the posology recommendations in the case of toxic epidermal necrolysis (TEN), add it as a new warning and add it to the list of adverse drug reactions (ADRs) with frequency uncommon and to update the frequency of interstitial lung disease (ILD) based on an internal safety information review. The Package Leaflet is updated accordingly. In addition, the MAH took the opportunity to introduce minor editorial changes to the PI."
Request for Supplementary Information adopted on 30.03.2023.

Tecvayli - teclistamab - EMEA/H/C/005865/II/0003

Janssen-Cilag International N.V., Rapporteur: Johanna Lähteenvuo, "Update of sections 4.2, 4.6 and 5.2 of the SmPC in order to revise the dosing schedule, amend recommendations on contraception and breast-feeding and to update pharmacokinetic information, based on the latest data available; the Package Leaflet is updated accordingly. In addition, the MAH took the opportunity to introduce minor editorial changes to the PI and update the list of local representatives in the Package Leaflet."
Request for Supplementary Information adopted on 25.05.2023.

TEPMETKO - tepotinib - EMEA/H/C/005524/II/0005

Merck Europe B.V., Rapporteur: Filip Josephson, "Update of sections 4.5 and 5.2 of the SmPC in order to remove interactions with 'CYP and P-gp inducers' and 'dual strong CYP3A and P-gp inhibitors, and P-gp inhibitors' and to update pharmacokinetic information based on final results from the drug-drug interaction (DDI) studies MS200095-0051 and MS200095-0053. Study MS200095-0051 is a phase 1, open-label, single-sequence, cross-over study to evaluate the effect of multiple doses of carbamazepine on single-dose tepotinib pharmacokinetics in healthy participants, while study MS200095-0053 is a phase 1, open-label, single-sequence, cross-over study to evaluate the effect of

multiple doses of itraconazole on single-dose tepotinib pharmacokinetics in healthy participants. The Package Leaflet is updated accordingly. In addition, the MAH took the opportunity to introduce minor changes to the PI."
Request for Supplementary Information adopted on 09.02.2023.

TUKYSA - tucatinib - EMEA/H/C/005263/II/0013

Seagen B.V., Rapporteur: Aaron Sosa Mejia, "Submission of the final report from study ONT-380-206 (HER2CLIMB) listed as a PAES in the Annex II of the Product Information. This is a phase 2 randomized, double-blinded, controlled study of tucatinib vs. placebo in combination with capecitabine and trastuzumab in patients with pretreated unresectable locally advanced or metastatic HER2+ breast carcinoma. The Annex II is updated accordingly."

Vargatef - nintedanib - EMEA/H/C/002569/II/0047/G

Boehringer Ingelheim International GmbH, Rapporteur: Aaron Sosa Mejia, "Grouped application containing:
C.I.4: Update of sections 4.2, 5.1 and 5.2 of the SmPC in order to update information regarding the paediatric population based on results from study 1199-0337; this is a double blind, randomised, placebo-controlled trial to evaluate the dose-exposure and safety of nintedanib on top of standard of care for 24 weeks, followed by open label treatment with nintedanib of variable duration, in children and adolescents (6 to 17 year-old) with clinically significant fibrosing interstitial lung disease. The Package Leaflet is updated accordingly.
C.I.4: Update of sections 4.2 and 5.2 of the SmPC in order to improve the recommendation for the administration of nintedanib based on food compatibility data. The Package Leaflet is updated accordingly.
In addition, the MAH took the opportunity to update the list of local representatives in the Package Leaflet and to introduce minor editorial changes to the PI."
Request for Supplementary Information adopted on 26.04.2023, 26.01.2023.

Vokanamet - canagliflozin / metformin - EMEA/H/C/002656/II/0067/G Janssen-Cilag International N.V., Rapporteur: Martina Weise, "C.I.4: Update of section 4.5 of the SmPC in order to add drug-drug interaction information with lithium, based on a safety review. In addition, the MAH took the opportunity to introduce minor editorial changes to the PI. C.I.4: Update of section 4.8 of the SmPC in order to update the frequency for 'vitamin B12 deficiency' in the list of adverse drug reactions (ADRs) to 'common', based on a safety review. The Package Leaflet is updated accordingly." Request for Supplementary Information adopted on 14.04.2023.	
Xultophy - insulin degludec / liraglutide - EMEA/H/C/002647/II/0049 Novo Nordisk A/S, Rapporteur: Kristina Dunder, "Update of section 4.8 of the SmPC in order to add dysgeusia to the list of adverse drug reactions (ADRs) with frequency uncommon based on post-marketing data and literature. The Package Leaflet is updated accordingly. In addition, the MAH took the opportunity to update section 4.9 of the SmPC to update overdose information and to amend Annex A." Opinion adopted on 01.06.2023.	Positive Opinion adopted by consensus on 01.06.2023.
Zokinvy - lonafarnib - EMEA/H/C/005271/II/0004, Orphan EigerBio Europe Limited, Rapporteur: Johann Lodewijk Hillege, "Update of sections 4.2, 4.4, 4.5 and 6.6 of the SmPC in order to update drug-drug interaction information based on final results from Drug-Drug Interaction study EIG-LNF-021. This is a phase I, single-center, two period, single sequence, study to assess the effects of lonafarnib autoinhibition following multiple-dose lonafarnib and the effects of a nonspecific CYP2C9 inhibitor on multiple-dose lonafarnib pharmacokinetics in healthy subjects. The Package Leaflet is updated accordingly."	
WS2368 **Invokana-** **EMEA/H/C/002649/WS2368/0061** **Vokanamet-** **EMEA/H/C/002656/WS2368/0066** Janssen-Cilag International N.V., Lead Rapporteur: Martina Weise, "To update section	

4.4 of the SmPC in order amend an existing warning on the diabetic ketoacidosis, to indicate that glucosuria may persist longer than expected and that DKA may be prolonged after discontinuation of canagliflozin in some patients based on a cumulative review of literature, MAH global safety database, preclinical and clinical pharmacology data, and clinical study data including cases reports." Request for Supplementary Information adopted on 12.05.2023, 08.12.2022.	
WS2407 **Efficib-EMEA/H/C/000896/WS2407/0110** **Janumet-EMEA/H/C/000861/WS2407/0109** **Ristfor-EMEA/H/C/001235/WS2407/0098** **Velmetia-EMEA/H/C/000862/WS2407/0115** Merck Sharp & Dohme B.V., Lead Rapporteur: Johann Lodewijk Hillege, "To include significant changes to sections 4.4 and 4.8 of the SmPC and section 4 of the Package Leaflet for the medicinal products Janumet, Velmetia, Ristfor and Efficib, containing the active substances Metformin hydrochloride and Sitagliptin phosphate in order to include a warning for vitamin B12 deficiency and to change the frequency of vitamin B12 deficiency from very rare to common following the assessment of the medicinal product Glucophage, which also contains the active substance metformin, assessed as part of a mutual recognition procedure FR/H/0181/001-3. The same wording is used for the combination product. The Package Leaflet is updated accordingly. In addition, the MAH took the opportunity to update the list of local representatives in the Package Leaflet for Janumet, Ristfor and Efficib and to improve the wording in section 2 of the Package Leaflet." Opinion adopted on 01.06.2023. Request for Supplementary Information adopted on 09.02.2023.	Positive Opinion adopted by consensus on 01.06.2023.
WS2467 **Adrovance-EMEA/H/C/000759/WS2467/0051** **FOSAVANCE-EMEA/H/C/000619/WS2467/0054** **VANTAVO-**	

EMEA/H/C/001180/WS2467/0041
Organon N.V., Lead Rapporteur: Christian Gartner, "Update of sections 4.4 and 4.8 the SmPC in order to include information on the risk of low-energy fractures in bones other than femur based on post-marketing case reports and the literature. The Package Leaflet and Labelling are updated accordingly. In addition, the MAH the opportunity to update the list of local representatives in the Package Leaflet, to bring the PI in line with the latest QRD template and to introduce editorial changes."

WS2488
Aluvia-EMEA/H/W/000764/WS2488/0118
Kaletra-EMEA/H/C/000368/WS2488/0197
Norvir-EMEA/H/C/000127/WS2488/0168
AbbVie Deutschland GmbH & Co. KG, Lead Rapporteur: Johann Lodewijk Hillege, "C.I.z - To update section 4.5, Interactions with other medicinal products and other forms of interaction, of the Norvir SmPC to align with the text in the Prezista product information, following the final Assessment Report for Norvir Post-Authorisation Measure LEG 033.12 (EMEA/H/C/000127/LEG 033.012) on 27 January 2023. The PL has been updated accordingly. The MAH is taking the opportunity to also update the Kaletra and Aluvia product information with the same information. Furthermore, the MAH has taken the opportunity to implement minor editorial changes to the Romanian and Norwegian Patient Information Leaflets."

WS2489/G
Kinzalmono-
EMEA/H/C/000211/WS2489/0119/G
Micardis-
EMEA/H/C/000209/WS2489/0127/G
Pritor-
EMEA/H/C/000210/WS2489/0132/G
Boehringer Ingelheim International GmbH, Lead Rapporteur: Armando Genazzani, "Grouped application consisting of:
C.I.4: Update of section 4.8 of the SmPC in order to include "hyponatremia" to the list of adverse drug reactions (ADRs) with frequency "rare", based on post-marketing data and literature;
C.I.z (Type IB unforeseen): Update of section

4.2 to include the possibility of using the combination of telmisartan and amlodipine for lowering blood pressure; C.I.z (Type IB unforeseen): Update of section 4.7 of the SmPC to replace the terms "dizziness" and "drowsiness" by "syncope" and "vertigo" to align with adverse reactions table in section 4.8 of SmPC. The Package Leaflet is updated accordingly. In addition, the MAH took the opportunity to update the list of local representatives in the Package Leaflet, bring the PI in line with the latest QRD template version 10.3, and to implement editorial changes to the SmPC."	
WS2492 **Glyxambi-** **EMEA/H/C/003833/WS2492/0052** **Synjardy-** **EMEA/H/C/003770/WS2492/0071** Boehringer Ingelheim International GmbH, Lead Rapporteur: Johann Lodewijk Hillege, "Submission of the final report from study PASS 1245-0137 listed as a category 3 study in the RMP. This is a multicentre international randomised parallel group double-blind placebo-controlled clinical trial of EMPAgliflozin once daily to assess cardiorenal outcomes in patients with chronic KIDNEY disease."	

B.5.3. CHMP-PRAC assessed procedures

Carbaglu - carglumic acid - **EMEA/H/C/000461/II/0045** Recordati Rare Diseases, Rapporteur: Fátima Ventura, PRAC Rapporteur: Ana Sofia Diniz Martins, "Update of sections 4.2, 4.4 and 5.2 of the SmPC in order to include a proposed dose adjustment for patients with impaired renal function based on final results from study RCD-P0-027; this is a Phase I, multicenter, open-label, parallel-group adaptive pharmacokinetic single dose study of oral Carbaglu in subjects with normal and varying degrees of impaired renal function. The Package Leaflet is updated accordingly. The RMP version 2.2 has also been submitted. In addition, the MAH took the opportunity to implement editorial changes in Annex II and Labelling, and to bring the PI in line with the latest QRD template version 10.3." Request for Supplementary Information adopted	

on 30.03.2023.	
Coagadex - human coagulation factor X - EMEA/H/C/003855/II/0046, Orphan BPL Bioproducts Laboratory GmbH, Rapporteur: Daniela Philadelphy, PRAC Rapporteur: Menno van der Elst, "Submission of the final report from study TEN06 - NCT03161626 (REC EMEA/H/C/003855). This is a non-interventional, multicentre, post-marketing registry study in three patients with moderate or severe hereditary FX deficiency, to assess Coagadex administered peri-operatively for haemostatic cover in major surgery during routine post-marketing use. The primary objective is to collect additional surgical data on the clinical effectiveness of Coagadex, in a post-marketing environment, for peri-operative haemostatic cover during major surgery in patients with moderate or severe hereditary factor X (FX) deficiency. The RMP version 3.0 has also been submitted." Opinion adopted on 08.06.2023. Request for Supplementary Information adopted on 16.03.2023.	Positive Opinion adopted by consensus on 08.06.2023.
Enhertu - trastuzumab deruxtecan - EMEA/H/C/005124/II/0031 Daiichi Sankyo Europe GmbH, Rapporteur: Aaron Sosa Mejia, PRAC Rapporteur: Inês Ribeiro-Vaz, "Update of sections 4.8, 5.1 and 5.2 of the SmPC in order to update safety, efficacy and pharmacokinetic information based on data from study DS8201-A-U301 and study DS8201-A-U302. Study U301was a Phase 3, randomized, 2-arm, open-label, multicenter study designed to compare the safety and efficacy of T-DXd vs TPC in HER2-positive, unresectable and/or metastatic BC subjects who were resistant or refractory to T-DM1. Study U302 was a Phase 3, multicenter, randomized, open-label, 2-arm, active-controlled study in subjects with unresectable and/or metastatic HER2-positive (IHC 3+ or ISH-positive) BC previously treated with trastuzumab plus taxane in the advanced/metastatic setting or who had progressed within 6 months after neoadjuvant or adjuvant treatment involving a regimen including trastuzumab plus taxane. The Package Leaflet and Annex II are updated accordingly. The updated RMP version 4.1 has also been	Request for supplementary information adopted with a specific timetable.

submitted." Request for Supplementary Information adopted on 08.06.2023.	
Fluenz Tetra - influenza vaccine (live attenuated, nasal) - EMEA/H/C/002617/II/0130 AstraZeneca AB, Rapporteur: Christophe Focke, PRAC Rapporteur: Jean-Michel Dogné, "Submission of the final report from study MA-VA-MEDI3250-1116 (A Case Control Study of the Effectiveness of Q/LAIV Versus Inactivated Influenza Vaccine and No Vaccine in Subjects 2 to 17 Years of Age) listed as a category 3 study in the RMP. This was an observational study. The objective of this study was to evaluate the effectiveness of Q/LAIV compared to IIV or no vaccine in community-dwelling subjects 2 to 17 years of age against laboratory-confirmed influenza. The RMP version 11.0 has also been submitted." Opinion adopted on 08.06.2023.	Positive Opinion adopted by consensus on 08.06.2023.
GAVRETO - pralsetinib - EMEA/H/C/005413/II/0010 Roche Registration GmbH, Rapporteur: Aaron Sosa Mejia, PRAC Rapporteur: Ulla Wändel Liminga, "Update of sections 4.8, 5.1 and 5.2 of the SmPC in order to update efficacy and safety information in the treatment of adult patients with RET fusion-positive advanced NSCLC based on final results (NSCLC indication) from study ARROW/BO42863, a Phase 1/2 Study of the Highly-selective RET Inhibitor, BLU 667, in Patients With Thyroid Cancer, Non-Small Cell Lung Cancer (NSCLC), and Other Advanced Solid Tumours listed as a specific obligation in the Annex II. The RMP version 1.5 has also been submitted." Request for Supplementary Information adopted on 25.05.2023, 23.02.2023.	
GAVRETO - pralsetinib - EMEA/H/C/005413/II/0012 Roche Registration GmbH, Rapporteur: Aaron Sosa Mejia, PRAC Rapporteur: Ulla Wändel Liminga, "Update of sections 4.2, 4.4 and 4.5 of the SmPC in order to amend posology recommendations, warnings and drug-drug interaction information regarding the co-administration with CYP3A4 inhibitors, P-gp inhibitors and CYP3A4 inducers based on final	

results from the DDI study GP43162, listed as a category 3 study in the RMP, as well as results from the physiologically based pharmacokinetic (PBPK) analyses summarised in the PBPK Report 1120689. Study GP43162 is a phase 1, open-label, fixed-sequence study to evaluate the effect of a single dose of cyclosporine on the single dose pharmacokinetics of pralsetinib in healthy subjects. The RMP version 1.6 has also been submitted." Request for Supplementary Information adopted on 30.03.2023.	
GIVLAARI - givosiran - EMEA/H/C/004775/II/0011/G, Orphan Alnylam Netherlands B.V., Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Martin Huber, "Update of section 5.3 of the SmPC based on final results from study AS1-GLP18-007 listed as a category 3 study in the RMP; This is a 104-week Subcutaneous Injection Carcinogenicity Study in Sprague Dawley Rats. Update of section 5.3 of the SmPC based on final results from study AS1-GLP18-004; This is a 26-week Subcutaneous Injection Carcinogenicity Study in TgRasH2 Mice. The RMP version 2.1 has also been submitted. In addition, the MAH took the opportunity to update the list of local representatives in the Package Leaflet." Opinion adopted on 08.06.2023. Request for Supplementary Information adopted on 14.04.2023, 12.01.2023, 29.09.2022.	Positive Opinion adopted by consensus on 08.06.2023.
Imnovid - pomalidomide - EMEA/H/C/002682/II/0047, Orphan Bristol-Myers Squibb Pharma EEIG, Rapporteur: Carolina Prieto Fernandez, PRAC Rapporteur: Monica Martinez Redondo, "Update of section 4.4 of the SmPC, Annex IID and Article 127a and the tools/documents included in the Educational Healthcare Professional Kit, in order to harmonise the terminology utilised in the RMP and PI documents relating to the safety concern of teratogenicity and its risk minimisation measure of the Pregnancy Prevention Plan across the 3 IMiDs. These proposed changes will only have a limited impact on the National Competent Authority (NCA)-approved content/text of the educational materials, and the key messages to the HCP	Positive Opinion adopted by consensus on 08.06.2023.

and patients. Furthermore, the regulatory obligations regarding the PPP will not be impacted. The updated RMP version 16 was provided." Opinion adopted on 08.06.2023. Request for Supplementary Information adopted on 14.04.2023, 09.02.2023, 27.10.2022.	
Piqray - alpelisib - EMEA/H/C/004804/II/0018 Novartis Europharm Limited, Rapporteur: Carolina Prieto Fernandez, PRAC Rapporteur: Menno van der Elst, "Update of sections 4.5 and 5.2 of the SmPC in order to update drug-drug interaction information, based on final results from study BYL719A2111; this is a phase 1, open-label, fixed-sequence, two-period drug-drug interaction (DDI) study evaluating the PK probe substrates for CYP3A4, CYP2B6, CYP2C8, CYP2C9, and CYP2C19 when administered either alone or in combination with repeated doses of alpelisib. The Annex II and Package Leaflet are updated accordingly. The RMP version 6.0 has also been submitted. In addition, the MAH took the opportunity to introduce minor editorial changes to the PI." Request for Supplementary Information adopted on 16.03.2023.	
Revlimid - lenalidomide - EMEA/H/C/000717/II/0123 Bristol-Myers Squibb Pharma EEIG, Rapporteur: Alexandre Moreau, PRAC Rapporteur: Tiphaine Vaillant, "Update of section 4.4 of the SmPC, Annex IID and Article 127a and the tools/documents included in the Educational Healthcare Professional Kit, in order to harmonise the terminology utilised in the RMP and PI documents relating to the safety concern of teratogenicity and its risk minimisation measure of the Pregnancy Prevention Plan across the 3 IMiDs. These proposed changes will only have a limited impact on the National Competent Authority (NCA)-approved content/text of the educational materials, and the key messages to the HCP and patients. Furthermore, the regulatory obligations regarding the PPP will not be impacted. The MAH is also taking the opportunity to update the RMP with PASS Protocol milestones. The updated RMP version 38.2 was provided."	Positive Opinion adopted by consensus on 08.06.2023.

Opinion adopted on 08.06.2023. Request for Supplementary Information adopted on 14.04.2023, 09.02.2023, 27.10.2022.	
Simponi - golimumab - EMEA/H/C/000992/II/0109 Janssen Biologics B.V., Rapporteur: Kristina Dunder, PRAC Rapporteur: Mari Thorn, "Update of the Package Leaflet in order to update the Instructions for Use (IFU) for the pre-filled pen." Request for Supplementary Information adopted on 26.04.2023, 26.01.2023.	
Thalidomide BMS - thalidomide - EMEA/H/C/000823/II/0076 Bristol-Myers Squibb Pharma EEIG, Rapporteur: Alexandre Moreau, PRAC Rapporteur: Tiphaine Vaillant, "Update of section 4.4 of the SmPC, Annex IID and Article 127a and the tools/documents included in the Educational Healthcare Professional Kit, in order to harmonise the terminology utilised in the RMP and PI documents relating to the safety concern of teratogenicity and its risk minimisation measure of the Pregnancy Prevention Plan across the 3 IMiDs. These proposed changes will only have a limited impact on the National Competent Authority (NCA)-approved content/text of the educational materials, and the key messages to the HCP and patients. Furthermore, the regulatory obligations regarding the PPP will not be impacted. The MAH is also taking the opportunity to update the RMP with PASS Protocol milestones, and to make some editorial changes in the labelling. The updated RMP version 20 was provided." Opinion adopted on 08.06.2023. Request for Supplementary Information adopted on 14.04.2023, 09.02.2023, 27.10.2022.	Positive Opinion adopted by consensus on 08.06.2023.
Vabysmo - faricimab - EMEA/H/C/005642/II/0002 Roche Registration GmbH, Rapporteur: Jayne Crowe, PRAC Rapporteur: Inês Ribeiro-Vaz, "Update of sections 4.2, 4.4, 4.8 and 5.1 of the SmPC in order to update efficacy and safety information and to update the warnings and the list of adverse drug reactions (ADRs), based on longer-term results from studies GR40306 (TENAYA) and GR40844 (LUCERNE); these are phase 3, multicenter, randomized, double-masked, active comparator-controlled, 112-	

week studies to evaluate the efficacy and safety of faricimab in patients with neovascular age-related macular degeneration (nAMD); the Package Leaflet is updated accordingly. The RMP version 2.0 has also been submitted. In addition, the MAH took the opportunity to introduce minor editorial changes to the PI."
Request for Supplementary Information adopted on 23.02.2023.

Veklury - remdesivir - EMEA/H/C/005622/II/0046

Gilead Sciences Ireland UC, Rapporteur: Janet Koenig, PRAC Rapporteur: Eva Jirsová, "Update of sections 4.6 and 5.1 of the SmPC in order to update information on pregnancy and breast-feeding based on final results from study IMPAACT 2032 listed as a category 3 study in the RMP; this is a phase 4, prospective, open-label, non-randomized study to address PK and safety of remdesivir in pregnant women. The Package Leaflet is updated accordingly. The RMP version 5.2 has also been submitted."
Request for Supplementary Information adopted on 16.03.2023.

Veklury - remdesivir - EMEA/H/C/005622/II/0050

Gilead Sciences Ireland UC, Rapporteur: Janet Koenig, PRAC Rapporteur: Eva Jirsová, "Update of sections 4.2, 4.4, 4.8 and 5.2 of the SmPC in order to address the safety of remdesivir and its metabolites in patients with hepatic impairment and to update information on hepatic and coagulation laboratory abnormalities based on final results from study GS US 540 9014: "A phase 1 open-label, adaptive, single-dose study to evaluate the pharmacokinetics of remdesivir and its metabolite(s) in subjects with normal hepatic function and hepatic impairment", listed as a category 3 study in the RMP, and on safety data from post-marketing and clinical trials experience.
The Package Leaflet is updated accordingly. The RMP version 5.4 has also been submitted. In addition, the MAH took the opportunity submit Minor Linguistic Amendments (MLA) for Veklury."

XOSPATA - gilteritinib - EMEA/H/C/004752/II/0013, Orphan

Astellas Pharma Europe B.V., Rapporteur: Ingrid

Wang, PRAC Rapporteur: Martin Huber, "Update of sections 4.2 and 5.2 in order to update the information on renal impairment based on final results from study 2215-CL-0114, listed as a category 3 study in the RMP. Study 2215-CL-0114 is a phase 1, single-dose, open-label study to investigate the effect of renal impairment on gilteritinib pharmacokinetics, safety and tolerability in 9 participants with severe renal impairment compared to 8 participants with normal renal function. The RMP version 4.0 has also been submitted. In addition, the MAH took the opportunity to introduce editorial changes."	

B.5.4. PRAC assessed procedures

PRAC Led **Aldurazyme - laronidase - EMEA/H/C/000477/II/0085** Sanofi B.V., PRAC Rapporteur: Nathalie Gault, PRAC-CHMP liaison: Alexandre Moreau, "To update section 4.2 of the SmPC in order to modify the administration instructions following the assessment of procedure PSUSA/00001830/202104 based on literature review. The Package Leaflet is updated accordingly. The RMP version 1.0 has also been submitted." Request for Supplementary Information adopted on 08.06.2023, 09.02.2023.	Request for supplementary information adopted with a specific timetable.
PRAC Led **CABOMETYX - cabozantinib - EMEA/H/C/004163/II/0033** Ipsen Pharma, PRAC Rapporteur: Menno van der Elst, PRAC-CHMP liaison: Johann Lodewijk Hillege, "Submission of the final report from study F-FR-60000-001 (CASSIOPE) listed as a category 3 study in the RMP. This is a prospective, non-imposed and non-interventional study of cabozantinib tablets in adults with advanced renal cell carcinoma (RCC) following prior vascular endothelial growth factor (VEGF)-targeted therapy. The RMP version 7.0 has also been submitted." Request for Supplementary Information adopted on 08.06.2023.	Request for supplementary information adopted with a specific timetable.
PRAC Led **Eurartesim - piperaquine tetraphosphate /**	Request for supplementary information adopted with a specific timetable.

artenimol - EMEA/H/C/001199/II/0040/G Alfasigma S.p.A., PRAC Rapporteur: Martin Huber, PRAC-CHMP liaison: Janet Koenig, "C.I.13: Submission of the final report from the effectiveness evaluation survey for Eurartesim (protocol no. 3366) listed as a category 3 study in the RMP. This is a European multi-centre online survey to assess physician understanding of the revised edition of the educational material. Consequential changes to RMP version 16.1 have been implemented. C.I.11.b: Submission of an updated RMP version 16.1 in order to delete "Severe Malaria" from the Missing Information." Request for Supplementary Information adopted on 08.06.2023.	
PRAC Led **EXJADE - deferasirox - EMEA/H/C/000670/II/0085** Novartis Europharm Limited, PRAC Rapporteur: Tiphaine Vaillant, PRAC-CHMP liaison: Alexandre Moreau, "Submission of an updated RMP version 21.0 in order to include the physician survey CICL670A2429 as a PASS category 3, based on the submission of a draft version of the protocol for the physician survey CICL670A2429. The Annex IID is updated to remove one sentence related to 'surveillance programme' and to introduce a minor correction." Request for Supplementary Information adopted on 08.06.2023.	Request for supplementary information adopted with a specific timetable.
PRAC Led **Fintepla - fenfluramine - EMEA/H/C/003933/II/0017, Orphan** UCB Pharma SA, Rapporteur: Thalia Marie Estrup Blicher, PRAC Rapporteur: Martin Huber, PRAC-CHMP liaison: Janet Koenig, "Submission of an updated RMP version 2.10 in order to implement a targeted follow-up questionnaire (FUQ) to further improve the collection of follow-up information on cases of vascular heart disease (VHD) and pulmonary arterial hypertension (PAH) suggested by PRAC following the assessment of procedure EMEA/H/C/PSUSA/00010907/202112." Request for Supplementary Information adopted on 08.06.2023, 14.04.2023, 12.01.2023.	Request for supplementary information adopted with a specific timetable.
PRAC Led **JCOVDEN - COVID-19 vaccine Janssen**	Positive Opinion adopted by consensus on

(Ad26.COV2.S) - EMEA/H/C/005737/II/0071/G Janssen-Cilag International N.V., PRAC Rapporteur: Ulla Wändel Liminga, PRAC-CHMP liaison: Kristina Dunder, "Grouped application consisting of: 1) Submission of the final study report of a clinical TTS characterisation study listed as a category 3 study in the RMP. This is a Test Pre- and Post-Vaccination Serum Across All Populations Using Clinical Samples From Ad26-based Company Vaccine Studies Other Than Ad26.COV2.S; 2) Submission of the Addendum to final CSR of the study VAC31518COV2001 listed as a category 3 study in the RMP. This is a randomized, double-blind, placebo-controlled Phase 2a study to evaluate a range of dose levels and vaccination intervals of Ad26.COV2.S in healthy adults aged 18 to 55 years, and adults aged 65 years and older. The RMP version 7.1 is submitted and updated accordingly. In addition, the MAH updated the milestones of several studies." Opinion adopted on 08.06.2023.	08.06.2023.
PRAC Led **Nexium Control - esomeprazole - EMEA/H/C/002618/II/0038** GlaxoSmithKline Dungarvan Ltd, PRAC Rapporteur: Rugile Pilviniene, PRAC-CHMP liaison: Vilma Petrikaite, "Submission of an updated RMP version 2.0 in order to update the list of safety concerns to meet the definition of important risk and missing information provided in GVP Module V Rev. 2" Request for Supplementary Information adopted on 08.06.2023.	Request for supplementary information adopted with a specific timetable.
PRAC Led **Olumiant - baricitinib - EMEA/H/C/004085/II/0039/G** Eli Lilly Nederland B.V., PRAC Rapporteur: Adam Przybylkowski, PRAC-CHMP liaison: Ewa Balkowiec Iskra, "Submission of the final reports from studies I4V-MC-B016 and I4V-MC-B011 (in terms of Objective 4 for RA cohort) listed as category 3 non-interventional PASS studies in the RMP. B016 is a drug utilisation study for the assessment of off-label use of baricitinib in the paediatric population in the United Kingdom.	Positive Opinion adopted by consensus on 08.06.2023.

B011 is a retrospective cohort study to assess the safety of baricitinib compared with other therapies used in the treatment of rheumatoid arthritis in Nordic countries. The RMP version 19.1 has also been submitted." Opinion adopted on 08.06.2023.	
PRAC Led **Ozurdex - dexamethasone - EMEA/H/C/001140/II/0044** AbbVie Deutschland GmbH & Co. KG, PRAC Rapporteur: Maria del Pilar Rayon, PRAC-CHMP liaison: Maria Concepcion Prieto Yerro, "Submission of an updated Annex II and RMP version 11 in order to remove additional risk minimisation measure: Patient guide, audio CD (where required)." Request for Supplementary Information adopted on 08.06.2023, 16.03.2023.	Request for supplementary information adopted with a specific timetable.
PRAC Led **Pradaxa - dabigatran etexilate - EMEA/H/C/000829/II/0144** Boehringer Ingelheim International GmbH, PRAC Rapporteur: Anette Kirstine Stark, PRAC-CHMP liaison: Thalia Marie Estrup Blicher, "Submission of the final report from the Human Factors Study (007-HFE-009035), listed as a category 3 study in the RMP; this is a non-interventional study to assess the effectiveness of a training video to mitigate potential medication errors during the reconstitution and dosing of the dabigatran etexilate paediatric oral solution." Opinion adopted on 08.06.2023.	Positive Opinion adopted by consensus on 08.06.2023.
PRAC Led **Revlimid - lenalidomide - EMEA/H/C/000717/II/0126** Bristol-Myers Squibb Pharma EEIG, PRAC Rapporteur: Tiphaine Vaillant, PRAC-CHMP liaison: Alexandre Moreau, "Submission of the final report from study CC-5013-MDS-010 listed as an obligation in the Annex II of the Product Information. This is a prospective non-interventional post-authorisation safety study (PASS), designed as a disease registry of patients with transfusion dependent IPSS low or intermediate-1-risk myelodysplastic syndromes (MDS) and isolated del(5q). Section D of the Annex II and the RMP (version 39) are updated accordingly."	Request for supplementary information adopted with a specific timetable.

Request for Supplementary Information adopted on 08.06.2023.	
PRAC Led **Sialanar - glycopyrronium - EMEA/H/C/003883/II/0026** Proveca Pharma Limited, PRAC Rapporteur: Zane Neikena, PRAC-CHMP liaison: Elita Poplavska, "Submission of an updated RMP version 3.1 in order to remove a Drug Utilisation Study (DUS). In addition, the safety concerns, pharmacovigilance activities and risk minimisation measures in the RMP of Sialanar are revised to be in line with the Guideline on good pharmacovigilance practices (GVP) Module V – Risk management systems (Rev 2). Furthermore, Annex II D "Conditions or restrictions with regard to the safe and effective use of the medicinal product" was revised to delete information about the utilisation study from the key elements of the physician educational material." Opinion adopted on 08.06.2023. Request for Supplementary Information adopted on 12.05.2023, 16.03.2023, 12.01.2023.	Positive Opinion adopted by consensus on 08.06.2023.
PRAC Led **Vaxzevria - COVID 19 vaccine (ChAdOx1 S [recombinant]) - EMEA/H/C/005675/II/0091** AstraZeneca AB, Rapporteur: Sol Ruiz, PRAC Rapporteur: Jean-Michel Dogné, PRAC-CHMP liaison: Christophe Focke, "Submission of the final report from study D8111R00020 listed as a category 3 study in the RMP. This is a systematic literature review of observational studies evaluating safety after vaccination with Vaxzevria in patients taking immunosuppressant medications and/or with primary immunodeficiency." Opinion adopted on 08.06.2023.	Positive Opinion adopted by consensus on 08.06.2023.
PRAC Led **WS2483 Lixiana-EMEA/H/C/002629/WS2483/0045 Roteas-EMEA/H/C/004339/WS2483/0032** Daiichi Sankyo Europe GmbH, Lead PRAC Rapporteur: Nathalie Gault, PRAC-CHMP liaison: Alexandre Moreau, "Submission of the final report from DSE-EDO-04-14-EU (Non-Interventional Study on Edoxaban Treatment in Routine Clinical Practice for Patients with Non-	Positive Opinion adopted by consensus on 08.06.2023.

Valvular Atrial Fibrillation, ETNA-AF Europe), listed as a category 3 study in the RMP (MEA 006). This is a multicentre, prospective, non-interventional, observational PASS. The RMP version 15.1 has also been submitted."
Opinion adopted on 08.06.2023.

PRAC Led
WS2487
Humalog-
EMEA/H/C/000088/WS2487/0199
Liprolog-
EMEA/H/C/000393/WS2487/0159
Eli Lilly Nederland B.V., Lead PRAC Rapporteur: Mari Thorn, PRAC-CHMP liaison: Kristina Dunder, "To remove severe hypoglycemia as a result of incorrect or incomplete data provided to a compatible software application which is listed as an important potential risk for the Tempo Pen and all associated risk minimisation measures, following PRAC assessment of F3Z-MC-B030 PASS protocol (EMA/PRAC/781358/2022, 29 September 2022)."

B.5.5. CHMP-CAT assessed procedures

Abecma - idecabtagene vicleucel - EMEA/H/C/004662/II/0032/G, Orphan, ATMP
Bristol-Myers Squibb Pharma EEIG, Rapporteur: Rune Kjeken, CHMP Coordinator: Ingrid Wang
Request for Supplementary Information adopted on 17.05.2023.

Abecma - idecabtagene vicleucel - EMEA/H/C/004662/II/0034, Orphan, ATMP
Bristol-Myers Squibb Pharma EEIG, Rapporteur: Rune Kjeken, CHMP Coordinator: Ingrid Wang

Breyanzi - lisocabtagene maraleucel / lisocabtagene maraleucel - EMEA/H/C/004731/II/0013/G, ATMP
Bristol-Myers Squibb Pharma EEIG, Rapporteur: Concetta Quintarelli, CHMP Coordinator: Armando Genazzani
Opinion adopted on 16.06.2023.
Request for Supplementary Information adopted on 17.02.2023.

Breyanzi - lisocabtagene maraleucel /

lisocabtagene maraleucel -
EMEA/H/C/004731/II/0018/G, ATMP
Bristol-Myers Squibb Pharma EEIG, Rapporteur: Concetta Quintarelli, CHMP Coordinator: Armando Genazzani

Breyanzi - lisocabtagene maraleucel / lisocabtagene maraleucel -
EMEA/H/C/004731/II/0019, ATMP
Bristol-Myers Squibb Pharma EEIG, Rapporteur: Concetta Quintarelli, CHMP Coordinator: Armando Genazzani

CARVYKTI - ciltacabtagene autoleucel -
EMEA/H/C/005095/II/0016, Orphan, ATMP
Janssen-Cilag International NV, Rapporteur: Jan Mueller-Berghaus, CHMP Coordinator: Jan Mueller-Berghaus,
Opinion adopted on 16.06.2023.

Imlygic - talimogene laherparepvec -
EMEA/H/C/002771/II/0062/G, ATMP
Amgen Europe B.V., Rapporteur: Maija Tarkkanen, CHMP Coordinator: Johanna Lähteenvuo
Request for Supplementary Information adopted on 17.05.2023.

Kymriah - tisagenlecleucel -
EMEA/H/C/004090/II/0069, Orphan, ATMP
Novartis Europharm Limited, Rapporteur: Rune Kjeken, CHMP Coordinator: Ingrid Wang
Request for Supplementary Information adopted on 17.05.2023.

Kymriah - tisagenlecleucel -
EMEA/H/C/004090/II/0070/G, Orphan, ATMP
Novartis Europharm Limited, Rapporteur: Rune Kjeken, CHMP Coordinator: Ingrid Wang
Opinion adopted on 16.06.2023.

Libmeldy - atidarsagene autotemcel -
EMEA/H/C/005321/II/0015, Orphan, ATMP
Orchard Therapeutics (Netherlands) B.V., Rapporteur: Carla Herberts, CHMP Coordinator: Johann Lodewijk Hillege

B.5.6. CHMP-PRAC-CAT assessed procedures

B.5.7. PRAC assessed ATMP procedures

B.5.8. Unclassified procedures and worksharing procedures of type I variations

WS2399/G **Mirapexin-** **EMEA/H/C/000134/WS2399/0104/G** **Sifrol-** **EMEA/H/C/000133/WS2399/0095/G** Boehringer Ingelheim International GmbH, Lead Rapporteur: Thalia Marie Estrup Blicher Opinion adopted on 08.06.2023. Request for Supplementary Information adopted on 30.03.2023, 19.01.2023.	Positive Opinion adopted by consensus on 08.06.2023.
WS2412 **Hexacima-** **EMEA/H/C/002702/WS2412/0144** **Hexyon-** **EMEA/H/C/002796/WS2412/0148** Sanofi Pasteur, Lead Rapporteur: Jan Mueller-Berghaus Opinion adopted on 01.06.2023. Request for Supplementary Information adopted on 30.03.2023.	Positive Opinion adopted by consensus on 01.06.2023.
WS2427/G **Silodosin Recordati-** **EMEA/H/C/004964/WS2427/0011/G** **Silodyx-** **EMEA/H/C/001209/WS2427/0051/G** **Urorec-** **EMEA/H/C/001092/WS2427/0054/G** Recordati Ireland Ltd, Generic, Generic of Urorec, Lead Rapporteur: Margareta Bego Request for Supplementary Information adopted on 08.06.2023.	Request for supplementary information adopted with a specific timetable.
WS2445 **Ambirix-** **EMEA/H/C/000426/WS2445/0127** **Cervarix-** **EMEA/H/C/000721/WS2445/0122** **Fendrix-** **EMEA/H/C/000550/WS2445/0082** **Infanrix hexa-** **EMEA/H/C/000296/WS2445/0329** **Synflorix-** **EMEA/H/C/000973/WS2445/0180**	

Twinrix Adult- **EMEA/H/C/000112/WS2445/0162** **Twinrix Paediatric-** **EMEA/H/C/000129/WS2445/0163** GlaxoSmithkline Biologicals SA, Lead Rapporteur: Kristina Dunder	
WS2456/G **Infanrix hexa-** **EMEA/H/C/000296/WS2456/0328/G** GlaxoSmithkline Biologicals SA, Lead Rapporteur: Christophe Focke Opinion adopted on 01.06.2023.	Positive Opinion adopted by consensus on 01.06.2023.
WS2458 **Juluca-EMEA/H/C/004427/WS2458/0051** **Tivicay-EMEA/H/C/002753/WS2458/0087** **Triumeq-** **EMEA/H/C/002754/WS2458/0112** ViiV Healthcare B.V., Lead Rapporteur: Janet Koenig Opinion adopted on 01.06.2023.	Positive Opinion adopted by consensus on 01.06.2023.
WS2464 **Aerius-EMEA/H/C/000313/WS2464/0105** **Azomyr-** **EMEA/H/C/000310/WS2464/0109** **Neoclarityn-** **EMEA/H/C/000314/WS2464/0103** Organon N.V., Duplicate, Duplicate of Allex (SRD), Azomyr, Opulis (SRD), Lead Rapporteur: Christophe Focke, "C.I.7.a - To delete the pharmaceutical form orodispersible tablets from the marketing authorisation of: - Aerius from EU/1/00/160/037 up to and including EU/1/00/160/060 - Azomyr from EU/1/00/157/035 up to and including EU/1/00/157/058 - Neoclarityn from EU/1/00/161/035 up to and including EU/1/00/161/058" Opinion adopted on 08.06.2023.	Positive Opinion adopted by consensus on 08.06.2023.
WS2466 **Fluenz Tetra-** **EMEA/H/C/002617/WS2466/0128** **Pandemic influenza vaccine H5N1 AstraZeneca-** **EMEA/H/C/003963/WS2466/0063** AstraZeneca AB, Lead Rapporteur: Christophe Focke Opinion adopted on 01.06.2023.	Positive Opinion adopted by consensus on 01.06.2023.

WS2468/G **Hexacima-** **EMEA/H/C/002702/WS2468/0148/G** **Hexyon-** **EMEA/H/C/002796/WS2468/0152/G** Sanofi Pasteur, Lead Rapporteur: Jan Mueller-Berghaus	
WS2469 **Hexacima-** **EMEA/H/C/002702/WS2469/0147** **Hexyon-** **EMEA/H/C/002796/WS2469/0151** **MenQuadfi-** **EMEA/H/C/005084/WS2469/0023** Sanofi Pasteur Europe, Duplicate, Duplicate of Hexacima, Lead Rapporteur: Jan Mueller-Berghaus	
WS2472 **Afinitor-** **EMEA/H/C/001038/WS2472/0085** **Votubia-** **EMEA/H/C/002311/WS2472/0081** Novartis Europharm Limited, Lead Rapporteur: Janet Koenig Opinion adopted on 01.06.2023.	Positive Opinion adopted by consensus on 01.06.2023.
WS2473 **ProQuad-** **EMEA/H/C/000622/WS2473/0161** **Zostavax-** **EMEA/H/C/000674/WS2473/0146** Merck Sharp & Dohme B.V., Lead Rapporteur: Jan Mueller-Berghaus	
WS2484 **Filgrastim Hexal-** **EMEA/H/C/000918/WS2484/0071** **Zarzio-EMEA/H/C/000917/WS2484/0072** Sandoz GmbH, Lead Rapporteur: Johann Lodewijk Hillege	

B.5.9. Information on withdrawn type II variation / WS procedure

B.5.10. Information on type II variation / WS procedure with revised timetable

B.6. START OF THE PROCEDURES TIMETABLES FOR INFORMATION

B.6.1. Start of procedure for New Applications: timetables for information

liquid ethanolic extract 30 per cent (w/w) of allium cepa fresh bulb and citrus limon fresh fruit / dry aqueous extract of paullinia cupana seed / dry hydroethanolic extract of theobroma cacao seed - EMEA/H/C/004155 treatment of alopecia areata in children and adolescents
fruquintinib - EMEA/H/C/005979 treatment of metastatic colorectal cancer
meningococcal group a, b, c, w and y vaccine - EMEA/H/C/006165 indicated for active immunisation to prevent invasive disease caused by Neisseria meningitidis groups A, B, C, W, and Y
ustekinumab - EMEA/H/C/005918 treatment of adult patients with moderately to severely active Crohn's disease and active ulcerative colitis.

B.6.2. Start of procedure for Extension application according to Annex I of Reg. 1234/2008): timetables for information

Menveo - meningococcal group A, C, W135 and Y conjugate vaccine - EMEA/H/C/001095/X/0119 GSK Vaccines S.r.l, Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Menno van der Elst, "Extension application to introduce a new pharmaceutical form (solution for injection). The RMP (version 11.0) is updated in accordance."
Vyepti - eptinezumab - EMEA/H/C/005287/X/0011 H. Lundbeck A/S, Rapporteur: Jan Mueller-Berghaus, "Line extension application to add a new strength (300 mg concentrate for solution for infusion)."

XALKORI - crizotinib -
EMEA/H/C/002489/X/0080/G
Pfizer Europe MA EEIG, Rapporteur: Alexandre Moreau, PRAC Rapporteur: Tiphaine Vaillant, "Extension application to introduce a new pharmaceutical form (granules in capsules for opening) associated with new strengths (20, 50 and 150 mg), grouped with a type II variation (C.I.6.a) to include the treatment of paediatric patients with relapsed or refractory, systemic ALK-positive ALCL or unresectable, recurrent, or refractory ALK-positive IMT to change the lower end of the age range from >=6 years to ≥1 year for Xalkori following the assessment of II/0072 based on final results from study ADVL0912. As a consequence, sections 4.1, 4.2, 4.4, 4.5, 4.8, 5.1, 5.2 and 5.3 of the SmPC are updated. The Package Leaflet is updated in accordance. Version 9.1 of the RMP has also been submitted."

B.6.3. Restart of procedure - responses received to Day 120 List of Questions timetables: for information

B.6.4. Annual Re-assessments: timetables for adoption

EVKEEZA - evinacumab -
EMEA/H/C/005449/S/0010
Ultragenyx Germany GmbH, Rapporteur: Johann Lodewijk Hillege, Co-Rapporteur: Alar Irs, PRAC Rapporteur: Mari Thorn

B.6.5. Renewals of Marketing Authorisations: timetables for adoption provided only if the validation has been completed

GHRYVELIN - macimorelin -
EMEA/H/C/004660/R/0020
Consilient Health Limited, Rapporteur: Martina Weise, Co-Rapporteur: Jean-Michel Race (FR) (MNAT with FR for Clinical Efficacy, FR for Clinical Pharmacology, FR for Clinical Safety, FR for Non-Clinical, FR for Coordination, CZ for Quality), PRAC Rapporteur: Liana Gross-Martirosyan

Miglustat Dipharma - miglustat -
EMEA/H/C/004904/R/0019
DIPHARMA Arzneimittel GmbH, Generic, Generic of Zavesca, Rapporteur: Frantisek Drafi, PRAC Rapporteur: Mari Thorn

Mulpleo - lusutrombopag - EMEA/H/C/004720/R/0018
Shionogi B.V., Rapporteur: Daniela Philadelphy, Co-Rapporteur: Ewa Balkowiec Iskra, PRAC Rapporteur: Mari Thorn

Rizmoic - naldemedine - EMEA/H/C/004256/R/0023
Shionogi B.V., Rapporteur: Thalia Marie Estrup Blicher, Co-Rapporteur: Christophe Focke, PRAC Rapporteur: Rhea Fitzgerald

Skyrizi - risankizumab - EMEA/H/C/004759/R/0039
AbbVie Deutschland GmbH & Co. KG, Rapporteur: Finbarr Leacy, PRAC Rapporteur: Liana Gross-Martirosyan

Zirabev - bevacizumab - EMEA/H/C/004697/R/0029
Pfizer Europe MA EEIG, Rapporteur: Eva Skovlund, Co-Rapporteur: Alexandre Moreau, PRAC Rapporteur: Anette Kirstine Stark

B.6.6. VARIATIONS – START OF THE PROCEDURE

Timetables for adoption provided that the validation has been completed.

B.6.7. Type II Variations scope of the Variations: Extension of indication

CARVYKTI - ciltacabtagene autoleucel - EMEA/H/C/005095/II/0021, Orphan, ATMP
Janssen-Cilag International NV, Rapporteur: Jan Mueller-Berghaus, CHMP Coordinator: Jan Mueller-Berghaus, PRAC Rapporteur: Jo Robays, "Extension of indication to include treatment of adult patients with relapsed and refractory multiple myeloma, who have received at least 1 prior therapy, including an IMiD and a PI, have demonstrated disease progression on or after the last therapy and are refractory to lenalidomide for CARVYKTI, based on interim results from study MMY3002 listed as a specific obligation (SOB/006) in the Annex II. This is an ongoing, Phase 3, randomized, open-label, multicentre study to determine whether treatment with cilta-cel provides an efficacy benefit compared to standard therapy in participants with relapsed and lenalidomide-refractory multiple myeloma. As a consequence, sections 4.1, 4.4, 4.5, 4.8, 5.1 and 5.2 of the

SmPC are updated. The Package Leaflet is updated in accordance. Version 4.1 of the RMP has also been submitted. In addition, the MAH took the opportunity to update Annex II of the PI. As part of the application the MAH is requesting a 1-year extension of the market protection." Request for 1 year of market protection for a new indication (Article 14(11) of Regulation (EC) 726/2004)	
CellCept - mycophenolate mofetil - EMEA/H/C/000082/II/0170/G Roche Registration GmbH, Rapporteur: Thalia Marie Estrup Blicher, "C.I.6.a: Extension of indication to include paediatric patients (3 months to 18 years of age) for hepatic and cardiac transplants and to extend the indication for renal transplants for paediatric patients starting from 3 months, based on pharmacokinetic data, published literature and the Roche Global Safety Database. As a consequence, sections 4.1, 4.2, 4.8 and 5.2 of the SmPC are updated. The Package Leaflet is updated accordingly. Type IB (C.I.z): To update section 4.2 of the SmPC for the CellCept 500 mg tablets formulation in order to be in line with the other three CellCept formulations. And for alignment with the current QRD guidance, the Package Leaflet was updated to cross reference section 2 in section 6 for sodium content. In addition, the MAH took the opportunity to introduce minor editorial changes to the PI and bring the PI in line with the latest QRD template version 10.3."	
Cibinqo - abrocitinib - EMEA/H/C/005452/II/0010 Pfizer Europe MA EEIG, Rapporteur: Kristina Dunder, PRAC Rapporteur: Nikica Mirošević Skvrce, "Extension of indication to include treatment of adolescents 12 to < 18 years of age with moderate to severe atopic dermatitis for CIBINQO based on final results from non-clinical study 00655292 [21GR211] and interim results from clinical study B7451015; this is a Phase III multi-center, long-term extension study investigating the efficacy and safety of abrocitinib, with or without topical medications, administered to subjects aged 12 years and	

older with moderate to severe atopic dermatitis. As a consequence, sections 4.1, 4.2, 4.8 and 5.1 of the SmPC are updated. The Package Leaflet is updated in accordance. Version 4.1 of the RMP has also been submitted."

EVKEEZA - evinacumab - EMEA/H/C/005449/II/0011

Ultragenyx Germany GmbH, Rapporteur: Johann Lodewijk Hillege, Co-Rapporteur: Alar Irs, PRAC Rapporteur: Mari Thorn, "Extension of indication to include the treatment of paediatric patients with homozygous familial hypercholesterolaemia (HoFH) aged 5 years and older for EVKEEZA, based on interim results from study R1500-CL-17100, as well as supportive information from an updated interim analysis of study R1500-CL-1719, and an extrapolation analysis (including population PK, population PK/PD, and simulation analyses). R1500-CL-17100 is an ongoing multicentre, three-part, single-arm, open-label study evaluating the efficacy, safety, and tolerability of evinacumab in paediatric patients aged ≥ 5 to 11 years with HoFH. As a consequence, sections 4.1, 4.2, 4.8, 5.1 and 5.2 of the SmPC are updated. The Package Leaflet is updated in accordance. Version 1.1 of the RMP has also been submitted. In addition, the marketing authorisation holder took the opportunity to introduce minor editorial changes to the PI. Furthermore, the PI is brought in line with the latest QRD template version 10.3."

Fexinidazole Winthrop - fexinidazole - EMEA/H/W/002320/II/0016

Sanofi Winthrop Industrie, Rapporteur: Fátima Ventura, PRAC Rapporteur: Liana Gross-Martirosyan, "Extension of indication to include treatment of both first stage (haemo-lymphatic) and second stage (meningo-encephalitic) of human African trypanosomiasis (HAT) due to Trypanosoma brucei rhodesiense for FEXINIDAZOLE WINTHROP based final results from study DNDI-FEX-07-HAT - Efficacy and safety of fexinidazole in patients with Human African Trypanosomiasis (HAT) due to Trypanosoma brucei rhodesiense: a multicentre, open-label clinical trial; this is a phase-II/III, multicenter, open-label, non-randomized, single-arm clinical trial to assess the efficacy

and safety of fexinidazole in patients with r-HAT. The primary objective of study DNDi-FEX-07-HAT, for which the final results are presented in this application, is: To show that the fatality rate (r HAT or treatment related death) at EOH in stage-2 patients treated with fexinidazole is smaller than a threshold of unacceptable rate of 8.5%. It is noteworthy that the primary objective of study DNDi-FEX-07-HAT is related to a safety parameter. As a consequence, sections 4.1, 4.4, 4.8, 5.1 and 5.2 of the SmPC are updated. The Package Leaflet is updated in accordance. Version 3.0 of the RMP has also been submitted."

Keytruda - pembrolizumab - EMEA/H/C/003820/II/0138

Merck Sharp & Dohme B.V., Rapporteur: Armando Genazzani, PRAC Rapporteur: Menno van der Elst, "Extension of indication to include KEYTRUDA in combination with gemcitabine-based chemotherapy for the first-line treatment of locally advanced unresectable or metastatic biliary tract carcinoma in adults, based on final results from study KEYNOTE-966; this is a Phase 3 randomized, double blind study of Pembrolizumab plus Gemcitabine/Cisplatin versus Placebo plus Gemcitabine/Cisplatin as first-line therapy in participants with advanced and/or unresectable biliary tract carcinoma. As a consequence, sections 4.1, 4.4 and 5.1 of the SmPC are updated. The Package Leaflet is updated in accordance. Version 43.1 of the RMP has also been submitted."

VeraSeal - human fibrinogen / human thrombin - EMEA/H/C/004446/II/0027

Instituto Grifols, S.A., Rapporteur: Daniela Philadelphy, Co-Rapporteur: Ewa Balkowiec Iskra, PRAC Rapporteur: Amelia Cupelli, "Extension of indication to include treatment of children for VeraSeal, based on final results from study IG1405; this is a prospective, randomized, active-controlled, single-blind, parallel group clinical trial to evaluate the safety and efficacy of VeraSeal as an adjunct to haemostasis during surgery in paediatric subjects. As a consequence, sections 4.1, 4.2 and 5.1 of the SmPC are updated. The Package Leaflet is updated in accordance. Version 6.0 of the RMP has also been submitted."

B.6.8. CHMP assessed procedures scope: Pharmaceutical aspects

COMIRNATY - COVID-19 mRNA vaccine (nucleoside-modified) - EMEA/H/C/005735/II/0182/G
BioNTech Manufacturing GmbH, Rapporteur:
Filip Josephson

B.6.9. CHMP assessed procedures scope: Non-Clinical and Clinical aspects

B.6.10. PRAC assessed procedures

B.6.11. CHMP-CAT assessed procedures

B.6.12. CHMP-PRAC-CAT assessed procedures

B.6.13. PRAC assessed ATMP procedures

B.6.14. Unclassified procedures and worksharing procedures of type I variations

B.7. DOCUMENTS TABLED IN MMD AFTER THE CHMP PLENARY

B.7.1. Yearly Line listing for Type I and II variations

B.7.2. Monthly Line listing for Type I variations

B.7.3. Opinion on Marketing Authorisation transfer (MMD only)

B.7.4. Notifications in accordance with Article 61(3) of Council Directive 2001/83/EC (MMD only)

B.7.5. Request for supplementary information relating to Notification of Type I variation (MMD only)

B.7.6. Notifications of Type I Variations (MMD only)

C. Annex C - Post-Authorisation Measures (PAMs), (Line listing of Post authorisation measures with a description of the PAM. Procedures starting in that given month with assessment timetabled)

D. Annex D - Post-Authorisation Measures (PAMs), (Details on PAMs including description and conclusion, for adoption by CHMP in that given month, or finalised ones with PRAC recommendation and no adoption by CHMP needed)

E. Annex E - EMA CERTIFICATION OF PLASMA MASTER FILES

Information related to plasma master files cannot be released at the present time as these contain commercially confidential information.

E.1. PMF Certification Dossiers:

E.1.1. Annual Update

E.1.2. Variations:

E.1.3. Initial PMF Certification:

E.2. Time Tables – starting & ongoing procedures: For information

PMF timetables starting and ongoing procedures	Tabled in MMD and sent by post mail (folder E).

F. ANNEX F - Decision of the Granting of a Fee Reduction/Fee Waiver

F.1. Parallel Distribution - Pursuant to Article 9 of Council Regulation (EC) No. 2743/98 of 14 December 1998, as amended

F.2. Request for scientific opinion on justification of exceptional circumstance and for imperative grounds of public health

G. ANNEX G

G.1. Final Scientific Advice (Reports and Scientific Advice letters):

Information related to Scientific Advice cannot be released at the present time as these contain commercially confidential information.

G.2. PRIME

Some information related to PRIME cannot be released at the present time as these contain commercially confidential information.

G.2.1. List of procedures concluding at 19-22 June 2023 CHMP plenary

G.2.2. List of procedures starting in June 2023 for July 2023 CHMP adoption of outcomes

H. ANNEX H - Product Shared Mailboxes – e-mail address