24 February 2020
EMA/CHMP/92703/2020
Inspections, Human Medicines Pharmacovigilance and Committees Division

Committee for medicinal products for human use (CHMP)

Agenda for the meeting on 24-27 February 2020

Chair: Harald Enzmann – Vice-Chair: Bruno Sepodes

24 February 2020, 13:00 – 19:30, room 1C

25 February 2020, 08:30 – 19:30, room 1C

26 February 2020, 08:30 – 19:30, room 1C

27 February 2020, 08:30 – 16:00, room 1C

Health and safety information

In accordance with the Agency's health and safety policy, delegates are to be briefed on health, safety and emergency information and procedures prior to the start of the meeting.

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 24-27 February 2020. See February 2020 CHMP minutes (to be published post March 2020 CHMP meeting).

1.2. Adoption of agenda

CHMP agenda for 24-27 February 2020.

1.3. Adoption of the minutes

CHMP minutes for 27-30 January 2020.

CHMP ORGAM minutes for 20 January 2020.

2. Oral Explanations

2.1. Pre-authorisation procedure oral explanations

2.1.1. rituximab - EMEA/H/C/005387

treatment of Non-Hodgkin's lymphoma (NHL), Chronic lymphocytic leukaemia (CLL)

Scope: Oral explanation

Action: Oral explanation to be held on Wednesday, 26 February 2020 at 09:00

List of Outstanding Issues adopted on 12.12.2019, 27.06.2019.

2.1.2. alpelisib - EMEA/H/C/004804

treatment of postmenopausal women and men, with hormone receptor (HR)-positive, human epidermal growth factor receptor2 (HER2)-negative, advanced breast cancer with a PIK3CA mutation in combination with fulvestrant after disease progression following an endocrine-based regimen.

Scope: Oral explanation

Action: Oral explanation to be held on Wednesday, 26 February 2020 at 15:30

List of Outstanding Issues adopted on 19.09.2019. List of Questions adopted on 29.05.2019.

2.1.3. rituximab - EMEA/H/C/004807

treatment of Non-Hodgkin's lymphoma (NHL), Chronic lymphocytic leukaemia (CLL) and Rheumatoid arthritis

Scope: Oral explanation

Action: Oral explanation to be held on Wednesday, 26 February 2020 at 09:00

List of Outstanding Issues adopted on 12.12.2019, 27.06.2019. List of Questions adopted on 18.10.2018.

2.2. Re-examination procedure oral explanations

2.2.1. Hopveus - sodium oxybate - EMEA/H/C/004962

D&A PHARMA; for the treatment of alcohol dependence

Scope: Oral explanation, Report from ad-hoc expert group meeting held on 11 February 2020

Action: Oral explanation to be held on Tuesday, 25 February 2020 at 14:00

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

Opinion adopted on 17.10.2019. List of Outstanding Issues adopted on 26.04.2019. List of Questions adopted on 15.11.2018.

See 3.5

2.3. Post-authorisation procedure oral explanations

2.3.1. WS1683, Elebrato Ellipta-EMEA/H/C/004781/WS1683/0012, Temybric Ellipta-EMEA/H/C/005254/WS1683/0001, Trelegy Ellipta-EMEA/H/C/004363/WS1683/0010

GlaxoSmithKline Trading Services Limited

Lead Rapporteur: Peter Kiely

Scope: "Update of SmPC in order to add information in section 5.1 on survival data from the IMPACT study"

Oral explanation

Action: Oral explanation to be held on Tuesday, 25 February 2020 at 16:00

Request for Supplementary Information adopted on 12.12.2019, 17.10.2019.

2.4. Referral procedure oral explanations

2.4.1. Ranitidine - EMEA/H/A-31/1491

MAH various

Rapporteur: Johann Lodewijk Hillege, Co-Rapporteur: Daniela Melchiorri

Scope: Oral explanations

Action: Oral explanations to be held on Tuesday, 25 February 2020 at 09:00, 10:00 and 11:00

Tests performed in a random selection of ranitidine API batches and finished products available in the EU have shown levels of NDMA which raise concerns.

European Commission triggered on 12 September 2019 a referral procedure under Article 31 of Directive 2001/83/EC to evaluate the relevance of these findings, the potential root causes and their impact on the benefit-risk balance of medicinal products containing ranitidine.

See 10.6

3. Initial applications

3.1. Initial applications; Opinions

3.1.1. cefiderocol - EMEA/H/C/004829

treatment of infections due to aerobic Gram-negative bacteria

Scope: Opinion

Action: For adoption

List of Outstanding Issues adopted on 14.11.2019. List of Questions adopted on 25.06.2019.

3.1.2. tigecycline - EMEA/H/C/005114

Treatment of soft tissue and intra-abdominal infections:
- complicated skin and soft tissue infections, excluding diabetic foot infections;
- complicated intra-abdominal infections.
should be used only in situations where it is known or suspected that other alternatives are not suitable

Scope: Opinion

Action: For adoption

List of Outstanding Issues adopted on 12.12.2019, 19.09.2019, 29.05.2019. List of Questions adopted on 13.12.2018.

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

3.2.1. cabazitaxel - EMEA/H/C/005178

treatment of adult patients with metastatic castration resistant prostate cancer previously treated with a docetaxel-containing regimen

Scope: List of outstanding issues

Action: For adoption

List of Questions adopted on 19.09.2019.

3.2.2. doxorubicin - EMEA/H/C/005194

treatment of breast cancer, ovarian cancer, progressive multiple myeloma and AIDS-related Kaposi's sarcoma

Scope: List of outstanding issues

Action: For adoption

List of Outstanding Issues adopted on 12.12.2019. List of Questions adopted on 19.09.2019.

3.2.3. glasdegib - Orphan - EMEA/H/C/004878

Pfizer Europe MA EEIG; treatment of newly diagnosed de novo or secondary acute myeloid leukaemia

Scope: List of outstanding issues

Action: For adoption

List of Questions adopted on 19.09.2019.

3.2.4. fingolimod - EMEA/H/C/005191

treatment of multiple sclerosis

Scope: List of outstanding issues

Action: For adoption

List of Questions adopted on 19.09.2019.

3.2.5. fingolimod - EMEA/H/C/005282

treatment of multiple sclerosis

Scope: List of outstanding issues

Action: For adoption

List of Questions adopted on 19.09.2019.

3.2.6. influenza vaccine (surface antigen, inactivated) - Article 28 - EMEA/H/C/004993

Active immunisation against influenza in the elderly (65 years of age and older) and in children 6 months to less than 6 years of age.

Scope: List of outstanding issues

Action: For adoption

List of Outstanding Issues adopted on 12.12.2019. List of Questions adopted on 25.07.2019.

3.2.7. imlifidase - Orphan - EMEA/H/C/004849

Hansa Biopharma AB; indicated for desensitization treatment of highly sensitized adult kidney transplant patients with positive crossmatch against an available deceased donor.

Scope: List of outstanding issues

Action: For adoption

List of Questions adopted on 27.06.2019.

3.2.8. insulin aspart - EMEA/H/C/005033

treatment of diabetes mellitus

Scope: List of outstanding issues

Action: For adoption

List of Questions adopted on 17.10.2019.

3.2.9. teriparatide - EMEA/H/C/005087

treatment of osteoporosis

Scope: List of outstanding issues

Action: For adoption

List of Questions adopted on 19.09.2019.

3.2.10. teriparatide - EMEA/H/C/005388

treatment of osteoporosis

Scope: List of outstanding issues

Action: For adoption

List of Questions adopted on 19.09.2019.

3.2.11. luspatercept - Orphan - EMEA/H/C/004444

Celgene Europe BV; - treatment of adult patients with very low- to intermediate-risk myelodysplastic syndromes (MDS)-associated anaemia; - the treatment of adult patients

with beta-thalassaemia (β-thalassaemia)-associated anaemia who require RBC transfusions.

Scope: List of outstanding issues

Action: For adoption

List of Questions adopted on 19.09.2019.

3.2.12. isatuximab - Orphan - EMEA/H/C/004977

sanofi-aventis groupe; For the treatment of patients with multiple myeloma (MM)

Scope: List of outstanding issues

Action: For adoption

List of Outstanding Issues adopted on 12.12.2019. List of Questions adopted on 19.09.2019.

3.2.13. ivosidenib - Orphan - EMEA/H/C/005056

Agios Netherlands B.V.; Treatment of adult patients (≥ 18 years old) with relapsed or refractory acute myeloid leukaemia (AML) with an isocitrate dehydrogenase-1 (IDH1) R132 mutation

Scope: List of outstanding issues

Action: For adoption

List of Questions adopted on 29.05.2019.

3.2.14. bupivacaine / meloxicam - EMEA/H/C/005205

for application into the surgical site to reduce postoperative pain

Scope: List of outstanding issues

Action: For adoption

List of Questions adopted on 25.07.2019.

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

3.3.1. acalabrutinib - Orphan - EMEA/H/C/005299

AstraZeneca AB; Treatment of adult patients with chronic lymphocytic leukaemia (CLL)/small lymphocytic lymphoma (SLL)

Scope: List of questions

Action: For adoption

3.3.2. insulin aspart - EMEA/H/C/004965

treatment of diabetes mellitus

Scope: List of questions

Action: For adoption

3.3.3. meningococcal group a, c, w135 and y conjugate vaccine - Article 28 - EMEA/H/C/005084

immunization against Neisseria meningitidis serogroups A, C, W-135 and Y

Scope: List of questions

Action: For adoption

3.3.4. ebola vaccine (rdna, replication-incompetent) - EMEA/H/C/005343

Accelerated assessment

is indicated for active immunization for prevention of disease caused by Ebola virus

Scope: List of questions

Action: For adoption

3.3.5. caffeine citrate - EMEA/H/C/005435

treatment of primary apnoea

Scope: List of questions

Action: For adoption

3.3.6. salmeterol xinafoate / fluticasone propionate - EMEA/H/C/004881

for treatment of asthma

Scope: List of questions

Action: For adoption

3.3.7. sunitinib - EMEA/H/C/005419

treatment of gastrointestinal stromal tumour (GIST) and metastatic renal cell carcinoma (MRCC) and pancreatic neuroendocrine tumours (pNET)

Scope: List of questions

Action: For adoption

3.3.8. influenza quadrivalent vaccine (rDNA) - EMEA/H/C/005159

prevention of influenza disease

Scope: List of questions

Action: For adoption

3.3.9. ebola vaccine (rdna, replication-incompetent) - EMEA/H/C/005337

Accelerated assessment

is indicated for active immunization for prevention of disease caused by Ebola virus (Zaire ebolavirus species)

Scope: List of questions

Action: For adoption

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

3.4.1. tagraxofusp - Orphan - EMEA/H/C/005031

TMC Pharma (EU) Limited; treatment of adult patients with blastic plasmacytoid dendritic cell neoplasm (BPDCN)

Scope: Draft list of questions for the SAG meeting

Action: For adoption

List of Outstanding Issues adopted on 25.06.2019. List of Questions adopted on 24.04.2019.

3.4.2. elexacaftor / tezacaftor / ivacaftor - Orphan - EMEA/H/C/005269

Vertex Pharmaceuticals (Ireland) Limited; treatment of cystic fibrosis

Scope: Letter from third party, draft response letter

Action: For information

List of Questions adopted on 30.01.2020.

3.4.3. arachis hypogaea allergens - EMEA/H/C/004917

immunotherapy (OIT) for patients aged 4 to 17 years with a confirmed diagnosis of peanut allergy

Scope: Letter from the applicant dated 17 February 2020 requesting an extension of clock-stop to respond to the list of questions adopted in November 2019.

List of Questions adopted on 14.11.2019

3.4.4. pexidartinib - Orphan - EMEA/H/C/004832

Daiichi Sankyo Europe GmbH; treatment of adult patients with symptomatic tenosynovial giant cell tumour (TGCT), also referred to as giant cell tumour of the tendon sheath (GCT-TS) or pigmented villonodular synovitis (PVNS)

Scope: Draft list of experts for SAG Oncology meeting

Action: For adoption

List of Outstanding Issues adopted on 12.12.2019. List of Questions adopted on 25.07.2019.

3.4.5. trastuzumab - EMEA/H/C/005066

treatment of metastatic and early breast cancer and metastatic gastric cancer (MGC)

Scope: Letter from the applicant dated 04 February 2020 requesting an extension of clock-stop to respond to the list of questions adopted in September 2019.

Action: For adoption

List of Questions adopted on 19.09.2019.

3.4.6. ozanimod - EMEA/H/C/004835

Treatment of multiple sclerosis

Scope: List of experts for the SAG Neurology meeting

Action: For adoption

List of Outstanding Issues adopted on 12.12.2019. List of Questions adopted on 25.07.2019.

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

3.5.1. Hopveus - sodium oxybate - EMEA/H/C/004962

D&A PHARMA; for the treatment of alcohol dependence

Scope: Oral explanation, Report from ad-hoc expert group meeting held on 11 February 2020

Action: For adoption

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

Opinion adopted on 17.10.2019. List of Outstanding Issues adopted on 26.04.2019. List of Questions adopted on 15.11.2018.

See 2.2

3.6. Initial applications in the decision-making phase

No items

3.7. Withdrawals of initial marketing authorisation application

No items

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. Emgality - galcanezumab - EMEA/H/C/004648/X/0004

Eli Lilly Nederland B.V.

Rapporteur: Daniela Melchiorri (IT) (MNAT with ES for Quality), Co-Rapporteur: Kristina Dunder, PRAC Rapporteur: Kirsti Villikka

Scope: "Extension application to add a new strength of 100 mg/ml solution for injection in pre-filled syringe for Emgality, associated with a new indication (episodic cluster headache)."

Action: For adoption

List of Outstanding Issues adopted on 17.10.2019. List of Questions adopted on 27.06.2019.

4.1.2. Entyvio - vedolizumab - EMEA/H/C/002782/X/0040

Takeda Pharma A/S

Rapporteur: Daniela Melchiorri, Co-Rapporteur: Ewa Balkowiec Iskra, PRAC Rapporteur: Adam Przybylkowski

Scope: "Extension application to introduce a new pharmaceutical form (solution for injection), associated with a new strength (108 mg) and a new route of administration (subcutaneous use)."

Action: For adoption

List of Outstanding Issues adopted on 12.12.2019. List of Questions adopted on 25.07.2019.

4.1.3. Suboxone - buprenorphine / naloxone - EMEA/H/C/000697/X/0042

Indivior Europe Limited

Rapporteur: Janet Koenig, PRAC Rapporteur: Martin Huber

Scope: "Extension application to introduce a new pharmaceutical form (sublingual film) associated with four new strengths (2/0.5, 4/1, 8/2, and 16/4 mg) and a new route of administration (either sublingual or buccal administration)"

Action: For adoption

List of Questions adopted on 25.07.2019.

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

4.2.1. Harvoni - ledipasvir / sofosbuvir - EMEA/H/C/003850/X/0081/G

Gilead Sciences Ireland UC

Rapporteur: Filip Josephson, PRAC Rapporteur: Ana Sofia Diniz Martins

Scope: "Extension application to introduce a new strength (45/200 mg film-coated tablets) and a new pharmaceutical form (oral granules) associated with new strengths (33.75/150 and 45/200 mg). The new presentations are indicated in combination with other medicinal products for the treatment of chronic hepatitis C (CHC) in patients aged 3 to < 12 years. The extension application is grouped with a type II variation (C.I.6.a) to include paediatric use in patients aged 3 to < 12 years who weigh greater than or equal to 35 kg to the existing presentations of 90/400 mg film-coated tablets. The RMP (version 6.1) is updated in accordance. In addition, the MAH took the opportunity to implement minor linguistic corrections throughout the Product Information."

Action: For adoption

List of Questions adopted on 17.10.2019.

4.2.2. Sovaldi - sofosbuvir - EMEA/H/C/002798/X/0059/G

Gilead Sciences Ireland UC

Rapporteur: Filip Josephson, PRAC Rapporteur: Ana Sofia Diniz Martins

Scope: "Extension application to introduce a new strength (200 mg film-coated tablets) and a new pharmaceutical form (oral granules) associated with new strengths (150 and 200 mg). The new presentations are indicated in combination with other medicinal products for the treatment of chronic hepatitis C (CHC) in patients aged 3 to < 12 years. The extension application is grouped with a type II variation (C.I.6.a) to include paediatric use in patients aged 3 to < 12 years who weigh greater than or equal to 35 kg to the existing presentations of 400 mg film-coated tablets. The RMP (version 9.1) is updated in accordance. In addition, the MAH took the opportunity to implement minor linguistic corrections throughout the Product Information."

Action: For adoption

List of Questions adopted on 17.10.2019.

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. Epclusa - sofosbuvir / velpatasvir - EMEA/H/C/004210/X/0043/G

Gilead Sciences Ireland UC

Rapporteur: Filip Josephson, PRAC Rapporteur: Ana Sofia Diniz Martins

Scope: "Extension application to introduce a new strength (200/50 mg film-coated tablets). The new formulation is indicated for the treatment of chronic hepatitis C (CHC) in patients aged 6 years and older.
The extension application is grouped with a type II variation (C.I.6.a) to include paediatric use in patients aged 6 to < 18 years who weigh greater than or equal to 35 kg to the existing presentation (400/100 mg film-coated tablets).
Sections 4.1, 4.2, 4.8, 5.1 and 5.2 of the SmPC and the Package Leaflet are updated to support the extended indication. The RMP (version 5.1) is updated in accordance."

Action: For adoption

4.3.2. Pemetrexed Hospira - pemetrexed - EMEA/H/C/003970/X/0021

Pfizer Europe MA EEIG

Rapporteur: Alar Irs

Scope: "Extension application to introduce a new pharmaceutical form with its associated strength (25 mg/ml concentrate for solution for infusion)."

Action: For adoption

4.3.3. Pradaxa - dabigatran etexilate - EMEA/H/C/000829/X/0122/G

Boehringer Ingelheim International GmbH

Rapporteur: Mark Ainsworth, Co-Rapporteur: Jean-Michel Race, PRAC Rapporteur: Anette Kirstine Stark

Scope: "Extension application to add two new pharmaceutical forms for Pradaxa (coated granules (20 mg, 30 mg, 40 mg, 50 mg, 110 mg, 150 mg) and powder and solvent for oral solution (6.25 mg/ml)), grouped with:
-A type II variation (C.I.6.a) - Extension of indication to include new indication for Pradaxa 75 mg, 110 mg, 150 mg capsules based on the paediatric trials 1160.106 and 1160.108. As a consequence, sections 4.1, 4.2, 4.3, 4.4, 4.5, 4.7, 4.8, 4.9, 5.1, 5.2 and 5.3 of the SmPC are updated. The Package Leaflet and Labelling are updated in accordance. In addition, the Marketing authorisation holder (MAH) took the opportunity to update the list of local representatives in the Package Leaflet. The RMP version 37.0 has also been submitted.
-Type IB (B.I.b.1.c)
-Type IA (B.I.b.1.b)
-Type IB (B.I.b.1.d)
-Type IA (B.I.b.2.a)
-Type IA (B.I.b.1.d)
-Type IA (B.I.d.1.a.1)
-Type IA (B.II.d.1.a)
-Type IB (B.II.d.1.d)
-Type IA (B.II.d.2.a)
-Type IA (B.II.c.1.c)."

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. Alunbrig - brigatinib - EMEA/H/C/004248/II/0003

Takeda Pharma A/S

Rapporteur: Sinan B. Sarac, PRAC Rapporteur: Marcia Sofia Sanches de Castro Lopes Silva

Scope: "Extension of indication to include first-line treatment of adult patients with anaplastic lymphoma kinase (ALK)-positive advanced non-small cell lung cancer (NSCLC) previously untreated with an ALK inhibitor for Alunbrig.
The addition of a new indication is supported by data from Study 301 (AP26113-13-301; ALTA 1L), a phase 3, randomized, open label, comparative, multicenter, international phase 3 study. As a consequence, sections 4.1, 4.8 and 5.1 of the SmPC are updated. The Package leaflet and labelling are updated in accordance. The RMP version 5.1 has also been submitted. Minor editorial corrections are also proposed." 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 19.09.2019.

5.1.2. Carmustine Obvius - carmustine - EMEA/H/C/004326/II/0002

Obvius Investment B.V

Rapporteur: Natalja Karpova, Co-Rapporteur: Andrea Laslop, PRAC Rapporteur: Jan Neuhauser

Scope: "carmustine with or without total body irradiation (TBI), as conditioning treatment prior to allogeneic or autologous haematopoietic progenitor cell transplantation (HPCT) in haematological diseases"

Action: For adoption

Request for Supplementary Information adopted on 17.10.2019.

5.1.3. Cosentyx - secukinumab - EMEA/H/C/003729/II/0057

Novartis Europharm Limited

Rapporteur: Tuomo Lapveteläinen, PRAC Rapporteur: Eva A. Segovia

Scope: "Extension of indication to include the treatment of moderate to severe plaque psoriasis in children and adolescents from the age of 6 years who are candidates for systemic therapy for Cosentyx; as a consequence, sections 4.1, 4.2, 4.4, 4.5, 4.8, 5.1 and 5.2 of the SmPC are updated. Section 6.6 of the SmPC for the solution for injection is also updated. The Package Leaflet is updated in accordance. In addition, the Marketing authorisation holder (MAH) took the opportunity to update the list of local representatives in the Package Leaflet. The RMP version 6.0 has also been submitted.
Furthermore, the Annex II is brought in line with the latest QRD template version 11.0."

Action: For adoption

5.1.4. Deltyba - delamanid - Orphan - EMEA/H/C/002552/II/0040

Otsuka Novel Products GmbH

Rapporteur: Koenraad Norga, PRAC Rapporteur: Jean-Michel Dogné

Scope: "Extension of indication to include adolescents and children above 6 years with a body weight of at least 30 kg. As a consequence, sections 4.1, 4.2, 5.1 and 5.2 of the SmPC and corresponding relevant sections of the PL are updated accordingly. The updated RMP version 3.2 has also been submitted. Furthermore, the PI is being brought in line with the latest QRD template."

Action: For adoption

5.1.5. Imfinzi - durvalumab - EMEA/H/C/004771/II/0014/G

AstraZeneca AB

Rapporteur: Sinan B. Sarac, Co-Rapporteur: Jorge Camarero Jiménez, PRAC Rapporteur: David Olsen

Scope: "Extension of indication to include the use of Imfinzi in combination with etoposide and either carboplatin or cisplatin for the first-line treatment of adults with extensive-stage small cell lung cancer (ES-SCLC). The proposed indication is supported by study D419QC00001 (CASPIAN), an ongoing Phase III randomised, multicentre, open-label, comparative study designed to determine the efficacy and safety of durvalumab, or durvalumab and tremelimumab, in combination with etoposide and platinum-based chemotherapy (EP) for the first-line treatment of patients with ES-SCLC.
In addition, the MAH proposes to update sections 4.4 and 4.8 of the SmPC to update the safety information based on the Durvalumab Pan-Tumour Pool, a safety dataset comprising of 9 clinical studies building on the existing safety database and summarising the safety information for durvalumab monotherapy characterised across tumour types in the durvalumab clinical program to date.
The Package Leaflet is updated in accordance. The RMP version 2.1 has also been

submitted."

Action: For adoption

5.1.6. Invokana - canagliflozin - EMEA/H/C/002649/II/0046

Janssen-Cilag International NV

Rapporteur: Martina Weise, Co-Rapporteur: Kristina Dunder, PRAC Rapporteur: Martin Huber

Scope: "Update of sections 4.1 , 4.2, 4.8 and 5.1 of the Summary of Product Characteristics to add a new therapeutic indication for Invokana (canagliflozin) for the treatment of stage 2 or 3 chronic kidney disease and albuminuria, as an adjunct to standard of care, in adults with type 2 diabetes mellitus. The proposed new indication is based upon new clinical efficacy and safety data from the Phase 3 study: Canagliflozin and Renal Events in Diabetes with Established Nephropathy Clinical Evaluation Trial (CREDENCE) (DNE3001).
The Package Leaflet is updated in accordance. The RMP version 8.1 has also been submitted. In addition, the Marketing authorisation holder (MAH) took the opportunity 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 14.11.2019.

5.1.7. Keytruda - pembrolizumab - EMEA/H/C/003820/II/0057

Merck Sharp & Dohme B.V.

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

Scope: "Extension of indication to include 1st line treatment of locally advanced or metastatic non-small cell lung cancer tumours expressing PD-L1 with a ≥ 1% tumour proportion score (TPS), based on data from study KEYNOTE-042; an international, randomized, open-label Phase 3 study investigating KEYTRUDA monotherapy compared to standard of care platinum-based chemotherapy in patients with locally advanced or metastatic PD-L1 positive (TPS ≥ 1%) NSCLC, and on supportive data from the final planned analysis of KEYNOTE-024; a Phase 3 randomized open-label study of KEYTRUDA monotherapy compared to platinum-based chemotherapy in metastatic NSCLC with PD-L1 TPS ≥50%. As a result, sections 4.1, 4.4, 4.8 and 5.1 of the SmPC have been updated. An updated RMP version 18.1 was provided as part of the application."

Action: For adoption

Request for Supplementary Information adopted on 19.09.2019, 28.03.2019, 18.10.2018.

5.1.8. OFEV - nintedanib - Orphan - EMEA/H/C/003821/II/0026

Boehringer Ingelheim International GmbH

Rapporteur: Peter Kiely, Co-Rapporteur: Ewa Balkowiec Iskra, PRAC Rapporteur: Nikica Mirošević Skvrce

Scope: "Extension of indication to include new indication for OFEV for the treatment of Systemic Sclerosis associated Interstitial Lung Disease (SSc-ILD).
As a consequence, sections 4.1, 4.2, 4.3, 4.4, 4.5, 4.6, 4.8, 5.1 and 5.2 of the SmPC are updated. The Package Leaflet is updated in accordance. In addition, the Marketing authorisation holder (MAH) took the opportunity to update the list of local representatives in the Package Leaflet. The MAH takes this opportunity to also introduce minor linguistic corrections to the Annexes for France and Sweden. The RMP version 7.0 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 14.11.2019, 27.06.2019.

5.1.9. Otezla - apremilast - EMEA/H/C/003746/II/0029

Celgene Europe BV

Rapporteur: Peter Kiely, Co-Rapporteur: Janet Koenig, PRAC Rapporteur: Eva A. Segovia

Scope: "Extension of indication to include treatment of adult patients with oral ulcers associated with Behçet's disease (BD) who are candidates for systemic therapy. As a consequence, sections 4.1, 4.2, 4.8 and 5.1 of the SmPC and sections 1, 2 and 4 of the PL are updated accordingly. The updated RMP version 12.0 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 17.10.2019.

5.1.10. Sivextro - tedizolid phosphate - EMEA/H/C/002846/II/0035

Merck Sharp & Dohme B.V.

Rapporteur: Bruno Sepodes, PRAC Rapporteur: Maria del Pilar Rayon

Scope: "Extension of indication (treatment of ABSSSI in adults) to include adolescent population from 12 years old and older for Sivextro; as a consequence, sections 4.1, 4.2, 4.8 and 5.2 of the SmPC are updated. Sections 1 and 2 of the Package Leaflet are updated in accordance. The updated RMP version 5.1 has also been submitted. In addition, the (MAH) took the opportunity to bring the PI in line with the latest QRD template version 10.1"

Action: For adoption

5.1.11. Vokanamet - canagliflozin / metformin - EMEA/H/C/002656/II/0051

Janssen-Cilag International NV

Rapporteur: Martina Weise, PRAC Rapporteur: Menno van der Elst

Scope: "Update of sections 4.1 , 4.2, 4.8 and 5.1 of the Summary of Product Characteristics to add a new therapeutic indication for Vokanamet (canagliflozin/metformin) for the treatment of stage 2 or 3 chronic kidney disease and albuminuria, as an adjunct to standard

of care, in adults with type 2 diabetes mellitus. The proposed new indication is based upon new clinical efficacy and safety data from the Phase 3 study: Canagliflozin and Renal Events in Diabetes with Established Nephropathy Clinical Evaluation Trial (CREDENCE) (DNE3001). The Package Leaflet is updated in accordance. The RMP version 8.1 has also been submitted. In addition, the Marketing authorisation holder (MAH) took the opportunity 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 14.11.2019.

5.1.12. Xolair - omalizumab - EMEA/H/C/000606/II/0101

Novartis Europharm Limited

Rapporteur: Kristina Dunder, PRAC Rapporteur: Annika Folin

Scope: "Extension of indication to include treatment of nasal polyps in adult patients with inadequate response to intranasal corticosteroids for Xolair; 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. In addition, the Marketing authorisation holder (MAH) took the opportunity to make minor editorial changes in section 4.2 of the SmPC and in the PL and to update the phone number of the NL local representative. The RMP version 16.0 has also been submitted. Furthermore, the PI is brought in line with the latest QRD template version 10.1."

Action: For adoption

5.1.13. Zavicefta - ceftazidime / avibactam - EMEA/H/C/004027/II/0019

Pfizer Ireland Pharmaceuticals

Rapporteur: Bjorg Bolstad, Co-Rapporteur: Romaldas Mačiulaitis

Scope: "Extension of indication to include bacteraemia (in association with, or suspected to be associated with, the currently approved indications for complicated intra-abdominal infection (cIAI), complicated urinary tract infection (cUTI) and hospital-acquired pneumonia, including ventilator-associated pneumonia (HAP/VAP)) for Zavicefta; as a consequence, sections 4.1 and 4.2 of the SmPC are updated in order to add this indication and the posology. The Package Leaflet is updated in accordance."

Action: For adoption

5.1.14. WS1737 Edistride - dapagliflozin - EMEA/H/C/004161/WS1737/0034 Forxiga - dapagliflozin - EMEA/H/C/002322/WS1737/0053

AstraZeneca AB

Lead Rapporteur: Kristina Dunder, Lead Co-Rapporteur: Martina Weise, PRAC Rapporteur: Annika Folin

Scope: "Update of sections 4.1, 4.2, 4.4, 4.8 and 5.1 of the SmPC for Edistride and Forxiga to add a new indication for the treatment of symptomatic heart failure with reduced ejection

fraction in adults. The Package Leaflet and Labelling are updated in accordance.
The RMP version 18 has also been submitted.
Furthermore, the PI is brought in line with the latest QRD template version 10.1, as well as editorial change (addition of SI unit for blood glucose).” Request for 1 year of market protection for a new indication (Article 14(11) of Regulation (EC) 726/2004)

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

No items

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

No items

6. Ancillary medicinal substances in medical devices

6.1. Ancillary medicinal substances in medical devices; Opinions/ Day 180 list of outstanding issues / Day 120 list of questions

No items

6.2. Update of Ancillary medicinal substances in medical devices

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. lonafarnib - H0005271

E34.8 Hutchinson-Gilford Progeria Syndrome and Progeroid Laminopathies

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

Action: For adoption

8.1.2. berotralstat - H0005138

Indicated for routine prophylaxis to prevent hereditary angioedema (HAE) attacks in adults and adolescents 12 years of age and older.

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

Action: For adoption

8.1.3. lumasiran - Orphan - H0005040

Alnylam Netherlands B.V.; Treatment of Primary Hyperoxaluria Type 1

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

Action: For adoption

8.2. Priority Medicines (PRIME)

Disclosure of 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. Bavencio - avelumab - EMEA/H/C/004338/II/0013

Merck Europe B.V.

Rapporteur: Filip Josephson, PRAC Rapporteur: Hans Christian Siersted

Scope: "Update of section 5.1 of the SmPC in order to update efficacy information following results from study EMR100070-003 Part B listed as a specific obligation in the Annex II; this is a Phase II, open-label, multicenter trial to investigate the clinical activity and safety of avelumab (MSB0010718C) in subjects with Merkel cell carcinoma. With this submission the company is also taking the opportunity to update annex-II proposing deletion of the specific obligation and proposing the switch from conditional to full marketing authorisation. The package leaflet and the RMP (version 2.1) are updated accordingly."

Action: For information

9.1.2. Benlysta - belimumab - EMEA/H/C/002015/II/0076

GlaxoSmithKline (Ireland) Limited

Rapporteur: Kristina Dunder, PRAC Rapporteur: Ulla Wändel Liminga

Scope: "Update of sections 4.4 and 4.8 of the SmPC in order to update the safety information based on the final results from study BEL115467 listed as a imposed PASS in the Annex II; this is a Randomized, Double-Blind, Placebo-Controlled 52-Week Study to Assess Adverse Events of Special Interest in Adults with Active, Autoantibody-Positive Systemic Lupus Erythematosus Receiving Belimumab; the Package Leaflet is updated accordingly.
The RMP version 36 has also been submitted. The MAH took the opportunity to make the following minor updates to the RMP:
- Updated information on transplant study, updated information on BEL116543, added information on paediatric continuation study in Module SIV.
- Updated exposure information and information for BEL116543 in Module SIV.2.
- Update data on revised rates of pregnancy and lactation in Module SIV.3.
- Correction of an error within Annex 3 and provision of the updated protocol Amendment 7 of a Phase 3, Randomized, Double-Blind, Placebo-Controlled Study to Evaluate the Efficacy and Safety of belimumab plus Standard of Care versus Placebo plus standard of Care in Adult Subjects with Active Lupus Nephritis (BEL114054).
This RMP also includes changes being reviewed in procedure II/0062, related to the use of Benlysta in paediatric patients, and BEL116027, evaluating the safety and efficacy of temporary discontinuation of belimumab.
In addition, the Marketing authorisation holder took the opportunity to make minor editorial changes to the Annex II and the label."

Action: For discussion

9.1.3. Cablivi - caplacizumab - EMEA/H/C/004426/II/0021, Orphan

Ablynx NV

Rapporteur: Filip Josephson

Scope: "Submission of the results of the study ALX-0681-MS-01, a Modelling/Simulation study performed for the paediatric population as part of the approved Paediatric Investigation Plan (EMEA-001157-PIP-01-11-M02) for Cablivi"

Action: For discussion

9.1.4. Juluca - dolutegravir / rilpivirine - EMEA/H/C/004427/II/0016 Dovato - dolutegravir / lamivudine - EMEA/H/C/004909/II/0001 Triumeq - dolutegravir / abacavir / lamivudine - EMEA/H/C/002754/II/0069 Tivicay - dolutegravir - EMEA/H/C/002753/II/0052

ViiV Healthcare B.V.

Lead Rapporteur: Filip Josephson

Scope: Update of section 4.6 of the SmPC in order to update the safety information regarding the occurrence of neural tube defects with the DTG-containing regimens based on interim analysis from Tsepamo study. This is a birth outcomes surveillance study being conducted in Botswana that was designed to evaluate adverse birth outcomes by HIV status and antiretroviral regimen, and to determine if there is an increased risk of neural tube defects among infants exposed to efavirenz at conception. This surveillance system captures all antiretroviral exposure including dolutegravir. The SmPC is updated accordingly. The RMP is not submitted.

List of experts for the SAG-HIV/viral diseases meeting adopted via written procedure on 05 February 2020, Report from SAG-HIV/viral diseases meeting held on 06 February 2020

Action: For information

9.1.5. OCALIVA - obeticholic acid – Orphan - EMEA/H/C/004093/R/0018

Intercept Pharma International Limited

Rapporteur: Jorge Camarero Jiménez, PRAC Rapporteur: Menno van der Elst

Scope: Renewal

Action: For discussion

Request for Supplementary Information adopted on 17.10.2019, 19.09.2019.

9.1.6. Qutenza - capsaicin - EMEA/H/C/000909/II/0048

Grunenthal GmbH

Rapporteur: Bruno Sepodes

Scope: "Update of sections 4.2 and 5.1 of the SmPC in order to amend the posology based on PACE and STRIDE studies. The Package Leaflet is updated accordingly."

Action: For discussion

Request for Supplementary Information adopted on 12.12.2019.

9.1.7. Zynteglo - autologous CD34+ cell enriched population that contains hematopoietic stem cells transduced with lentiglobin BB305 lentiviral vector encoding the beta-A-T87Q-globin gene - EMEA/H/C/003691/R/0005, Orphan, ATMP

bluebird bio (Netherlands) B.V

Rapporteur: Carla Herberts, Co-Rapporteur: Violaine Closson Carella, CHMP Coordinator Rapporteur: Paula Boudewina van Hennik, CHMP Coordinator Co-Rapporteur: Alexander Moreau

Scope: Opinion

Action: For discussion

9.1.8. Lifmior – etanercept - EMEA/H/C/004167

Pfizer Europe MA EEIG

Rapporteur: Maria Concepcion Prieto Yerro, PRAC Rapporteur: Eva A. Segovia

Scope: The marketing authorisation of Lifmior expired on 16 February 2020 due to end of the sunset clause

Action: For information

10. Referral procedures

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

10.1.1. Yondelis - EMEA/H/C/0773/A-20/0060

MAH: Pharma Mar S.A.

Rapporteur: Sinan B. Sarac, Co-Rapporteur: Jorge Camarero Jiménez

Scope: Start of procedure, List of Questions, Timetable, Appointment of Rapporteurs

Action: For adoption

European Commission triggered a referral procedure under Article 20 of Regulation (EC) No 726/2004 to request CHMP to assess study OVC-3006, which failed to meet its endpoints in the indication of ovarian cancer, and its impact on the benefit risk balance for the centrally authorised medicinal product(s) Yondelis (trabectedin)

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

10.2.1. Direct Oral Anticoagulants (DOAC) - EMEA/H/A-5(3)/1478

MAHs: various

Referral Rapporteur: Martina Weise, Referral Co-Rapporteurs: Kristina Dunder

Rapporteurs for involved CAPs:

Eliquis (APIXABAN):
Rapporteur: Johann Lodewijk Hillege, Co-Rapporteur: Bart van der Schueren;

Pradaxa (DABIGATRAN ETEXILATE):
Rapporteur: Mark Ainsworth, Co-Rapporteur: Jean-Michel Race;

Xarelto (RIVAROXABAN):
Rapporteur: Kristina Dunder, Co-Rapporteur: Martina Weise

Scope: Opinion

Action: For adoption

10.2.2. Presence of nitrosamine impurities in human medicinal products containing chemically synthesised active pharmaceutical ingredients EMEA/H/A-5(3)/1490

MAHs: various

Referral Rapporteur: Martina Weise, Referral Co-Rapporteurs: Kristina Dunder

Scope: Preparation for the ad-hoc expert group meeting

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

No items

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

10.6.1. Ranitidine - EMEA/H/A-31/1491

MAH various

Rapporteur: Johann Lodewijk Hillege, Co-Rapporteur: Daniela Melchiorri

Scope: Oral explanations

Action: Oral explanations to be held on Tuesday, 25 February 2020 at 09:00, 10:00 and 11:00

Tests performed in a random selection of ranitidine API batches and finished products available in the EU have shown levels of NDMA which raise concerns.

European Commission triggered on 12 September 2019 a referral procedure under Article 31 of Directive 2001/83/EC to evaluate the relevance of these findings, the potential root causes and their impact on the benefit-risk balance of medicinal products containing ranitidine.

See 2.4

10.6.2. Panexcell Clinical Laboratories Priv. Ltd - Multiple NAPs (EMEA/H/A-31/1494)

MAH : various

Referral Rapporteur: TBC, Referral Co-Rapporteur :TBC

Scope: Start of procedure, List of Questions, Timetable, Appointment of Rapporteurs

Action: For adoption

Article 31 procedure triggered by the German Federal Institute of Drugs and Medical Devices (BfArM) concerning the contract research organisation (CRO) Panexcell Clinical Laboratories Priv. Ltd. located in Navi Mumbai 400 701, India.

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

February 2020 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

Action: For information

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

No items

14.2. Coordination with EMA Scientific Committees

14.2.1. Pharmacovigilance Risk Assessment Committee (PRAC)

Summary of recommendations and advice of PRAC meeting held on 10-13 February 2020

Action: For information

List of Union Reference Dates and frequency of submission of Periodic Safety Update Reports (EURD list) for February 2020

Action: For adoption

14.2.2. Committee for Advanced Therapies (CAT)

CAT draft minutes of meeting held on 19-21 February 2020

Action: For information

14.2.3. Paediatric Committee (PDCO)

PIPs reaching D30 at February 2020 PDCO

Action: For information

Report from the PDCO meeting held on 25-28 February 2020

Action: For information

14.2.4. Committee for Orphan Medicinal Products (COMP)

Report from the COMP meeting held on 18-20 February 2020

Action: For information

14.2.5. Coordination Group for Mutual Recognition and Decentralised Procedures – Human (CMDh)

Report from the Coordination Group for Mutual Recognition and Decentralised Procedures – Human (CMDh) on the meeting held on 25-27 February 2020

Action: For information

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

14.3.1. Ad-hoc Influenza Working Group

Scope EU Strain selection for the Influenza Vaccines for the Season 2020/2021: Report from the Ad Hoc Influenza working group to the BWP

Action: For adoption

Scope: EU Recommendation for the Seasonal Influenza Vaccine Composition for the Season 2020/2021

Action: For adoption

14.3.2. Biologics Working Party (BWP)

Chair: TBC

Reports from BWP February 2020 meeting to CHMP:

- 11 reports on products in scientific advice and protocol assistance
- 14 reports on products in pre-authorisation procedures
- 5 reports on products in post-authorisation procedures
- 3 reports on products in plasma master file

Action: For adoption

Election of the new BWP chair

Action: For adoption

Nomination(s) received

14.3.3. Central Nervous System Working Party (CNSWP)

Chair: Karl Broich/André Elferink

CNSWP response to CMDh question on bioequivalence requirements for Cmax for carbamazepine as NTI drug

Action: For adoption

14.3.4. Scientific Advice Working Party (SAWP)

Report from the SAWP meeting held on 10-13 February 2020. Table of conclusions

Action: For information

Scientific advice letters:

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

14.4. Cooperation within the EU regulatory network

No items

14.5. Cooperation with International Regulators

14.5.1. International Council on Harmonisation (ICH)

ICH M13 – Bioequivalence for immediate-release dosage forms: nomination of expert

Action: For adoption

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. Oncology Training

Action: For information

15.1.2. UK withdrawal from the EU – update

Action: For discussion

16. 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)*

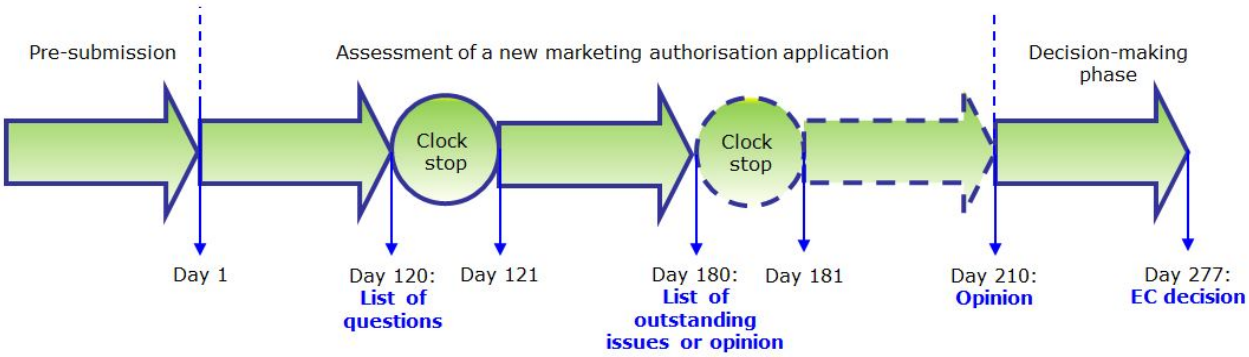
The items listed in this section are those for which marketing authorisation holders (MAHs) or applicants have been invited to the CHMP plenary meeting to address questions raised by the Committee. Oral explanations normally relate to on-going applications (section 3, 4 and 5) or referral procedures (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

24 February 2020
EMA/CHMP/92695/2020

Annex to 24-27 February 2020 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 February 2020: **For adoption**

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

Final Outcome of Rapporteurship allocation for February 2020: **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

Defitelio - defibrotide -
EMEA/H/C/002393/S/0045, Orphan
Gentium S.r.l., Rapporteur: Kristina Dunder, PRAC Rapporteur: Ulla Wändel Liminga

Naglazyme - galsulfase -
EMEA/H/C/000640/S/0078
BioMarin International Limited, Rapporteur: Fátima Ventura, PRAC Rapporteur: Ana Sofia Diniz Martins
Request for Supplementary Information adopted on 14.11.2019.

Obizur - susoctocog alfa -
EMEA/H/C/002792/S/0028
Baxalta Innovations GmbH, Rapporteur: Andrea Laslop, PRAC Rapporteur: Brigitte Keller-Stanislawski

Orphacol - cholic acid -
EMEA/H/C/001250/S/0033, Orphan

Laboratoires CTRS, Rapporteur: Konstantinos Markopoulos, PRAC Rapporteur: Sofia Trantza

Raxone - idebenone -
EMEA/H/C/003834/S/0019, Orphan
Santhera Pharmaceuticals (Deutschland) GmbH, Rapporteur: John Joseph Borg, PRAC Rapporteur: Amelia Cupelli
Request for Supplementary Information adopted on 30.01.2020.

Vedrop - tocofersolan -
EMEA/H/C/000920/S/0035
Recordati Rare Diseases, Rapporteur: Melinda Sobor, PRAC Rapporteur: Melinda Palfi

B.2. RENEWALS OF MARKETING AUTHORISATIONS OUTCOMES

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

Lumark - lutetium (177Lu) chloride -
EMEA/H/C/002749/R/0014
I.D.B. Holland B.V., Rapporteur: Jean-Michel Race, Co-Rapporteur: Maria Concepcion Prieto Yerro, PRAC Rapporteur: Ronan Grimes
Request for Supplementary Information adopted on 12.12.2019.

B.2.2. Renewals of Marketing Authorisations for unlimited validity

Bortezomib Accord - bortezomib -
EMEA/H/C/003984/R/0022
Accord Healthcare S.L.U., Generic, Generic of VELCADE, Rapporteur: Milena Stain, PRAC Rapporteur: Amelia Cupelli
Request for Supplementary Information adopted on 30.01.2020.

Farydak - panobinostat -
EMEA/H/C/003725/R/0020, Orphan
Secura Bio Limited, Rapporteur: Paula Boudewina van Hennik, Co-Rapporteur: Filip Josephson, PRAC Rapporteur: Sofia Trantza

IKERVIS - ciclosporin -
EMEA/H/C/002066/R/0017
Santen Oy, Rapporteur: Peter Kiely, Co-Rapporteur: Agnes Gyurasics, PRAC Rapporteur: Jan Neuhauser
Request for Supplementary Information adopted

on 19.09.2019.

Kanuma - sebelipase alfa - EMEA/H/C/004004/R/0025, Orphan
Alexion Europe SAS, Rapporteur: Bart Van der Schueren, Co-Rapporteur: Fátima Ventura, PRAC Rapporteur: Ulla Wändel Liminga

OPDIVO - nivolumab - EMEA/H/C/003985/R/0074
Bristol-Myers Squibb Pharma EEIG, Rapporteur: Jorge Camarero Jiménez, Co-Rapporteur: Paula Boudewina van Hennik, PRAC Rapporteur: Brigitte Keller-Stanislawski
Request for Supplementary Information adopted on 12.12.2019.

Respreeza - human alpha1-proteinase inhibitor - EMEA/H/C/002739/R/0036
CSL Behring GmbH, Rapporteur: Kristina Dunder, Co-Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Maria del Pilar Rayon

Strensiq - asfotase alfa - EMEA/H/C/003794/R/0044, Orphan
Alexion Europe SAS, Rapporteur: Daniela Melchiorri, Co-Rapporteur: Jayne Crowe, PRAC Rapporteur: Rhea Fitzgerald
Request for Supplementary Information adopted on 30.01.2020.

Zerbaxa - ceftolozane / tazobactam - EMEA/H/C/003772/R/0026
Merck Sharp & Dohme B.V., Rapporteur: Bjorg Bolstad, Co-Rapporteur: Ewa Balkowiec Iskra, PRAC Rapporteur: Adam Przybylkowski

B.2.3. Renewals of Conditional Marketing Authorisations

Deltyba - delamanid - EMEA/H/C/002552/R/0041, Orphan
Otsuka Novel Products GmbH, Rapporteur: Koenraad Norga, PRAC Rapporteur: Jean-Michel Dogné
Request for Supplementary Information adopted on 30.01.2020.

Natpar - parathyroid hormone - EMEA/H/C/003861/R/0022, Orphan
Shire Pharmaceuticals Ireland Limited, Rapporteur: Bart Van der Schueren, PRAC Rapporteur: Rhea Fitzgerald
Request for Supplementary Information adopted

on 12.12.2019.	
OCALIVA - obeticholic acid - EMEA/H/C/004093/R/0018, Orphan Intercept Pharma International Limited, Rapporteur: Jorge Camarero Jiménez, PRAC Rapporteur: Liana Gross-Martirosyan Request for Supplementary Information adopted on 19.09.2019.	See 9.1
Rubraca - rucaparib - EMEA/H/C/004272/R/0016 Clovis Oncology Ireland Limited, Rapporteur: Jorge Camarero Jiménez, PRAC Rapporteur: Annika Folin Request for Supplementary Information adopted on 30.01.2020.	
Zynteglo - autologous CD34+ cell enriched population that contains hematopoietic stem cells transduced with lentiglobin BB305 lentiviral vector encoding the beta-A-T87Q-globin gene - EMEA/H/C/003691/R/0005, Orphan, ATMP bluebird bio (Netherlands) B.V, Rapporteur: Carla Herberts, Co-Rapporteur: Violaine Closson Carella, CHMP Coordinators: Paula Boudewina van Hennik and Alexandre Moreau, PRAC Rapporteur: Brigitte Keller-Stanislawski Request for Supplementary Information adopted on 24.01.2020.	See 9.1

B.3. POST-AUTHORISATION PHARMACOVIGILANCE OUTCOMES

Signal detection

PRAC recommendations on signals adopted at the PRAC meeting held on 10-13 February 2020
PRAC:

Signal of haemophagocytic lumphohistiocytosis:

OPDIVO - nivolumab – EMEA/H/C/003985

Rapporteur: Jorge Camarero Jiménez, Co-Rapporteur: Paula Boudewina van Hennik

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 February 2020 meeting:

EMEA/H/C/PSUSA/00000234/201907
(aripiprazole)
CAPS:
Abilify (EMEA/H/C/000471) (aripiprazole), Otsuka Pharmaceutical Netherlands B.V., Rapporteur: Bruno Sepodes
Abilify Maintena (EMEA/H/C/002755) (aripiprazole), Otsuka Pharmaceutical Netherlands B.V., Rapporteur: Bruno Sepodes
Aripiprazole Sandoz (EMEA/H/C/004008) (aripiprazole), Sandoz GmbH, Rapporteur: John Joseph Borg
NAPS:
ARIPIPRAZOL HEUMANN - HEUMANN PHARMA GMBH CO GENERICA KG
PRAC Rapporteur: Ana Sofia Diniz Martins, "From: 17/07/2018 To: 16/07/2019"

EMEA/H/C/PSUSA/00002127/201908
(natalizumab)
CAPS:
Tysabri (EMEA/H/C/000603) (natalizumab), Biogen Netherlands B.V., Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Brigitte Keller-Stanislawski, "Period Covered From: 06/08/2018 To: 06/08/2019"

EMEA/H/C/PSUSA/00009255/201907
(perampanel)
CAPS:
Fycompa (EMEA/H/C/002434) (perampanel), Eisai GmbH, Rapporteur: Alexandre Moreau, PRAC Rapporteur: Ghania Chamouni, "Period Covered From: 21/07/2018 To: 21/07/2019"

EMEA/H/C/PSUSA/00010457/201907
(pegaspargase (centrally authorised product))
CAPS:
Oncaspar (EMEA/H/C/003789) (pegaspargase), Les Laboratoires Servier, Rapporteur: Alexandre Moreau, PRAC Rapporteur: Annika Folin, "From: 14/07/2018 To: 14/07/2019"

EMEA/H/C/PSUSA/00010609/201907
(sarilumab)
CAPS:
Kevzara (EMEA/H/C/004254) (sarilumab), sanofi-aventis groupe, Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Eva A. Segovia, "From: 12/07/2018 To: 12/07/2019"

B.4. EPARs / WPARs

Arsenic trioxide Mylan - arsenic trioxide - EMEA/H/C/005235 Mylan Ireland Limited, treatment of relapsed acute promyelocytic leukaemia (APL), Generic, Generic of TRISENOX, Generic application (Article 10(1) of Directive No 2001/83/EC)	For information only. Comments can be sent to the PL in case necessary.
Azacitidine betapharm - azacitidine - EMEA/H/C/005075 betapharm Arzneimittel GmbH, Treatment of myelodysplastic syndromes (MDS), chronic myelomonocytic leukemia (CMML) and acute myeloid leukemia (AML), Generic, Generic of Vidaza, Generic application (Article 10(1) of Directive No 2001/83/EC)	For information only. Comments can be sent to the PL in case necessary.
Azacitidine Mylan - azacitidine - EMEA/H/C/004984 Mylan Ireland Limited, treatment of adult patients who are not eligible for haematopoietic stem cell transplantation (HSCT), Generic, Generic of Vidaza, Generic application (Article 10(1) of Directive No 2001/83/EC)	For information only. Comments can be sent to the PL in case necessary.
Budesonide/Formoterol Teva Pharma B.V. - budesonide / formoterol fumarate dihydrate - EMEA/H/C/004882 Teva Pharma B.V., treatment of asthma and COPD, Hybrid application (Article 10(3) of Directive No 2001/83/EC)	For information only. Comments can be sent to the PL in case necessary.
Cinacalcet Accordpharma - cinacalcet - EMEA/H/C/005236 Accord Healthcare S.L.U., treatment of secondary hyperparathyroidism and hypercalcaemia, Generic, Generic of Mimpara, Generic application (Article 10(1) of Directive No 2001/83/EC)	For information only. Comments can be sent to the PL in case necessary.
Evenity - romosozumab - EMEA/H/C/004465 UCB Pharma S.A., Treatment of osteoporosis, 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.
GIVLAARI - givosiran - EMEA/H/C/004775, Orphan Alnylam Netherlands B.V., Treatment of acute hepatic porphyria (AHP) in adults and adolescents aged 12 years and older., 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.

Isturisa - osilodrostat - EMEA/H/C/004821, Orphan Novartis Europharm Limited, treatment of Cushing's syndrome, 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.
Liumjev - insulin lispro - EMEA/H/C/005037 Eli Lilly Nederland B.V., Treatment of diabetes mellitus in adults, Known active substance (Article 8(3) of Directive No 2001/83/EC)	For information only. Comments can be sent to the PL in case necessary.
Nilemdo - bempedoic acid - EMEA/H/C/004958 FGK Representative Service GmbH, treatment of primary hypercholesterolaemia or mixed dyslipidaemia, 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.
NUBEQA - darolutamide - EMEA/H/C/004790 Bayer AG, treatment of non-metastatic castration resistant prostate cancer (nmCRPC), 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.
Nustendi - bempedoic acid / ezetimibe - EMEA/H/C/004959 FGK Representative Service GmbH, treatment of primary hypercholesterolaemia or mixed dyslipidaemia, 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.
Ruxience - rituximab - EMEA/H/C/004696 Pfizer Europe MA EEIG, treatment of non-Hodgkin’s lymphoma (NHL), chronic lymphocytic leukaemia (CLL), rheumatoid arthritis (RA), granulomatosis with polyangiitis (GPA) and microscopic polyangiitis (MPA), and Pemphigus vulgaris (PV), Similar biological application (Article 10(4) of Directive No 2001/83/EC)	For information only. Comments can be sent to the PL in case necessary.
Rybelsus - semaglutide - EMEA/H/C/004953 Novo Nordisk A/S, treatment of type 2 diabetes mellitus, Known active substance (Article 8(3) of Directive No 2001/83/EC)	For information only. Comments can be sent to the PL in case necessary.
Staquis - crisaborole - EMEA/H/C/004863 Pfizer Europe MA EEIG, treatment of mild to moderate atopic dermatitis, New active substance (Article 8(3) of Directive No	For information only. Comments can be sent to the PL in case necessary.

2001/83/EC)	
Trepulmix - treprostinil sodium - EMEA/H/C/005207, Orphan SciPharm Sarl, treatment of thromboembolic pulmonary hypertension (CTEPH), Hybrid application (Article 10(3) of Directive No 2001/83/EC)	For information only. Comments can be sent to the PL in case necessary.
Vaxchora - cholera vaccine, oral, live - EMEA/H/C/003876 Emergent Netherlands B.V., indicated for active immunisation against disease caused by Vibrio cholerae serogroup O1 in adults and children aged 6 years and older, 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

Adenuric - febuxostat - EMEA/H/C/000777/II/0056 Menarini International Operations Luxembourg S.A., Rapporteur: Andrea Laslop Request for Supplementary Information adopted on 13.02.2020.	Request for supplementary information adopted with a specific timetable.
BeneFIX - nonacog alfa - EMEA/H/C/000139/II/0161/G Pfizer Europe MA EEIG, Rapporteur: Jan Mueller-Berghaus Request for Supplementary Information adopted on 05.12.2019.	
Brineura - cerliponase alfa - EMEA/H/C/004065/II/0019, Orphan BioMarin International Limited, Rapporteur: Martina Weise Request for Supplementary Information adopted on 06.02.2020.	Request for supplementary information adopted with a specific timetable.
Buvidal - buprenorphine - EMEA/H/C/004651/II/0005 Camurus AB, Rapporteur: Peter Kiely Opinion adopted on 13.02.2020.	Positive Opinion adopted by consensus on 13.02.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Cimzia - certolizumab pegol - EMEA/H/C/001037/II/0085/G UCB Pharma S.A., Rapporteur: Kristina Dunder	Positive Opinion adopted by consensus on 13.02.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP

Opinion adopted on 13.02.2020. Request for Supplementary Information adopted on 12.12.2019.	recommendation.
CRYSVITA - burosumab - EMEA/H/C/004275/II/0007/G, Orphan Kyowa Kirin Holdings B.V., Rapporteur: Kristina Dunder Request for Supplementary Information adopted on 21.11.2019, 12.09.2019.	
Elaprase - idursulfase - EMEA/H/C/000700/II/0082 Shire Human Genetic Therapies AB, Rapporteur: Johann Lodewijk Hillege Request for Supplementary Information adopted on 12.12.2019, 12.09.2019.	
Eylea - aflibercept - EMEA/H/C/002392/II/0058 Bayer AG, Rapporteur: Alexandre Moreau Opinion adopted on 13.02.2020. Request for Supplementary Information adopted on 14.11.2019.	Positive Opinion adopted by consensus on 13.02.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
IKERVIS - ciclosporin - EMEA/H/C/002066/II/0018 Santen Oy, Rapporteur: Peter Kiely Opinion adopted on 20.02.2020. Request for Supplementary Information adopted on 16.01.2020.	Positive Opinion adopted by consensus on 20.02.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Ilumetri - tildrakizumab - EMEA/H/C/004514/II/0010/G Almirall S.A, Rapporteur: Jan Mueller-Berghaus Opinion adopted on 20.02.2020. Request for Supplementary Information adopted on 28.11.2019.	Positive Opinion adopted by consensus on 20.02.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Inflectra - infliximab - EMEA/H/C/002778/II/0081/G Pfizer Europe MA EEIG, Duplicate, Duplicate of Remsima, Rapporteur: Outi Mäki-Ikola Opinion adopted on 06.02.2020. Request for Supplementary Information adopted on 28.11.2019.	Positive Opinion adopted by consensus on 06.02.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Natpar - parathyroid hormone - EMEA/H/C/003861/II/0020/G, Orphan Shire Pharmaceuticals Ireland Limited, Rapporteur: Bart Van der Schueren Request for Supplementary Information adopted on 07.11.2019.	

Onpattro - patisiran - EMEA/H/C/004699/II/0011/G, Orphan Alnylam Netherlands B.V., Rapporteur: Kristina Dunder Request for Supplementary Information adopted on 06.02.2020.	Request for supplementary information adopted with a specific timetable.
Ozempic - semaglutide - EMEA/H/C/004174/II/0011 Novo Nordisk A/S, Rapporteur: Johann Lodewijk Hillege Request for Supplementary Information adopted on 13.02.2020.	Request for supplementary information adopted with a specific timetable.
Remicade - infliximab - EMEA/H/C/000240/II/0225/G Janssen Biologics B.V., Rapporteur: Kristina Dunder Opinion adopted on 13.02.2020.	Positive Opinion adopted by consensus on 13.02.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Remsima - infliximab - EMEA/H/C/002576/II/0075/G Celltrion Healthcare Hungary Kft., Rapporteur: Outi Mäki-Ikola Opinion adopted on 06.02.2020. Request for Supplementary Information adopted on 28.11.2019.	Positive Opinion adopted by consensus on 06.02.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
RoActemra - tocilizumab - EMEA/H/C/000955/II/0093/G Roche Registration GmbH, Rapporteur: Jan Mueller-Berghaus Request for Supplementary Information adopted on 13.02.2020.	Request for supplementary information adopted with a specific timetable.
Rotarix - rotavirus vaccine (live, oral) - EMEA/H/C/000639/II/0116/G GlaxoSmithKline Biologicals S.A., Rapporteur: Bart Van der Schueren Opinion adopted on 13.02.2020.	Positive Opinion adopted by consensus on 13.02.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Skilarence - dimethyl fumarate - EMEA/H/C/002157/II/0019 Almirall S.A, Rapporteur: Janet Koenig Request for Supplementary Information adopted on 13.02.2020.	Request for supplementary information adopted with a specific timetable.
Somavert - pegvisomant - EMEA/H/C/000409/II/0091 Pfizer Europe MA EEIG, Rapporteur: Jean-Michel Race Opinion adopted on 06.02.2020. Request for Supplementary Information adopted	Positive Opinion adopted by consensus on 06.02.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

on 05.12.2019.	
Trumenba - meningococcal group B vaccine (recombinant, adsorbed) - EMEA/H/C/004051/II/0020/G Pfizer Europe MA EEIG, Rapporteur: Johann Lodewijk Hillege Opinion adopted on 06.02.2020. Request for Supplementary Information adopted on 05.12.2019.	Positive Opinion adopted by consensus on 06.02.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
VIZAMYL - flutemetamol (^{18}F) - EMEA/H/C/002557/II/0022/G GE Healthcare AS, Rapporteur: Maria Concepcion Prieto Yerro Opinion adopted on 06.02.2020.	Positive Opinion adopted by consensus on 06.02.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Ziextenzo - pegfilgrastim - EMEA/H/C/004802/II/0005/G Sandoz GmbH, Rapporteur: Andrea Laslop Request for Supplementary Information adopted on 06.02.2020.	Request for supplementary information adopted with a specific timetable.
WS1524 **HyQvia-EMEA/H/C/002491/WS1524/0048** **Kiovig-EMEA/H/C/000628/WS1524/0090** Takeda Manufacturing Austria AG, Lead Rapporteur: Jan Mueller-Berghaus Opinion adopted on 20.02.2020. Request for Supplementary Information adopted on 19.09.2019, 14.03.2019.	Positive Opinion adopted by consensus on 20.02.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1720/G **Ambirix-EMEA/H/C/000426/WS1720/0104/G** **Twinrix Adult-EMEA/H/C/000112/WS1720/0139/G** **Twinrix Paediatric-EMEA/H/C/000129/WS1720/0140/G** GlaxoSmithkline Biologicals SA, Lead Rapporteur: Bart Van der Schueren Request for Supplementary Information adopted on 16.01.2020.	
WS1744/G **Hexacima-EMEA/H/C/002702/WS1744/0095/G** **Hexaxim-EMEA/H/W/002495/WS1744/0100/G** **Hexyon-EMEA/H/C/002796/WS1744/0099/G** Sanofi Pasteur, Lead Rapporteur: Jan Mueller-Berghaus	Positive Opinion adopted by consensus on 06.02.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

Opinion adopted on 06.02.2020.	

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

Avamys - fluticasone furoate - EMEA/H/C/000770/II/0040 GlaxoSmithKline (Ireland) Limited, Rapporteur: Ewa Balkowiec Iskra, "Update of section 4.8 of the SmPC in order to add bronchospasm and dyspnoea to the list of adverse drug reactions with a frequency unknown based on post-marketing experience. The package leaflet is updated accordingly. In addition, the MAH took the opportunity to update the list of local representatives in the package leaflet." Request for Supplementary Information adopted on 06.02.2020.	Request for supplementary information adopted with a specific timetable.
Cablivi - caplacizumab - EMEA/H/C/004426/II/0021, Orphan Ablynx NV, Rapporteur: Filip Josephson, , "Submission of the results of the study ALX-0681-MS-01, a Modelling/Simulation study performed for the paediatric population as part of the approved Paediatric Investigation Plan (EMEA-001157-PIP-01-11-M02) for Cablivi"	See agenda 9.1
Dengvaxia - dengue tetravalent vaccine (live, attenuated) - EMEA/H/C/004171/II/0007/G Sanofi Pasteur, Rapporteur: Bart Van der Schueren, "C.I.13: Submission of the final report from studies CYD63 and CYD64 listed as a category 3 study in the RMP. These are booster studies to evaluate the safety and immunogenicity of a booster dose of dengue vaccine administered in a subset of subjects who received third dose of dengue vaccine 4-5 years before, in Phase II studies." Opinion adopted on 13.02.2020.	Positive Opinion adopted by consensus on 13.02.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Dovato - dolutegravir / lamivudine - EMEA/H/C/004909/II/0008 ViiV Healthcare B.V., Rapporteur: Filip Josephson, "Update of section 5.1 of the SmPC in order to update the safety and efficacy information following the week 48 results from TANGO study (204862); TANGO (204862) is an on-going 200-week, Phase III, randomized, open-label, active controlled, multicenter,	Positive Opinion adopted by consensus on 13.02.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

parallel-group study, evaluating the efficacy, safety, and tolerability of switching to Dovato in HIV-1 infected adults who are virologically suppressed. The RMP version has not been submitted." Opinion adopted on 13.02.2020.	
Dovato - dolutegravir / lamivudine - EMEA/H/C/004909/II/0009 ViiV Healthcare B.V., Rapporteur: Filip Josephson, "Update of section 5.1 in order to update the safety and efficacy following the week 96 results from 204861 (GEMINI-1) and 205543 (GEMINI-2) studies listed as a specific category 3 study in the RMP; these are two identical pivotal ongoing, randomized, double-blind, parallel group, 148-week, phase III studies to evaluate the efficacy, safety and tolerability of dolutegravir plus 3TC compared to dolutegravir plus tenofovir/emtricitabine in HIV-1-infected treatment naïve patients. The RMP version has not been submitted." Opinion adopted on 13.02.2020.	Positive Opinion adopted by consensus on 13.02.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Feraccru - ferric maltol - EMEA/H/C/002733/II/0024 Norgine B.V., Rapporteur: Maria Concepcion Prieto Yerro, "To update sections 4.4 and 5.1 of the SmPC in order to update safety and efficacy information based on final results from study ST 10-01-304 this is a phase 3b, randomized, controlled, multicentre study with oral ferric maltol (Feraccru) or intravenous iron (ferric carboxymaltose; FCM), for the treatment of iron deficiency anaemia in subjects with inflammatory bowel disease."	
Imbruvica - ibrutinib - EMEA/H/C/003791/II/0058, Orphan Janssen-Cilag International NV, Rapporteur: Filip Josephson, PRAC Rapporteur: Nikica Mirošević Skvrce, "Update of sections 4.5 and 4.6 of the SmPC following the results from study CLL1017 (MEA 012.2). The PL is updated accordingly." Request for Supplementary Information adopted on 13.02.2020.	Request for supplementary information adopted with a specific timetable.
Jivi - damoctocog alfa pegol - EMEA/H/C/004054/II/0004 Bayer AG, Rapporteur: Sinan B. Sarac, "Submission of the final Clinical Study Report PH-40657 for the pharmacokinetic study (study 19096) comparing pharmacokinetic parameters	Positive Opinion adopted by consensus on 06.02.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

of Jivi vs. Elocta." Opinion adopted on 06.02.2020. Request for Supplementary Information adopted on 05.12.2019.	
Kisqali - ribociclib - EMEA/H/C/004213/II/0020 Novartis Europharm Limited, Rapporteur: Filip Josephson, "Update of sections 4.4 and 4.8 of the SmPC in order to add a warning and include safety information on toxic epidermal necrolysis. The Package Leaflet is updated accordingly." Opinion adopted on 13.02.2020.	Positive Opinion adopted by consensus on 13.02.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Kisqali - ribociclib - EMEA/H/C/004213/II/0022 Novartis Europharm Limited, Rapporteur: Filip Josephson, "Update of section 5.1 of the SmPC based on the final OS analysis from study CLEE011F2301 (MONALEESA-3), a randomised double-blind, placebo-controlled, multicentre phase III clinical study in the treatment of men and postmenopausal women with hormone receptor positive, HER2-negative advanced breast cancer who had received no or only one line of prior endocrine treatment, in combination with fulvestrant versus fulvestrant alone." Request for Supplementary Information adopted on 13.02.2020.	Request for supplementary information adopted with a specific timetable.
MabThera - rituximab - EMEA/H/C/000165/II/0169 Roche Registration GmbH, Rapporteur: Sinan B. Sarac, "Update of the SmPC sections 4.8, 5.1 and 5.2. with the results of the Post-authorisation efficacy study (PAES) randomised phase 3 study (PEMPHIX WA29330) which further investigated the efficacy of Mabthera in the subgroup of patients with established PV as well as characterised its long term efficacy and safety on disease progression. Annex II and PL are updated accordingly." Request for Supplementary Information adopted on 12.12.2019.	
Maviret - glecaprevir / pibrentasvir - EMEA/H/C/004430/II/0031 AbbVie Deutschland GmbH & Co. KG, Rapporteur: Jean-Michel Race, "Submission of the final clinical study report from the Phase 3b study M16-156 (A Multicenter, Open-Label Study to Evaluate the Efficacy and Safety of Glecaprevir (GLE)/Pibrentasvir (PIB) in Treatment-Naïve	Positive Opinion adopted by consensus on 13.02.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

Adults in Brazil with Chronic Hepatitis C Virus (HCV) Genotype 1 – 6 Infection).” Opinion adopted on 13.02.2020.	
Nivestim - filgrastim - EMEA/H/C/001142/II/0061 Pfizer Europe MA EEIG, Rapporteur: Outi Mäki-Ikola, “To update section 4.4 of the SmPC to add a warning on the content of a derivative of natural rubber latex in the needle cover formulation. The Package Leaflet are updated accordingly.” Request for Supplementary Information adopted on 06.02.2020.	Request for supplementary information adopted with a specific timetable.
Perjeta - pertuzumab - EMEA/H/C/002547/II/0047 Roche Registration GmbH, Rapporteur: Sinan B. Sarac, “Submission of the final report from study W020698 (CLEOPATRA), a phase III, randomized, double blind, placebo-controlled clinical trial to evaluate the efficacy and safety of pertuzumab + trastuzumab + docetaxel vs placebo + trastuzumab + docetaxel in previously untreated HER2-positive metastatic breast cancer.” Request for Supplementary Information adopted on 06.02.2020.	Request for supplementary information adopted with a specific timetable.
PREVYMIS - letermovir - EMEA/H/C/004536/II/0013, Orphan Merck Sharp & Dohme B.V., Rapporteur: Filip Josephson, “Update of section 5.1 of the SmPC in order to update the viral resistance profile that may be associated with a change in susceptibility to letermovir considering new intro pharmacology data based on the analysis of the patients’ samples included in the study MK-8228. This study is a Phase III Randomized, Placebo-Controlled Clinical Trial to Evaluate the Safety and Efficacy of MK-8228 (Letermovir) for the Prevention of Clinically Significant Human Cytomegalovirus (CMV) Infection in Adult, CMV Seropositive Allogeneic Hematopoietic Stem Cell. This variation follows the recommendation dated 9th November 2017 that asked for the submission when available of the results to update the CMV phenotypic resistance analyses of all clinical isolates for subjects failing letermovir treatment and to explore the possibility to obtain additional pre-failure CMV genotypic data from available samples.”	

Request for Supplementary Information adopted on 12.12.2019, 17.10.2019.	
PREVYMIS - letermovir - EMEA/H/C/004536/II/0014, Orphan Merck Sharp & Dohme B.V., Rapporteur: Filip Josephson, "Update of section 4.5 of the SmPC in order to update the safety information with regard to the drug interaction information following the results from study MK-8228-039, a clinical pharmacology trial entitled "A Study to Assess the Effect of P-gp/BCRP Inhibition, following Multiple Oral Doses of Itraconazole, on the Steady-State Pharmacokinetics of MK-8228 in Healthy Adult Subjects" listed as a category 3 study in the RMP. The RMP version has not been submitted. In addition, the Marketing authorisation holder (MAH) took the opportunity to submit the protein binding report as part of the rifampin study MK-8228-038 as it was requested within the previous type II variation (EMEA/H/C/004536/II/0011) ." Opinion adopted on 13.02.2020. Request for Supplementary Information adopted on 05.12.2019.	Positive Opinion adopted by consensus on 13.02.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Qutenza - capsaicin - EMEA/H/C/000909/II/0048 Grunenthal GmbH, Rapporteur: Bruno Sepodes, "Update of sections 4.2 and 5.1 of the SmPC in order to amend the posology based on PACE and STRIDE studies. The Package Leaflet is updated accordingly." Request for Supplementary Information adopted on 12.12.2019.	See agenda 9.1
Resolor - prucalopride - EMEA/H/C/001012/II/0049/G Shire Pharmaceuticals Ireland Limited, Rapporteur: Kristina Dunder, "Update of section 4.8 of the SmPC in order to update the safety information following the final results from study SHP555-802 (a cohort Study of the Relative Incidence of Major Cardiovascular Events) and based on an analysis of all potential major adverse cardiovascular events (MACE) from completed Phase 2/4 clinical studies in adult subjects. In addition, the Marketing authorisation holder (MAH) took the opportunity to update typographical errors in sections 4.4 and 5.1" Request for Supplementary Information adopted	

on 31.10.2019.	
Revatio - sildenafil - EMEA/H/C/000638/II/0086 Pfizer Europe MA EEIG, Rapporteur: Johann Lodewijk Hillege, "Update of sections 4.2 and 5.1 of the SmPC based on the final results from study A1481316; this is a multi-centre, randomised, placebo-controlled, double-blind, two-armed, parallel group study to evaluate efficacy and safety of iv sildenafil in the treatment of neonates with persistent pulmonary hypertension of the newborn (PPHN) or hypoxic respiratory failure and at risk for PPHN, with a long-term follow-up investigation of developmental progress 12 and 24 months after completion of study treatment. In addition some minor editorial updates to the Product Information are being proposed." Request for Supplementary Information adopted on 16.01.2020, 05.12.2019.	
RoActemra - tocilizumab - EMEA/H/C/000955/II/0091 Roche Registration GmbH, Rapporteur: Jan Mueller-Berghaus, "Update of sections 4.1, 4.2, 4.4, 4.8, 5.1 and 5.2 of the SmPC for RoActemra 162 mg solution for injection in pre-filled pen in order to align with the approved indications for RoActemra 162 mg solution for injection in pre-filled syringe to include active systemic juvenile idiopathic arthritis (sJIA) and juvenile idiopathic polyarthritis; the Package Leaflet is updated accordingly. In addition, the Marketing authorisation holder took the opportunity to make minor editorial changes in sections 3, 4.2, 4.4 and 5.1 of the SmPC for RoActemra 162 mg solution for injection in pre-filled syringe and the Annex II."	
Shingrix - herpes zoster vaccine (recombinant, adjuvanted) - EMEA/H/C/004336/II/0021 GlaxoSmithkline Biologicals SA, Rapporteur: Bart Van der Schueren, "C.I.13: Submission of the final report from study Zoster-063, listed as a category 3 study in the RMP version 2.0. The study was conducted to investigate the impact of reactogenicity on health-related quality of life in subjects ≥50 YOA following Shingrix vaccination." Request for Supplementary Information adopted on 13.02.2020.	Request for supplementary information adopted with a specific timetable.

Skyrizi - risankizumab - EMEA/H/C/004759/II/0008 AbbVie Deutschland GmbH & Co. KG, Rapporteur: Peter Kiely, "Update of SmPC 5.1 'Pharmacodynamic Properties' of the Skyrizi SmPC. The change pertains to the addition of information on retreatment after withdrawal of risankizumab to the summary of the IMMhance clinical study (M15-992)." Request for Supplementary Information adopted on 13.02.2020.	Request for supplementary information adopted with a specific timetable.
Soliris - eculizumab - EMEA/H/C/000791/II/0111, Orphan Alexion Europe SAS, Rapporteur: Jorge Camarero Jiménez, "C.I.4 Update of sections 4.2, 4.4, 4.5, 4.8, 5.1 and 5.2 (only editorial change) of the SmPC in order to update information on posology, warnings on infusion reactions, immunogenicity and concomitant use of other medicinal products, interactions and pharmacodynamics following procedures EMEA/H/C/000791/II/0105 and EMEA/H/C/000791/II/0107 displaying interim and final results from study ECU-NMO-302 and ECU-MG-302, respectively which are open label extension phase of pivotal RCT ECU-NMO-301 and ECU-MG-301 supporting indications for NMOSD AQP4+ and gMG respectively. Annex IID (to be aligned with RMP 19.3 approved in the EMEA/H/C/000791/II/0105) and the Package Leaflet have been updated accordingly."	
TAGRISSO - osimertinib - EMEA/H/C/004124/II/0032 AstraZeneca AB, Rapporteur: Jorge Camarero Jiménez, "Update of sections 4.4 and 4.8 of the SmPC in order to include erythema multiforme as an adverse drug reaction following the review of the MAH internal safety data. The Package Leaflet is updated accordingly. In addition, the MAH took the opportunity of this procedure to add the event frequency of Stevens-Johnson syndrome to align with the approved text in the SmPC." Request for Supplementary Information adopted on 13.02.2020.	Request for supplementary information adopted with a specific timetable.
Tecentriq - atezolizumab - EMEA/H/C/004143/II/0034 Roche Registration GmbH, Rapporteur: Sinan B. Sarac, "Update of section 5.1 of the SmPC in	Positive Opinion adopted by consensus on 13.02.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

order to include updated overall survival data from study IMvigor 211 (GO29294), a phase III study to investigate the efficacy and safety of atezolizumab compared with chemotherapy in patients with locally advanced or metastatic urothelial bladder cancer after failure of platinum-containing chemotherapy." Opinion adopted on 13.02.2020.	
Tecentriq - atezolizumab - EMEA/H/C/004143/II/0035 Roche Registration GmbH, Rapporteur: Sinan B. Sarac, "Update of sections 4.2, 4.8, 5.1 and 5.2 of the SmPC in order to reflect the results of study GO29664, (iMATRIX) evaluate the safety and pharmacokinetics of Tecentriq in paediatric (<18, n=69) and young adult patients (18-30 years, n=18) with relapsed or progressive solid tumours as well as with Hodgkin's and non-Hodgkin's lymphoma. This study was agreed under the Paediatric Investigational Plan EMEA-001638-PIP01-14-M02 (EMA decision: P/0207/2019). The Package Leaflet is updated accordingly." Opinion adopted on 13.02.2020.	Positive Opinion adopted by consensus on 13.02.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Vaxelis - diphtheria, tetanus, pertussis (acellular, component), hepatitis B (rDNA), poliomyelitis (inact.) and haemophilus type b conjugate vaccine (adsorbed) - EMEA/H/C/003982/II/0054 MCM Vaccine B.V., Rapporteur: Bart Van der Schueren, "Update of section 4.8 of the Vaxelis SmPC in order to add Hypotonic Hyporesponsive Episode to the list of post-marketing adverse events, based on a cumulative assessment of post-marketing data from the Marketing authorisation holder (MAH) global safety database. The Package Leaflet is updated accordingly. In addition, the MAH made minor editorial changes to the product information." Opinion adopted on 13.02.2020.	Positive Opinion adopted by consensus on 13.02.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Vemlidy - tenofovir alafenamide - EMEA/H/C/004169/II/0023 Gilead Sciences Ireland UC, Rapporteur: Janet Koenig, "Update of sections 4.4, 4.8, 5.1 and 5.2 of the SmPC based on interim analysis data at Week 24 from study GS-US-320-4035. This is a Phase 2, open-label study evaluating the efficacy and safety of tenofovir alafenamide (TAF) in virologically suppressed chronic hepatitis B	Request for supplementary information adopted with a specific timetable.

subjects with renal and/or hepatic impairment who switch from tenofovir disoproxil fumarate and/or other oral antiviral agents to TAF 25 mg once daily. Supportive safety and pharmacokinetic data are also provided from Study GS-US-292-1825 (phase 3b trial in which Elvitegravir/cobicistat/emtricitabine/tenofovir alafenamide an fixed-dose TAF containing combination indicated for the treatment of HIV-1 infection, was evaluated in HIV-1 infected subjects with end stage renal disease). Study GS-US-320-4035 is included in the RMP version 4.2 under additional PhV activities." Request for Supplementary Information adopted on 06.02.2020.	
VEYVONDI - vonicog alfa - EMEA/H/C/004454/II/0010 Baxalta Innovations GmbH, Rapporteur: Jan Mueller-Berghaus, "Variation to add hypersensitivity reactions (including anaphylaxis) in sections 4.4 and 4.8 of the SmPC." Request for Supplementary Information adopted on 16.01.2020.	
VITRAKVI - larotrectinib - EMEA/H/C/004919/II/0001 Bayer AG, Rapporteur: Filip Josephson, "Update of section 4.5 of the SmPC in order to remove the warning related to drug-drug interactions between larotrectinib and CYP2 substrates based on the results of study PH-40955 investigating the inductive potential of Larotrectinib on the expression of cytochrome P450 (CYP) enzymes. The Package Leaflet is updated accordingly." Request for Supplementary Information adopted on 13.02.2020.	Request for supplementary information adopted with a specific timetable.
Xaluprine - mercaptopurine - EMEA/H/C/002022/II/0022, Orphan Nova Laboratories Ireland Limited, Rapporteur: Filip Josephson, "Update of sections 4.4, 4.8 and 4.9 of the SmPC to add further information on hepatic toxicity. The MAH took the opportunity to implement minor editorial changes to the SmPC and PIL." Request for Supplementary Information adopted on 13.02.2020, 21.11.2019, 12.09.2019.	Request for supplementary information adopted with a specific timetable.
Zebinix - eslicarbazepine acetate - EMEA/H/C/000988/II/0074 Bial - Portela & Cª, S.A., Rapporteur: Martina	Request for supplementary information adopted with a specific timetable.

Weise, "Update of sections 4.2 and 5.2 of the SmPC in order to add the possibility to crush the tablets for patients unable to swallow whole tablets based on final results from study EP093-155; this is an interventional study to investigate and compare the relative bioavailability and bioequivalence of a crushed tablet and an intact tablet of ESL (800 mg); The Package Leaflet is updated accordingly." Request for Supplementary Information adopted on 13.02.2020.	
WS1683 **Elebrato Ellipta-EMEA/H/C/004781/ WS1683/0012** **Temybric Ellipta-EMEA/H/C/005254/ WS1683/0001** **Trelegy Ellipta-EMEA/H/C/004363/ WS1683/0010** GlaxoSmithKline Trading Services Limited, Lead Rapporteur: Peter Kiely, "Update of SmPC in order to add information in section 5.1 on survival data from the IMPACT study" Request for Supplementary Information adopted on 12.12.2019, 17.10.2019.	
WS1749 **AZILECT-EMEA/H/C/000574/WS1749/ 0084** **Rasagiline ratiopharm-EMEA/H/C/003957/ WS1749/0016** Teva B.V., Lead Rapporteur: Bruno Sepodes, Lead PRAC Rapporteur: Ana Sofia Diniz Martins, "Submission of the final report from study TV1030-CNS-50024 listed as a category 3 study in the RMP. This is a non-interventional retrospective cohort study which was conducted using the United States Medicare research database to assess the potential risk of melanoma associated with the use of rasagilline mesylate in patients with Parkinson's disease."	
WS1762 **Dovato-EMEA/H/C/004909/WS1762/0007** **Juluca-EMEA/H/C/004427/WS1762/0018** **Tivicay-EMEA/H/C/002753/WS1762/0055** **Triumeq-EMEA/H/C/002754/WS1762/ 0076** ViiV Healthcare B.V., Lead Rapporteur: Filip Josephson, "Update of sections 4.3 and 4.5 of the SmPC in order to add a new contraindication in relation to the co-administration of dolutegravir	Positive Opinion adopted by consensus on 06.02.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

with medicinal products with narrow therapeutic windows that are substrates of organic cation transporter 2 (OCT2), including but not limited to fampridine (also known as dalfampridine). The Package Leaflet is updated accordingly. The RMP has not been submitted. In addition, the Marketing authorisation holder (MAH) took the opportunity to remove the drug-drug interactions for products no longer authorised in the EU (boceprevir, dofetilide, nelfinavir), update the local MAH contacts in Belgium/Luxembourg, remove the inverted triangle for additional monitoring and add the date of first authorisation in the case of Dovato and add the date of last marketing authorisation renewal for Triumeq only." Opinion adopted on 06.02.2020.	

B.5.3. CHMP-PRAC assessed procedures

Baraclude - entecavir - EMEA/H/C/000623/II/0064 Bristol-Myers Squibb Pharma EEIG, Rapporteur: Filip Josephson, PRAC Rapporteur: Ulla Wändel Liminga, "Update of sections 4.8 and 5.1 of the SmPC in order to reflect the completion of the two paediatric studies AI463028 (Evaluation of the pharmacokinetics, safety, tolerability and efficacy of Entecavir (ETV) in pediatric subjects with chronic hepatitis B virus (HBV) infection who are HBeAg-Positive) and AI463189 (A Comparative Study of the Antiviral Efficacy and Safety of Entecavir (ETV) versus Placebo in Pediatric Subjects with Chronic Hepatitis B Virus (HBV) Infection who are HBeAg-Positive) and section 5.3 to reflect the outcome of study AI463080 (Randomized, Observational Study of Entecavir to Assess Long-term Outcomes Associated with Nucleoside/Nucleotide Monotherapy for Patients with Chronic HBV Infection: The REALM Study). Section 5.2 of the SmPC is also updated to remove information on the pharmacokinetics of entecavir in lamivudine-experienced paediatric patients, at the request of the CHMP. The RMP version 15 has also been submitted, which implements Revision 2 of the EU-RMP template. In addition, the Marketing authorisation holder (MAH) took the opportunity to bring the PI in line with the latest QRD template version 10.1 and to make minor editorial changes to the PI."	Request for supplementary information adopted with a specific timetable.

Request for Supplementary Information adopted on 13.02.2020.	
Bavencio - avelumab - EMEA/H/C/004338/II/0013 Merck Europe B.V., Rapporteur: Filip Josephson, PRAC Rapporteur: Hans Christian Siersted, "Update of section 5.1 of the SmPC in order to update efficacy information following results from study EMR100070-003 Part B listed as a specific obligation in the Annex II; this is a Phase II, open-label, multicenter trial to investigate the clinical activity and safety of avelumab (MSB0010718C) in subjects with Merkel cell carcinoma. With this submission the company is also taking the opportunity to update annex-II proposing deletion of the specific obligation and proposing the switch from conditional to full marketing authorisation. The package leaflet and the RMP (version 2.1) are updated accordingly."	See agenda 9.1
Benlysta - belimumab - EMEA/H/C/002015/II/0076 GlaxoSmithKline (Ireland) Limited, Rapporteur: Kristina Dunder, PRAC Rapporteur: Ulla Wändel Liminga, "Update of sections 4.4 and 4.8 of the SmPC in order to update the safety information based on the final results from study BEL115467 listed as an imposed PASS in the Annex II; this is a Randomized, Double-Blind, Placebo-Controlled 52-Week Study to Assess Adverse Events of Special Interest in Adults with Active, Autoantibody-Positive Systemic Lupus Erythematosus Receiving Belimumab; the Package Leaflet is updated accordingly. The RMP version 36 has also been submitted. The MAH took the opportunity to make the following minor updates to the RMP: - Updated information on transplant study, updated information on BEL116543, added information on paediatric continuation study in Module SIV. - Updated exposure information and information for BEL116543 in Module SIV.2. - Update data on revised rates of pregnancy and lactation in Module SIV.3. - Correction of an error within Annex 3 and provision of the updated protocol Amendment 7 of a Phase 3, Randomized, Double-Blind, Placebo-Controlled Study to Evaluate the Efficacy and Safety of belimumab plus Standard of Care versus Placebo plus standard of Care in Adult	See agenda 9.1

Subjects with Active Lupus Nephritis (BEL114054). This RMP also includes changes being reviewed in procedure II/0062, related to the use of Benlysta in paediatric patients, and BEL116027, evaluating the safety and efficacy of temporary discontinuation of belimumab. In addition, the Marketing authorisation holder took the opportunity make minor editorial changes to the Annex II and the label."	
Biktarvy - bictegravir / emtricitabine / tenofovir alafenamide - EMEA/H/C/004449/II/0027 Gilead Sciences Ireland UC, Rapporteur: Jean-Michel Race, PRAC Rapporteur: Liana Gross-Martirosyan, "Update of sections 4.8 and 5.1 of the Biktarvy SmPC to reflect pooled efficacy and safety data from the final clinical study reports of two antiretroviral therapy-naive adult studies through 144 weeks of treatment, GS-US-380-1489 (A Phase 3, Randomized, Double-Blind Study to Evaluate the Safety and Efficacy of GS-9883/Emtricitabine/ Tenofovir Alafenamide Versus Abacavir [ABC]/Dolutegravir [DTG]/Lamivudine [3TC] in HIV-1 Infected, Antiretroviral Treatment-Naive Adults) and GS-US-380-1490 (A Phase 3, Randomized, Double-Blinded Study to Evaluate the Safety and Efficacy of GS-9883/ Emtricitabine/Tenofovir Alafenamide Versus Dolutegravir + Emtricitabine/Tenofovir Alafenamide in HIV-1 Infected, Antiretroviral Treatment-Naive Adults). Both studies are listed as Category 3 studies in the RMP and this submission therefore fulfils MEA 001 and MEA 002. The RMP version 3.0 has also been submitted. In addition, the Marketing authorisation holder (MAH) took the opportunity to make some minor editorial changes to the PI and update Annex II with regards to PSUR requirements. Furthermore, the PI is brought in line with the latest QRD template version 10.1." Opinion adopted on 13.02.2020.	Positive Opinion adopted by consensus on 13.02.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Descovy - emtricitabine / tenofovir alafenamide - EMEA/H/C/004094/II/0044 Gilead Sciences Ireland UC, Rapporteur: Bruno Sepodes, PRAC Rapporteur: Ana Sofia Diniz Martins, "Submission of the final report from Study GS-US-311-1717 "A Phase 3b, Randomized, Double-Blind, Switch Study to	

Evaluate F/TAF in HIV-1 Infected Subjects who are Virologically Suppressed on Regimens Containing ABC/3TC" , listed as additional pharmacovigilance activity in the Descovy EU Risk Management Plan (RMP). This submission provides efficacy, clinical virology and safety data for virologically suppressed HIV-infected, who switch to regimens containing F/TAF from regimens containing abacavir (ABC)/lamivudine (3TC). No amendments are proposed to the Summary of Products Characteristics, product labelling and Patient Information Leaflet for the product. The RMP version 4.1 has been submitted." Request for Supplementary Information adopted on 12.12.2019.	
Gazyvaro - obinutuzumab - EMEA/H/C/002799/II/0036, Orphan Roche Registration GmbH, Rapporteur: Sinan B. Sarac, PRAC Rapporteur: Annika Folin, "Update of sections 4.8 and 5.1 of the SmPC based on data from the final CSR of the pivotal study GA04753g/GO01297/GADOLIN to fulfil a Category 3 PAM (MEA 006). The PL and RMP are updated accordingly." Opinion adopted on 13.02.2020. Request for Supplementary Information adopted on 28.11.2019.	Positive Opinion adopted by consensus on 13.02.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Gazyvaro - obinutuzumab - EMEA/H/C/002799/II/0038, Orphan Roche Registration GmbH, Rapporteur: Sinan B. Sarac, PRAC Rapporteur: Annika Folin, "Submission of final CSR for study MO28543/GREEN to fulfil the post authorization commitment [MEA] 005, the RMP is updated with the deletion of the study under PhV plan, (RMP version 6.1)" Request for Supplementary Information adopted on 13.02.2020.	Request for supplementary information adopted with a specific timetable.
Herceptin - trastuzumab - EMEA/H/C/000278/II/0158 Roche Registration GmbH, Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Brigitte Keller-Stanislawski, "Submission of the final report from study BO29159 (MetaPHER) a post-authorization safety measure Category 3 Non-Imposed Post-Approval Safety Study (NI-PASS), following approval of the Herceptin SC line extension procedure	Request for supplementary information adopted with a specific timetable.

EMEA/H/C/278/X/60 to generate and evaluate additional safety and tolerability data for the approved triplet regimen (Herceptin+Perjeta+docetaxel) in the advanced breast cancer setting. In addition, bioanalytical supportive studies are presented. An updated version of the Herceptin Risk Management Plan (version 21) has also been submitted." Request for Supplementary Information adopted on 13.02.2020.	
Incresync - alogliptin / pioglitazone - EMEA/H/C/002178/II/0029 Takeda Pharma A/S, Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Menno van der Elst, "Incresync RMP (version 10.1) includes the following updates: (i) MAH's proposal for removal of additional risk minimisation measures and consequently the Drug Utilisation Study along with removal of relevant commitment (EMEA/H/C/002182/LEG/009). (ii) RMP updated in the new template in order to implement the GVP Module V Revision 2 template along with revising/removal of the safety concerns as summarised below: · The safety concerns for the alogliptin component have been updated. · The safety concerns for the pioglitazone component have been aligned with the consolidated Pioglitazone family RMP (RMP V27.0 as part of the worksharing variation procedure [EMEA/H/C/XXXX/WS/1680]). · The safety concerns for alogliptin/pioglitazone FDC have been updated. (iii) Targeted Adverse Event (AE) Follow-up Questionnaires related to AEs of severe hypersensitivity skin reactions, hepatic events, pancreatitis, bladder cancer, bone fractures, and macular oedema were also removed. (iv) RMP has also been updated to reflect the removal of the inverted black triangle as agreed as part of alogliptin renewal procedure (EMEA/H/C/002182/R/0019). Annex II of the PI has been updated accordingly, as well as local representative for Poland." Opinion adopted on 13.02.2020. Request for Supplementary Information adopted on 28.11.2019.	Positive Opinion adopted by consensus on 13.02.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Kisqali - ribociclib - EMEA/H/C/004213/II/0021	

Novartis Europharm Limited, Rapporteur: Filip Josephson, PRAC Rapporteur: Hans Christian Siersted, "Update of sections 4.2 and 4.4 of the SmPC in order to add a warning on ILD/pneumonitis and related dose modification recommendations. The Package Leaflet is updated accordingly. The RMP version 4.0 has also been submitted."	
Lonsurf - trifluridine / tipiracil - EMEA/H/C/003897/II/0016 Les Laboratoires Servier, Rapporteur: Paula Boudewina van Hennik, PRAC Rapporteur: Annika Folin, "Update of sections 4.2, 4.4 and 5.2 of the SmPC in order to update information on patients with severe renal impairment based on final results from study TO-TAS-102-107 (A Phase 1, Open-label Study to Evaluate the Safety, Tolerability, and Pharmacokinetics of TAS-102 in Patients With Advanced Solid Tumors and Varying Degrees of Renal Impairment). The Package Leaflet is updated accordingly. The updated RMP version 6.3 has also been submitted. In addition, the Marketing authorisation holder (MAH) took the opportunity to bring the RMP in line with the template revision 2 of the good Pharmacovigilance practice module V guideline." Request for Supplementary Information adopted on 12.12.2019, 19.09.2019.	
Symkevi - tezacaftor / ivacaftor - EMEA/H/C/004682/II/0016, Orphan Vertex Pharmaceuticals (Ireland) Limited, Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Rhea Fitzgerald, "Update of sections 4.8 and 5.1 of the SmPC in order to update the information based on final results from study VX14-661-110 listed as a category 3 study in the RMP; this is a Phase 3, multicenter, open label, rollover study for Studies 103, 106, 107, 108, 109, 111, 112, and 114 designed to evaluate the long-term safety and tolerability of TEZ/IVA treatment for 96 weeks in cystic fibrosis (CF) subjects 12 years and older, homozygous or heterozygous for the F508del CFTR mutation. In addition, the Marketing authorisation holder (MAH) took the opportunity to bring the PI in line with the latest QRD template version 10.1. The RMP version 2.2 has also been submitted." Request for Supplementary Information adopted	Request for supplementary information adopted with a specific timetable.

on 13.02.2020.

Tamiflu - oseltamivir - EMEA/H/C/000402/II/0142

Roche Registration GmbH, Rapporteur: Outi Mäki-Ikola, PRAC Rapporteur: Kirsti Villikka, "Update of sections 4.2, 4.4, 4.8 and 5.1 of the SmPC following completion of the paediatric studies NV25719 and NV20234 and downstream population PK and PK/PD analysis, listed in the approved Tamiflu Paediatric Investigation Plan (PIP) (EMEA-000365-PIP01-08-M10); the study NV25719 was a prospective, open-label, randomized study which investigated PK and PD of two weight adjusted oseltamivir doses for the treatment of influenza-infected immunocompromised (IC) children less than 13 years of age. The study NV20234 was a prospective, double-blind, randomized trial which investigated safety and viral resistance to oseltamivir treatment in influenza-infected IC adults, adolescents and children. The purpose of this variation is to establish a dose recommendation for the treatment of paediatric IC patients.
The Package Leaflet and Labelling are updated accordingly. The updated RMP version 19 has also been submitted."
Request for Supplementary Information adopted on 17.10.2019.

Xultophy - insulin degludec / liraglutide - EMEA/H/C/002647/II/0034

Novo Nordisk A/S, Rapporteur: Kristina Dunder, PRAC Rapporteur: Menno van der Elst, "Update of section 4.2 of the SmPC in order to change the wording "transfer from basal insulin" to "transfer from any insulin regimen", based on data from study NN9068-4184 (DUAL II Japan: A double-blinded trial comparing the efficacy and safety of insulin degludec/liraglutide and insulin degludec both in combination with metformin in Japanese subjects with type 2 diabetes mellitus inadequately controlled with basal or pre-mix/combination insulin therapy and oral anti-diabetic drugs) as well as data from the post-marketing setting. In addition, the MAH took the opportunity to make a minor correction in SmPC section 5.1 and to implement changes in the annexes in accordance with QRD template 10.1.
The MAH provided an updated RMP version 9.0 as

part of the application."	
WS1704 **Alimta-EMEA/H/C/000564/WS1704/0058** **Pemetrexed Lilly-EMEA/H/C/004114/ WS1704/0010** Eli Lilly Nederland B.V., Lead Rapporteur: Alexandre Moreau, Lead PRAC Rapporteur: Ghania Chamouni, "Worksharing to update section 4.8 of the SmPC as requested by CHMP following the assessment of the PSUR covering the period between 05 February 2015 and 04 February 2018. To comply with SmPC guideline and latest QRD update, the Alimta and Pemetrexed Lilly SmPCs are updated combining multiple tables of ADRs into two tables: one for the ADRs reported in the pivotal registration trials and one for ADRs from the postmarketing period (both clinical trials and spontaneous reporting), organized by SOC with the respective frequency categories. The Package Leaflet is updated accordingly. In addition an updated RMP version 6.1 has been submitted to implement the revised GVP Module V (Rev 2) format as requested by CHMP following the assessment of the PSUR covering the period between 05 February 2015 and 04 February 2018." Request for Supplementary Information adopted on 28.11.2019.	

B.5.4. PRAC assessed procedures

PRAC Led **Ameluz - 5-aminolevulinic acid - EMEA/H/C/002204/II/0040** Biofrontera Bioscience GmbH, Rapporteur: Janet Koenig, PRAC Rapporteur: Martin Huber, PRAC-CHMP liaison: Janet Koenig, "To update the RMP for Ameluz to version 11.1 based on the new RMP template (GVP module V, rev.2), as well as the implementation of changes assessed and agreed by PRAC in the recently finalised PSUSA procedure (EMEA/H/C/002204/PSUSA/ 00010006/20180614)." Request for Supplementary Information adopted on 13.02.2020.	Request for supplementary information adopted with a specific timetable.
PRAC Led **Betmiga - mirabegron - EMEA/H/C/002388/II/0033** Astellas Pharma Europe B.V., Rapporteur: Maria	Request for supplementary information adopted with a specific timetable.

Concepcion Prieto Yerro, PRAC Rapporteur: Maria del Pilar Rayon, PRAC-CHMP liaison: Maria Concepcion Prieto Yerro, "Final study report of the PASS Study 178-CL-114 – An evaluation of Cardiovascular Events in Users of Mirabegron and Other Treatments for Overactive Bladder." Request for Supplementary Information adopted on 13.02.2020.	
PRAC Led **Cimzia - certolizumab pegol - EMEA/H/C/001037/II/0086** UCB Pharma S.A., Rapporteur: Kristina Dunder, PRAC Rapporteur: Ulla Wändel Liminga, PRAC-CHMP liaison: Kristina Dunder, "Submission of the final report from study (UP0038) listed as a category 3 study in the RMP. This is a non-interventional post-authorisation safety study with the aim to evaluate the effectiveness of Cimzia risk minimisation educational materials for healthcare professionals and patients." Opinion adopted on 13.02.2020.	Positive Opinion adopted by consensus on 13.02.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
PRAC Led **Colobreathe - colistimethate sodium - EMEA/H/C/001225/II/0044/G** Teva B.V., Rapporteur: Ewa Balkowiec Iskra, PRAC Rapporteur: Adam Przybylkowski, PRAC-CHMP liaison: Ewa Balkowiec Iskra, "Submission of the final Post-authorisation safety study report for CLB-MD-05: An observational safety study of Colobreathe (colistimethate sodium dry powder for inhalation) compared with other inhaled anti-pseudomonal antibiotics in cystic fibrosis patients using cystic fibrosis registries. The MAH is also providing an updated RMP, reflecting results from CLB-MD-05 but also the results form CLB-MD-08 that had been provided previously." Opinion adopted on 13.02.2020. Request for Supplementary Information adopted on 31.10.2019.	Positive Opinion adopted by consensus on 13.02.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
PRAC Led **Constella - linaclotide - EMEA/H/C/002490/II/0043** Allergan Pharmaceuticals International Limited, Rapporteur: Martina Weise, PRAC Rapporteur: Martin Huber, PRAC-CHMP liaison: Martina Weise, "Submission of the final report from study "Linaclotide Utilization Study in Selected	Request for supplementary information adopted with a specific timetable.

European Populations" listed as a category 3 study in the RMP. This is a Drug Utilisation Study (DUS) address following safety concerns - The potential for off-label use and abuse/excessive use - Extent of use in pregnancy and lactation, and male patients - Assess the extent of off-label use and the extent of use in males and in pregnant females" Request for Supplementary Information adopted on 13.02.2020, 28.11.2019.	
PRAC Led **Cubicin - daptomycin - EMEA/H/C/000637/II/0074** Merck Sharp & Dohme B.V., PRAC Rapporteur: Pernille Harg, PRAC-CHMP liaison: Bjorg Bolstad, "Submission of an updated RMP version 12.0 in order to delete all risks and additional risk minimisation measures, in line with GVP module V revision 2. Annex II of the Product Information is updated accordingly. In addition, the MAH took the opportunity to align the Product Information with the QRD template version 10.1 and update the list of local representatives." Opinion adopted on 13.02.2020. Request for Supplementary Information adopted on 31.10.2019.	Positive Opinion adopted by consensus on 13.02.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
PRAC Led **Fampyra - fampridine - EMEA/H/C/002097/II/0046** Biogen Netherlands B.V., Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Liana Gross-Martirosyan, PRAC-CHMP liaison: Johann Lodewijk Hillege, "Update of sections 4.2, 4.3, 4.4, 4.8 and 5.1 of the SmPC in order to update the existing contraindication for renal impaired patients, update the frequency of seizure to uncommon and reflect safety information based on final results from study 218MS401 (LIBERATE) listed as category 3 study in the RMP; Study 218MS401 was a Phase IV prospective, noninterventional, multicentre, observational study in MS patients who began Fampyra treatment in the postmarketing setting. The Package Leaflet is updated accordingly. An updated RMP version 13.1 has also been submitted to align with the RMP Rev. 2 template." Request for Supplementary Information adopted	Request for supplementary information adopted with a specific timetable.

on 13.02.2020.	
PRAC Led **Firmagon - degarelix - EMEA/H/C/000986/II/0035** Ferring Pharmaceuticals A/S, Rapporteur: Alexandre Moreau, PRAC Rapporteur: Ghania Chamouni, PRAC-CHMP liaison: Alexandre Moreau, "Submission of the FE 200486 CS39 Post Authorisation Safety Study (PASS) report; this was a Prospective Observational Safety Study in Patients with Advanced Prostate Cancer Treated with FIRMAGON (Degarelix) or a GnRH Agonist." Opinion adopted on 13.02.2020.	Positive Opinion adopted by consensus on 13.02.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
PRAC Led **Rebif - interferon beta-1a - EMEA/H/C/000136/II/0144** Merck Europe B.V., Rapporteur: Filip Josephson, PRAC Rapporteur: Ulla Wändel Liminga, PRAC-CHMP liaison: Filip Josephson, "C.I.11 for RMP: Submission of an updated RMP version 11 in order to comply with the Good Pharmacovigilance Practices (GVP) Module 5 RMP revision 2 requirements, and to ensure the appropriate time needed for effective review and analysis of all RMP Sections" Opinion adopted on 13.02.2020.	Positive Opinion adopted by consensus on 13.02.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
PRAC Led **Retacrit - epoetin zeta - EMEA/H/C/000872/II/0094** Pfizer Europe MA EEIG, Rapporteur: Martina Weise, PRAC Rapporteur: Martin Huber, PRAC-CHMP liaison: Martina Weise, "Pfizer's biosimilar epoetin zeta list of safety concerns has been aligned to the Innovator's Eprex (reference product, INN epoetin alfa)." Request for Supplementary Information adopted on 13.02.2020.	Request for supplementary information adopted with a specific timetable.
PRAC Led **Saxenda - liraglutide - EMEA/H/C/003780/II/0025** Novo Nordisk A/S, Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Menno van der Elst, PRAC-CHMP liaison: Johann Lodewijk Hillege, "Submission of the final report from study NN8022-4241, a retrospective drug utilisation study (DUS) undertaken to investigate patterns of use of liraglutide containing drugs in routine clinical practice, listed as a category 3 study in the RMP. An updated RMP version 31 was agreed	Positive Opinion adopted by consensus on 13.02.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

during the procedure." Opinion adopted on 13.02.2020.	
PRAC Led **SIRTURO - bedaquiline - EMEA/H/C/002614/II/0038, Orphan** Janssen-Cilag International NV, Rapporteur: Filip Josephson, PRAC Rapporteur: Ulla Wändel Liminga, PRAC-CHMP liaison: Filip Josephson, "To update the RMP (version 5.1) to revise the Summary of Safety Concerns for Sirturo in response to a request received from PRAC/CHMP in the context of the assessment of the Renewal (EMEA/H/C/002614/R/0035) of the Conditional Marketing Authorisation of SIRTURO. As requested by the PRAC/CHMP, data on co-administration of bedaquiline and HIV-protease inhibitors are also summarised. No changes are proposed to the Product Information of SIRTURO." Opinion adopted on 13.02.2020.	Positive Opinion adopted by consensus on 13.02.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
PRAC Led **Teysuno - tegafur / gimeracil / oteracil - EMEA/H/C/001242/II/0042** Nordic Group B.V., Rapporteur: Paula Boudewina van Hennik, PRAC Rapporteur: Menno van der Elst, PRAC-CHMP liaison: Johann Lodewijk Hillege, "Submission of an updated RMP version 9 in order to update safety specifications (re-classifying and removing risks from the list of important safety concerns as outlined in PSUSA/2875/201801)." Request for Supplementary Information adopted on 13.02.2020.	Request for supplementary information adopted with a specific timetable.
PRAC Led **VELCADE - bortezomib - EMEA/H/C/000539/II/0093** Janssen-Cilag International NV, Rapporteur: Daniela Melchiorri, PRAC Rapporteur: Amelia Cupelli, PRAC-CHMP liaison: Daniela Melchiorri, "Update of the RMP (finally agreed version 30.2) in line with the latest RMP template revision 2; as a consequence, Annex II of the PI is updated to reflect the removal of the additional risk minimisation activities (educational materials). In addition, the MAH took the opportunity to update the list of local representatives in the PL. Furthermore, the PI is being brought in line with the latest QRD template (version 10.1)." Opinion adopted on 13.02.2020.	Positive Opinion adopted by consensus on 13.02.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

Request for Supplementary Information adopted on 31.10.2019.	
PRAC Led **WS1747** **Enbrel-EMEA/H/C/000262/WS1747/0231** **LIFMIOR-EMEA/H/C/004167/WS1747/0025** Pfizer Europe MA EEIG, Lead Rapporteur: Maria Concepcion Prieto Yerro, Lead PRAC Rapporteur: Eva A. Segovia, PRAC-CHMP liaison: Maria Concepcion Prieto Yerro, "Submission of an updated RMP (version 7.0) in line with the latest GVP Module V RMP template (rev. 2) and revision of the list of safety concerns. In addition, the MAH took the opportunity to implement outcomes of previous procedures (type II variation EMEA/H/C/WS/1270 and PSUR EMEA/H/C/PSUSA/001295/201902) to remove or consolidate risks. The MAH is also removing the addendum to RMP version 6.3., making clinical and post-marketing data updates in the RMP and reflecting the completion of post-authorisation studies." Request for Supplementary Information adopted on 13.02.2020.	Request for supplementary information adopted with a specific timetable.

B.5.5. CHMP-CAT assessed procedures

Alofisel - darvadstrocel - EMEA/H/C/004258/II/0009, Orphan, ATMP Takeda Pharma A/S, Rapporteur: Lisbeth Barkholt, CHMP Coordinator: Kristina Dunder Request for Supplementary Information adopted on 08.11.2019.	
Alofisel - darvadstrocel - EMEA/H/C/004258/II/0010/G, Orphan, ATMP Takeda Pharma A/S, Rapporteur: Lisbeth Barkholt, CHMP Coordinator: Kristina Dunder Request for Supplementary Information adopted on 06.12.2019.	
Imlygic - talimogene laherparepvec - EMEA/H/C/002771/II/0036, ATMP Amgen Europe B.V., Rapporteur: Olli Tenhunen, CHMP Coordinator: Tuomo Lapveteläinen	
Imlygic - talimogene laherparepvec - EMEA/H/C/002771/II/0037, ATMP	

Amgen Europe B.V., Rapporteur: Olli Tenhunen, CHMP Coordinator: Tuomo Lapveteläinen
Yescarta - axicabtagene ciloleucel - EMEA/H/C/004480/II/0015, Orphan, ATMP Kite Pharma EU B.V., Rapporteur: Jan Mueller-Berghaus, CHMP Coordinator: Jan Mueller-Berghaus

B.5.6. CHMP-PRAC-CAT assessed procedures

Kymriah - tisagenlecleucel - EMEA/H/C/004090/II/0013/G, Orphan, ATMP Novartis Europharm Limited, Rapporteur: Rune Kjeken, CHMP Coordinator: Ingrid Wang, PRAC Rapporteur: Brigitte Keller-Stanislawski, "Submission of a group of 3 type II variations (C.I.4) to include: - Long-term efficacy and safety of Kymriah in relapsed/refractory DLBCL based on the 24 months follow-up results from study CCTL019C2201 (update of sections 4.4, 4.8, 5.1 and 5.2 of the SmPC) - Interim results from study CCTL019B2202 (update of sections 4.4, 4.8, 5.1 and 5.2 of the SmPC) - Interim results from study CCTL019B2205J (update section 5.2 of the SmPC) The Annex II and the Package Leaflet are updated accordingly. In addition, the Marketing authorisation holder (MAH) took the opportunity to clarify the wording of the indication to include patients over 25 years of age and to introduce some minor editorial corrections throughout the SmPC and the Package Leaflet. The RMP version 2.0 has also been submitted." Request for Supplementary Information adopted on 06.12.2019.

B.5.7. PRAC assessed ATMP procedures

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

WS1699 **Hexacima-EMEA/H/C/002702/WS1699/0093** **Hexaxim-EMEA/H/W/002495/WS1699/0098**

Hexyon-EMEA/H/C/002796/WS1699/0097 Sanofi Pasteur, Lead Rapporteur: Jan Mueller-Berghaus Request for Supplementary Information adopted on 16.01.2020.	
WS1729 **Juluca-EMEA/H/C/004427/WS1729/0020** **Tivicay-EMEA/H/C/002753/WS1729/0056** **Triumeq-EMEA/H/C/002754/WS1729/0077** ViiV Healthcare B.V., Lead Rapporteur: Filip Josephson Opinion adopted on 13.02.2020.	Positive Opinion adopted by consensus on 13.02.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1734 **Riarify-EMEA/H/C/004836/WS1734/0005** **Trimbow-EMEA/H/C/004257/WS1734/0010** **Trydonis-EMEA/H/C/004702/WS1734/0005** Chiesi Farmaceutici S.p.A., Lead Rapporteur: Janet Koenig, "To provide an updated Enviromental Risk Assessment (ERA) report." Request for Supplementary Information adopted on 12.12.2019.	
WS1738/G **Dovato-EMEA/H/C/004909/WS1738/0011/G** **Juluca-EMEA/H/C/004427/WS1738/0021/G** **Tivicay-EMEA/H/C/002753/WS1738/0057/G** **Triumeq-EMEA/H/C/002754/WS1738/0078/G** ViiV Healthcare B.V., Lead Rapporteur: Filip Josephson Opinion adopted on 20.02.2020.	Positive Opinion adopted by consensus on 20.02.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1746 **Biktarvy-EMEA/H/C/004449/WS1746/0026** **Descovy-EMEA/H/C/004094/WS1746/0045** **Genvoya-EMEA/H/C/004042/WS1746/0067** **Odefsey-EMEA/H/C/004156/WS1746/0044** **Vemlidy-EMEA/H/C/004169/WS1746/0022** Gilead Sciences Ireland UC, Lead Rapporteur:	Positive Opinion adopted by consensus on 06.02.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

Jean-Michel Race Opinion adopted on 06.02.2020.	
WS1751/G **Keppra-EMEA/H/C/000277/WS1751/0185/G** UCB Pharma S.A., Lead Rapporteur: Koenraad Norga	
WS1759/G **Blitzima-EMEA/H/C/004723/WS1759/0030/G** **Ritemvia-EMEA/H/C/004725/WS1759/0030/G** **Truxima-EMEA/H/C/004112/WS1759/0033/G** Celltrion Healthcare Hungary Kft., Lead Rapporteur: Sol Ruiz Opinion adopted on 06.02.2020.	Positive Opinion adopted by consensus on 06.02.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

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

bevacizumab - EMEA/H/C/005556
treatment of metastatic carcinoma of the colon or rectum, metastatic breast cancer and recurrence of platinum-sensitive epithelial ovarian, fallopian tube or primary peritoneal cancer. first-line treatment of patients with unresectable advanced, metastatic or recurrent non-small cell lung cancer. first line treatment of patients with advanced and/or metastatic renal cell cancer.

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

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

SCENESSE - afamelanotide -

EMEA/H/C/002548/S/0032, Orphan
Clinuvel Europe Limited, Rapporteur: Janet Koenig, PRAC Rapporteur: Martin Huber

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

Cotellic - cobimetinib - EMEA/H/C/003960/R/0019
Roche Registration GmbH, Rapporteur: Filip Josephson, Co-Rapporteur: Koenraad Norga, PRAC Rapporteur: Menno van der Elst

Entresto - sacubitril / valsartan - EMEA/H/C/004062/R/0031
Novartis Europharm Limited, Rapporteur: Johann Lodewijk Hillege, Co-Rapporteur: Kristina Dunder, PRAC Rapporteur: Anette Kirstine Stark

Hetlioz - tasimelteon - EMEA/H/C/003870/R/0018, Orphan
Vanda Pharmaceuticals Germany GmbH, Rapporteur: Jayne Crowe, Co-Rapporteur: Martina Weise, PRAC Rapporteur: Adam Przybylkowski

Imlygic - talimogene laherparepvec - EMEA/H/C/002771/R/0039, ATMP
Amgen Europe B.V., Rapporteur: Olli Tenhunen, Co-Rapporteur: Rune Kjeken, CHMP Coordinators: Tuomo Lapveteläinen and Ingrid Wang, PRAC Rapporteur: Brigitte Keller-Stanislawski

Kyprolis - carfilzomib - EMEA/H/C/003790/R/0044, Orphan
Amgen Europe B.V., Rapporteur: Jorge Camarero Jiménez, Co-Rapporteur: Alexandre Moreau, PRAC Rapporteur: Nikica Mirošević Skvrce

Nucala - mepolizumab - EMEA/H/C/003860/R/0031
GlaxoSmithKline Trading Services Limited, Rapporteur: Peter Kiely, PRAC Rapporteur: Brigitte Keller-Stanislawski

Omidria - phenylephrine / ketorolac - EMEA/H/C/003702/R/0015
Omeros Ireland Limited, Rapporteur: Jayne Crowe, Co-Rapporteur: Agnes Gyurasics, PRAC Rapporteur: Jan Neuhauser

Pemetrexed medac - pemetrexed - EMEA/H/C/003905/R/0008

medac Gesellschaft fur klinische Spezialpraparate mbH, Generic, Generic of Alimta, Rapporteur: Bart Van der Schueren, PRAC Rapporteur: Ghania Chamouni

Pemetrexed Sandoz - pemetrexed - EMEA/H/C/004011/R/0008
Sandoz GmbH, Generic, Generic of Alimta, Rapporteur: Bjorg Bolstad, PRAC Rapporteur: Ghania Chamouni

RAVICTI - glycerol phenylbutyrate - EMEA/H/C/003822/R/0034, Orphan
lmmedica Pharma AB, Rapporteur: Sinan B. Sarac, Co-Rapporteur: Jayne Crowe, PRAC Rapporteur: Ilaria Baldelli

Votubia - everolimus - EMEA/H/C/002311/R/0065, Orphan
Novartis Europharm Limited, Rapporteur: Janet Koenig, Co-Rapporteur: Kristina Dunder, PRAC Rapporteur: Martin Huber

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

Dupixent - dupilumab - EMEA/H/C/004390/II/0027
sanofi-aventis groupe, Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Kimmo Jaakkola, "Extension of indication to include atopic dermatitis patients from 6 years to 11 years. Consequently, the sections 4.1, 4.2, 4.8, 5.1 and 5.2 are updated. The PL is updated accordingly."

Imbruvica - ibrutinib - EMEA/H/C/003791/II/0059, Orphan
Janssen-Cilag International NV, Rapporteur: Filip Josephson, PRAC Rapporteur: Nikica Mirošević Skvrce, "Extension of indication in chronic lymphocytic leukaemia (CLL) to add combination with rituximab as follows: In combination with rituximab or obinutuzumab for the treatment of adult patients with previously untreated CLL.
This extension of the approved CLL indication is based on results from the Phase 3 Eastern Cooperative Oncology Group-American College of Radiology Imaging Network (ECOG ACRIN) Study

E1912 (also referred to as PCYC-1126e-CA).
The SmPC is revised to include information related to the new indication. The PL has been revised accordingly. Minor editorial changes have been implemented in Annex II and Annex IIIA. An updated RMP has been submitted."

Nordimet - methotrexate - EMEA/H/C/003983/II/0016

Nordic Group B.V., Rapporteur: Bruno Sepodes, PRAC Rapporteur: Martin Huber, "Extension of indication to include the treatment of mild to moderate Crohn's disease either alone or in combination with corticosteroids in patients refractory or intolerant to thiopurines for Nordimet;
as a consequence, sections 4.1, 4.2 and 5.1 of the SmPC are updated. The Package Leaflet is updated in accordance. The RMP version 5.0 has also been submitted. The MAH took the opportunity to update the RMP with changes related to GVP V version 2 template and the outcome of MTX referral."

NovoThirteen - catridecacog - EMEA/H/C/002284/II/0026/G

Novo Nordisk A/S, Rapporteur: Jean-Michel Race, PRAC Rapporteur: Ghania Chamouni, "Extension of indication to include treatment of bleeding episodes in patients with congenital factor XIII A-subunit deficiency as well as minor surgery based on the results of study NN1841-3868 and the PRO-RBDD registry. As a consequence, sections 4.1, 4.2, 4.4, 4.6, 5.1, 5.2 of the SmPC and the RMP version 15 has been submitted. Annex IID and the package leaflet have been updated accordingly.
Furthermore, the PI is brought in line with the latest QRD template version. Minor editorial updates have also been made."
Request for 1 year of market protection for a new indication (Article 14(11) of Regulation (EC) 726/2004)

OPDIVO - nivolumab - EMEA/H/C/003985/II/0080

Bristol-Myers Squibb Pharma EEIG,
Co-Rapporteur: Paula Boudewina van Hennik, PRAC Rapporteur: Brigitte Keller-Stanislawski, "Extension of indication to include treatment of adult patients with unresectable advanced, recurrent or metastatic oesophageal squamous

cell carcinoma (OSCC) after prior fluoropyrimidine- and platinum-based chemotherapy. As a consequence, sections 4.1, 4.2, 4.4, 4.8 and 5.1 of the SmPC are updated. The Package Leaflet is updated in accordance. Version 16.0 of the RMP has also been submitted."

Remsima - infliximab - EMEA/H/C/002576/II/0082

Celltrion Healthcare Hungary Kft., Rapporteur: Outi Mäki-Ikola, PRAC Rapporteur: Kimmo Jaakkola, "Extension of indication to add Crohn's disease, ulcerative colitis, ankylosing spondylitis, psoriatic arthritis and psoriasis to the Remsima SC pharmaceutical form to be in line with the IV formulation."

Shingrix - herpes zoster vaccine (recombinant, adjuvanted) - EMEA/H/C/004336/II/0022

GlaxoSmithkline Biologicals SA, Rapporteur: Bart Van der Schueren, Co-Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Sonja Hrabcik, "Extension of indication to include a new population for Shingrix; as a consequence, sections 4.1, 4.2, 4.4, 4.8 and 5.1 of the SmPC are updated in order to delete a warning and to add new safety and efficacy information. The Package Leaflet is updated in accordance. The RMP version 2.1 has also been submitted."

Spravato - esketamine - EMEA/H/C/004535/II/0001/G

Janssen-Cilag International N.V., Rapporteur: Martina Weise, Co-Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Kirsti Villikka, "C.I.6(a): Extension of indication to include a new indication for Spravato for the rapid reduction of depressive symptoms in adult patients with a moderate to severe depressive episode of MDD who have current suicidal ideation with intent. As a consequence, sections 4.1, 4.2, 4.4, 4.8 and 5.1 the SmPC are updated. The RMP version 2.1 has also been submitted.
B.II.e.5.a.2: Addition of a new pack size (multipack) of 24 nasal spray devices (multipack of 8 packs of 3 nasal spray devices) corresponding to 4 weeks of treatment in the new indication.
The Package Leaflet and labelling are updated in accordance. In addition, the Marketing

authorisation holder (MAH) took the opportunity to clarify the wording in Annex II.D."

WS1769
Iscover-EMEA/H/C/000175/WS1769/0140
Plavix-EMEA/H/C/000174/WS1769/0138
sanofi-aventis groupe, Lead Rapporteur: Bruno Sepodes, Lead PRAC Rapporteur: Marcia Sofia Sanches de Castro Lopes Silva, "Extension of indication to include adult patients with high risk Transient Ischemic Attack (TIA) (ABCD2 score ≥4) or minor Ischemic Stroke (IS) (NIHSS ≤3) within 24 hours of either the TIA or IS event. The new indication is based on the results of two double-blind, randomised, placebo-controlled phase III trials (studies POINT & CHANCE); as a consequence, sections 4.1, 4.2, 4.4 and 5.1 of the SmPC are updated, the PL is updated accordingly. Version 1.0 of the RMP has also been submitted."

WS1782
Lacosamide UCB-EMEA/H/C/005243/WS1782/0006
Vimpat-EMEA/H/C/000863/WS1782/0088
UCB Pharma S.A., Lead Rapporteur: Filip Josephson, Lead Co-Rapporteur: Daniela Melchiorri, Lead PRAC Rapporteur: Ulla Wändel Liminga, "Extension of indication to include the treatment as adjunctive therapy of primary generalised tonic-clonic seizures in adults, adolescents and children from 4 years of age with idiopathic generalised epilepsy for Lacosamide UCB and Vimpat; consequently, sections 4.1, 4.2, 4.8 and 5.1 of the SmPC are updated. The Package Leaflet is updated in accordance. The RMP version 15.0 has also been submitted. Furthermore, the PI is brought in line with the latest QRD template version 10.1. The MAH also takes the opportunity to align the PI of Lacosamide UCB with the PI of Vimpat and to implement some corrections in BG, CS, DA, FR, DE, HU, PL and ES."

B.6.8. CHMP assessed procedures scope: Pharmaceutical aspects

HyQvia - human normal immunoglobulin - EMEA/H/C/002491/II/0055
Baxalta Innovations GmbH, Rapporteur: Jan Mueller-Berghaus

Kiovig - human normal immunoglobulin - EMEA/H/C/000628/II/0098
Takeda Manufacturing Austria AG, Rapporteur: Jan Mueller-Berghaus

Mepsevii - vestronidase alfa - EMEA/H/C/004438/II/0013/G, Orphan
Ultragenyx Germany GmbH, Rapporteur: Johann Lodewijk Hillege

Ogivri - trastuzumab - EMEA/H/C/004916/II/0013
Mylan S.A.S, Rapporteur: Koenraad Norga

Ondexxya - andexanet alfa - EMEA/H/C/004108/II/0007
Portola Netherlands B.V., Rapporteur: Jan Mueller-Berghaus

Ondexxya - andexanet alfa - EMEA/H/C/004108/II/0010/G
Portola Netherlands B.V., Rapporteur: Jan Mueller-Berghaus

Orencia - abatacept - EMEA/H/C/000701/II/0137/G
Bristol-Myers Squibb Pharma EEIG, Rapporteur: Outi Mäki-Ikola

Privigen - human normal immunoglobulin - EMEA/H/C/000831/II/0157/G
CSL Behring GmbH, Rapporteur: Jan Mueller-Berghaus

Puregon - follitropin beta - EMEA/H/C/000086/II/0106/G
Merck Sharp & Dohme B.V., Rapporteur: Peter Kiely

Ritonavir Mylan - ritonavir - EMEA/H/C/004549/II/0007/G
Mylan S.A.S, Generic, Generic of Norvir, Rapporteur: John Joseph Borg

Shingrix - herpes zoster vaccine (recombinant, adjuvanted) - EMEA/H/C/004336/II/0026
GlaxoSmithkline Biologicals SA, Rapporteur: Bart Van der Schueren

Shingrix - herpes zoster vaccine (recombinant, adjuvanted) - EMEA/H/C/004336/II/0027/G
GlaxoSmithkline Biologicals SA, Rapporteur: Bart Van der Schueren

Vaxelis - diphtheria, tetanus, pertussis (acellular, component), hepatitis B (rDNA), poliomyelitis (inact.) and haemophilus type b conjugate vaccine (adsorbed) - EMEA/H/C/003982/II/0055
MCM Vaccine B.V., Rapporteur: Bart Van der Schueren

Zaltrap - aflibercept - EMEA/H/C/002532/II/0055/G
sanofi-aventis groupe, Rapporteur: Filip Josephson

WS1736/G
Elebrato Ellipta-EMEA/H/C/004781/ WS1736/0015/G
Temybric Ellipta-EMEA/H/C/005254/ WS1736/0003/G
Trelegy Ellipta-EMEA/H/C/004363/ WS1736/0013/G
GlaxoSmithKline Trading Services Limited, Lead Rapporteur: Peter Kiely

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

AUBAGIO - teriflunomide - EMEA/H/C/002514/II/0028
sanofi-aventis groupe, Rapporteur: Martina Weise, "To update section 4.6 of the SmPC with additional information in relation to human experience of use of teriflunomide during pregnancy, from an analysis of the data recorded in the global safety database and available sources (clinical trial cases, registries and cohort studies, literature and post-marketing pregnancy reports).
The MAH also took the opportunity to update sections 2 and 4.4 of the SmPC to align with the updated annex of the guideline excipients with regards to sodium.
The Labelling and Package Leaflet are updated accordingly."

Bridion - sugammadex - EMEA/H/C/000885/II/0036
Merck Sharp & Dohme B.V., Rapporteur: Outi Mäki-Ikola, "update of sections 4.2, 4.4, 4.8, 5.1 and 5.2 of the SmPC with information on morbidly obese patients (based on study report P146MK8616 - a phase 4 randomized, active-comparator controlled trial to study the efficacy and safety of sugammadex (MK-8616)

for the reversal of neuromuscular blockade induced by either rocuronium bromide or vecuronium bromide in morbidly obese subjects) and information related to the excipient sodium in accordance with the revised Annex to the EC guideline on excipients. The Patient Leaflet is updated accordingly. The MAH also took the opportunity to include the changes related to the new EMA QRD template version 10.1 and to implement some editorial changes."	
Brintellix - vortioxetine - EMEA/H/C/002717/II/0025 H. Lundbeck A/S, Rapporteur: Bart Van der Schueren, "Update of sections 4.2, 4.4, 4.8, 5.1 and 5.2 of the SmPC to reflect the outcomes of the paediatric clinical study 12710A (a paediatric efficacy and safety study in adolescent MDD patients) and the study 12708A (paediatric pharmacokinetics and tolerability study in children and adolescent patients with DSM-IV diagnosis of depressive and anxiety disorder). In addition, the MAH took the opportunity to propose minor amendments to the labelling and to update the list of local representatives in the Package Leaflet."	
Esmya - ulipristal acetate - EMEA/H/C/002041/II/0048 Gedeon Richter Plc., Rapporteur: Kristina Dunder, "Update of sections 4.4 and 4.8 of the SmPC in order to add a warning and update the safety information based on a new case of liver transplantation following exposure to Esmya; the Package Leaflet and Labelling are updated accordingly"	See 9.1
Faslodex - fulvestrant - EMEA/H/C/000540/II/0068 AstraZeneca AB, Rapporteur: Filip Josephson, "Update of sections 4.8 and 5.1 of the SmPC in order to update the safety and efficacy information based on the results from study Phase 3 Study A5481023 (PALOMA-3) a randomized controlled study of fulvestrant and palbociclib combination. In addition, the MAH took the opportunity to make a number of editorial changes to the PI to comply with the new QRD template v10.1 and the addition of the respective strength and pharmaceutical form to the corresponding Marketing Authorisation	

Number."

Gardasil 9 - human papillomavirus vaccine [types 6, 11, 16, 18, 31, 33, 45, 52, 58] (recombinant, adsorbed) - EMEA/H/C/003852/II/0037/G

MSD Vaccins, Rapporteur: Kristina Dunder, "Update of section 5.1 of the SmPC based on the 2nd interim report from studies V503-002-20 (MEA 005) and V503-021 (MEA 004) listed as a category 3 in the RMP and on final results from study V501-015-21-01 (qHPV); these are effectiveness and immunogenicity long-term follow-up (LTFU) studies from the 9-valent HPV and 4-valent HPV (qHPV) vaccines programs in women 16-26YOA. In addition, the Marketing authorisation holder (MAH) took the opportunity to update the list of local representatives in the Package Leaflet and to bring the PI in line with the latest QRD template version 10.1. In addition, one linguistic comment received from the Czech NCA on the PI during procedure EMEA/H/C/003852/II/033 will be implemented as well."

Herceptin - trastuzumab - EMEA/H/C/000278/II/0160

Roche Registration GmbH, Rapporteur: Jan Mueller-Berghaus, "Update of section 4.7 of the SmPC in order to add "dizziness and somnolence" to the recommendations on the effects on the patient's ability to drive and use machines. Update of section 4.8 of the SmPC to remove Herpes zoster, Erysipelas, Cellulitis Common, Sepsis, Thinking abnormal, Ataxia, Paresis, Brain oedema, Pericarditis, Bradycardia and Hepatic failure as adverse drug reactions. An update of the frequencies of adverse reactions is proposed in accordance to a change in the company core datasheet (CDS) for Herceptin: Anaphylactic reaction and Anaphylactic shock is changed to frequency Rare, Wheezing is changed to frequency Uncommon, Pneumonitis is changed to frequency Uncommon and Palpitation is changed to frequency Common. The MAH is taking the opportunity to update section 2 of the Herceptin PL to ensure compliance with the guidance on Excipients in the Labelling and Package Leaflet of medicinal products for Human Use (SANTE 2017-11668). The Package Leaflet is updated accordingly."

Kyprolis - carfilzomib - EMEA/H/C/003790/II/0043, Orphan

Amgen Europe B.V., Rapporteur: Jorge Camarero Jiménez, "Update of section 4.8 of the SmPC in order to include cardiomyopathy as a new adverse drug reaction with uncommon frequency following a signal evaluation triggered by a request from the Therapeutic Goods Administration (TGA) Australian authority. The RMP version 11.0 has also been submitted. In addition, the MAH took the opportunity to make some minor editorial changes to the PI."

Opsumit - macitentan - EMEA/H/C/002697/II/0035/G, Orphan

Janssen-Cilag International N.V., Rapporteur: Maria Concepcion Prieto Yerro, "Update of section 4.5 of the SmPC in order to update the drug-drug interaction information of macitentan with Breast cancer resistance protein (BCRP) substrate drugs based on final results from studies AC-055-122 and AC-055-123 ; these are single-center, open-label, one-sequence, two-treatment studies investigating the effect of macitentan at steady state on the pharmacokinetics of rosuvastatin and riociguat respectively in healthy male subjects. In addition, a minor editorial change was introduced in section 5.1."

Perjeta - pertuzumab - EMEA/H/C/002547/II/0048

Roche Registration GmbH, Rapporteur: Sinan B. Sarac, "Update of sections 4.2 and 4.4 of the SmPC in order to add safety information on elderly patients based on a safety review. The Package Leaflet is updated accordingly. In addition, the MAH took the opportunity to make minor amendments to section 4.7 of the SmPC and to update the PL in accordance with the excipient guideline and in line with the SmPC."

Tecentriq - atezolizumab - EMEA/H/C/004143/II/0040

Roche Registration GmbH, Rapporteur: Sinan B. Sarac, "Update of section 4.8 of the SmPC to include the adverse drug reactions hyperthyroidism and hypertension, identified in study IMvigor130. The package leaflet is proposed to be updated accordingly."

Xyrem - sodium oxybate - EMEA/H/C/000593/II/0088

UCB Pharma S.A., Rapporteur: Bruno Sepodes, "Update of sections 4.4. and 4.8 of the SmPC in order to update the safety information to add choking sensation; the Package Leaflet is updated accordingly."

WS1718
Eviplera-EMEA/H/C/002312/WS1718/0101
Odefsey-EMEA/H/C/004156/WS1718/0045
Gilead Sciences Ireland UC, Lead Rapporteur: Bruno Sepodes, "Update of section 4.6 of the Eviplera and Odefsey SmPCs in order to reflect rilpivirine data from the Antiretroviral Pregnancy Registry (APR) Interim Report issued in December 2019. The Eviplera Package Leaflet is updated in accordance. In addition, the Worksharing applicant (WSA) took the opportunity to bring the PI in line with the latest QRD template version 10.1 and to update the PI in line with the Annex to the European Commission guideline on `Excipients in the labelling and package leaflet of medicinal products for human use' (EMA/CHMP/302620/2017 Rev.1) for both products."

B.6.10. CHMP-PRAC assessed procedures

Orkambi - lumacaftor / ivacaftor - EMEA/H/C/003954/II/0055
Vertex Pharmaceuticals (Ireland) Limited, Rapporteur: Daniela Melchiorri, PRAC Rapporteur: Rhea Fitzgerald, "Update of SmPC section 4.8 following results of safety study in children. Additionally, the applicant took the opportunity to update the SmPC in line with the latest version of the QRD template v10.1. The PL is updated accordingly.
In addition, the RMP is updated and version 7,1 is submitted."

Palynziq - pegvaliase - EMEA/H/C/004744/II/0007/G, Orphan
BioMarin International Limited, Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Rhea Fitzgerald, "Update of sections 4.4, 4.8 and 5.1 of the SmPC based on final results from studies 1655-003, a long-term extension of a Phase 2, open-label, dose-finding

study and 165-302 a Phase 3, randomised, double-blind, placebo-controlled, four-arm, discontinuation study which are listed in the RMP as category 3 studies. The RMP version 2.0 has also been submitted. In addition, the SmPC was amended with minor editorial changes."	
Tremfya - guselkumab - EMEA/H/C/004271/II/0020 Janssen-Cilag International N.V., Rapporteur: Melinda Sobor, PRAC Rapporteur: Brigitte Keller-Stanislawski, "Update of sections 4.4 and 4.8 to include anaphylactic reactions. Additionally the RMP is updated"	

B.6.11. PRAC assessed procedures

PRAC Led **BeneFIX - nonacog alfa - EMEA/H/C/000139/II/0163** Pfizer Europe MA EEIG, PRAC Rapporteur: Brigitte Keller-Stanislawski, PRAC-CHMP liaison: Jan Mueller-Berghaus, "Update of the RMP to remove LETE (Less than therapeutic effect) as an important identified risk. In addition, in the light of GVP Module V Revision 2,1 the MAH proposes to remove patient populations that were previously identified as Missing information."	
PRAC Led **Ceplene - histamine dihydrochloride - EMEA/H/C/000796/II/0040** Noventia Pharma Srl, Rapporteur: Jayne Crowe, PRAC Rapporteur: Rhea Fitzgerald, PRAC-CHMP liaison: Jayne Crowe, "Submission of an updated RMP version 8.1 in order to include the information about the termination of the non-interventional study (NIS) of category 3 Ceplene-3290 and of the PAES Ceplene Cohort Study 3306. -RMP has been adapted to the new RMP template (GVP V rev.2, mandatory since 31 March 2018). -The safety concerns (Part II Modules SVII and SVIII, Part V, Part VI) have been changed. A new important potential risk, named "Drug effect decreased as a consequence of drug interaction", has been added to the list of safety concerns. -Part III and Part IV have been updated to include information about the termination of the non-interventional study (NIS) of category 3 Ceplene-3290 and of the PAES Ceplene Cohort	

Study 3306. -updated information about the other ongoing studies included in the Pharmacovigilance Plan, "Ceplene-3292" and "Ceplene-3298", have been included in Part III and related parts/modules. -Details about the Marketing Authorization Holder have been updated accordingly upon implementation of transfer from previous Marketing Authorisation Holder, Meda AB, (positive decision received 08 December 2017)"	
PRAC Led **Lemtrada - alemtuzumab - EMEA/H/C/003718/II/0031** Sanofi Belgium, Duplicate, Duplicate of Lemtrada (WD), Rapporteur: Mark Ainsworth, PRAC Rapporteur: Anette Kirstine Stark, PRAC-CHMP liaison: Sinan B. Sarac, "Submission of an update the RMP (version 7.0) incorporating all amendments and additional activities defined in the Article 20 referral procedure (EMEA/H/A-20/1483/C/3718/0028)."	
PRAC Led **Siklos - hydroxycarbamide - EMEA/H/C/000689/II/0045** Addmedica S.A.S., Rapporteur: Koenraad Norga, PRAC Rapporteur: Laurence de Fays, PRAC-CHMP liaison: Bart Van der Schueren, "Update of sections 4.2, 4.3, 4.4, 4.5, 4.6, 4.8 and 4.9 of the SmPC as a consequence of the final study results of the non-interventional cohort study ESCORT-HU (European Sickle Cell Disease Cohort – Hydroxyurea) and harmonisation with other HU products. In addition, Annex II is amended to delete the paragraph about the treatment guide for physicians. The PIL is updated in accordance with the changes to the SmPC. The RMP is updated to reflect the finalisation of the ESCORT-HU study."	
PRAC Led **Taxotere - docetaxel - EMEA/H/C/000073/II/0136/G** Sanofi Mature IP, Rapporteur: Alexandre Moreau, PRAC Rapporteur: Ghania Chamouni, PRAC-CHMP liaison: Alexandre Moreau, "C.I.3: Update of sections 4.4 and 4.8 of the SmPC to add a warning and safety information about tumour lysis syndrome based on a cumulative safety review requested as part of the last PSUR; The Package Leaflet is updated accordingly. In	

addition, the MAH took the opportunity to make minor corrections to the SmPC and update the list of local representatives in the Package Leaflet.
C.I.3: Update of section 4.8 of the SmPC to add safety information about myositis based on cumulative safety review requested as part of the last PSUR; the Package Leaflet is updated accordingly."

PRAC Led
WS1773
Exelon-EMEA/H/C/000169/WS1773/0128
Prometax-EMEA/H/C/000255/WS1773/0128
Novartis Europharm Limited, Lead Rapporteur: Alexandre Moreau, Lead PRAC Rapporteur: Ghania Chamouni, PRAC-CHMP liaison: Alexandre Moreau, "Submission of an updated RMP v 10.0 to reflect the results of the Drug Utilisation Study CENA713D2409 (submitted and assessed in variation WS-1557, opinion adopted in July 2019) and to reassess all important risks in accordance of GVP revision 2. In addition, as requested by the PRAC following the assessment of the PSUSA/00002654/201901, some safety concerns have been removed."

PRAC Led
WS1775
Renagel-EMEA/H/C/000254/WS1775/0114
Renvela-EMEA/H/C/000993/WS1775/0051
Sevelamer carbonate Winthrop-EMEA/H/C/003971/WS1775/0024
Genzyme Europe BV, Lead Rapporteur: Bart Van der Schueren, Lead PRAC Rapporteur: Laurence de Fays, PRAC-CHMP liaison: Bart Van der Schueren, "Submission of an updated RMP version 10 in order to remove the important potential risk "sevelamer crystals associated with serious gastrointestinal disorders" from the list of safety concerns in the RMP of sevelamer hydrochloride/carbonate products, as agreed by the CHMP during the procedure for the renewal of the marketing authorization for Sevelamer Carbonate Winthrop (EMEA/H/C/003971/R/0022)."

B.6.12. CHMP-CAT assessed procedures

B.6.13. CHMP-PRAC-CAT assessed procedures

B.6.14. PRAC assessed ATMP procedures

B.6.15. Unclassified procedures and worksharing procedures of type I variations

WS1770/G
Infanrix hexa-EMEA/H/C/000296/
WS1770/0271/G
GlaxoSmithkline Biologicals SA, Lead
Rapporteur: Bart Van der Schueren

WS1789/G
AZILECT-EMEA/H/C/000574/WS1789/
0086/G
Rasagiline ratiopharm-EMEA/H/C/003957/
WS1789/0018/G
Teva B.V., Lead Rapporteur: Bruno Sepodes

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. Timetables – 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.

Qualification of Biomarkers:

HTA:

G.2. Ongoing procedures

G.3. PRIME

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

G.3.1. List of procedures concluding at 24-27 February 2020 CHMP plenary:

G.3.2. List of procedures starting in February 2020 for March 2020 CHMP adoption of outcomes

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