11 November 2019
EMA/CHMP/606044/2019
Inspections, Human Medicines Pharmacovigilance and Committees Division

Committee for medicinal products for human use (CHMP)

Agenda for the meeting on 11-14 November 2019

Chair: Harald Enzmann – Vice-Chair: Bruno Sepodes

11 November 2019, 13:00 – 19:30, room 1C

12 November 2019, 08:30 – 19:30, room 1C

13 November 2019, 08:30 – 19:30, room 1C

14 November 2019, 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 11-14 November 2019. See November 2019 CHMP minutes (to be published post December 2019 CHMP meeting).

1.2. Adoption of agenda

CHMP agenda for 11-14 November 2019

1.3. Adoption of the minutes

CHMP minutes for 14 – 17 October 2019.

ORGAM Minutes from 4 November 2019

2. Oral Explanations

2.1. Pre-authorisation procedure oral explanations

2.1.1. clopidogrel / acetylsalicylic acid - EMEA/H/C/004996

indicated for the secondary prevention of atherothrombotic events

Scope: Possible oral explanation

Action: Possible oral explanation to be held on Wednesday 13 November 2019 at 11:00

List of Outstanding Issues adopted on 19.09.2019, 27.06.2019. List of Questions adopted on 31.01.2019.

2.1.2. enasidenib - Orphan - EMEA/H/C/004324

Celgene Europe BV; treatment of acute myeloid leukaemia (AML)

Scope: Oral explanation

Action: Oral explanation to be held on Tuesday 12 November 2019 at 11:00

List of Outstanding Issues adopted on 19.09.2019, 26.04.2019. List of Questions adopted on 18.10.2018.

2.1.3. plazomicin - EMEA/H/C/004457

treatment of complicated urinary tract infection (cUTI), including pyelonephritis; treatment of bloodstream infection (BSI); treatment of infections due to Enterobacteriaceae

Scope: Possible oral explanation

Action: Possible oral explanation to be held on Tuesday 12 November 2019 at 09:00

List of Outstanding Issues adopted on 25.07.2019. List of Questions adopted on 28.02.2019.

2.2. Re-examination procedure oral explanations

No items

2.3. Post-authorisation procedure oral explanations

2.3.1. Keytruda - pembrolizumab - EMEA/H/C/003820/II/0072

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 a new indication for Keytruda as monotherapy for the treatment of recurrent locally advanced or metastatic oesophageal cancer in adults whose tumours express PD L1 with a CPS ≥ 10 and who have received prior systemic therapy; as a consequence, sections 4.1, 4.2, and 5.1 of the SmPC, and section 1 of the PL are updated accordingly. The updated RMP version 25.1 has also been submitted."

Possible oral Explanation

Action: Possible oral explanation to be held on Wednesday 13 November 2019 at 09:00

Request for Supplementary Information adopted on 19.09.2019, 29.05.2019.

See 5.1

2.4. Referral procedure oral explanations

2.4.1. Lemtrada - Alemtuzumab - EMEA/H/A-20/1483

Sanofi Belgium

PRAC led Referral

PRAC Rapporteur: Brigitte Keller-Stanislawski; PRAC Co-rapporteur: Ulla Wändel Liminga

CHMP Rapporteurs Lemtrada: Rapporteur: Mark Ainsworth, Co-Rapporteur: Filip Josephson

Scope: Review of the benefit-risk balance following notification by European Commission of a referral under Article 20 of Regulation (EC) No 726/2004, based on pharmacovigilance data

Oral Explanation, PRAC Recommendation, CHMP opinion

Action: Oral explanation to be held on Tuesday 12 November 2019 at 14:15

See 10.1

3. Initial applications

3.1. Initial applications; Opinions

3.1.1. deferasirox - EMEA/H/C/005156

treatment of chronic iron overload

Scope: Opinion

Action: For adoption

List of Outstanding Issues adopted on 19.09.2019. List of Questions adopted on 28.03.2019.

3.1.2. osilodrostat - Orphan - EMEA/H/C/004821

Novartis Europharm Limited; treatment of Cushing's syndrome

Scope: Opinion

Action: For adoption

List of Outstanding Issues adopted on 17.10.2019, 25.07.2019. List of Questions adopted on 28.03.2019.

3.1.3. siponimod - EMEA/H/C/004712

treatment of secondary progressive multiple sclerosis (SPMS)

Scope: Opinion, SAG Neurology report

The list of experts for the SAG Neurology meeting held on 7 November was adopted via written procedure on 6 November 2019.

Action: For adoption

List of Outstanding Issues adopted on 19.09.2019, 29.05.2019. List of Questions adopted on 31.01.2019.

3.1.4. polatuzumab vedotin - Orphan - EMEA/H/C/004870

Roche Registration GmbH; treatment of mature B cell lymphomas

Scope: Opinion

Action: For adoption

List of Outstanding Issues adopted on 19.09.2019, 25.07.2019, 25.06.2019. List of Questions adopted on 24.04.2019.

3.1.5. solriamfetol - EMEA/H/C/004893

indicated to improve wakefulness in patients with narcolepsy or obstructive sleep apnoea.

Scope: Opinion

Action: For adoption

List of Outstanding Issues adopted on 25.07.2019. List of Questions adopted on 28.03.2019.

3.1.6. fostamatinib - EMEA/H/C/005012

indicated for the treatment of thrombocytopenia

Scope: Opinion

Action: For adoption

List of Outstanding Issues adopted on 19.09.2019, 27.06.2019. List of Questions adopted on 31.01.2019.

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

3.2.1. azacitidine - EMEA/H/C/005147

Treatment of myelodysplastic syndromes (MDS), chronic myelomonocytic leukemia (CMML) and acute myeloid leukemia (AML) and AML with >30% marrow blasts according to the WHO classification.

Scope: List of outstanding issues

Action: For adoption

List of Outstanding Issues adopted on 17.10.2019. List of Questions adopted on 29.05.2019.

3.2.2. cefiderocol - EMEA/H/C/004829

Treatment of infections due to aerobic Gram-negative bacteria

Scope: List of outstanding issues

Action: For adoption

List of Questions adopted on 25.06.2019.

3.2.3. insulin lispro - EMEA/H/C/005037

Treatment of diabetes mellitus in adults

Scope: List of outstanding issues

Action: For adoption

List of Questions adopted on 25.07.2019.

3.2.4. bempedoic acid - EMEA/H/C/004958

treatment of primary hypercholesterolaemia or mixed dyslipidaemia

Scope: List of outstanding issues

Action: For adoption

List of Questions adopted on 27.06.2019.

3.2.5. bempedoic acid / ezetimibe - EMEA/H/C/004959

treatment of primary hypercholesterolaemia or mixed dyslipidaemia

Scope: List of outstanding issues

Action: For adoption

List of Questions adopted on 27.06.2019.

3.2.6. darolutamide - EMEA/H/C/004790

treatment of non-metastatic castration resistant prostate cancer (nmCRPC)

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. abicipar pegol - EMEA/H/C/005103

treatment of neovascular (wet) age-related macular degeneration (AMD)

Scope: List of questions

Action: For adoption

3.3.2. amikacin - Orphan - EMEA/H/C/005264

Insmed Netherlands B.V.; treatment of lung infection as part of combination antibacterial drug regiment in adults

Scope: List of questions

Action: For adoption

3.3.3. avapritinib - Orphan - EMEA/H/C/005208

Blueprint Medicines (Netherlands) B.V.; treatment of gastrointestinal stromal tumours

Scope: List of questions

Action: For adoption

3.3.4. bevacizumab - EMEA/H/C/005106

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.

Scope: List of questions

Action: For adoption

3.3.5. ioflupane (^{123}I) - EMEA/H/C/005135

is indicated for detecting loss of functional dopaminergic neuron terminals in the striatum

Scope: List of questions

Action: For adoption

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

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

Scope: List of questions

Action: For adoption

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

3.4.1. trastuzumab - EMEA/H/C/005066

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

Scope: Letter from the applicant dated 08 October 2019 requesting an extension of clock stop to respond to the List of Questions adopted on 19.09.2019

Action: For adoption

List of Questions adopted on 19.09.2019.

3.4.2. selinexor - Orphan - EMEA/H/C/005127

Karyopharm Europe GmbH; treatment of patients with relapsed refractory multiple myeloma (RRMM)

Scope: Letter from the applicant dated 29 October 2019 requesting an extension to the clock stop to respond to the List of Outstanding Issues adopted on 19.09.2019

Action: For adoption

List of Outstanding Issues adopted on 19.09.2019. List of Questions adopted on 24.04.2019.

3.4.3. crizanlizumab - Orphan - EMEA/H/C/004874

Novartis Europharm Limited; Treatment of sickle cell disease

Scope: Letter from the applicant dated 04 November 2019 requesting an extension to the clock stop to respond to the List of Questions adopted on 17.10.2019

Action: For adoption

List of Questions adopted on 17.10.2019.

3.4.4. idebenone - Orphan - EMEA/H/C/005123

Santhera Pharmaceuticals (Deutschland) GmbH; treatment of respiratory dysfunction in patients with Duchenne muscular dystrophy (DMD) not using glucocorticoids

Scope: Letter from the applicant dated 05 November 2019 requesting an extension to the clock stop to respond to the List of Questions adopted on 17.10.2019

Action: For adoption

List of Questions adopted on 17.10.2019.

3.4.5. methylthioninium chloride - EMEA/H/C/002776

is indicated as an aid for the enhanced visualization and detection of colorectal lesions in adult patients undergoing screening / surveillance colonoscopy for colorectal cancer.

Scope: Letter from the applicant dated 06 November 2019 requesting an extension to the clock stop to respond to the List of Questions adopted on 27.06.2019.

Action: For adoption

List of Questions adopted on 27.06.2019.

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

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

Scope: Letter from the applicant dated 30 October 2019 requesting an extension to the clock stop to respond to the List of Questions adopted on 25.06.2019.

Action: For adoption

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

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; medium to long-term maintenance of alcohol abstinence and treatment of mild to moderate alcohol withdrawal syndrome

Scope: Appointment of re-examination Rapporteurs, draft timetable

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.

3.6. Initial applications in the decision-making phase

No items

3.7. Withdrawals of initial marketing authorisation application

3.7.1. ciprofloxacin - EMEA/H/C/004394

treatment of non-cystic fibrosis bronchiectasis (NCFBE) patients with chronic lung infection with Pseudomonas aeruginosa (P. aeruginosa)

Scope: Letter from the applicant dated 29 October 2019 informing about the withdrawal of marketing authorisation application

Action: For information

List of Outstanding Issues adopted on 28.03.2019. List of Questions adopted on 26.07.2018.

3.7.2. viable T-cells - Orphan - ATMP - EMEA/H/C/002397

Kiadis Pharma Netherlands B.V.; Adjunctive treatment in haemotopoietic stem cell transplantation (HSCT) for a malignant disease

Scope: Letter from the applicant dated 06 November 2019 informing about the withdrawal of marketing authorisation application

Action: For information

Oral explanation held on 11.10.2019. List of Outstanding Issues adopted on 21.06.2019, 14.09.2018, 25.05.2018. List of Questions adopted on 08.09.2017.

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. Akynzeo - fosnetupitant / netupitant / palonosetron - EMEA/H/C/003728/X/0018

Helsinn Birex Pharmaceuticals Limited

Rapporteur: Peter Kiely, Co-Rapporteur: Jean-Michel Race, PRAC Rapporteur: Ilaria Baldelli

Scope: “Extension application to introduce the new pharmaceutical form ‘powder for concentrate for solution for infusion’ and a new strength for the fixed combination of fosnetupitant (pro-drug of netupitant) and palonosetron of 235 mg/0.25 mg, to be administered intravenously (new route of administration).”

Action: For adoption

List of Outstanding Issues adopted on 19.09.2019. List of Questions adopted on 28.03.2019.

4.1.2. Humalog - insulin lispro - EMEA/H/C/000088/X/0169

Eli Lilly Nederland B.V.

Rapporteur: Kristina Dunder, PRAC Rapporteur: Annika Folin

Scope: Quality

Action: For adoption

List of Outstanding Issues adopted on 19.09.2019. List of Questions adopted on 26.04.2019.

4.1.3. Liprolog - insulin lispro - EMEA/H/C/000393/X/0130

Eli Lilly Nederland B.V.

Rapporteur: Kristina Dunder, PRAC Rapporteur: Annika Folin

Scope: “Extension application to change process steps. The RMP is updated (version 9.3) accordingly and in line with revision 2 of GVP module V on ‘Risk management systems’.”

Action: For adoption

List of Outstanding Issues adopted on 19.09.2019. List of Questions adopted on 26.04.2019.

4.1.4. Pemetrexed Fresenius Kabi - pemetrexed - EMEA/H/C/003895/X/0009

Fresenius Kabi Deutschland GmbH

Rapporteur: Bjorg Bolstad, PRAC Rapporteur: Ghania Chamouni

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

Action: For adoption

List of Outstanding Issues adopted on 19.09.2019. List of Questions adopted on 31.01.2019.

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

No items

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

4.3.1. Halimatoz - adalimumab - EMEA/H/C/004866/X/0013

Sandoz GmbH

Rapporteur: Milena Stain, PRAC Rapporteur: Ulla Wändel Liminga

Scope: "Extension application to add a new strength of 20mg (20mg/0.4ml) for Halimatoz solution for injection in pre-filled syringe.
The RMP (version 2.0) is updated in accordance. The MAH took also the opportunity to consolidate the RMP with changes approved in two other procedures (WS1565 and IB/11) and to align the PI with the latest QRD template (v.10.1)."

Action: For adoption

4.3.2. Hefiya - adalimumab - EMEA/H/C/004865/X/0013

Sandoz GmbH

Rapporteur: Milena Stain, PRAC Rapporteur: Ulla Wändel Liminga

Scope: "Extension application to add a new strength of 20mg (20mg/0.4ml) for Hefiya solution for injection in pre-filled syringe.
The RMP (version 2.0) is updated in accordance. The MAH took also the opportunity to consolidate the RMP with changes approved in two other procedures (WS1565 and IB/10) and to align the PI with the latest QRD template (v.10.1)."

Action: For adoption

4.3.3. Hyrimoz - adalimumab - EMEA/H/C/004320/X/0013

Sandoz GmbH

Rapporteur: Milena Stain, PRAC Rapporteur: Ulla Wändel Liminga

Scope: “Extension application to add a new strength of 20mg (20mg/0.4ml) for Hyrimoz solution for injection in pre-filled syringe.
The RMP (version 2.0) is updated in accordance. The MAH took also the opportunity to consolidate the RMP with changes approved in two other procedures (WS1565 and IB/11) and to align the PI with the latest QRD template (v.10.1).”

Action: For adoption

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

4.4.1. 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).”

List of Questions to the SAG Neurology

Action: For adoption

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

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. BLINCYTO - blinatumomab - Orphan - EMEA/H/C/003731/II/0030

Amgen Europe B.V.

Rapporteur: Alexandre Moreau, Co-Rapporteur: Daniela Melchiorri, PRAC Rapporteur: Eva Jirsová

Scope: “To modify the approved therapeutic indication to include the treatment of Philadelphia chromosome positive CD19 positive B-cell precursor acute lymphoblastic leukaemia (ALL) in adult and paediatric patients with relapsed or refractory ALL and adult patients in first or second complete remission with minimal residual disease (MRD) greater than or equal to 0.1%. As a consequence, sections 4.1, 4.2, 4.4, 4.8, 5.1 and 5.2 of the SmPC and the PL are updated accordingly. The updated RMP version 10.0 has also been submitted.”

Action: For adoption

5.1.2. Brilique - ticagrelor - EMEA/H/C/001241/II/0047/G

AstraZeneca AB

Rapporteur: Johann Lodewijk Hillege, Co-Rapporteur: Maria Concepcion Prieto Yerro, PRAC Rapporteur: Menno van der Elst

Scope: "Extension of indication to include, in co administration with acetylsalicylic acid (ASA), the prevention of atherothrombotic events in adult patients with coronary artery disease (CAD) and type 2 diabetes mellitus (T2DM) without a history of myocardial infarction who have undergone percutaneous coronary intervention (PCI) based on the final results of study D513BC00001 (THEMIS), a phase III multinational, randomised, double-blind, placebo controlled study to evaluate the effect of ticagrelor twice daily on the incidence of cardiovascular death, myocardial infarction or stroke in patients with T2DM; as a consequence, sections 4.1, 4.2, 4.4, 4.8 and 5.1 of the SmPC are updated.
Update of section 4.8 of the SmPC with new safety information on traumatic haemorrhages based on the final results from study D513BC00001 (THEMIS) and data from the ticagrelor clinical development programme and post-marketing data.
The Package Leaflet is updated in accordance. The RMP version 12 has also been submitted."

Action: For adoption

5.1.3. 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

5.1.4. Kadcyla - trastuzumab emtansine - EMEA/H/C/002389/II/0045

Roche Registration GmbH

Rapporteur: Sinan B. Sarac, PRAC Rapporteur: Hans Christian Siersted

Scope: "Extension of indication to include the adjuvant treatment of adult patients with HER2-positive early breast cancer, 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. In addition, the Applicant took the opportunity to introduce editorial changes." 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, 29.05.2019.

5.1.5. Keytruda - pembrolizumab - EMEA/H/C/003820/II/0072

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 a new indication for Keytruda as monotherapy for the treatment of recurrent locally advanced or metastatic oesophageal cancer in adults whose tumours express PD L1 with a CPS ≥ 10 and who have received prior systemic therapy; as a consequence, sections 4.1, 4.2, and 5.1 of the SmPC, and section 1 of the PL are updated accordingly. The updated RMP version 25.1 has also been submitted."

Action: Oral explanation to be held on Wednesday 13 November 2019 at 09:00

Request for Supplementary Information adopted on 19.09.2019, 29.05.2019.

See 2.3

5.1.6. 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 27.06.2019.

5.1.7. Revlimid - lenalidomide - Orphan - EMEA/H/C/000717/II/0107

Celgene Europe BV

Rapporteur: Alexandre Moreau, Co-Rapporteur: Filip Josephson, PRAC Rapporteur: Ghania Chamouni

Scope: "Extension of indication to include Revlimid in combination with rituximab (anti-CD20 antibody) for the treatment of adult patients with previously treated follicular lymphoma or marginal zone lymphoma.
As a consequence, sections 4.1, 4.2, 4.4, 4.8 and 5.1 of the SmPC are updated; the PL is updated in accordance. An updated EU RMP (version 36.2) has also been submitted."

Action: For adoption

Request for Supplementary Information adopted on 19.09.2019, 29.05.2019.

5.1.8. Trumenba - meningococcal group B vaccine (recombinant, adsorbed) - EMEA/H/C/004051/II/0013

Pfizer Europe MA EEIG

Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Jean-Michel Dogné

Scope: "Extension of indication for Trumenba to include active immunisation of children 1-9 years old. Sections 4.1, 4.2, 4.8 and 5.1 of the SmPC are updated in parallel based on the results from the two pivotal studies B1971017 and B1971035. The Package Leaflet is updated in accordance. The RMP version 2.0 has also been submitted.
In addition, the Marketing authorisation holder (MAH) took the opportunity to submit a corrected version of the final report of study B1971016, which was included in the initial marketing authorisation application."

Action: For adoption

Request for Supplementary Information adopted on 25.07.2019, 28.02.2019.

5.1.9. Ultomiris - ravulizumab - EMEA/H/C/004954/II/0002

Alexion Europe SAS

Rapporteur: Jorge Camarero Jiménez, Co-Rapporteur: Agnes Gyurasics, PRAC Rapporteur: Kimmo Jaakkola

Scope: "Extension of indication to include the treatment of patients with atypical haemolytic uremic syndrome (aHUS) for Ultomiris; as a consequence, sections 4.1, 4.2, 4.4, 4.8, 5.1, 5.2 and 6.6 of the SmPC are updated. The Package Leaflet is updated in accordance. In addition, Annex II.D is proposed to be updated to include the risk of thrombotic microangiopathy (TMA) with the new indication in the educational materials. The RMP version 1.6 has also been submitted."

Action: For adoption

5.1.10. 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

5.1.11. WS1372
OPDIVO - nivolumab - EMEA/H/C/003985/WS1372/0053
Yervoy - ipilimumab - EMEA/H/C/002213/WS1372/0057

Bristol-Myers Squibb Pharma EEIG

Lead Rapporteur: Jorge Camarero Jiménez, Lead Co-Rapporteur: Paula Boudewina van Hennik, PRAC Rapporteur: Brigitte Keller-Stanislawski

Scope: "Extension of indication to include first-line treatment of adult patients with metastatic Non-Small Cell Lung Carcinoma (NSCLC) for Opdivo and Yervoy; as a consequence, sections 4.1, 4.2, 4.4, 4.8, 5.1 and 5.2 of the SmPC are updated in order to add information from the pivotal study CA209227 (an open-label, randomised phase 3 trial of nivolumab, or nivolumab plus ipilimumab, or nivolumab plus platinum doublet chemotherapy versus platinum doublet chemotherapy in subjects with chemotherapy-naïve stage IV or recurrent NSCLC). The Package Leaflet and RMP (version 14.0 for Opdivo and version 21.0 for Yervoy) are updated in accordance. In addition, the MAH has taken the opportunity to introduce minor editorial and formatting revisions in the PI."

Action: For adoption

Request for Supplementary Information adopted on 13.12.2018, 26.07.2018.

5.1.12. WS1542
Bretaris Genuair - aclidinium - EMEA/H/C/002706/WS1542/0040
Eklira Genuair - aclidinium - EMEA/H/C/002211/WS1542/0040

AstraZeneca AB

Lead Rapporteur: Ewa Balkowiec Iskra, Lead Co-Rapporteur: Peter Kiely

Scope: "Extension of indication to include reduction of COPD exarcerbations for Eklira Genuair and Bretaris Genuair; as a consequence, sections 4.1, 4.4, 4.8 and 5.3 of the SmPC are updated. The Package Leaflet is updated in accordance.
In addition, the Worksharing applicant (WSA) took the opportunity to update the list of local representatives in the Package Leaflet for Bretaris Genuair and to implement minor editorial changes in section 4.4, 4.6, 5.3 of the SmPC and section 2 of the PL for both Eklira Genuair and Bretaris Genuair."

Action: For adoption

Request for Supplementary Information adopted on 19.09.2019, 26.04.2019.

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

5.2.1. Suliqua - insulin glargine / lixisenatide - EMEA/H/C/004243/II/0011

sanofi-aventis groupe

Rapporteur: Kristina Dunder, PRAC Rapporteur: Menno van der Elst

Scope: "Extension of indication to include "treatment of adults with insufficiently controlled

type 2 diabetes mellitus to improve glycaemic control as an adjunct to diet and exercise in addition to other oral medicinal products for the treatment of diabetes" based on the phase 3 Study EFC13794; a 26-week randomized, open-label, active controlled, parallel-group, study assessing the efficacy and safety of the insulin glargine/lixisenatide fixed ratio combination in adults with Type 2 Diabetes inadequately controlled on GLP-1 receptor agonist and metformin (alone or with pioglitazone and/or SGLT2 inhibitors), followed by a fixed ratio combination single-arm 26-week extension period.
As a consequence, sections 4.1, 4.2, 4.4, 4.8 and 5.1 of the SmPC are updated and the Package Leaflet is updated in accordance. In addition, the MAH took the opportunity to update the contact details of the local representatives in Denmark, the Netherlands and the UK in the Package Leaflet and to implement minor editorial changes in the annexes. An updated RMP version 4.0 was provided as part of the application."

Request by the applicant for an extension to the clock stop to respond to the Request for Supplementary Information adopted on 17.10.2019.

The CHMP agreed to the request by the applicant for an extension to the clock stop and adopted the new timetable by written procedure on 08 November 2019.

Action: For information

Request for Supplementary Information adopted on 17.10.2019, 27.06.2019.

5.2.2. Opsumit - macitentan - Orphan - EMEA/H/C/002697/II/0029

Janssen-Cilag International N.V.

Rapporteur: Maria Concepcion Prieto Yerro, Co-Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Eva A. Segovia

Scope: "Extension of indication to include treatment of patients with inoperable chronic thromboembolic pulmonary hypertension (CTEPH), based on the pivotal study MERIT-1 (AC-055E201), together with 6 months of efficacy and safety data (cut-off date 17 October 2017) from its ongoing open-label extension study MERIT-2 (AC-055E202), as well as a drug-drug interaction (DDI) study (AC-055-122) of macitentan and rosuvastatine, a DDI study (AC-055-123) of macitentan and riociguat, and observational data from the OPUS Registry (OPsumit USers Registry; cut-off date of 17 April 2018).
As a consequence, sections 4.1, 4.2, 4.4, 4.5, 4.8 and 5.1 are being updated and the Package Leaflet is being updated accordingly. In addition, the MAH took the opportunity to implement editorial changes and to align the annexes with the latest QRD template and to update the contact details of the local representatives in the Package Leaflet.
An updated RMP version 9.2 was provided as part of the application."

Letter from the MAH dated 8 November 2019 informing about the withdrawal of the extension of indication application

Action: For information

Request for Supplementary Information adopted on 28.03.2019, 13.12.2018.

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. belantamab mafodotin - H0004935

Treatment of Multiple Myeloma

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

Action: For adoption

8.1.2. fostemsavir - H0005011

indicated in combination with other antiretroviral agents for the treatment of HIV-1 infection in heavily treatment-experienced adults.

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. WS1587/G Abasaglar-EMEA/H/C/002835/WS1587/0028/G Humalog-EMEA/H/C/000088/WS1587/0178/G

Applicant: Eli Lilly Nederland B.V.

Lead rapporteur: Kristina Dunder

Scope: Type II variation. B.IV.z. to introduce an additional prefilled pen presentation for Abasaglar, solution for injection (EU/1/14/944/007, EU/1/14/944/008, EU/1/14/944/012, EU/1/14/944/013), Humalog, solution for injection (EU/1/96/007/002, EU/1/96/007/004, EU/1/96/007/020, EU/1/96/007/021 EU/1/96/007/023), Humalog Kwikpen solution for injection (EU/1/96/007/031, EU/1/96/007/032, EU/1/96/007/039, EU/1/96/007/040, EU/1/96/007/041, EU/1/96/007/042) and Humalog Junior Kwikpen, solution for injection (EU/1/96/007/043, EU/1/96/007/044, EU/1/96/007/045). The pack contains 5 pre-filled pens. Type IAIN B. II.e.5.a.1 to request the 2x5 multipack.
As a consequence, the following sections were updated 1, 4.2, 4.4, 6.2, 6.4, 6.5, 6.6, 8 of the SmPC in order to add new pre-filled pen presentation; the Package Leaflet and Labelling are updated accordingly. In addition, the Worksharing applicant (WSA) took the opportunity to make an editorial change (removing comma in SK address in the PL).

Action: For adoption

Request for Supplementary Information adopted on 19.09.2019

9.1.2. Tyverb - lapatinib - EMEA/H/C/000795/II/0059

Applicant: Novartis Europharm Limited

Rapporteur: Filip Josephson

Scope: Update of section 5.1 of the SmPC in order to update Table 8 based on updated/corrected results from study EGF114299/LAP016A2307, an interventional study with progression free survival rate as primary objective, original report submitted during procedure EMEA/H/C/00795/II/0051.

Action: For discussion

Request for Supplementary Information adopted on 26.04.2019.

9.1.3. ZOELY - Nomegestrol acetate, estradiol - EMEA/H/C/001213/II/0050

Applicant: Theramex Ireland Limited

CHMP Rapporteur: Jean-Michel Race; PRAC Rapporteur: Adrien Inoubli

Scope: Update of sections 4.3 and 4.4 of the SmPC in order to add a new contraindication and a new warning regarding meningioma, upon request by PRAC following the assessment of Post-authorisation measure “LEG 014”. The package leaflet is being updated accordingly. In addition, the MAH took the opportunity to update the contact details of the local representatives in the Netherlands and Portugal in the package leaflet.

CHMP request for PRAC advice

Action: For adoption

9.1.4. Cufence - trientine dihydrochloride - EMEA/H/C/004111/II/0002/G

Univar BV

Rapporteur: Milena Stain

Scope: “B.II.a.3.b.2) (type II)
B.II.b.4.b) (type IA)
B.II.b.3.a) (type IB)
B.II.a.1.a) (type IB)
B.II.d.1.a) (type IA)
B.II.f.1.d) (type IB
C.1.4. Update of sections updates of sections 4.5 and 5.2 of the SmPC in order to add information on food interaction and pk based on results from study TR-003 PK are proposed. In addition, the Marketing authorisation holder (MAH) took the opportunity to bring the PI in line with the latest QRD template.”

Action: For adoption

9.1.5. Zydelig - idelalisib - EMEA/H/C/003843/II/0047

Gilead Sciences Ireland UC

Rapporteur: Filip Josephson, PRAC Rapporteur: Martin Huber

Scope: “submission of the final clinical study report for study 101-09, A Phase 2 Study to Assess the Efficacy and Safety of Idelalisib in Subjects with Indolent BCell Non-Hodgkin Lymphomas Refractory to Rituximab and Alkylating Agents. This submission is an Annex II postauthorisation measure (ANX 002) and a category I commitment in the Zydelig Risk Management Plan (RMP).This submission also includes an update to the PI”
Request for Supplementary Information adopted on 19.09.2019.

Action: For adoption

9.1.6. Increlex - Mecasermin - EMEA/H/C/000704/II/0060

Ipsen Pharma

Rapporteur: Outi Mäki-Ikola, PRAC Rapporteur: Kirsti Villikka

Scope: Update of sections 4.1, 4.2, 4.3, 4.4, 4.8 and 4.9 of the SmPC in order to update the safety information on benign or malignant neoplasia based on the EU Registry Study, the Ipsen global safety database and literature review. The Package Leaflet is updated accordingly. The MAH also submitted the updated RMP version 11.
In addition, the MAH took the opportunity to update the list of local representatives in the Package Leaflet

Action: For adoption

Request for Supplementary Information adopted on 19.09.2019.

10. Referral procedures

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

10.1.1. Lemtrada - Alemtuzumab - EMEA/H/A-20/1483

Applicant: Sanofi Belgium

PRAC led Referral

PRAC Rapporteur: Brigitte Keller-Stanislawski; PRAC Co-rapporteur: Ulla Wändel Liminga

CHMP Rapporteurs Lemtrada: Rapporteur: Mark Ainsworth, Co-Rapporteur: Filip Josephson

Scope: Review of the benefit-risk balance following notification by European Commission of a referral under Article 20 of Regulation (EC) No 726/2004, based on pharmacovigilance data

Oral Explanation, PRAC Recommendation, CHMP opinion

Action: Oral explanation to be held on Tuesday 12 November 2019 at 14:15

See 2.4

10.1.2. Xeljanz - Tofacitinib - EMEA/H/A-20/1485

Applicant: Pfizer Europe MA EEIG

PRAC led Referral

PRAC Rapporteur: Liana Gross-Martirosyan; PRAC Co-rapporteur: Amelia Cupelli

CHMP Rapporteurs Xeljanz: Rapporteur: Daniela Melchiorri, Co-Rapporteur: Johann Lodewijk Hillege

Scope: Review of the benefit-risk balance following notification by the European Commission (EC) of a referral under Article 20 of Regulation (EC) No 726/2004, based on pharmacovigilance data

PRAC Recommendation, CHMP opinion

Action: For adoption

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

No items

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

No items

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

No items

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

No items

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

No items

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

No items

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

No items

11. Pharmacovigilance issue

11.1. Early Notification System

November 2019 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

Information related to briefing meetings taking place with applicants cannot be released at the present time as it is deemed to contain commercially confidential information

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 28-31 October 2019

Action: For information

List of Union Reference Dates and frequency of submission of Periodic Safety Update Reports (EURD list) for November 2019

Action: For adoption

14.2.2. Committee for Advanced Therapies (CAT)

CAT draft minutes of meeting held on 06-08 November 2019

Action: For information

14.2.3. Paediatric Committee (PDCO)

PIPs reaching D30 at November 2019 PDCO

Action: For information

Report from the PDCO meeting held on 12-15 November 2019

Action: For information

14.2.4. Committee for Orphan Medicinal Products (COMP)

Report from the COMP meeting held on 05-07 November 2019

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 12-14 November 2019

Action: For information

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

14.3.1. Scientific Advice Working Party (SAWP)

Chair: Anja Schiel

Report from the SAWP meeting held on 28-31 October 2019. 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.

14.3.2. Biologics Working Party (BWP)

Chair: Sol Ruiz/Nanna Aaby Kruse

Reports from BWP November 2019 meeting to CHMP for adoption:

- 14 reports on products in scientific advice and protocol assistance
- 4 reports on products in pre-authorisation procedures
- 4 reports on products in plasma master file

Action: For adoption

14.3.3. Antimicrobial Advice Ad Hoc Expert Group (AMEG)

Scope: Scientific advice on the AMEG categorisation of antimicrobials in the European Union; overview of comments

Background information: request from the EC for the update of the AMEG advice on the impact on public health and animal health of the use of antibiotics in animals (link); further extension for the deadline to submit the advice

Action: For discussion

14.3.4. Pharmacokinetics Working Party (PKWP)

Chair(s): TBC/Henrike Potthast

Election of PKWP chair

Jan Welink's second 3-year term expired in September 2019.

14.4. Cooperation within the EU regulatory network

No items

14.5. Cooperation with International Regulators

No items

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

No items

14.7. CHMP work plan

No items

14.8. Planning and reporting

No items

14.9. Others

No items

15. Any other business

15.1. AOB topic

15.1.1. Future-proofing EMA

Update on the Future-proofing EMA

Action: For information

15.1.2. Oncology Training

Action: For information

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)*

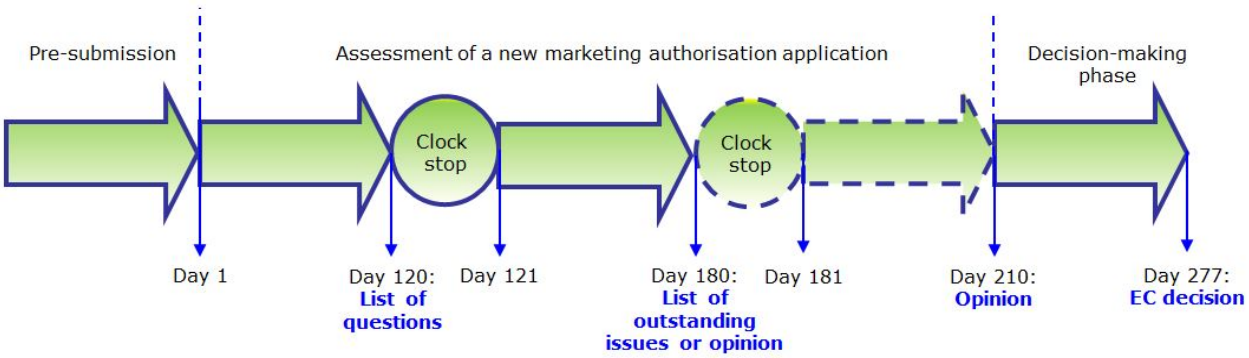
The items listed in this section are those for which marketing authorisation holders (MAHs) or applicants have been invited to the CHMP plenary meeting to address questions raised by the Committee. Oral explanations normally relate to on-going applications (section 3, 4 and 5) or referral procedures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

11 November 2019
EMA/CHMP/606074/2019

Annex to 11-14 November 2019 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 November 2019: **For adoption**

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

Final Outcome of Rapporteurship allocation for November 2019: **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

B.2. RENEWALS OF MARKETING AUTHORISATIONS OUTCOMES

Brineura - cerliponase alfa - EMEA/H/C/004065/S/0018, Orphan
BioMarin International Limited, Rapporteur: Martina Weise, PRAC Rapporteur: Ulla Wändel Liminga

IMVANEX - smallpox vaccine (live modified vaccinia virus Ankara) - EMEA/H/C/002596/S/0041
Bavarian Nordic A/S, Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Brigitte Keller-Stanislawski

Lojuxta - lomitapide - EMEA/H/C/002578/S/0036
Amryt Pharmaceuticals DAC, Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Menno van der Elst

Mepsevii - vestronidase alfa - EMEA/H/C/004438/S/0007, Orphan
Ultragenyx Germany GmbH, Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Eva A. Segovia

Naglazyme - galsulfase -
EMEA/H/C/000640/S/0078
BioMarin International Limited, Rapporteur: Fátima Ventura, PRAC Rapporteur: Ana Sofia Diniz Martins

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

Sivextro - tedizolid phosphate -
EMEA/H/C/002846/R/0031
Merck Sharp & Dohme B.V., Rapporteur: Bruno Sepodes, Co-Rapporteur: Filip Josephson, PRAC Rapporteur: Maria del Pilar Rayon
Request for Supplementary Information adopted on 19.09.2019.

B.2.2. Renewals of Marketing Authorisations for unlimited validity

Akynzeo - fosnetupitant / netupitant / palonosetron - EMEA/H/C/003728/R/0024
Helsinn Birex Pharmaceuticals Limited, Rapporteur: Peter Kiely, Co-Rapporteur: Jean-Michel Race, PRAC Rapporteur: Ilaria Baldelli
Request for Supplementary Information adopted on 17.10.2019.

Gardasil 9 - human papillomavirus vaccine [types 6, 11, 16, 18, 31, 33, 45, 52, 58] (recombinant, adsorbed) -
EMEA/H/C/003852/R/0035
MSD Vaccins, Rapporteur: Kristina Dunder, Co-Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Jean-Michel Dogné

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.

Jinarc - tolvaptan -
EMEA/H/C/002788/R/0027
Otsuka Pharmaceutical Netherlands B.V., Rapporteur: Daniela Melchiorri, Co-Rapporteur: Romaldas Mačiulaitis, PRAC Rapporteur: Amelia Cupelli

Mysimba - naltrexone hydrochloride /

bupropion hydrochloride - EMEA/H/C/003687/R/0033
Orexigen Therapeutics Ireland Limited, Rapporteur: Mark Ainsworth, Co-Rapporteur: Andrea Laslop, PRAC Rapporteur: Martin Huber
Request for Supplementary Information adopted on 19.09.2019.

Orbactiv - oritavancin - EMEA/H/C/003785/R/0027
Menarini International Operations Luxembourg S.A., Rapporteur: Janet Koenig, Co-Rapporteur: Kristina Dunder, PRAC Rapporteur: Adam Przybylkowski
Request for Supplementary Information adopted on 19.09.2019.

Prolia - denosumab - EMEA/H/C/001120/R/0082
Amgen Europe B.V., Rapporteur: Kristina Dunder, Co-Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Ulla Wändel Liminga

Synjardy - empagliflozin / metformin - EMEA/H/C/003770/R/0044
Boehringer Ingelheim International GmbH, Rapporteur: Johann Lodewijk Hillege, Co-Rapporteur: Daniela Melchiorri, PRAC Rapporteur: Eva A. Segovia

B.2.3. Renewals of Conditional Marketing Authorisations

Caprelsa - vandetanib - EMEA/H/C/002315/R/0041
Genzyme Europe BV, Rapporteur: Alexandre Moreau, PRAC Rapporteur: Ghania Chamouni
Request for Supplementary Information adopted on 17.10.2019.

Cometriq - cabozantinib - EMEA/H/C/002640/R/0032, Orphan
Ipsen Pharma, Rapporteur: Paula Boudewina van Hennik, Co-Rapporteur: Bjorg Bolstad, PRAC Rapporteur: Menno van der Elst

Holoclar - ex vivo expanded autologous human corneal epithelial cells containing stem cells - EMEA/H/C/002450/R/0026, Orphan, ATMP
Chiesi Farmaceutici S.p.A., Rapporteur: Egbert Flory, Co-Rapporteur: Paolo Gasparini, CHMP Coordinators: Jan Mueller-Berghaus and Daniela

Melchiorri, PRAC Rapporteur: Rhea Fitzgerald
Request for Supplementary Information adopted on 11.10.2019.

SIRTURO - bedaquiline - EMEA/H/C/002614/R/0035, Orphan
Janssen-Cilag International NV, Rapporteur: Filip Josephson, PRAC Rapporteur: Ulla Wändel Liminga

B.3. POST-AUTHORISATION PHARMACOVIGILANCE OUTCOMES

Signal detection

PRAC recommendations on signals adopted at the PRAC meeting held on 28-31 October 2019 PRAC

PSUR procedures for which PRAC adopted a recommendation for variation of the terms of the MA at its November 2019 meeting:

EMEA/H/C/PSUSA/00001751/201904
(insulin glargine)
CAPS:
Abasaglar (EMEA/H/C/002835) (insulin glargine), Eli Lilly Nederland B.V., Rapporteur: Kristina Dunder
Lantus (EMEA/H/C/000284) (insulin glargine), Sanofi-Aventis Deutschland GmbH, Rapporteur: Johann Lodewijk Hillege
Semglee (EMEA/H/C/004280) (insulin glargine), Mylan S.A.S, Rapporteur: Martina Weise
Toujeo (EMEA/H/C/000309) (insulin glargine), Sanofi-Aventis Deutschland GmbH, Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Menno van der Elst, “From: 20/04/2018 To: 20/04/2019”

EMEA/H/C/PSUSA/00002892/201903
(tenofovir disoproxil)
CAPS:
Tenofovir disoproxil Mylan
(EMEA/H/C/004049) (tenofovir disoproxil),
Mylan S.A.S, Rapporteur: Romaldas Mačiulaitis
Tenofovir disoproxil Zentiva
(EMEA/H/C/004120) (tenofovir disoproxil),
Zentiva k.s., Rapporteur: John Joseph Borg
Viread (EMEA/H/C/000419) (tenofovir disoproxil), Gilead Sciences Ireland UC, Rapporteur: Jean-Michel Race
NAPS:
TENOFOVIR FARMOZ - FARMOZ - SOCIEDADE TÉCNICO MEDICINAL, S.A.
PRAC Rapporteur: Adrien Inoubli, "31/03/2018 To: 30/03/2019"

EMEA/H/C/PSUSA/00010077/201903
(canagliflozin, canagliflozin / metformin)
CAPS:
Invokana (EMEA/H/C/002649) (canagliflozin), Janssen-Cilag International NV, Rapporteur: Martina Weise
Vokanamet (EMEA/H/C/002656) (canagliflozin / metformin), Janssen-Cilag International NV, Rapporteur: Martina Weise, PRAC Rapporteur: Martin Huber, "Period covered by the PSUR: 29 March 2018 to 28 March 2019"

EMEA/H/C/PSUSA/00010143/201903
(dimethyl fumarate (multiple sclerosis))
CAPS:
TECFIDERA (EMEA/H/C/002601) (dimethyl fumarate), Biogen Netherlands B.V., Rapporteur: Martina Weise, PRAC Rapporteur: Martin Huber, "25/03/2017 To: 25/03/2019"

EMEA/H/C/PSUSA/00010703/201904
(axicabtagene ciloleucel)
CAPS:
YESCARTA (EMEA/H/C/004480) (axicabtagene ciloleucel), Kite Pharma EU B.V., Rapporteur: Jan Mueller-Berghaus, CHMP Coordinator: Jan Mueller-Berghaus, PRAC Rapporteur: Anette Kirstine Stark, "From: 18/10/2018 To: 17/04/2019"

B.4. EPARs / WPARs

BAQSIMI - glucagon - EMEA/H/C/003848 For information only. Comments can be sent to Eli Lilly Nederland B.V., treatment of severe

hypoglycaemia, Known active substance (Article 8(3) of Directive No 2001/83/EC)	the PL in case necessary.
Ervebo - recombinant vesicular stomatitis virus - zaire ebolavirus vaccine (live) - EMEA/H/C/004554 Merck Sharp & Dohme B.V., Ebola Vaccine, 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.
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.
Pegfilgrastim Mundipharma - pegfilgrastim - EMEA/H/C/005312 Mundipharma Biologics S.L., treatment of neutropenia, 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.
Quofenix - delafloxacin - EMEA/H/C/004860 A. Menarini Industrie Farmaceutiche Riunite s.r.l., treatment of Acute Bacterial Skin and Skin Structure Infections (ABSSSI) in adults, 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.
Rinvoq - upadacitinib - EMEA/H/C/004760 AbbVie Deutschland GmbH & Co. KG, treatment of moderate to severe active rheumatoid arthritis, 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.
Spravato - esketamine - EMEA/H/C/004535 Janssen-Cilag International N.V., treatment-resistant depression, 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.
VANFLYTA - quizartinib - EMEA/H/C/004468, Orphan Daiichi Sankyo Europe GmbH, treatment of acute myeloid leukaemia, 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

AJOVY - fremanezumab - EMEA/H/C/004833/II/0002 TEVA GmbH, Rapporteur: Jan Mueller-Berghaus Opinion adopted on 24.10.2019. Request for Supplementary Information adopted on 25.07.2019.	Positive Opinion adopted by consensus on 24.10.2019. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Aripiprazole Mylan Pharma - aripiprazole - EMEA/H/C/003803/II/0012 Mylan S.A.S, Generic, Generic of Abilify, Rapporteur: Bjorg Bolstad Opinion adopted on 07.11.2019. Request for Supplementary Information adopted on 12.09.2019.	Positive Opinion adopted by consensus on 07.11.2019. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
CellCept - mycophenolate mofetil - EMEA/H/C/000082/II/0149/G Roche Registration GmbH, Rapporteur: Sinan B. Sarac	
Coagadex - human coagulation factor X - EMEA/H/C/003855/II/0023, Orphan BPL Bioproducts Laboratory GmbH, Rapporteur: Andrea Laslop Request for Supplementary Information adopted on 24.10.2019.	Request for supplementary information adopted with a specific timetable.
Eylea - aflibercept - EMEA/H/C/002392/II/0055/G Bayer AG, Rapporteur: Alexandre Moreau Request for Supplementary Information adopted on 07.11.2019.	Request for supplementary information adopted with a specific timetable.
Eylea - aflibercept - EMEA/H/C/002392/II/0058 Bayer AG, Rapporteur: Alexandre Moreau	
Kalydeco - ivacaftor - EMEA/H/C/002494/II/0080, Orphan Vertex Pharmaceuticals (Ireland) Limited, Rapporteur: Maria Concepcion Prieto Yerro Request for Supplementary Information adopted on 24.10.2019.	Request for supplementary information adopted with a specific timetable.
LIBTAYO - cemiplimab - EMEA/H/C/004844/II/0003 Regeneron Ireland Designated Activity Company (DAC), Rapporteur: Sinan B. Sarac Request for Supplementary Information adopted on 24.10.2019.	Request for supplementary information adopted with a specific timetable.
Natpar - parathyroid hormone - EMEA/H/C/003861/II/0020/G, Orphan	Request for supplementary information adopted with a specific timetable.

Shire Pharmaceuticals Ireland Limited, Rapporteur: Bart Van der Schueren Request for Supplementary Information adopted on 07.11.2019.	
Natpar - parathyroid hormone - EMEA/H/C/003861/II/0021, Orphan Shire Pharmaceuticals Ireland Limited, Rapporteur: Bart Van der Schueren	
NeuroBloc - botulinum toxin type B - EMEA/H/C/000301/II/0104/G Sloan Pharma S.a.r.l, Rapporteur: Bruno Sepodes Request for Supplementary Information adopted on 31.10.2019, 26.09.2019.	Request for supplementary information adopted with a specific timetable.
Ongentys - opicapone - EMEA/H/C/002790/II/0009 Bial - Portela & Cª, S.A., Rapporteur: Nithyanandan Nagercoil Opinion adopted on 24.10.2019. Request for Supplementary Information adopted on 14.02.2019, 13.09.2018, 26.04.2018.	Positive Opinion adopted by consensus on 24.10.2019. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Palynziq - pegvaliase - EMEA/H/C/004744/II/0002, Orphan BioMarin International Limited, Rapporteur: Johann Lodewijk Hillege	
Pelgraz - pegfilgrastim - EMEA/H/C/003961/II/0013/G Accord Healthcare S.L.U., Rapporteur: Sol Ruiz Request for Supplementary Information adopted on 31.10.2019.	Request for supplementary information adopted with a specific timetable.
Pemetrexed Hospira - pemetrexed - EMEA/H/C/003970/II/0020/G Pfizer Europe MA EEIG, Generic, Generic of Alimta, Rapporteur: Alar Irs Request for Supplementary Information adopted on 24.10.2019, 12.09.2019.	Request for supplementary information adopted with a specific timetable.
Praluent - alirocumab - EMEA/H/C/003882/II/0049/G sanofi-aventis groupe, Rapporteur: Johann Lodewijk Hillege	
Prevenar 13 - pneumococcal polysaccharide conjugate vaccine (13-valent, adsorbed) - EMEA/H/C/001104/II/0180/G Pfizer Europe MA EEIG, Rapporteur: Kristina Dunder Request for Supplementary Information adopted	Request for supplementary information adopted with a specific timetable.

on 07.11.2019, 12.09.2019.	
RotaTeq - rotavirus vaccine (live, oral) - EMEA/H/C/000669/II/0079/G MSD Vaccins, Rapporteur: Kristina Dunder Request for Supplementary Information adopted on 12.09.2019.	
Synflorix - pneumococcal polysaccharide conjugate vaccine (adsorbed) - EMEA/H/C/000973/II/0141 GlaxoSmithkline Biologicals SA, Rapporteur: Kristina Dunder Opinion adopted on 07.11.2019.	Positive Opinion adopted by consensus on 07.11.2019. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Tepadina - thiotepa - EMEA/H/C/001046/II/0034, Orphan ADIENNE S.r.l., Rapporteur: Alexandre Moreau Request for Supplementary Information adopted on 19.09.2019.	
Tepadina - thiotepa - EMEA/H/C/001046/II/0035/G, Orphan ADIENNE S.r.l., Rapporteur: Alexandre Moreau Request for Supplementary Information adopted on 19.09.2019.	
Xofigo - radium-223 - EMEA/H/C/002653/II/0037 Bayer AG, Rapporteur: Janet Koenig Opinion adopted on 31.10.2019.	Positive Opinion adopted by consensus on 31.10.2019. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Zessly - infliximab - EMEA/H/C/004647/II/0009/G Sandoz GmbH, Rapporteur: Bjorg Bolstad Opinion adopted on 24.10.2019.	Positive Opinion adopted by consensus on 24.10.2019. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1587/G **Abasaglar-EMEA/H/C/002835/WS1587/0028/G** **Humalog-EMEA/H/C/000088/WS1587/0178/G** Eli Lilly Nederland B.V., Lead Rapporteur: Kristina Dunder"Type II variation. B.IV.z. to introduce an additional prefilled pen presentation for Abasaglar, solution for injection (EU/1/14/944/007, EU/1/14/944/008, EU/1/14/944/012, EU/1/14/944/013), Humalog, solution for injection (EU/1/96/007/002, EU/1/96/007/004, EU/1/96/007/020, EU/1/96/007/021 EU/1/96/007/023), Humalog Kwikpen solution for injection (EU/1/96/007/031, EU/1/96/007/032, EU/1/96/007/039, EU/1/96/007/040, EU/1/96/007/041,	See 9.1

EU/1/96/007/042) and Humalog Junior Kwikpen, solution for injection (EU/1/96/007/043, EU/1/96/007/044, EU/1/96/007/045). The pack contains 5 pre-filled pens. Type IAIN B. II.e.5.a.1 to request the 2x5 multipack. As a consequence, the following sections were updated 1, 4.2, 4.4, 6.2, 6.4, 6.5, 6.6, 8 of the SmPC in order to add new pre-filled pen presentation; the Package Leaflet and Labelling are updated accordingly. In addition, the Worksharing applicant (WSA) took the opportunity to make an editorial change (removing comma in SK address in the PL)" Request for Supplementary Information adopted on 19.09.2019.	
WS1612/G **Herceptin-EMEA/H/C/000278/WS1612/0155/G** **Kadcyla-EMEA/H/C/002389/WS1612/0047/G** Roche Registration GmbH, Lead Rapporteur: Jan Mueller-Berghaus Opinion adopted on 07.11.2019. Request for Supplementary Information adopted on 12.09.2019.	Positive Opinion adopted by consensus on 07.11.2019. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1678 **Rixathon-EMEA/H/C/003903/WS1678/0027** **Riximyo-EMEA/H/C/004729/WS1678/0028** Sandoz GmbH, Lead Rapporteur: Jan Mueller-Berghaus Request for Supplementary Information adopted on 19.09.2019.	
WS1700/G **Humalog-EMEA/H/C/000088/WS1700/0180/G** **Liprolog-EMEA/H/C/000393/WS1700/0141/G** Eli Lilly Nederland B.V., Informed Consent of Humalog, Lead Rapporteur: Kristina Dunder Request for Supplementary Information adopted on 24.10.2019.	Request for supplementary information adopted with a specific timetable.

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

Abraxane - paclitaxel - EMEA/H/C/000778/II/0097	

Celgene Europe BV, Rapporteur: Paula Boudewina van Hennik, “Update of sections 4.2, 4.5, 4.8, 5.1 and 5.2 of the SmPC based on the results of study ABI-007-PST-001. This was a phase 1/2, multicenter, open-label, dose-finding study to assess the safety, tolerability and efficacy of weekly abraxane in paediatric patients with recurrent or refractory solid tumours, listed in the PIP, submitted in order to fulfil Article 46.”	
Advagraf - tacrolimus - EMEA/H/C/000712/II/0054 Astellas Pharma Europe B.V., Rapporteur: Jayne Crowe, “Update of section 4.2 of the SmPC, to reinforce the existing wording regarding the switching between different oral formulations of tacrolimus with different release characteristics.” Opinion adopted on 24.10.2019. Request for Supplementary Information adopted on 12.09.2019, 14.06.2019.	Positive Opinion adopted by consensus on 24.10.2019. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Afstyla - lonoctocog alfa - EMEA/H/C/004075/II/0024 CSL Behring GmbH, Rapporteur: Jan Mueller-Berghaus, “Update of sections 4.4 and 4.8 of the SmPC with safety information regarding the development of factor VIII inhibitors in patients treated with Afstyla based on clinical trial and post-marketing data reviewed recently with data lock point 03 January 2019. The PL is updated accordingly. Additionally, local representatives' details for Bulgaria and Croatia have been updated.” Request for Supplementary Information adopted on 10.10.2019.	
Aptivus - tipranavir - EMEA/H/C/000631/II/0083/G Boehringer Ingelheim International GmbH, Rapporteur: Jean-Michel Race, “C.1.4 (type II) - Update section 5.3 of the SmPC in order to update the preclinical safety information based on the outcome of ICH M7 assessment which identified 5-trifluoro-2-methylpyridinol (5-TFMP), a class 2 mutagen. B.I.b.1.h (type IB) B.II.b.2.a (type IA) B.II.d.1.g (type IB) B.II.f.1.a.1 (type IB) B.II.f.1.d (type IB) B.II.f.1.e (type IB) . The Package Leaflet and Labelling are updated	

accordingly.”	
Brintellix - vortioxetine - EMEA/H/C/002717/II/0022/G H. Lundbeck A/S, Rapporteur: Bart Van der Schueren, “Update of sections 4.8 and 5.1 of the SmPC in order to describe effects of vortioxetine on treatment-emergent sexual dysfunction based on the outcome of 2 prospective clinical studies (Studies 318 and 4001). Update of sections 4.4 and 5.2 of the SmPC in order to reflect the outcome of study 401 in subjects with severe hepatic impairment.” Request for Supplementary Information adopted on 27.06.2019.	
Cufence - trientine dihydrochloride - EMEA/H/C/004111/II/0002/G Univar BV, Rapporteur: Milena Stain “B.II.a.3.b.2) (type II) B.II.b.4.b) (type IA) B.II.b.3.a) (type IB) B.II.a.1.a) (type IB) B.II.d.1.a) (type IA) B.II.f.1.d) (type IB C.1.4. Update of sections updates of sections 4.5 and 5.2 of the SmPC in order to add information on food interaction and pk based on results from study TR-003 PK are proposed. In addition, the Marketing authorisation holder (MAH) took the opportunity to bring the PI in line with the latest QRD template.”	See 9.1
Deltyba - delamanid - EMEA/H/C/002552/II/0037, Orphan Otsuka Novel Products GmbH, Rapporteur: Koenraad Norga, “C.I.13 MIC report as amendment to CSR 242-09-213.” Opinion adopted on 24.10.2019. Request for Supplementary Information adopted on 11.07.2019.	Positive Opinion adopted by consensus on 24.10.2019. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Dovato - dolutegravir / lamivudine - EMEA/H/C/004909/II/0001 ViiV Healthcare B.V., Rapporteur: Filip Josephson, “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 dolutegavir. The SmPC is updated accordingly. The RMP is not submitted."	
Eurartesim - piperaquine tetraphosphate / artenimol - EMEA/H/C/001199/II/0036 Alfasigma S.p.A., Rapporteur: Janet Koenig, "Changes to sections 4.2, 4.4 and 4.6 of the SmPC with reference to the posology and the recommendation during pregnancy; sections 2 and 3 of the leaflet (PL) are amended accordingly and reference to the pregnancy register deleted from Annex II." Request for Supplementary Information adopted on 07.11.2019, 19.09.2019.	Request for supplementary information adopted with a specific timetable.
Faslodex - fulvestrant - EMEA/H/C/000540/II/0067 AstraZeneca AB, Rapporteur: Filip Josephson, "To update a warning in section 4.6 of the SmPC following an overview of non-clinical data, clinical pharmacology simulation/modelling data, supporting documentation and safety data. The Package Leaflet is updated accordingly. In addition, the applicant has taken the opportunity to correct a minor mistake in the address of one of the manufacturers responsible for batch release in Annex II and PL." Opinion adopted on 07.11.2019.	Positive Opinion adopted by consensus on 07.11.2019. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Feraccru - ferric maltol - EMEA/H/C/002733/II/0022 Norgine B.V., Rapporteur: Maria Concepcion Prieto Yerro, "Update of sections 4.2, 4.4 and 5.2 of the SmPC to include information on patients with chronic kidney disease, following the submission of the final study report of study ST10-01-303." Request for Supplementary Information adopted on 12.09.2019.	
Herceptin - trastuzumab - EMEA/H/C/000278/II/0157 Roche Registration GmbH, Rapporteur: Jan Mueller-Berghaus, "To submit the results from biopharmaceutic studies and clinical pharmacology studies on the improved sensitivity of the assay developed and validated to assess rHuPH20, included as a recommendation."	Positive Opinion adopted by consensus on 31.10.2019. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

Opinion adopted on 31.10.2019.	
IBRANCE - palbociclib - EMEA/H/C/003853/II/0016 Pfizer Europe MA EEIG, Rapporteur: Filip Josephson, "Update of section 5.1 of the SmPC in order to update with information following submission of the final results from the pivotal study A5481023 "A double blind, Phase 3 trial of fulvestrant with or without palbociclib in pre- and postmenopausal women with hormone receptor positive, HER2-negative metastatic breast cancer that progressed on prior endocrine therapy" listed as a recommendation at the time of initial MA." Request for Supplementary Information adopted on 26.09.2019, 25.07.2019, 02.05.2019, 31.01.2019.	
IBRANCE - palbociclib - EMEA/H/C/003853/II/0024 Pfizer Europe MA EEIG, Rapporteur: Filip Josephson, "Update of section 5.1 of the SmPC based on the final report from a non-clinical study (PD-0332991) evaluating the correlation of palbociclib response to RB1 status." Opinion adopted on 31.10.2019. Request for Supplementary Information adopted on 12.09.2019.	Positive Opinion adopted by consensus on 31.10.2019. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
IDELVION - albutrepenonacog alfa - EMEA/H/C/003955/II/0034, Orphan CSL Behring GmbH, Rapporteur: Jan Mueller-Berghaus, "Submission of a variation to update the dosing regimen as follows: -21-day prophylaxis regimen with rIX-FP at a dose of 100 IU/kg body weight for patients ≥ 12 years who are well controlled on a 14-day prophylaxis regimen. -10- or 14-day prophylaxis regimen with rIX-FP at a dose of 75 IU/kg body weight for patients < 12 years who are well controlled on a 7-day prophylaxis regimen. This submission also updates the existing population PK model with additional intravenous and subcutaneous (SC) data from the PTPs in the PTP arm of study CSL654_3003 and re-evaluates the covariates that are possible determinants of PK variability."	
Juluca - dolutegravir / rilpivirine - EMEA/H/C/004427/II/0016 ViiV Healthcare B.V., Rapporteur: Janet Koenig,	

"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 dolutegavir. The SmPC is updated accordingly. The RMP is not submitted."	
Kisqali - ribociclib - EMEA/H/C/004213/II/0018 Novartis Europharm Limited, Rapporteur: Filip Josephson, "Update of section 5.2 of the SmPC to include updated information about the use of Kisqali in patients with mild or moderate renal impairment based on the results of Study CLEE011A2116 Part II and additional data from breast cancer patients with mild or moderate renal impairment."	
Kolbam - cholic acid - EMEA/H/C/002081/II/0028, Orphan Retrophin Europe Ltd, Rapporteur: Konstantinos Markopoulos, "Submission of the final report from study CAC-002-01, listed as a category 3 study in the RMP. This is a Phase 3, open-label, single arm, non-randomized study investigating cholic acid in the treatment of subjects with inborn errors of bile acid metabolism. The study was a continuation study that included eligible subjects who had previously received cholic acid in studies CAC-91-10-10 or CAC-001-01 as well as newly diagnosed subjects." Request for Supplementary Information adopted on 19.09.2019, 29.05.2019, 28.02.2019.	
Maviret - glecaprevir / pibrentasvir - EMEA/H/C/004430/II/0027 AbbVie Deutschland GmbH & Co. KG, Rapporteur: Jean-Michel Race, "Submission of the final clinical study report from study M16-133, this is a phase 3b, single Arm, open label, multicenter study aimed to evaluate the efficacy and safety of glecaprevir (GLE)/pibrentasvir (PIB) in treatment of naïve adults with chronic	Request for supplementary information adopted with a specific timetable.

Hepatitis C Virus (HCV) Genotypes 1 – 6 infection and aspartate aminotransferase to platelet ratio index (APRI) ≤ 1." Request for Supplementary Information adopted on 24.10.2019.	
Mimpara - cinacalcet - EMEA/H/C/000570/II/0065 Amgen Europe B.V., Rapporteur: Kristina Dunder, "Update of section 4.8 of the SmPC in order to the new ADR 'chondrocalcinosis pyrophosphate' with a frequency of unknown. In addition, the MAH took the opportunity to bring the PI in line with the latest QRD template version 10.1 and to implement a minor correction to the List of Excipients in section 6.1 of the SmPC."	
Ongentys - opicapone - EMEA/H/C/002790/II/0020 Bial - Portela & Cª, S.A., Rapporteur: Martina Weise, "Update of sections 4.5 and 5.2 of the SmPC to add information on drug interaction and pharmacokinetic properties of opicapone based on final results from drug interaction studies NBI-OPC-1708 and NBI-OPC-1707. Study NBI-OPC-1708 is a phase 1, open-label, one-sequence crossover, drug-interaction study to evaluate and compare the pharmacokinetics of repaglinide when administered alone and concomitantly with opicapone. Study NBI-OPC-1707 is a Phase 1, randomized, open-label, 2-period crossover drug interaction study of the effect of administration of single dose of quinidine on the pharmacokinetics of opicapone. In addition, the marketing authorisation holder took the opportunity to delete the local representative for UK from the PL, according to the guidance provided on UK's withdrawal from the EU regarding medicinal products for human and veterinary use within the framework of the Centralised Procedure" Request for Supplementary Information adopted on 24.10.2019.	Request for supplementary information adopted with a specific timetable.
OPDIVO - nivolumab - EMEA/H/C/003985/II/0073 Bristol-Myers Squibb Pharma EEIG, Rapporteur: Jorge Camarero Jiménez, "To update sections 4.8 and 5.1 of the SmPC based on the final results from two studies: CA209017 (Open-label	Positive Opinion adopted by consensus on 24.10.2019. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

Randomized Phase III Trial comparing Nivolumab Versus Docetaxel in Previously Treated Advanced or Metastatic Squamous Cell Non-small Cell Lung Cancer) and CA209057 (Open-label Randomized Phase III Trial comparing Nivolumab Versus Docetaxel in Previously Treated Metastatic Non-Squamous Non-small Cell Lung Cancer)."
Opinion adopted on 24.10.2019.

Pradaxa - dabigatran etexilate - EMEA/H/C/000829/II/0118/G

Boehringer Ingelheim International GmbH, Rapporteur: Mark Ainsworth, "Update of section 4.5 of the SmPC in order to add a warning regarding the interaction between Pradaxa and the fixed-dose combination of the P-gp inhibitors glecaprevir and pibrentasvir based on the phase I drug-drug interaction study results. The Package Leaflet was updated accordingly.
Update of section 4.8 of the SmPC with new safety information regarding adverse reaction alopecia following the confirmation of signal "alopecia associated with dabigatran" by the EMA and the cumulative review of cases of alopecia and related terms that was provided in PSUR submitted by 27 May 2019. In addition small editorial corrections under "Adverse reaction" Table 2 were made additionally to highlight that information on some side effects was obtained from post-marketing data. The Package Leaflet was updated accordingly."
Request for Supplementary Information adopted on 12.09.2019.

Prevenar 13 - pneumococcal polysaccharide conjugate vaccine (13-valent, adsorbed) - EMEA/H/C/001104/II/0181

Pfizer Europe MA EEIG, Rapporteur: Kristina Dunder, "Update of section 5.1 of the SmPC in order to update efficacy information based on results from a public health analysis and publication of data from the CAPiTA (Community-Acquired Pneumonia Immunization Trial in Adults), a double-blind, randomized, placebo-controlled efficacy trial of 13-valent pneumococcal conjugate vaccine (PCV13)."
Request for Supplementary Information adopted on 19.09.2019.

Qtern - saxagliptin / dapagliflozin - EMEA/H/C/004057/II/0024

AstraZeneca AB, Rapporteur: Johann Lodewijk

Hillege, PRAC Rapporteur: Amelia Cupelli, "Update of sections 4.2, 4.4 and 5.1 of the SmPC with information on the glycaemic efficacy and renal safety of dapagliflozin in patients with Type 2 Diabetes Mellitus and moderate renal impairment (CKD 3A) based on final results from study D1690C00024 (DERIVE) (dapagliflozin), and to reflect a change in renal cut-off value for saxagliptin. The package leaflet is updated accordingly. The RMP version 4.1 has also been submitted. In addition, the MAH took the opportunity to update SmPC sections 2, 4.8, 5.2 and Annex II to include the required excipient information in relation to sodium levels and lactose following the update to the Annex to the European Commission guideline on "Excipients in the labelling and package leaflet of medicinal products for human use", as well as to bring the PI in line with EMA guidance ("Compilation of QRD decisions on stylistic matters in product information", EMA/25090/2002 Rev.18, published 08 December 2017)." Request for Supplementary Information adopted on 27.06.2019.	
Repatha - evolocumab - EMEA/H/C/003766/II/0038 Amgen Europe B.V., Rapporteur: Johann Lodewijk Hillege, "Update of sections 4.2, 4.4 and 5.2 of the SmPC in order to update the safety information based on final results from study 20140213. This is a phase 1 open-label interventional study to evaluate the safety, pharmacokinetics, and pharmacodynamics of evolocumab after a single 140 mg subcutaneous dose in subjects with normal renal function or severe renal insufficiency or end stage renal disease receiving haemodialysis. The Package Leaflet is updated accordingly."	
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	Request for supplementary information adopted with a specific timetable.

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.

Rezolsta - darunavir / cobicistat - EMEA/H/C/002819/II/0035

Janssen-Cilag International NV, Rapporteur: Johann Lodewijk Hillege, “C.I.4 Update of sections 4.8 and 5.1 of the SmPC to update the efficacy and safety information of Rezolsta following results from study TMC114FD2HTX3001 (AMBER); this is an ongoing Phase 3, randomized, active-controlled, double-blind study to evaluate efficacy and safety of darunavir/cobicistat/emtricitabine/tenofovir alafenamide (D/C/F/TAF) once daily fixed dose combination (FDC) regimen versus a regimen consisting of darunavir/cobicistat (DRV/COBI) FDC co-administered with emtricitabine/tenofovir disoproxil fumarate FDC in antiretroviral treatment-naïve human immunodeficiency virus type 1 infected subjects.
The applicant takes the opportunity to update section 4.5 to remove the interaction with simeprevir, following the withdrawal of Olysio Marketing Authorization. In addition, the MAH has implemented some minor administrative updates throughout the Product Information. The Package Leaflet is updated accordingly.”

Rizmoic - naldemedine - EMEA/H/C/004256/II/0004

Shionogi B.V., Rapporteur: Mark Ainsworth, “Submission of the final report from non-clinical study S-297995-PF-360-N as agreed in letter of recommendation to CHMP: In-vitro data determining whether naldemedine inhibits in a time dependent manner the OATP1B1, OATP1B3, OAT1 and OAT3 transporters.”

RoActemra - tocilizumab - EMEA/H/C/000955/II/0089

Roche Registration GmbH, Rapporteur: Jan Mueller-Berghaus, “Submission of the final report from study WA28119. This is a Phase III, multicenter, randomized, double-blind, placebo-controlled study to assess the efficacy and safety of tocilizumab in subjects with giant cell arteritis.).”

RXULTI - brexpiprazole - EMEA/H/C/003841/II/0003 Otsuka Pharmaceutical Netherlands B.V., Rapporteur: Daniela Melchiorri, “To update section 4.4 of the SmPC (paragraph “Impulse-control disorders”) based on the Company Core Data Sheet of brexpiprazole. In addition, the applicant has taken the opportunity to update the section 4.2 of the SmPC requested by EMA (see annex to cover letter) and to perform additional changes, i.e. editorial changes in the SmPC and Package Leaflet.” Request for Supplementary Information adopted on 12.09.2019.	
Shingrix - herpes zoster vaccine (recombinant, adjuvanted) - EMEA/H/C/004336/II/0016 GlaxoSmithkline Biologicals SA, Rapporteur: Bart Van der Schueren, “Update of section 4.5 of the SmPC in order to reflect information related to coadministration based on the final results from studies ZOSTER-035 and ZOSTER-042; these are immunogenicity and safety studies in which Shingrix was co-administered either with Merck's 23-valent pneumococcal polysaccharide vaccine (Pneumovax 23; ZOSTER-035) or with GSK's reduced-antigen-content diphtheria and tetanus toxoids and acellular pertussis (dTpa) vaccine (Boostrix; ZOSTER-042); the Package Leaflet is updated accordingly.” Opinion adopted on 24.10.2019. Request for Supplementary Information adopted on 19.09.2019, 25.07.2019.	Positive Opinion adopted by consensus on 24.10.2019. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
SIMBRINZA - brinzolamide / brimonidine - EMEA/H/C/003698/II/0018/G Novartis Europharm Limited, Rapporteur: Maria Concepcion Prieto Yerro, “Update of section 5.1 of the SmPC in order to update the safety information with adjunctive use of BID Simbrinza with a PGA therapy based on final results from study CQVJ499A2401; this is a phase 4, multicenter, randomized, double-masked, parallel-group study. Update of section 5.1 of the SmPC in order to update the safety information with adjunctive use of BID Simbrinza with a PGA/beta-blocker combination therapy based on final results from study CQVJ499A2402; this is a phase 4, multicenter, randomized, double-masked,	

parallel-group study." Request for Supplementary Information adopted on 12.09.2019.	
Sivextro - tedizolid phosphate - EMEA/H/C/002846/II/0032 Merck Sharp & Dohme B.V., Rapporteur: Bruno Sepodes, "To update the Marketing Authorization for Sivextro with the final report from Phase 3 study for the treatment of Hospital-Acquired Bacterial Pneumonia and Ventilator-Associated Bacterial Pneumonia (HABP/VABP) MK-1986-002; protocol TR701-132." Opinion adopted on 31.10.2019.	Positive Opinion adopted by consensus on 31.10.2019. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Tivicay - dolutegravir - EMEA/H/C/002753/II/0052 ViiV Healthcare B.V., Rapporteur: Filip Josephson, "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 dolutegavir. The SmPC is updated accordingly. The RMP is not submitted."	
Translarna - ataluren - EMEA/H/C/002720/II/0053/G, Orphan PTC Therapeutics International Limited, Rapporteur: Johann Lodewijk Hillege, "C.I.4: Update of section 5.3 of the SmPC in order to update the safety information based on final results Charles River 9001126 Three-month juvenile toxicology and toxicokinetic study planned in neonatal dogs listed as category 3 study in the RMP (MEA-005). C.I.13 Submission of the final report from study WIL-523008 listed as category 3 study in the RMP (MEA/003). This is a Seven-day tolerability and pharmacokinetic study in neonatal dogs. C.I.13 Submission of the final report from study WIL-523009 listed as category 3 study in the RMP (MEA/004). This is a One-month juvenile dose range-finding toxicology and toxicokinetic study planned in neonatal dogs age correlating with	Positive Opinion adopted by consensus on 24.10.2019. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

dosing in newborn paediatric patients to 2 years of age. C.I.13 Submission of the final report from study (Charles River 5700755 listed as category 3 study in the RMP (MEA/0024). This is a 28-day investigational toxicology and toxicokinetic study of ataluren in juvenile beagle dogs with an 8-week recovery period – Category 3." Opinion adopted on 24.10.2019. Request for Supplementary Information adopted on 18.07.2019.	
Triumeq - dolutegravir / abacavir / lamivudine - EMEA/H/C/002754/II/0069 ViiV Healthcare B.V., Rapporteur: Filip Josephson, "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 dolutegavir. The SmPC is updated accordingly. The RMP is not submitted."	
Truvada - emtricitabine / tenofovir disoproxil - EMEA/H/C/000594/II/0161 Gilead Sciences Ireland UC, Rapporteur: Bruno Sepodes, "Submission of the final clinical study report for the non-interventional study GS-US-276-0103, 'A Prospective, Observational Study of Individuals Who Seroconvert While Taking Truvada for Pre-Exposure Prophylaxis (PrEP)', listed as a Category 3 study in the Truvada RMP." Request for Supplementary Information adopted on 12.09.2019.	
Tyverb - lapatinib - EMEA/H/C/000795/II/0059 Novartis Europharm Limited, Rapporteur: Filip Josephson, "Update of section 5.1 of the SmPC in order to update Table 8 based on updated/corrected results from study EGF114299/LAP016A2307, an interventional study with progression free survival rate as primary objective, original report submitted	See 9.1

during procedure EMEA/H/C/00795/II/0051." Request for Supplementary Information adopted on 26.04.2019.	
Verzenios - abemaciclib - EMEA/H/C/004302/II/0006 Eli Lilly Nederland B.V., Rapporteur: Filip Josephson, "Update of section 4.8 of the SmPC in order to add interstitial lung disease (ILD)-like events (including pneumonitis) as a new adverse drug reaction. The Package Leaflet is updated accordingly. In addition, the Marketing authorisation holder (MAH) took the opportunity to update the list of local representatives in the Package Leaflet." Request for Supplementary Information adopted on 19.09.2019.	
XALKORI - crizotinib - EMEA/H/C/002489/II/0064 Pfizer Europe MA EEIG, Rapporteur: Alexandre Moreau, "Update of section 5.1 of the SmPC in order to reflect updated efficacy data from Study A8081001 in patients with ROS1-positive NSCLC. In addition, the Marketing authorisation holder (MAH) took the opportunity to bring the PI in line with the latest QRD template version 10.1." Opinion adopted on 31.10.2019.	Positive Opinion adopted by consensus on 31.10.2019. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Xermelo - telotristat ethyl - EMEA/H/C/003937/II/0014, Orphan Ipsen Pharma, Rapporteur: Martina Weise, "To update sections 4.2 and 5.2 of the SmPC following final results from study LX1606-111; this is a Phase 1, open-label, parallel-group study to evaluate the single-dose pharmacokinetics of Telotristat Ethyl in Male and Female Subjects with Severe Hepatic Impairment and Matched Subjects with Normal Function; the Package Leaflet is updated accordingly." Request for Supplementary Information adopted on 19.09.2019, 11.07.2019.	
Zoely - nomegestrol acetate / estradiol - EMEA/H/C/001213/II/0050 Theramex Ireland Limited, Rapporteur: Jean-Michel Race, "Update of sections 4.3 and 4.4 of the SmPC in order to add a new contraindication and a new warning regarding meningioma, upon request by PRAC following the assessment of Post-authorisation measure "LEG 014". The Package Leaflet is being updated accordingly. In addition, the MAH took the	

opportunity to update the contact details of the local representatives in the Netherlands and Portugal in the Package Leaflet." Request for Supplementary Information adopted on 19.09.2019.	
PRAC Led **WS1601** **Glyxambi-EMEA/H/C/003833/WS1601/0022** **Jentadueto-EMEA/H/C/002279/WS1601/0051** **Trajenta-EMEA/H/C/002110/WS1601/0038** Boehringer Ingelheim International GmbH, Lead Rapporteur: Johann Lodewijk Hillege, Lead PRAC Rapporteur: Menno van der Elst, PRAC-CHMP liaison: Johann Lodewijk Hillege, "Update of sections 4.2 and 5.1 of the Trajenta SmPC, update of sections 4.2, 4.4 and 5.1 of the Jentadueto SmPC and section 5.1 of the Glyxambi SmPC, based on the final results from study 1218.74 (CAROLINA study) listed as a category 3 study in the RMP of Jentadueto and Trajenta, in order to fulfil Trajenta MEA 008.1 and Jentadueto MEA 001.1; this is a phase III randomized, parallel group, double blind study to evaluate Cardiovascular safety of linagliptin versus glimepiride in patients with type 2 diabetes mellitus at high cardiovascular risk. The Package Leaflet for Trajenta is updated accordingly. The RMP version 13.0 for Jentadueto and Trajenta and version 5.0 for Glyxambi have also been submitted. In addition, the Worksharing applicant (WSA) took the opportunity to make corrections throughout the product information for Glyxambi and Jentadueto and to make corrections to the Bulgarian, French, Swedish translations for Glyxambi." Opinion adopted on 31.10.2019. Request for Supplementary Information adopted on 11.07.2019.	Positive Opinion adopted by consensus on 31.10.2019. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1605 **Lyrica-EMEA/H/C/000546/WS1605/0097** **Pregabalin Pfizer-EMEA/H/C/003880/WS1605/0027** Pfizer Europe MA EEIG, Lead Rapporteur: Johann Lodewijk Hillege, "Addition in the SmPC section 4.5 of the wording on the risk of death, including in patients who are substance abusers." Request for Supplementary Information adopted	

on 12.09.2019, 23.05.2019.	
WS1701 **Epclusa-EMEA/H/C/004210/WS1701/0040** **Vosevi-EMEA/H/C/004350/WS1701/0032** Gilead Sciences Ireland UC, Lead Rapporteur: Filip Josephson, "Update of section 4.8 of the SmPC in order to add new safety information on rash and angioedema following a cumulative review of hypersensitivity with Epclusa and Vosevi, prompted by routine pharmacovigilance and signal detection activities. The Package Leaflet is updated accordingly. In addition, the MAH took the opportunity to introduce minor editorial changes throughout the Product Information."	
WS1705 **Aluvia-EMEA/H/W/000764/WS1705/0111** **Kaletra-EMEA/H/C/000368/WS1705/0180** AbbVie Deutschland GmbH & Co. KG, Lead Rapporteur: Jean-Michel Race, "Change of section 4.8 of the SmPC to update the safety information following a cumulative safety review of the incidence rate of Stevens-Johnson syndrome, erythema multiforme and jaundice during clinical trials. This variation closes LEG 110. The Package Leaflet is updated accordingly." Opinion adopted on 31.10.2019.	Positive Opinion adopted by consensus on 31.10.2019. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

B.5.3. CHMP-PRAC assessed procedures

Aranesp - darbepoetin alfa - EMEA/H/C/000332/II/0150 Amgen Europe B.V., Rapporteur: Martina Weise, PRAC Rapporteur: Martin Huber, "Update of the SmPC sections 4.4, 4.8, 5.1 based on the study data from Study 20070782 - a phase 3, randomized, double-blind, placebo-controlled, noninferiority study in subjects with chemotherapy-induced anemia receiving multi-cycle chemotherapy for the treatment of advanced stage nonsmall-cell lung cancer (NSCLC); study of epoetin alfa in metastatic breast cancer (EPO-ANE-3010) and the Company Core Data Sheet. In addition, the section 4.6 has been revised based on the recommendation from last Periodic Safety Update Report Number 33 dated 15 January 2018. Furthermore, the MAH took the	

opportunity to introduce minor editorial changes, update the information on local representatives and align the PI with the requirements of the QRD template 10.0. The PL is updated accordingly. The revised RMP version 9.3 has been also submitted." Request for Supplementary Information adopted on 25.07.2019, 28.03.2019.	
Avastin - bevacizumab - EMEA/H/C/000582/II/0110 Roche Registration GmbH, Rapporteur: Sinan B. Sarac, PRAC Rapporteur: Hans Christian Siersted, "Submission of the final report from study NEJ026 listed as an obligation in the Annex II of the Product Information. This is an open-label, randomized, Phase III study conducted in Japan to compare erlotinib + bevacizumab combination therapy versus erlotinib monotherapy as first-line therapies for patients with NSCLC with EGFR gene mutations (exon 19 deletion or exon 21 L858R substitution). The RMP version 30.0 has also been submitted. In addition, the Package leaflet is updated to reflect information on sodium content in compliance with the revised Annex to the European Commission guideline on "Excipients in the labelling and package leaflet of medicinal products for human use"." Request for Supplementary Information adopted on 31.10.2019, 05.09.2019.	Request for supplementary information adopted with a specific timetable.
Brinavess - vernakalant - EMEA/H/C/001215/II/0035 Correvio, Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Menno van der Elst, "Update of sections 4.4 and 4.8 of the SmPC in order to add a warning and update the safety information following updates to the Company Core Safety Datasheet (CCDS) based on the results of an integrated safety analysis performed on data of existing clinical studies with a stronger emphasis on treatment-related ADRs and an incidence rate above one percent. The Package Leaflet was updated accordingly. The RMP version 7.0 has also been submitted and incorporates the results from the new safety analysis. In addition the results from completed observational Cohort SPECTRUM study (A Prospective Observational Registry Study to Characterise Normal Conditions of Use, Dosing and Safety	Positive Opinion adopted by consensus on 31.10.2019. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

Following Administration of Vernakalant IV Sterile Concentrate: PASS Protocol 6621-049) currently under assessment within Brinavess II/34 were included in the updated RMP. In addition, the Marketing authorisation holder (MAH) took the opportunity to update sections 4.2, 4.4, 4.6, 4.7, 4.8, 5.1, 5.2, 5.3, 6.4 of the SmPC, Annex II, Labelling sections 3 and 5, Package Leaflet sections 2, 4, 5 and 6 to include editorial changes, to correct typographical errors and to bring the PI in line with the latest QRD template version 10 as well as to update statements related to the excipients in the SmPC and Package Leaflet in line with the EC Guideline on "Excipients in the Labelling and Package Leaflet of medicinal products for human use" of March 2018 and the EMA Annex to the EC Guideline of October 2017 (SANTE-2017-11668)." Opinion adopted on 31.10.2019. Request for Supplementary Information adopted on 05.09.2019, 14.06.2019.	
Increlex - mecasermin - EMEA/H/C/000704/II/0060 Ipsen Pharma, Rapporteur: Outi Mäki-Ikola, PRAC Rapporteur: Kirsti Villikka, "Update of sections 4.1, 4.2, 4.3, 4.4, 4.8 and 4.9 of the SmPC in order to update the safety information on benign or malignant neoplasia based on the EU Registry Study, the Ipsen global safety database and literature review. The Package Leaflet is updated accordingly. The MAH also submitted the updated RMP version 11. In addition, the MAH took the opportunity to update the list of local representatives in the Package Leaflet" Request for Supplementary Information adopted on 19.09.2019.	See 9.1
Mircera - methoxy polyethylene glycol-epoetin beta - EMEA/H/C/000739/II/0068 Roche Registration GmbH, Rapporteur: Maria Concepcion Prieto Yerro, PRAC Rapporteur: Eva A. Segovia, "Update of sections 4.4 and 4.8 of the SmPC to include information on the availability of anti-erythropoietin antibody (AEAB) testing and to amend the frequency of adverse drug reactions, respectively, based on the final report of study BH21260, listed as a category 3 study in	Positive Opinion adopted by consensus on 31.10.2019. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

the RMP (MEA008.5); this is a randomized, controlled, open-label, multicenter, parallel-group study to assess all-cause mortality and cardiovascular morbidity in patients with chronic kidney disease on dialysis and those not on renal replacement therapy under treatment with Mircera or reference ESAs. In addition, reference to the educational materials related to AEAB testing is removed from the Annex II. The RMP (version 12.3) is updated accordingly and transitioned to the new EU RMP template in line with the revised Good Pharmacovigilance Practice (GVP) Module V (Revision 2) guideline." Opinion adopted on 31.10.2019. Request for Supplementary Information adopted on 14.06.2019, 17.01.2019, 04.10.2018.	
NovoEight - turoctocog alfa - EMEA/H/C/002719/II/0030/G Novo Nordisk A/S, Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Brigitte Keller-Stanislawski, "Update of sections 4.2, 4.4, 4.8, 5.1 and 5.2 of the SmPC based on results of the Guardian 4 (NN7008-3809) Clinical Trial in Previously Untreated Patients (PUPs) and the Guardian 9 (NN7008-4239) PK Clinical Trial. The MAH has also updated the SmPC to align with the 'EMA Core SmPC for human plasma derived and recombinant coagulation factor VIII products, revision 3' and Annex to the European Commission guideline on `Excipients in the labelling and package leaflet of medicinal products for human use'. Further, some administrative updates have also been applied." Opinion adopted on 31.10.2019. Request for Supplementary Information adopted on 05.09.2019.	Positive Opinion adopted by consensus on 31.10.2019. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Odomzo - sonidegib - EMEA/H/C/002839/II/0024 Sun Pharmaceutical Industries Europe B.V., Rapporteur: Paula Boudewina van Hennik, PRAC Rapporteur: Željana Margan Koletić, "To submit the final report of study CLDE225X2116, listed as a category 3 study in the RMP. This is an interventional Phase Ib/II, open-label, multi-center, dose-finding study to assess the safety and efficacy of the oral combination of LDE225 and INC424 (Ruxolitinib) in subjects with myelofibrosis. The RMP version 7.1 has also been submitted." Request for Supplementary Information adopted	Request for supplementary information adopted with a specific timetable.

on 31.10.2019.	
Orkambi - lumacaftor / ivacaftor - EMEA/H/C/003954/II/0049 Vertex Pharmaceuticals (Ireland) Limited, Rapporteur: Daniela Melchiorri, PRAC Rapporteur: Rhea Fitzgerald, "Updated of section 4.8 of the SmPC with the safety data from the Phase 3, open-label, rollover study for Studies 109 and 011 Part B (Study 011B) designed to evaluate the long-term safety and tolerability of Orkambi treatment for 96 weeks in patients with cystic fibrosis, 6 years of age and older, homozygous for F508del." Request for Supplementary Information adopted on 31.10.2019, 05.09.2019.	Request for supplementary information adopted with a specific timetable.
Raxone - idebenone - EMEA/H/C/003834/II/0018, Orphan Santhera Pharmaceuticals (Deutschland) GmbH, Rapporteur: John Joseph Borg, PRAC Rapporteur: Amelia Cupelli, "Submission of the final report from study SNT-EAP-001 listed as a Specific Obligation (SOB11, former SOB4) in the Annex II of the Product Information. This is a follow-up study of patients in the Expanded Access Program (SNT-EPA-001) for Raxone in the treatment of patients with Leber's Hereditary Optic Neuropathy (LHON). The goal is to collect further long-term real-world efficacy and safety data. Annex II is modified accordingly. An updated RMP version 1.10 submitted accordingly."	
TECFIDERA - dimethyl fumarate - EMEA/H/C/002601/II/0058 Biogen Netherlands B.V., Rapporteur: Martina Weise, PRAC Rapporteur: Martin Huber, "Submission of CSR of study 109MS310, an open-label study to assess the effects of Tecfidera on lymphocyte subsets in subjects with relapsing remitting multiple sclerosis, listed as category 3 study in the RMP. The RMP (version 10.1) has been updated as a consequence of the completion of this study. The revised RMP also includes updates to reflect safety information available through to the data lock point of 24 January 2019 and to align with the EU RMP Module V (revision 2.01)." Request for Supplementary Information adopted on 31.10.2019, 11.07.2019.	Request for supplementary information adopted with a specific timetable.

TECFIDERA - dimethyl fumarate - EMEA/H/C/002601/II/0062 Biogen Netherlands B.V., Rapporteur: Martina Weise, PRAC Rapporteur: Martin Huber, "Update of sections 4.4 and 4.8 of the SmPC to add a warning on the risk of herpes zoster based on cumulative review data submitted in the ongoing PSUSA/00010143/201903. The Package Leaflet is updated accordingly." Request for Supplementary Information adopted on 19.09.2019.	
UDENYCA - pegfilgrastim - EMEA/H/C/004413/II/0003 ERA Consulting GmbH, Rapporteur: Martina Weise, PRAC Rapporteur: Menno van der Elst, "To update section 4.6 of the SmPC to remove reference to the pregnancy and lactation registry listed as a category 3 study in the RMP. The Package Leaflet is updated accordingly. The updated RMP version 1.5 has also been submitted." Opinion adopted on 31.10.2019. Request for Supplementary Information adopted on 05.09.2019.	Positive Opinion adopted by consensus on 31.10.2019. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
VeraSeal - human fibrinogen / human thrombin - EMEA/H/C/004446/II/0006/G Instituto Grifols, S.A., Rapporteur: Andrea Laslop, PRAC Rapporteur: Amelia Cupelli"B.IV.1.a.3 – Type II - To add a new CE marked applicator tip as a replacement for the current application cannula which allows the application of the product both by dripping and spraying without gas assistance. The safety concern of air or gas embolism that is currently included in the Risk Management Plan (RMP) is no longer applicable, consequently the RMP has been updated accordingly (version 4.0) and is provided in Module 1.8.2. RMP version 4.0 has been restructured in order to adapt to the new format of GVP Module V. B.II.e.6.a – Type 1B B.II.e.6.a – Type 1B B.II.b.3.a – Type 1A B.II.b.3.a – Type 1A Request for Supplementary Information adopted on 17.10.2019.	
Zelboraf - vemurafenib - EMEA/H/C/002409/II/0054 Roche Registration GmbH, Rapporteur: Filip	Positive Opinion adopted by consensus on 31.10.2019. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP

Josephson, PRAC Rapporteur: Annika Folin, "Update of sections 4.4 and 4.5 of the SmPC in order to add information and a precaution regarding concomitant strong CYP3A4 inhibitors based on final results from study GO29475 (MEA-011), a category 3 study in the RMP; the Package Leaflet is updated accordingly. The RMP version 12.0 has also been submitted. In addition, the Marketing authorisation holder (MAH) took the opportunity to bring the PL in line with the excipients guideline (EMA/CHMP/302620/2017) by adding information about the product's sodium content." Opinion adopted on 31.10.2019. Request for Supplementary Information adopted on 05.09.2019.	recommendation.
Zydelig - idelalisib - EMEA/H/C/003843/II/0047 Gilead Sciences Ireland UC, Rapporteur: Filip Josephson, PRAC Rapporteur: Martin Huber, "submission of the final clinical study report for study 101-09, A Phase 2 Study to Assess the Efficacy and Safety of Idelalisib in Subjects with Indolent BCell Non-Hodgkin Lymphomas Refractory to Rituximab and Alkylating Agents. This submission is an Annex II postauthorisation measure (ANX 002) and a category I commitment in the Zydelig Risk Management Plan (RMP).This submission also includes an update to the PI" Request for Supplementary Information adopted on 19.09.2019.	See 9.1
WS1690 Clopidogrel Zentiva-EMEA/H/C/000975/WS1690/0066 DuoPlavin-EMEA/H/C/001143/WS1690/0053 Iscover-EMEA/H/C/000175/WS1690/0136 Plavix-EMEA/H/C/000174/WS1690/0133 Sanofi Clir SNC, Lead Rapporteur: Bruno Sepodes, Lead PRAC Rapporteur: Marcia Sofia Sanches de Castro Lopes Silva, "To modify the Product Information (PI) in section 4.5 "Interaction with other medicinal products and other forms of interaction" of the SmPC and the corresponding section of the PL to add the signal of interaction of clopidogrel with boosted antiviral HIV therapy leading to insufficient inhibition of platelet aggregation in line with EPITT 19325.	

The MAH has made minor adjustments to the wording."	

B.5.4. PRAC assessed procedures

PRAC Led **Adempas - riociguat - EMEA/H/C/002737/II/0030, Orphan** Bayer AG, Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Kimmo Jaakkola, PRAC-CHMP liaison: Tuomo Lapveteläinen, "Submission of the final report from risk management plan (RMP) category 3 study 16657, EXPERT (EXPosurE Registry RiociguaT in patients with pulmonary hypertension) to collect information about the long term use of Adempas in real clinical practice. The RMP version 7.1 has also been submitted." Request for Supplementary Information adopted on 31.10.2019.	Request for supplementary information adopted with a specific timetable.
PRAC Led **AUBAGIO - teriflunomide - EMEA/H/C/002514/II/0025** sanofi-aventis groupe, Rapporteur: Martina Weise, PRAC Rapporteur: Martin Huber, PRAC-CHMP liaison: Martina Weise, "Submission of the patients and HCPs final survey reports to assess the effectiveness of the education materials; the survey reports are part of the additional pharmacovigilance activities in the RMP (category 3 studies). Within this submission the MAH is proposing a revised patient card with the following revisions: the patient card was restructured (general guidance, possible side effects, pregnancy), details related to the Accelerated Elimination Procedure were deleted and symptoms related to liver and infections are described." Request for Supplementary Information adopted on 31.10.2019, 05.09.2019.	Request for supplementary information adopted with a specific timetable.
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	Request for supplementary information adopted with a specific timetable.

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." Request for Supplementary Information adopted on 31.10.2019.	
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 11.1 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." Request for Supplementary Information adopted on 31.10.2019.	Request for supplementary information adopted with a specific timetable.
PRAC Led **Flixabi - infliximab - EMEA/H/C/004020/II/0039** Samsung Bioepis NL B.V., Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Ulla Wändel Liminga, PRAC-CHMP liaison: Kristina Dunder, "Update of the RMP to replace the current registries with one company-sponsored initiated registry (PERFUSE) and three IBD registries (CEDUR, CREDIT, and DREAM)" Opinion adopted on 31.10.2019. Request for Supplementary Information adopted on 11.07.2019, 11.04.2019.	Positive Opinion adopted by consensus on 31.10.2019. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
PRAC Led **Invokana - canagliflozin - EMEA/H/C/002649/II/0045/G** Janssen-Cilag International NV, Rapporteur: Martina Weise, PRAC Rapporteur: Martin Huber, PRAC-CHMP liaison: Martina Weise, "Submission of the final report from 3 non-interventional studies (listed as category 3 studies in the RMP): - Study RRA-21410, an Epidemiology Study to characterize the risk of LLA in subjects in the overall T2DM population and in a subpopulation with established CVD.	Positive Opinion adopted by consensus on 31.10.2019. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

- NAP4001, a Meta-Analysis from CANVAS, CANVAS-R and CREDENCE Studies to characterize the risk of LLA in subjects at high risk for CV events and/or progression of kidney disease. - Meta-Analysis from CANVAS, CANVAS-R and CREDENCE to evaluate the incidence of bladder cancer in the canagliflozin group compared to the placebo group." Opinion adopted on 31.10.2019.	
PRAC Led **Praluent - alirocumab - EMEA/H/C/003882/II/0050/G** sanofi-aventis groupe, Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Brigitte Keller-Stanislawski, PRAC-CHMP liaison: Jan Mueller-Berghaus, "Submission of an updated RMP version 5.0 in order to amend the list of safety concerns (removing 'cataract (in the context of very low LDL-C)' as important potential risk; 'long-term use (>5years)' and 'clinical impact of very low LDL-C for extended period of time' as missing information; and consequentially to remove the following additional Pharmacovigilance activities (category 3 studies in the RMP) from the RMP: study R727-CL-1609 (MEA 016), study OBS14697 (MEA 019) and study ALIROC07997 (MEA 017) based on a review of data since the MA was granted including the 1st interim report for study OBS14697, a drug utilisation study (DUS) of alirocumab in Europe to assess the effectiveness of the dosing recommendation to avoid very low LDL-C levels, in order to fulfil MEA 019.4." Request for Supplementary Information adopted on 31.10.2019.	Request for supplementary information adopted with a specific timetable.
PRAC Led **Revlimid - lenalidomide - EMEA/H/C/000717/II/0110, Orphan** Celgene Europe BV, Rapporteur: Alexandre Moreau, PRAC Rapporteur: Ghania Chamouni, PRAC-CHMP liaison: Alexandre Moreau, "Submission of the CC-5013-PASS-001 final study report dated 2 Nov 2018; this is a non-interventional post-authorisation safety study (PASS) to further characterise the safety profile of lenalidomide plus dexamethasone in the treatment of relapsed and/or refractory (R/R) MM in a real-world setting." Opinion adopted on 31.10.2019.	Positive Opinion adopted by consensus on 31.10.2019. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

Request for Supplementary Information adopted on 14.06.2019.	
PRAC Led **Selincro - nalmefene - EMEA/H/C/002583/II/0025** H. Lundbeck A/S, Rapporteur: Janet Koenig, PRAC Rapporteur: Martin Huber, PRAC-CHMP liaison: Janet Koenig, "submission for the Final Study Reports for the PASS 15649A: Use of Nalmefene (Selincro) in European databases: Cohort design using longitudinal electronic medical records or claims databases and PASS 14910A a non-interventional multicountry prospective cohort study to investigate the pattern of use of Selincro and frequency of selected adverse reactions in routine clinical practice." Request for Supplementary Information adopted on 31.10.2019, 05.09.2019.	Request for supplementary information adopted with a specific timetable.
PRAC Led **SIMBRINZA - brinzolamide / brimonidine - EMEA/H/C/003698/II/0019** Novartis Europharm Limited, Rapporteur: Maria Concepcion Prieto Yerro, PRAC Rapporteur: Rhea Fitzgerald, PRAC-CHMP liaison: Peter Kiely, "Submission of an updated RMP version 3.0 in order to remove metabolic acidosis/renal impairment as an important potential risk from the list of safety concerns and in addition update the Risk management plan to comply with the new GVP module V rev 2 RMP template." Opinion adopted on 31.10.2019. Request for Supplementary Information adopted on 05.09.2019.	Positive Opinion adopted by consensus on 31.10.2019. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
PRAC Led **Slenyto - melatonin - EMEA/H/C/004425/II/0010** RAD Neurim Pharmaceuticals EEC SARL, Rapporteur: Kristina Dunder, PRAC Rapporteur: Ana Sofia Diniz Martins, PRAC-CHMP liaison: Bruno Sepodes, "Submission of the Annual report #3 of the French RTU with data collected from 01 October 2015 to 01 October 2018. Update of the RMP to version 1.5 to reflect the interim RTU result and commit to submit final results when available." Opinion adopted on 31.10.2019. Request for Supplementary Information adopted on 05.09.2019.	Positive Opinion adopted by consensus on 31.10.2019. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

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, "Submission of an updated RMP version 30.1 in order to revise the list of safety concerns. This revision has been triggered by the PRAC recommendation received in outcome of the EU-PSUR covering the period from 26 April 2017 to 25 April 2018. As a consequence, the Annex II of the PI has been updated to reflect the removal of the additional risk minimisation activities. In addition, the applicant 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)." Request for Supplementary Information adopted on 31.10.2019.	Request for supplementary information adopted with a specific timetable.
PRAC Led **Vokanamet - canagliflozin / metformin - EMEA/H/C/002656/II/0050/G** Janssen-Cilag International NV, Rapporteur: Martina Weise, PRAC Rapporteur: Menno van der Elst, PRAC-CHMP liaison: Johann Lodewijk Hillege, "Submission of the final report from 3 non-interventional studies (listed as category 3 studies in the RMP): - Study RRA-21410, an Epidemiology Study to characterize the risk of LLA in subjects in the overall T2DM population and in a subpopulation with established CVD. - NAP4001, a Meta-Analysis from CANVAS, CANVAS-R and CREDENCE Studies to characterize the risk of LLA in subjects at high risk for CV events and/or progression of kidney disease. - Meta-Analysis from CANVAS, CANVAS-R and CREDENCE to evaluate the incidence of bladder cancer in the canagliflozin group compared to the placebo group." Opinion adopted on 31.10.2019.	Positive Opinion adopted by consensus on 31.10.2019. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
PRAC Led **WS1654** **Enbrel-EMEA/H/C/000262/WS1654/0228** **LIFMIOR-EMEA/H/C/004167/WS1654/0022** Pfizer Europe MA EEIG, Lead Rapporteur: Maria Concepcion Prieto Yerro, Lead PRAC Rapporteur:	Positive Opinion adopted by consensus on 31.10.2019. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

Eva A. Segovia, PRAC-CHMP liaison: Maria Concepcion Prieto Yerro, “C.I.13: Submission of the final report from study (B1801311 - BADBIR) listed as a category 3 study in the RMP. This is a prospective cohort study that compared patients treated with biologic interventions (etanercept, adalimumab, and ustekinumab) and patients with similar disease characteristics but exposed only to conventional non-biologic systemic therapies.”
Opinion adopted on 31.10.2019.

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

Kymriah - tisagenlecleucel - EMEA/H/C/004090/II/0014, Orphan, ATMP
Novartis Europharm Limited, Rapporteur: Rune Kjeken, CHMP Coordinator: Ingrid Wang

B.5.6. CHMP-PRAC-CAT assessed procedures

B.5.7. PRAC assessed ATMP procedures

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

WS1656/G
Copalia-EMEA/H/C/000774/WS1656/0108/G
Copalia HCT-EMEA/H/C/001159/WS1656/0079/G
Dafiro-EMEA/H/C/000776/WS1656/0111/G
Dafiro HCT-EMEA/H/C/001160/WS1656/0081/G
Exforge HCT-EMEA/H/C/001068/WS1656/0078/G
Novartis Europharm Limited, Lead Rapporteur: Mark Ainsworth
Request for Supplementary Information adopted on 19.09.2019.

WS1661/G
Entresto-EMEA/H/C/004062/WS1661/0026/G
Neparvis-EMEA/H/C/004343/WS1661/

0025/G Novartis Europharm Limited, Lead Rapporteur: Johann Lodewijk Hillege	
WS1669 **Ryzodeg-EMEA/H/C/002499/WS1669/0035** **Tresiba-EMEA/H/C/002498/WS1669/0042** **Xultophy-EMEA/H/C/002647/WS1669/0032** Novo Nordisk A/S, Lead Rapporteur: Kristina Dunder Request for Supplementary Information adopted on 12.09.2019.	
WS1673 **Infanrix hexa-EMEA/H/C/000296/WS1673/0263** GlaxoSmithkline Biologicals SA, Lead Rapporteur: Bart Van der Schueren	
WS1685 **Juluca-EMEA/H/C/004427/WS1685/0017** **Tivicay-EMEA/H/C/002753/WS1685/0053** **Triumeq-EMEA/H/C/002754/WS1685/0072** ViiV Healthcare B.V., Lead Rapporteur: Filip Josephson	
WS1692/G **Edistride-EMEA/H/C/004161/WS1692/0033/G** **Forxiga-EMEA/H/C/002322/WS1692/0052/G** AstraZeneca AB, Lead Rapporteur: Kristina Dunder	
WS1706 **Enurev Breezhaler-EMEA/H/C/002691/WS1706/0030** **Seebri Breezhaler-EMEA/H/C/002430/WS1706/0030** **Tovanor Breezhaler-EMEA/H/C/002690/WS1706/0034** Novartis Europharm Limited, Duplicate, Duplicate of Seebri Breezhaler, Lead Rapporteur: Mark Ainsworth, “To introduce a modification of the Instructions for Use (IFU) for the Breezhaler devices affecting the labeling (SmPC and PL) through simplification of its content and layout - condensing textual instructions, optimizing illustrations, and clustering of content to aid comprehension of the procedure of use.	Request for supplementary information adopted with a specific timetable.

In addition both the packaging and IFU in the SmPC and PL have included device-specific information in precise locations in order to meet the MDR requirements. This is because the device does not have its own packaging or leaflet." Request for Supplementary Information adopted on 07.11.2019.	
WS1707 **Ultibro Breezhaler-EMEA/H/C/002679/WS1707/0031** **Ulunar Breezhaler-EMEA/H/C/003875/WS1707/0032** **Xoterna Breezhaler-EMEA/H/C/003755/WS1707/0035** Novartis Europharm Limited, Lead Rapporteur: Mark Ainsworth, "To add onto the packaging and IFU in the SmPC and PL device-specific information in precise locations in order to meet the MDR requirements. This is because the device does not have its own packaging or leaflet." Opinion adopted on 07.11.2019.	Negative Opinion adopted by consensus on 07.11.2019. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1708 **Hirobriz Breezhaler-EMEA/H/C/001211/WS1708/0055** **Onbrez Breezhaler-EMEA/H/C/001114/WS1708/0053** **Oslif Breezhaler-EMEA/H/C/001210/WS1708/0053** Novartis Europharm Limited, Lead Rapporteur: Mark Ainsworth, "To introduce a modification of the Instructions for Use (IFU) for the Breezhaler devices affecting the labeling (SmPC and PL) through simplification of its content and layout - condensing textual instructions, optimizing illustrations, and clustering of content to aid comprehension of the procedure of use. In addition both the packaging and IFU in the SmPC and PL have included device-specific information in precise locations in order to meet the MDR requirements. This is because the device does not have its own packaging or leaflet. Finally, as notified to the Agency, the MAH took this opportunity to remove unnecessary details from the quality module 3.2.P.7 currently registered for Onbrez/ Hirobriz/ Oslif Breezhaler." Request for Supplementary Information adopted on 07.11.2019.	Request for supplementary information adopted with a specific timetable.

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

acalabrutinib - EMEA/H/C/005299, Orphan AstraZeneca AB, Treatment of adult patients with chronic lymphocytic leukaemia (CLL)/small lymphocytic lymphoma (SLL)	
bulevirtide - EMEA/H/C/004854, Orphan MYR GmbH, indicated for the treatment of chronic hepatitis delta virus (HDV) infection in adult patients with compensated liver disease.	**Accelerated review**
elexacaftor / tezacaftor / ivacaftor - EMEA/H/C/005269, Orphan Vertex Pharmaceuticals (Ireland) Limited, treatment of cystic fibrosis	**Accelerated review**
insulin aspart - EMEA/H/C/004965 treatment of diabetes mellitus	
Meningococcal group A, C, W135 and Y conjugate vaccine - EMEA/H/C/005084, Article 28 immunization against Neisseria meningitidis serogroups A, C, W-135 and Y	
caffeine citrate - EMEA/H/C/005435 treatment of primary apnoea	
salmeterol xinafoate / fluticasone propionate - EMEA/H/C/004881 for treatment of asthma	
sunitinib - EMEA/H/C/005419 treatment of gastrointestinal stromal tumour (GIST) and metastatic renal cell carcinoma (MRCC) and pancreatic neuroendocrine tumours (pNET)	
influenza quadrivalent vaccine (rDNA) - EMEA/H/C/005159 prevention of influenza disease	

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

Epclusa - sofosbuvir / velpatasvir - EMEA/H/C/004210/X/0043/G Gilead Sciences Ireland UC, Rapporteur: Filip Josephson, PRAC Rapporteur: Ana Sofia Diniz	

Martins, “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.”

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

Pfizer Europe MA EEIG, Generic, Generic of Alimta, Rapporteur: Alar Irs, “Extension application to introduce a new pharmaceutical form with its associated strength (25 mg/ml concentrate for solution for infusion).”

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, “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)

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

Increlex - mecasermin -
EMEA/H/C/000704/S/0061
Ipsen Pharma, Rapporteur: Outi Mäki-Ikola, PRAC Rapporteur: Kirsti Villikka

Myalepta - metreleptin -
EMEA/H/C/004218/S/0009, Orphan
Aegerion Pharmaceuticals B.V., Rapporteur: Bart Van der Schueren, PRAC Rapporteur: Adam Przybylkowski

Raxone - idebenone -
EMEA/H/C/003834/S/0019, Orphan
Santhera Pharmaceuticals (Deutschland) GmbH, Rapporteur: John Joseph Borg, PRAC Rapporteur: Amelia Cupelli

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

Aripiprazole Zentiva - aripiprazole -
EMEA/H/C/003899/R/0012
Zentiva, k.s., Generic, Generic of Abilify, Rapporteur: John Joseph Borg, PRAC Rapporteur: Ana Sofia Diniz Martins

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

Daxas - roflumilast -
EMEA/H/C/001179/R/0039
AstraZeneca AB, Rapporteur: Maria Concepcion Prieto Yerro, Co-Rapporteur: Jayne Crowe, PRAC Rapporteur: Maria del Pilar Rayon

Ivabradine Anpharm - ivabradine -
EMEA/H/C/004187/R/0014
ANPHARM Przedsiebiorstwo Farmaceutyczne S.A., Rapporteur: Johann Lodewijk Hillege, Co-Rapporteur: Outi Mäki-Ikola, PRAC

Rapporteur: Menno van der Elst

Odomzo - sonidegib -
EMEA/H/C/002839/R/0028
Sun Pharmaceutical Industries Europe B.V.,
Rapporteur: Paula Boudewina van Hennik,
Co-Rapporteur: Daniela Melchiorri, PRAC
Rapporteur: Željana Margan Koletić

Pregabalin Sandoz - pregabalin -
EMEA/H/C/004010/R/0012
Sandoz GmbH, Generic, Generic of Lyrica,
Rapporteur: Tomas Radimersky, PRAC
Rapporteur: Liana Gross-Martirosyan

Pregabalin Sandoz GmbH - pregabalin -
EMEA/H/C/004070/R/0013
Sandoz GmbH, Generic, Duplicate, Generic of Lyrica, Duplicate of Pregabalin Sandoz,
Rapporteur: Tomas Radimersky, PRAC
Rapporteur: Liana Gross-Martirosyan

Pregabalin Zentiva - pregabalin -
EMEA/H/C/003900/R/0021
Zentiva k.s., Generic, Generic of Lyrica,
Rapporteur: Alar Irs, PRAC Rapporteur: Liana Gross-Martirosyan

Repatha - evolocumab -
EMEA/H/C/003766/R/0040
Amgen Europe B.V., Rapporteur: Johann Lodewijk Hillege, Co-Rapporteur: Alar Irs, PRAC
Rapporteur: Kimmo Jaakkola

Strensiq - asfotase alfa -
EMEA/H/C/003794/R/0044, Orphan
Alexion Europe SAS, Rapporteur: Daniela Melchiorri, Co-Rapporteur: Jayne Crowe, PRAC
Rapporteur: Rhea Fitzgerald

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

CRYSVITA - burosumab -
EMEA/H/C/004275/II/0010/G, Orphan
Kyowa Kirin Holdings B.V., Rapporteur: Kristina Dunder, Co-Rapporteur: Jayne Crowe, PRAC
Rapporteur: Brigitte Keller-Stanislawski,
“Extension of indication to include treatment of adults with X-linked hypophosphataemia (XLH),

and modification of the currently approved indication in children and adolescents, by removing the qualification 'with growing skeletons', in order to include treatment in all children with radiographic evidence of bone disease.
The application provides new week-48 data from Study UX023-CL304; a randomized, double-blind, placebo-controlled, phase 3 study with open-label extension to assess the efficacy and safety of KRN23 in adults with XLH.
As a consequence, sections 4.1, 4.2, 4.4, 4.8, 5.1 and 5.2 of the SmPC are updated and the Package Leaflet is updated in accordance.
Furthermore, the PI is brought in line with the latest QRD template version 10.1.
The updated RMP version 2.0 has also been submitted."

INTELENCE - etravirine - EMEA/H/C/000900/II/0058

Janssen-Cilag International NV, Rapporteur: Jean-Michel Race, Co-Rapporteur: Bruno Sepodes, PRAC Rapporteur: Adrien Inoubli, "To extend the approved therapeutic indication of Intelence in order to include patient population from 2 to 6 years of age based on the 48 week study results from study TMC125-C234/P1090 (A Phase I/II, Open-label Trial to Evaluate the Safety, Tolerability, Pharmacokinetics and Antiviral Activity of Etravirine (ETR) in Antiretroviral (ARV) Treatment-experienced HIV-1 Infected Infants and Children, Aged ≥2 Months to <6 Years). As a consequence, sections 4.1, 4.2, 4.8, 5.1 and 5.2 of the SmPC and sections 1, 2 and 3 of the PL are updated accordingly. The updated RMP version 13.1 has also been submitted.
The RMP (version 13.1) of the product has been updated to remove the completed additional pharmacovigilance activities (TMC125-C234/P1090 (Week 48) and TMC125-EPPICC) from the pharmacovigilance plan of the RMP). The RMP has also been updated to meet the requirements and updated definitions in the European Medicines Agency (EMA) Guideline on good pharmacovigilance practices (GVP) Module V Revision 2 (EMA/838713/2011; Rev 2) and Guidance on the format of the RMP in the European Union (EMA/164014/2018 Rev 2.0.1) including proposed removal of safety

concerns.
The MAH took the opportunity to include some typographic changes in Annex II C and D."

Kalydeco - ivacaftor -
EMEA/H/C/002494/II/0082, Orphan
Vertex Pharmaceuticals (Ireland) Limited, Rapporteur: Maria Concepcion Prieto Yerro, PRAC Rapporteur: Maria del Pilar Rayon, "Extension of indication to include new population for Kalydeco 150 mg tables to extend the use to patients with cystic fibrosis (CF) aged 6 years and older and weighing 25 kg or more who have an R117H mutation in the CFTR gene and for Kalydeco granules 75 mg and 50 mg, to add patients with CF aged 12 months and older and weighing 7 kg to less than 25 kg who have an R117H mutation in the CFTR gene. This is based on a clinical trial and literature data, and post-marketing experience with Kalydeco. As a consequence, sections 4.1, 4.2, 4.4 and 5.1 of the SmPC are updated. The Package Leaflet is updated in accordance. The RMP version 8.5 has also been submitted."

Kineret - anakinra -
EMEA/H/C/000363/II/0070
Swedish Orphan Biovitrum AB (publ), Rapporteur: Mark Ainsworth, Co-Rapporteur: Fátima Ventura, PRAC Rapporteur: Hans Christian Siersted, "Extension of indication to include the treatment of Familial Mediterranean Fever (FMF) for Kineret, to be given in combination with colchicine, if appropriate; as a consequence, sections 4.1, 4.2, 4.8, 5.1 of the SmPC are updated. The Package Leaflet is updated in accordance. The RMP version 5.0 has also been submitted."

Taltz - ixekizumab -
EMEA/H/C/003943/II/0031
Eli Lilly Nederland B.V., Rapporteur: Kristina Dunder, Co-Rapporteur: Peter Kiely, PRAC Rapporteur: Brigitte Keller-Stanislawski, "Extension of indication to include the treatment of moderate to severe plaque psoriasis in children from the age of 6 years and adolescents who are candidates for systemic therapy for Taltz; as a consequence, sections 4.1, 4.2, 4.4, 4.8, 5.1 and 5.2 of the SmPC are updated with new safety and efficacy information. 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 7.1 has also been submitted."

Tremfya - guselkumab - EMEA/H/C/004271/II/0017

Janssen-Cilag International N.V., Rapporteur: Melinda Sobor, Co-Rapporteur: Peter Kiely, PRAC Rapporteur: Brigitte Keller-Stanislawski, "Addition of a new indication for the treatment of active psoriatic arthritis in adult patients who have had an inadequate response or who have been intolerant to a prior disease-modifying antirheumatic drug (DMARD) therapy. Consequently sections 4.1, 4.2, 4.5, 4.8, 5.1 and 5.2 of the SmPC are updated. The Package leaflet is updated accordingly.
Additionally minor QRD changes are introduced in annex II."

WS1695
Braftovi-EMEA/H/C/004580/WS1695/0008
Mektovi-EMEA/H/C/004579/WS1695/0007

Pierre Fabre Medicament, Lead Rapporteur: Janet Koenig, Lead Co-Rapporteur: Alar Irs, Lead PRAC Rapporteur: Rugile Pilviniene, "Extension of indication to include encorafenib in combination with binimetinib and cetuximab, for the treatment of adult patients with metastatic colorectal cancer (CRC) with a BRAF V600E mutation, who have received prior systemic therapy, as a consequence, sections 4.1, 4.2, 4.4, 4.5, 4.8, 5.1, 5.2, 5.3 of the SmPC are updated. The Package Leaflet is updated in accordance. The RMP version 1.1 has also been submitted. Furthermore, the PI is brought in line with the latest QRD template version 10.1."
Request for 1 year of market protection for a new indication (Article 14(11) of Regulation (EC) 726/2004)

B.6.8. CHMP assessed procedures scope: Pharmaceutical aspects

Ruconest - conestat alfa - EMEA/H/C/001223/II/0052

Pharming Group N.V, Rapporteur: Andrea Laslop

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

B.6.10. CHMP-PRAC assessed procedures

B.6.11. PRAC assessed procedures

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

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 - EMEA CERTIFICATION OF PLASMA MASTER FILES

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

E.1. PMF Certification Dossiers:

E.1.1. Annual Update

E.1.2. Variations:

E.1.3. Initial PMF Certification:

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

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

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

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

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

G. ANNEX G

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

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

G.2. 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 11-14 November 2019 CHMP plenary:

G.3.2. List of procedures starting in November 2019 for December 2019 CHMP adoption of outcomes

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