11 December 2017
EMA/CHMP/741899/2017
Inspections, Human Medicines Pharmacovigilance and Committees Division

Committee for medicinal products for human use (CHMP)

Draft agenda of the meeting on 11-14 December 2017

Chair: Tomas Salmonson – Vice-Chair: Harald Enzmann

11 December 2017, 13:00 – 19:30, room 3A

12 December 2017, 08:30 – 19:30, room 3A

13 December 2017, 08:30 – 19:30, room 3A

14 December 2017, 08:30 – 15:00, room 3A

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 December 2017. See December 2017 CHMP minutes (to be published post January 2018 CHMP meeting).

1.2. Adoption of agenda

CHMP agenda for 11-14 December 2017

1.3. Adoption of the minutes

CHMP minutes for 6-9 November 2017

2. Oral Explanations

2.1. Pre-authorisation procedure oral explanations

2.1.1. plitidepsin - Orphan - EMEA/H/C/004354

Pharma Mar, S.A.; treatment of multiple myeloma

Scope: Oral explanation

Action: Oral explanation to be held on 12 December 2017 at time 11:00

Oral explanation was held on 07.11.2017. List of Outstanding Issues adopted on 14.09.2017. List of Questions adopted on 23.02.2017.

See 3.1

2.1.2. enclomifene - EMEA/H/C/004198

treatment of hypogonadotrophic hypogonadism

Scope: Oral explanation

Action: Oral explanation to be held on 13 December 2017 at time 09:00

List of Outstanding Issues adopted on 12.10.2017. List of Questions adopted on 26.01.2017.

2.1.3. velmanase alfa - Orphan - EMEA/H/C/003922

Chiesi Farmaceutici S.p.A.; indicated for long-term enzyme replacement therapy in patients with alpha-mannosidosis

Scope: Oral explanation

Action: Oral explanation to be held on 12 December 2017 at time 14:00

List of Outstanding Issues adopted on 14.09.2017. List of Questions adopted on 26.01.2017.

See 3.1

2.1.4. human herpesvirus 3 - EMEA/H/C/004336

prevention of herpes zoster (HZ) and HZ-related complications

Scope: Oral explanation

Action: Oral explanation to be held on 12 December 2017 at time 16:00

List of Outstanding Issues adopted on 14.09.2017. List of Questions adopted on 21.04.2017.

2.1.5. abaloparatide - EMEA/H/C/004157

treatment of osteoporosis

Scope: Oral explanation

Action: Oral explanation to be held on 13 December 2017 at time 11:30

List of Outstanding Issues adopted on 20.07.2017, 15.12.2016. List of Questions adopted on 01.04.2016.

2.2. Re-examination procedure oral explanations

No items

2.3. Post-authorisation procedure oral explanations

2.3.1. Opdivo - nivolumab - EMEA/H/C/003985/II/0030

Bristol-Myers Squibb Pharma EEIG

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

Scope: Oral explanation/Opinion

Action: Oral explanation to be held on 11 December 2017 at time 16:00

Request for Supplementary Information adopted on 22.06.2017, 23.03.2017.

See 5.1

2.4. Referral procedure oral explanations

No items

3. Initial applications

3.1. Initial applications; Opinions

3.1.1. hydrocortisone - PUMA - EMEA/H/C/004416

treatment of adrenal insufficiency

Scope: Opinion

Action: For adoption

List of Outstanding Issues adopted on 12.10.2017. List of Questions adopted on 21.04.2017.

3.1.2. expanded human allogeneic mesenchymal adult stem cells extracted from adipose tissue - Orphan - ATMP - EMEA/H/C/004258

Tigenix, S.A.U.; treatment of complex perianal fistula(s)

Scope: Opinion

Action: For adoption

List of Outstanding Issues adopted on 17.02.2017. List of Questions adopted on 15.07.2016.

3.1.3. anagrelide - EMEA/H/C/004585

reduction of elevated platelet counts in at risk essential thrombocythaemia patients

Scope: Opinion

Action: For adoption

List of Outstanding Issues adopted on 14.09.2017. List of Questions adopted on 21.04.2017.

3.1.4. plitidepsin - Orphan - EMEA/H/C/004354

Pharma Mar, S.A.; treatment of multiple myeloma

Scope: Opinion/Oral explanation

Action: For adoption

Oral explanation was held on 07.11.2017. List of Outstanding Issues adopted on 14.09.2017. List of Questions adopted on 23.02.2017.

See 2.1

3.1.5. burosumab - Orphan - EMEA/H/C/004275

Kyowa Kirin Limited; treatment of X-linked hypophosphataemia (XLH)

Scope: Opinion

Action: For adoption

List of Outstanding Issues adopted on 12.10.2017. List of Questions adopted on 21.04.2017.

3.1.6. trastuzumab - EMEA/H/C/002575

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

Scope: Opinion

Action: For adoption

List of Outstanding Issues adopted on 09.11.2017. List of Questions adopted on 23.02.2017.

3.1.7. velmanase alfa - Orphan - EMEA/H/C/003922

Chiesi Farmaceutici S.p.A.; indicated for long-term enzyme replacement therapy in patients with alpha-mannosidosis

Scope: Opinion

Action: For adoption

List of Outstanding Issues adopted on 14.09.2017. List of Questions adopted on 26.01.2017.

See 2.1.

3.1.8. semaglutide - EMEA/H/C/004174

to improve glycaemic control in adults with type 2 diabetes and to prevent cardiovascular events

Scope: Opinion

Action: For adoption

List of Outstanding Issues adopted on 09.11.2017, 14.09.2017. List of Questions adopted on 21.04.2017.

3.1.9. rucaparib - Orphan - EMEA/H/C/004272

Clovis Oncology UK Ltd; treatment of ovarian cancer

Scope: Opinion

Action: For adoption

Oral explanation held on 08.11.2017. List of Outstanding Issues adopted on 09.11.2017, 14.09.2017. List of Questions adopted on 23.03.2017.

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

3.2.1. glibenclamide - Orphan - EMEA/H/C/004379

Ammtek; treatment of neonatal diabetes

Scope: 2nd 180 list of outstanding issue

Action: For adoption

List of Outstanding Issues adopted on 22.06.2017. List of Questions adopted on 24.01.2017.

3.2.2. betrixaban - EMEA/H/C/004309

treatment of prophylaxis of venous thromboembolism (VTE)

Scope: 2nd 180 list of outstanding issue, Report from SAG

Action: For adoption

List of Outstanding Issues adopted on 12.10.2017. List of Questions adopted on 21.04.2017.

3.2.3. efavirenz / emtricitabine / tenofovir disoproxil - EMEA/H/C/004274

treatment of HIV-1 infection

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 14.09.2017.

3.2.4. eteplirsen - Orphan - EMEA/H/C/004355

AVI Biopharma International Ltd; treatment of Duchenne muscular dystrophy

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 21.04.2017.

3.2.5. emicizumab - EMEA/H/C/004406

Accelerated assessment

routine prophylaxis to prevent bleeding or reduce the frequency of bleeding episodes in patients with haemophilia A (congenital factor VIII deficiency) with factor VIII inhibitors.

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 10.10.2017.

3.2.6. metreleptin - Orphan - EMEA/H/C/004218

Aegerion Pharmaceuticals Limited; treatment of leptin deficiency (lipodystrophy)

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 18.05.2017.

3.2.7. gemtuzumab ozogamicin - Orphan - EMEA/H/C/004204

Pfizer Limited; combination therapy with daunorubicin (DNR) and cytarabine (AraC) for the treatment of adult patients with previously untreated, de novo acute myeloid leukaemia (AML)

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 21.04.2017.

3.2.8. prasugrel - EMEA/H/C/004644

prevention of atherothrombotic events

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 14.09.2017.

3.2.9. pegfilgrastim - EMEA/H/C/004413

treatment of neutropenia

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 23.03.2017.

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

3.3.1. binimetinib - EMEA/H/C/004579

in combination with encorafenib is indicated for the treatment of adult patients with

unresectable or metastatic melanoma with a BRAF V600 mutation

Scope: Day 120 list of questions

Action: For adoption

3.3.2. encorafenib - EMEA/H/C/004580

in combination with binimetinib is indicated for the treatment of adult patients with unresectable or metastatic melanoma with a BRAF V600 mutation

Scope: Day 120 list of questions

Action: For adoption

3.3.3. gefitinib - EMEA/H/C/004826

treatment of non-small cell lung cancer

Scope: Day 120 list of questions

Action: For adoption

3.3.4. vigabatrin - PUMA - EMEA/H/C/004534

Treatment in monotherapy of infantile spasms (West's syndrome) and resistant partial epilepsy in infants and children

Scope: Day 120 list of questions

Action: For adoption

3.3.5. lenalidomide - EMEA/H/C/004857

treatment of multiple myeloma

Scope: Day 120 list of questions

Action: For adoption

3.3.6. voretigene neparvovec - Orphan - ATMP - EMEA/H/C/004451

Spark Therapeutics Ireland Ltd; treatment of patients with vision loss due to Leber congenital amaurosis or retinitis pigmentosa inherited retinal dystrophy

Scope: Day 120 list of questions

Action: For information

3.3.7. mexiletine hydrochloride - Orphan - EMEA/H/C/004584

Lupin (Europe) Limited; Treatment of myotonic disorders

Scope: Day 120 list of questions

Action: For adoption

3.3.8. paclitaxel - EMEA/H/C/004441

treatment of metastatic breast cancer

Scope: Day 120 list of questions

Action: For adoption

3.3.9. tezacaftor / ivacaftor - Orphan - EMEA/H/C/004682

Vertex Pharmaceuticals (Europe) Ltd.; treatment of cystic fibrosis

Scope: Day 120 list of questions

Action: For adoption

3.3.10. abemaciclib - EMEA/H/C/004302

treatment of hormone receptor (HR) positive, human epidermal growth factor receptor 2 (HER2) negative locally advanced or metastatic breast cancer

Scope: Day 120 list of questions

Action: For adoption

3.3.11. volanesorsen - Orphan - EMEA/H/C/004538

Akcea Therapeutics UK Ltd.; indicated as an adjunct to diet for the treatment of patients with familial chylomicronemia syndrome (FCS).

Scope: Day 120 list of questions

Action: For adoption

3.3.12. eravacycline - EMEA/H/C/004237

treatment of complicated intra-abdominal infections (cIAI) in adults

Scope: Day 120 list of questions

Action: For adoption

3.3.13. axicabtagene ciloleucel - Orphan - ATMP - EMEA/H/C/004480

Accelerated assessment

Kite Pharma EU B.V.; treatment of diffuse large B-cell lymphoma (DLBCL), primary mediastinal B-cell lymphoma (PMBCL) and transformed follicular lymphoma (TFL)

Scope: Day 120 list of questions

Action: For information

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

3.4.1. carmustine - EMEA/H/C/004326

treatment of brain tumors, multiple myeloma, Hodgkin's disease and non-Hodgkin's lymphomas

Scope: Request for extension of clock stop to respond to List of outstanding issue adopted on 12.10.2017

Action: For adoption

List of Outstanding Issues adopted on 12.10.2017, 20.07.2017. List of Questions adopted on 13.10.2016.

3.4.2. adalimumab - EMEA/H/C/004429

treatment of rheumatoid arthritis, psoriatic arthritis and ankylosing spondylitis

Scope: Request for extension of clock stop to respond to Day 120 list of questions adopted on 14.09.2017

Action: For adoption

List of Questions adopted on 14.09.2017

3.4.3. tildrakizumab - EMEA/H/C/004514

treatment of adults with moderate-to-severe plaque psoriasis

Scope: Request for extension of clock stop to respond to List of outstanding issue adopted on 27.07.2017

Action: For information

List of Questions adopted on 27.07.2017.

The CHMP adopted the extension of clock stop by written procedure on 22.11.2017.

3.4.4. rotigotine - EMEA/H/C/004286

treatment of idiopathic Restless Legs Syndrome and Parkinson's disease

Scope: Request by the applicant dated 30 November 2017 requesting an extension of clock stop to respond to the List of Questions adopted on 21 April 2017.

Action: For adoption

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

No items

3.6. Initial applications in the decision-making phase

No items

3.7. Withdrawals of initial marketing authorisation application

No items

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

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

No items

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

4.2.1. Lynparza - olaparib - Orphan - EMEA/H/C/003726/X/0016/G

AstraZeneca AB

Rapporteur: Alexandre Moreau, Co-Rapporteur: Bart Van der Schueren, PRAC Rapporteur: Carmela Macchiarulo

Scope: “Extension application to add a new pharmaceutical form associated with a new strength (100mg and 150 mg film-coated tablets) including an extension of the indication to treat patients with platinum-sensitive relapsed ovarian tumours. The extension application is grouped with a type II variation to align the PI for the currently authorised capsule licence with the safety updates proposed for the tablet formulation.”

Action: For adoption

List of Questions adopted on 14.09.2017.

4.2.2. Nuwiq - simoctocog alfa - EMEA/H/C/002813/X/0020

Octapharma AB

Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Ulla Wändel Liminga

Scope: “Extension application to add new strengths of 2500 IU, 3000 IU, 4000 IU for Nuwiq, powder and solvent for solution for injection.
The RMP (version 5.4) is updated accordingly.”

Action: For adoption

List of Questions adopted on 14.09.2017.

4.2.3. Daliresp - roflumilast - EMEA/H/C/002398/X/0031

AstraZeneca AB

Rapporteur: Concepcion Prieto Yerro, Co-Rapporteur: Jayne Crowe, PRAC Rapporteur: Dolores Montero Corominas

Scope: “Extension application to add a new strength of 250 µg in a PVC/PVDC/Alu blister of 28 tablets.”

Action: For adoption

List of Questions adopted on 20.07.2017.

4.2.4. Daxas - roflumilast - EMEA/H/C/001179/X/0035

AstraZeneca AB

Rapporteur: Concepcion Prieto Yerro, Co-Rapporteur: Jayne Crowe, PRAC Rapporteur: Dolores Montero Corominas

Scope: “Extension application to add a new strength of 250 µg in a PVC/PVDC/Alu blister of 28 tablets.”

Action: For adoption

List of Questions adopted on 20.07.2017.

4.2.5. Libertek - roflumilast - EMEA/H/C/002399/X/0032

AstraZeneca AB

Rapporteur: Concepcion Prieto Yerro, Co-Rapporteur: Jayne Crowe, PRAC Rapporteur: Dolores Montero Corominas

Scope: “Extension application to add a new strength of 250 µg in a PVC/PVDC/Alu blister of 28 tablets.”

Action: For adoption

List of Questions adopted on 20.07.2017.

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. Bosulif - bosutinib - Orphan - EMEA/H/C/002373/X/0026

Pfizer Limited

Rapporteur: Harald Enzmann

Scope: “Extension application to add a new strength of 400mg film-coated tablets.”

Action: For adoption

4.3.2. Inhixa - enoxaparin sodium - EMEA/H/C/004264/X/0018

Techdow Europe AB

Rapporteur: Andrea Laslop, PRAC Rapporteur: Menno van der Elst

Scope: "Extension application to add two new strengths of 12,000 IU (120 mg)/0.8 mL and 15,000 IU (150 mg)/1 mL for enoxaparin sodium solution for injection in pre-filled syringe, for subcutaneous, extracorporeal and intravenous administration."

Action: For adoption

4.3.3. Xeljanz - tofacitinib - EMEA/H/C/004214/X/0005/G

Pfizer Limited

Rapporteur: Robert James Hemmings, PRAC Rapporteur: Sabine Straus

Scope: "Extension application to introduce a new strength (10 mg film coated tablets). In addition, the MAH proposed a type II variation (C.I.6.a) to extend the indication to include 'the induction and maintenance of treatment of adult patients with moderately to severely active ulcerative colitis (UC) who have had an inadequate response, lost response, or were intolerant to either conventional therapy or a biologic agent"

Action: For adoption

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

No items

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

No items

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

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

5.1.1. Blincyto - blinatumomab - Orphan - EMEA/H/C/003731/II/0011

Amgen Europe B.V.

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

Scope: “Extension of Indication to include the treatment of adults with minimal residual disease (MRD) positive B-cell precursor acute lymphoblastic leukaemia (ALL)for BLINCYTO; as a consequence, sections 4.1, 4.2, 4.8, 5.1 and 5.2 of the SmPC are updated in order to add the new indication and its relevant posology, and to update the safety information. The Labelling is updated in accordance.
RMP version 4.0 is included in this submission”. Report from SAG Oncology.

Action: For adoption

Request for Supplementary Information adopted on 22.06.2017.

5.1.2. Cabometyx - cabozantinib - EMEA/H/C/004163/II/0003

Ipsen Pharma

Rapporteur: Robert James Hemmings, Co-Rapporteur: Bjorg Bolstad, PRAC Rapporteur: Sabine Straus

Scope: “Extension of indication to include for the treatment of advanced renal cell carcinoma the ‘treatment-naïve adults with intermediate or poor risk per IMDC criteria’ for CABOMETYX; as a consequence, sections 4.1, 4.4, 4.8 and 5.1 of the SmPC are updated in order to add a warning on dose reductions and dose interruptions and to update the safety information. The final report of the randomised phase II study comparing cabozantinib with commercially supplied sunitinib in subjects with previously untreated locally advanced or metastatic renal cell carcinoma (study A031203) is submitted in support of this application. The Package Leaflet is updated accordingly. The risk management plan (version 3.0) is also submitted.
In addition, the Marketing authorisation holder (MAH) took the opportunity to make some editorial changes in the product information.”

Action: For adoption

5.1.3. Feraccru - ferric maltol - EMEA/H/C/002733/II/0010

Shield TX (UK) Ltd

Rapporteur: Concepcion Prieto Yerro, Co-Rapporteur: Harald Enzmann, PRAC Rapporteur: Adam Przybylkowski

Scope: “Extension of indication to widen the indication for Feraccru from the treatment “in adults with Iron deficiency anaemia in patients with IBD” to the treatment of “adults with Iron deficiency”; As a consequence, sections 4.1, 4, 5.1 and 5.2 of the SmPC are updated. The Package Leaflet has been updated accordingly.
In addition, the Marketing authorisation holder (MAH) took the opportunity to update the list of local representatives in the Package Leaflet.”

Action: For adoption

5.1.4. Hizentra - human normal immunoglobulin - EMEA/H/C/002127/II/0087

CSL Behring GmbH

Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Brigitte Keller-Stanislawski

Scope: “Extension of Indication to include immunomodulatory therapy for the treatment of patients with chronic inflammatory demyelinating polyneuropathy (CIDP) as maintenance therapy to prevent relapse of neuromuscular disability and impairment. As a consequence, sections 4.1, 4.2, 4.8, 5.1 and 5.2 of the SmPC are updated. The Package Leaflet is updated in accordance. The RMP is updated (v. 4.0)”

Action: For adoption

Request for Supplementary Information adopted on 12.10.2017.

5.1.5. Inovelon - rufinamide - Orphan - EMEA/H/C/000660/II/0045

Eisai Ltd

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

Scope: “Extension of indication to include the treatment of seizures associated with Lennox Gastaut syndrome in patients 1 year of age and older as adjunctive therapy. As a consequence sections 4.1, 4.2, 4.5, 5.1 and 5.2. The Package Leaflet and the RMP (version 10.0) are updated accordingly. In addition the Marketing Authorisation Holder (MAH) took the opportunity to make small corrections with the Product Information and to update the name and contact details of the local representative in Belgium and Luxembourg. Furthermore, the Product Information is brought in line with the latest QRD template version 10.”

Action: For adoption

5.1.6. Kalydeco - ivacaftor - Orphan - EMEA/H/C/002494/II/0063/G

Vertex Pharmaceuticals (Europe) Ltd.

Rapporteur: Concepcion Prieto Yerro, PRAC Rapporteur: Dolores Montero Corominas

Scope: “1) C.I.6.a (type II) - Extension of Indication to include the combination regimen of the ivacaftor 150 mg evening dose and Symkevi (tezacaftor/ivacaftor);
2) B.II.e.5.a.2 (type IB) - to add a blister card pack presentation containing 28-tablets for the 150 mg film-coated tablets (EU/1/12/782/005);
3) B.II.e.5.a.2 (type IB) - to add a blister pack presentation containing 28-tablets for the 150 mg film-coated tablets (EU/1/12/782/006).
As a consequence, section4.1, 4.2, 4.4, 4.5, 4.8, 6.5 and 8 of the SmPC are updated. Annex A, the Package Leaflet and Labelling are updated in accordance.
An updated RMP (version 6.0) is included.”

Action: For adoption

5.1.7. Opdivo - nivolumab - EMEA/H/C/003985/II/0030

Bristol-Myers Squibb Pharma EEIG

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

Scope: “Extension of indication to include treatment of adults with mismatch repair deficient

(dMMR) or microsatellite instability high (MSI-H) metastatic colorectal cancer after prior fluoropyrimidine based therapy for Opdivo.
As a consequence, sections 4.1, 4.4, 4.8 and 5.1 of the SmPC are updated in order to add the new indication and update the safety information. The Package Leaflet is updated in accordance.
RMP version 9.0 is submitted with this application"

Report from SAG Oncology held 21 November 2017

Action: For adoption

Request for Supplementary Information adopted on 22.06.2017, 23.03.2017.

See 2.3

5.1.8. Opdivo - nivolumab - EMEA/H/C/003985/II/0039

Bristol-Myers Squibb Pharma EEIG

Rapporteur: Jorge Camarero Jiménez, PRAC Rapporteur: Brigitte Keller-Stanislawski

Scope: "Extension of Indication to include treatment of adult patients with advanced or recurrent gastric or gastroesophageal junction (GEJ) cancer after two or more prior systemic therapies, based on data from study ONO-4538-12. As a consequence, sections 4.1, 4.4, 4.8, and 5.1 of the SmPC are updated. The Annex II and Package Leaflet are updated in accordance. The RMP version version 11.0 has also been submitted."

Action: For adoption

5.1.9. Perjeta - pertuzumab - EMEA/H/C/002547/II/0034

Roche Registration Limited

Rapporteur: Sinan B. Sarac, Co-Rapporteur: Daniela Melchiorri, PRAC Rapporteur: Doris Stenver

Scope: "Extension of Indication for Perjeta in combination with trastuzumab and chemotherapy for the adjuvant treatment of adult patients with HER2-positive early breast cancer. The submission is based on the primary analysis of efficacy and safety data from the pivotal Phase III study BIG-4-11/BO25126/TOC4939g (APHINITY). With the submission of the APHINITY data, the MAH also aims to fulfil the Annex IID obligation from the approval of the neoadjuvant indication of Perjeta granted in 2015. Sections 4.2, 4.2, 4.4, 4.8, 5.1 and 5.2 of the SmPC are. Annex II and the Package Leaflet have been updated accordingly. The RMP version 10.0 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

5.1.10. Repatha - evolocumab - EMEA/H/C/003766/II/0017/G

Amgen Europe B.V.

Rapporteur: Johann Lodewijk Hillege, Co-Rapporteur: Alar Irs, PRAC Rapporteur: Kimmo Jaakkola

Scope: “Extension of Indication to include reduction of atherosclerotic cardiovascular disease risk in adults with high cardiovascular risk for Repatha based on the results from Study 20110118 (a category 3 PV activity in the Risk Management Plan, MEA 004); as a consequence, sections 4.1, 4.2, 4.8 and 5.1 of the SmPC are updated. The Package Leaflet is updated in accordance.

In addition, the Marketing authorisation holder (MAH) took the opportunity to update section 5.1 of the SmPC to include important mechanistic information for healthcare professionals based on Study 20120153 (a category 3 PV activity, MEA 006).
Submission of an updated RMP version 2.0 in order to add two category 3 studies in the RMP (Study 20160250 and Study 20150338), as well as to update the milestones of five category 3 studies (20110110, 20110271, 20120138, 20130286, 20130295)”

Action: For adoption

Request for Supplementary Information adopted on 14.09.2017.

5.1.11. Taltz - ixekizumab - EMEA/H/C/003943/II/0009

Eli Lilly Nederland B.V.

Rapporteur: Kristina Dunder, Co-Rapporteur: Greg Markey, PRAC Rapporteur: Brigitte Keller-Stanislawski

Scope: “Extension of Indication to include alone or in combination with conventional disease-modifying anti-rheumatic drug (cDMARD), the treatment of active psoriatic arthritis in adult patients who have responded inadequately to, or who are intolerant to one or more DMARD therapies. As a consequence, sections 4.1, 4.2, 4.4, 4.5, 4.8, 5.1, and 5.2 of the SmPC are updated to reflect the new safety and efficacy information. The Package Leaflet and RMP have been updated accordingly.”, Request for 1 year of market protection for a new indication (Article 14(11) of Regulation (EC) 726/2004)

Action: For adoption

Request for Supplementary Information adopted on 14.09.2017.

5.1.12. Translarna - ataluren - Orphan - EMEA/H/C/002720/II/0037

PTC Therapeutics International Limited

Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Sabine Straus

Scope: “Extension of Indication to include a new population (children from 2 to less than 5 years of age) for Translarna; as a consequence, sections 4.1, 4.2, 4.8, 5.1 and 5.2 of the SmPC are updated. The Package Leaflet and RMP (version 7.1) is updated in accordance.”

Action: For adoption

5.1.13. Truvada - emtricitabine / tenofovir disoproxil - EMEA/H/C/000594/II/0135

Gilead Sciences International Limited

Rapporteur: Greg Markey, Co-Rapporteur: Alexandre Moreau, PRAC Rapporteur: Julie Williams

Scope: “Extension of Indication to include pre-exposure prophylaxis of HIV in adolescents aged 12 to < 18 years at high risk; as a consequence, sections 4.1, 4.2, 4.4, 4.8, 5.1 and 5.2 of the SmPC are updated based on extrapolation of data for emtricitabine, tenofovir disoproxil fumarate, and Truvada in HIV-infected and uninfected subjects.

The Package Leaflet and Risk Management Plan (v.15) are updated in accordance.

The requested variation proposed amendments to the Summary of Product Characteristics, Package Leaflet and Risk Management Plan (RMP).

In addition, the Marketing authorisation holder (MAH) took the opportunity to make minor linguistic amendments”

Action: For adoption

Request for Supplementary Information adopted on 14.09.2017, 18.05.2017.

5.1.14. Xeljanz - tofacitinib - EMEA/H/C/004214/II/0006

Pfizer Limited

Rapporteur: Robert James Hemmings, Co-Rapporteur: Daniela Melchiorri, PRAC Rapporteur: Sabine Straus

Scope: “Extension of Indication to include treatment of adult patients with active psoriatic arthritis who have had an inadequate response or who have been intolerant to a prior disease-modifying anti-rheumatic drug (DMARD) therapy, based on data from studies A3921091, A3921092, A3921125. As a consequence, sections 4.1, 4.2, 4.4, 4.5, 4.8, 5.1 and 5.2 of the SmPC are updated. The Package Leaflet is updated in accordance. In addition, the Marketing authorisation holder (MAH) took the opportunity to update the Annex II with minor editorial changes. The RMP version 3.0 has also been submitted.”

Action: For adoption

5.1.15. Yervoy - ipilimumab - EMEA/H/C/002213/II/0044

Bristol-Myers Squibb Pharma EEIG

Rapporteur: Paula Boudewina van Hennik, PRAC Rapporteur: Sabine Straus

Scope: “Extension of indication to include the treatment of advanced (unresectable or metastatic) melanoma in children and adolescents 12 years of age and older for Yervoy. As a consequence sections 4.1, 4.2, 4.4, 4.5, 4.8, 5.1 and 5.2 of the SmPC are updated. The Package Leaflet and the RMP (version 15) are updated in accordance.”

Action: For adoption

Request for Supplementary Information adopted on 14.09.2017, 22.06.2017.

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

No items

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

5.3.1. Raxone - idebenone - Orphan - EMEA/H/C/003834/II/0003

Santhera Pharmaceuticals (Deutschland) GmbH

Rapporteur: John Joseph Borg, Co-Rapporteur: Andrea Laslop, PRAC Rapporteur: Carmela Macchiarulo

Scope: List of questions for the SAG

Action: For adoption

Opinion adopted on 14.09.2017

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

No items

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. Blincyto - blinatumomab - Orphan - EMEA/H/C/003731/II/0009

Amgen Europe B.V.

Rapporteur: Alexandre Moreau

Scope: "Update of sections 4.4, 4.8, 5.1 and 5.2 of the SmPC in order to update the safety information with the data from the study 103311. This study is fulfilling the specific obligation for the conditional MA. The SO is removed from annex II. The Package Leaflet is updated accordingly.
The MAH takes this opportunity to amend the format of the preparation instructions to improve clarity. The content is not impacted."

Action: For adoption

Request for Supplementary Information adopted on 20.07.2017, 26.01.2017.

9.1.2. Blincyto - blinatumomab - Orphan - EMEA/H/C/003731/R/0013

Amgen Europe B.V.

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

Scope: "Renewal"

Action: For adoption

9.1.3. Enbrel – etanercept - EMEA/H/C/000262 – WS1190 - 0210/G and Lifmior - EMEA/H/C/004167 - WS1190 - 0009/G

Pfizer Limited

Lead Rapporteur: Robert James Hemmings, Lead PRAC Rapporteur: Patrick Batty

Action: For discussion

Request for Supplementary Information adopted on 30.11.2017, 28.09.2017.

10. Referral procedures

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

No items

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

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

December 2017 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

Impact of Brexit on manufacturing and supply

Action: For information

12.2. GCP inspections

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

GCP Inspection Programme 2018-2019

Action: For adoption

12.3. Pharmacovigilance inspections

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

12.4. GLP inspections

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

13. Innovation Task Force

13.1. Minutes of Innovation Task Force

No items

13.2. Innovation Task Force briefing meetings

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

13.4. Nanomedicines activities

No items

14. Organisational, regulatory and methodological matters

14.1. Mandate and organisation of the CHMP

14.1.1. Area of expertise of co-opted member

Jean-Louis Robert will step down as CHMP co-opted member by end of the year 2017 (expertise in Quality (non-biologicals)).

Scope: Discussion on area of expertise

The discussion and agreement on the area of expertise is planned for the December 2017 Plenary. Proposals should be sent by **11 December 2017**.

Action: For adoption

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 27-30 November 2017

Action: For information

List of Union Reference Dates and frequency of submission of Periodic Safety Update

Reports (EURD list) for December 2017

Action: For adoption

14.2.2. Committee for Advanced Therapies (CAT)

CAT draft minutes of meeting held on 6-8 December 2017
Action: For information

14.2.3. Committee for Herbal Medicinal Products (HMPC)

Report from the HMPC meeting held on 20-21 November 2017

Action: For information

14.2.4. Paediatric Committee (PDCO)

PIPs reaching D30 at December 2017 PDCO

Action: For information

Report from the PDCO meeting held on 12-15 December 2017

Action: For information

Joint CHMP/PDCO session

Agenda for joint session

Action: For discussion

14.2.5. Committee for Orphan Medicinal Products (COMP)

Report from the COMP meeting held on 5-7 December 2017
Action: For information

14.2.6. 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 11-13 December 2017

Action: For information

14.2.7. Procedural advice on the evaluation of advanced therapy medicinal product in accordance with Article 8 of Regulation (EC) No 1394/2007

Scope: Present and adopt the final updated procedural advice for ATMP

Action: For adoption

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

14.3.1. Scientific Advice Working Party (SAWP)

Report from the SAWP meeting held on 27-30 November 2017. 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.

Re re-nomination of the SAWP members

Action: For adoption

14.3.2. Name Review Group (NRG)

Table of Decisions of the NRG meeting held on 22 November 2017.

Action: For adoption

14.3.3. Biologics Working Party (BWP)

Chair: Sol Ruiz/Nanna Aaby Kruse

Reports from BWP December 2017 meeting to CHMP

Action: For adoption

Work plan for the Biologics Working Party (BWP) for 2018 (EMA/CHMP/BWP/400117/2017)

Action: For adoption

14.3.4. Cardiovascular Working Party (CVSWP)

Chair: Pieter de Graeff/Kristina Dunder

Election of CVSWP Chair following resignation of Pieter de Graeff

Nominations should be sent to the CVSWP Secretariat by **7 December 2017.**

Action: For adoption

Work plan for the Cardiovascular Working Party (CVSWP) for 2018 (EMA/CHMP/365586/2017)

Action: For adoption

14.3.5. Biostatistics Working Party (BSWP)

Chair: Anja Schiel/Thomas Lang

Call for nominations for BSWP Vice-Chair.

Nominations should be sent by **15th January 2018**. Elections will take place at the January CHMP Plenary meeting.

Action: For information

Work plan for the Biostatistics Working Party (BSWP) for 2018 (EMA/CHMP/333668/2017)

Action: For adoption

14.3.6. Safety Working Party (SWP)

Chair: Jan Willem Van der Laan

Work plan for the Safety Working Party (SWP) for 2018 (EMA/CHMP/SWP/420217/2017)

Action: For adoption

14.3.7. Quality Working Party (QWP)

Chair: Keith Pugh

Work plan for the Quality Working Party (QWP) for 2018 (EMA/CHMP/CVMP/QWP/504882/2017)

Action: For adoption

14.3.8. Biosimilar Medicinal Product Working Party (BMWP)

Chair: Elena Wolff-Holz/Niklas Ekman

Work plan for the Biosimilar Medicinal Product Working Party (BMWP) for 2018 (EMA/CHMP/BMWP/420509/2017)

Action: For adoption

14.3.9. Vaccines Working Party (VWP)

Chair: Mair Powell/Svein Rune Andersen

Work plan for the Vaccines Working Party (VWP) for 2018 (EMA/CHMP/VWP/515395/2017)

Action: For adoption

14.3.10. Blood Products Working Party (BPWP)

Chair: Jacqueline Kerr

Work plan for the Blood Products Working Party (BPWP) for 2018 (EMA/CHMP/BPWP/399637/2017)

Action: For adoption

14.3.11. Pharmacogenomics Working Party (PGWP)

Chair: Krishna Prasad/Markus Paulmichl

Work plan for the Pharmacogenomics Working Party (PGWP) for 2018 (EMA/CHMP/370931/2017)

Action: For adoption

14.3.12. Infectious Diseases Working Party (IDWP)

Chair: Maria Jesus Fernandez Cortizo

Work plan for the Infectious Diseases Working Party (IDWP) for 2018 (EMA/517878/2017)

Action: For adoption

14.3.13. Pharmacokinetics Working Party (PKWP)

Chair: Jan Welink/Henrike Potthast

Work plan for the Pharmacokinetics Working Party (PKWP) for 2018 (EMA/CHMP/365756/2017)

Action: For adoption

Reflection paper on investigation of pharmacokinetics and pharmacodynamics in the obese population

Action: For adoption for 6 months public consultation

14.3.14. Rheumatology/Immunology Working Party (RIWP)

Chair: Jan Mueller-Berghaus

Work plan for the Rheumatology/Immunology Working Party (RIWP) for 2018 (EMA/799178/2017)

Action: For adoption

Guideline on clinical investigation of medicinal products for the treatment of rheumatoid arthritis (CPMP/EWP/556/95 Rev. 2)

Action: For adoption

14.3.15. Gastroenterology Drafting Group (GDG)

Chair: Elmer Schabel

Work Plan for the Gastroenterology Drafting Group (GDG) for 2018 (EMA/CHMP/350919/2017)

Action: For adoption

14.3.16. Respiratory Drafting Group (RDG)

Chair: Karolina Törneke

Work plan for the Respiratory Drafting Group (RDG) for 2018 (EMA/CHMP/383892/2017)

Action: For adoption

14.3.17. Excipients Drafting Group (ExcpDG)

Chair: Dominique Masset

Work plan for the CHMP Excipients Drafting Group (ExcpDG) for the revision of the EC guideline 'Excipients in the labelling and package leaflet of medicinal products for human use' for 2018 (EMA/CHMP/672198/2017)

Action: For adoption

14.4. Cooperation within the EU regulatory network

14.4.1. Draft Commission Regulation amending Regulation (EC) No 847/2000 as regards the definition of the concept 'similar medicinal product'

Letter from the European Commission

Action: For information

14.5. Cooperation with International Regulators

14.5.1. International Council for Harmonisation of Technical Requirements for Pharmaceuticals for Human Use (ICH)

ICH report to CHMP

Action: For information

Scope: ICH guideline E17 - Step 5 -: general principles for planning and design of multi-regional clinical trials (EMA/CHMP/ICH/453276/2016)

Action: For adoption

Scope: S3A Q&A - Step 5 - Note for guidance on toxicokinetics the assessment of systemic exposure in toxicity studies - questions and answers (EMA/CHMP/ICH/320985/2016)

Action: For adoption

Scope: Q12 draft Guideline 'Technical and regulatory considerations for pharmaceutical product lifecycle management' – Step 2 - (EMA/804273/2017)

Action: For adoption

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

None

14.7. CHMP work plan

14.7.1. CHMP 2018 Work Plan

Action: For adoption

14.8. Planning and reporting

None

14.9. Others

None

15. Any other business

15.1. AOB topic

15.1.1. Preparedness of the system and capacity increase

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)*

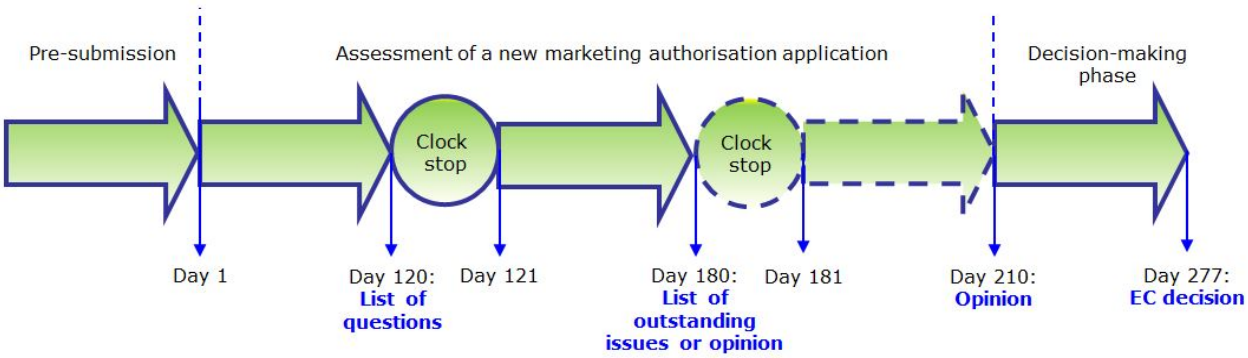
The items listed in this section are those for which marketing authorisation holders (MAHs) or applicants have been invited to the CHMP plenary meeting to address questions raised by the Committee. Oral explanations normally relate to on-going applications (section 3, 4 and 5) or referral procedures (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 December 2017
EMA/CHMP/741900/2017

Annex to 11-14 December 2017 CHMP Agenda

Pre submission and post authorisations issues

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Send a question via our website www.ema.europa.eu/contact

A. PRE SUBMISSION ISSUES

A.1. ELIGIBILITY REQUESTS

Report on Eligibility to Centralised Procedure for December 2017: **For adoption**

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

Final Outcome of Rapporteurship allocation for December 2017: **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

Strensiq - asfotase alfa - EMEA/H/C/003794/S/0024, Orphan
MAH: Alexion Europe SAS, Rapporteur: Greg Markey, PRAC Rapporteur: Almath Spooner

B.2. RENEWALS OF MARKETING AUTHORISATIONS OUTCOMES

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

B.2.2. Renewals of Marketing Authorisations for unlimited validity

Iclusig - ponatinib - EMEA/H/C/002695/R/0042, Orphan
MAH: Incyte Biosciences UK Ltd, Rapporteur: Greg Markey, Co-Rapporteur: Filip Josephson, PRAC Rapporteur: Patrick Batty

Memantine ratiopharm - memantine - EMEA/H/C/002671/R/0011
MAH: ratiopharm GmbH, Generic, Generic of Ebixa, Rapporteur: Bart Van der Schueren, PRAC Rapporteur: Dolores Montero Corominas

Mycamine - micafungin - EMEA/H/C/000734/R/0034

MAH: Astellas Pharma Europe B.V., Rapporteur: Harald Enzmann, Co-Rapporteur: Koenraad Norga, PRAC Rapporteur: Martin Huber
Request for Supplementary Information adopted on 12.10.2017.

Spedra - avanafil - EMEA/H/C/002581/R/0029
MAH: Menarini International Operations Luxembourg S.A., Rapporteur: Concepcion Prieto Yerro, Co-Rapporteur: Daniela Melchiorri, PRAC Rapporteur: Dolores Montero Corominas

Stribild - elvitegravir / cobicistat / emtricitabine / tenofovir disoproxil - EMEA/H/C/002574/R/0086
MAH: Gilead Sciences International Limited, Rapporteur: Robert James Hemmings, Co-Rapporteur: Joseph Emmerich, PRAC Rapporteur: Julie Williams

Thalidomide Celgene - thalidomide - EMEA/H/C/000823/R/0054, Orphan
MAH: Celgene Europe Limited, Rapporteur: Alexandre Moreau, Co-Rapporteur: Filip Josephson, PRAC Rapporteur: Ghania Chamouni
Request for Supplementary Information adopted on 12.10.2017.

Voriconazole Accord - voriconazole - EMEA/H/C/002669/R/0017
MAH: Accord Healthcare Limited, Generic, Generic of Vfend, Rapporteur: John Joseph Borg, PRAC Rapporteur: Menno van der Elst

B.2.3. Renewals of Conditional Marketing Authorisations

Blincyto - blinatumomab - EMEA/H/C/003731/R/0013, Orphan
See 9.1
MAH: Amgen Europe B.V., Rapporteur: Alexandre Moreau, Co-Rapporteur: Daniela Melchiorri, PRAC Rapporteur: Eva Jirsová
Request for Supplementary Information adopted on 20.07.2017.

Bosulif - bosutinib - EMEA/H/C/002373/R/0027, Orphan
MAH: Pfizer Limited, Rapporteur: Harald Enzmann, PRAC Rapporteur: Martin Huber

Caprelsa - vandetanib - EMEA/H/C/002315/R/0027
MAH: Genzyme Europe BV, Rapporteur:

Alexandre Moreau, PRAC Rapporteur: Ghania Chamouni
Request for Supplementary Information adopted on 09.11.2017.

B.3. POST-AUTHORISATION PHARMACOVIGILANCE OUTCOMES

Signal detection

PRAC recommendations on signals adopted at the PRAC meeting held on 27-30 November 2017 PRAC:

PSUR procedures for which PRAC adopted a recommendation for variation of the terms of the MA at its November 2017 meeting:

EMEA/H/C/PSUSA/00001353/201704
(febuxostat)
CAPS:
Adenuric (EMEA/H/C/000777) (febuxostat), MAH: Menarini International Operations Luxembourg S.A., Rapporteur: Andrea Laslop, PRAC Rapporteur: Jan Neuhauser, “21 Apr 2016 - 20 Apr 2017”

EMEA/H/C/PSUSA/00002029/201705
(methylthioninium chloride)
CAPS:
Methylthioninium chloride Proveblue (EMEA/H/C/002108) (methylthioninium chloride), MAH: Provepharm SAS, Rapporteur: Kristina Dunder, PRAC Rapporteur: Qun-Ying Yue, “06/05/2016 - 05/05/2017”

EMEA/H/C/PSUSA/00002772/201703
(somatropin)
CAPS:
NutropinAq (EMEA/H/C/000315) (somatropin), MAH: Ipsen Pharma, Rapporteur: Hanne Lomholt Larsen
Omnitrope (EMEA/H/C/000607) (somatropin), MAH: Sandoz GmbH, Rapporteur: Johann Lodewijk Hillege
Somatropin Biopartners (EMEA/H/C/002196) (somatropin), MAH: BioPartners GmbH, Rapporteur: Martina Weise
NAPS:
NAPS - EU
, PRAC Rapporteur: Doris Stenver, “01/10/2015 - 31/03/2017”

EMEA/H/C/PSUSA/00009261/201705

(pixantrone)
CAPS:
Pixuvri (EMEA/H/C/002055) (pixantrone), MAH: CTI Life Sciences Limited, Rapporteur: Greg Markey, PRAC Rapporteur: Patrick Batty, "11-Nov-2016 to 10-May-2017"

EMEA/H/C/PSUSA/00010186/201705
(vedolizumab)
CAPS:
Entyvio (EMEA/H/C/002782) (vedolizumab), MAH: Takeda Pharma A/S, Rapporteur: Greg Markey, PRAC Rapporteur: Adam Przybylkowski, "20Nov2016 – 19May2017"

EMEA/H/C/PSUSA/00010255/201705
(simeprevir)
CAPS:
OLYSIO (EMEA/H/C/002777) (simeprevir), MAH: Janssen-Cilag International NV, Rapporteur: Jorge Camarero Jiménez, PRAC Rapporteur: Julie Williams, "22 November 2016 – 21 May 2017"

EMEA/H/C/PSUSA/00010301/201705
(ibrutinib)
CAPS:
Imbruvica (EMEA/H/C/003791) (ibrutinib), MAH: Janssen-Cilag International NV, Rapporteur: Filip Josephson, PRAC Rapporteur: Patrick Batty, "13 May 2016 to 12 November 2016"

EMEA/H/C/PSUSA/00010455/201705
(lumacaftor / ivacaftor)
CAPS:
Orkambi (EMEA/H/C/003954) (lumacaftor / ivacaftor), MAH: Vertex Pharmaceuticals (Europe) Ltd., Rapporteur: Nithyanandan Nagercoil, PRAC Rapporteur: Almath Spooner, "20 Nov 16 to 19 May 2017"

EMEA/H/C/PSUSA/00010550/201705
(mycophenolate mofetil, mycophenolic acid)
CAPS:
CellCept (EMEA/H/C/000082) (mycophenolate mofetil), MAH: Roche Registration Limited, Rapporteur: Greg Markey
Myclausen (EMEA/H/C/001218) (mycophenolate mofetil), MAH: Passauer Pharma GmbH, Rapporteur: Andrea Laslop
Mycophenolate mofetil Teva (EMEA/H/C/000882) (mycophenolate mofetil),

MAH: Teva B.V., Rapporteur: Greg Markey

Myfenax (EMEA/H/C/000884) (mycophenolate mofetil), MAH: Teva B.V., Rapporteur: Greg Markey

NAPS:

Micofenolato De Mofetilo Cinfa - LABORATORIOS CINFA, S.A.

PRAC Rapporteur: Patrick Batty, “3 May 2016 - 2 May 2017”

B.4. EPARs / WPARs

Adynovi - rurioctocog alfa pegol - EMEA/H/C/004195

Applicant: Baxalta Innovations GmbH, treatment of haemophilia A, New active substance (Article 8(3) of Directive No 2001/83/EC)

Darunavir Krka - darunavir - EMEA/H/C/004273

Applicant: KRKA, d.d., Novo mesto, treatment of HIV-1 infection, Generic, Generic of Prezista, Generic application (Article 10(1) of Directive No 2001/83/EC)

Darunavir Krka d.d. - darunavir - EMEA/H/C/004891

Applicant: KRKA, d.d., Novo mesto, treatment of HIV-1 infection, Generic, Duplicate, Generic of Prezista, Duplicate of Darunavir Krka, Generic application (Article 10(1) of Directive No 2001/83/EC)

Fanaptum - iloperidone - EMEA/H/C/004149

Applicant: Vanda Pharmaceuticals Ltd., treatment of schizophrenia, New active substance (Article 8(3) of Directive No 2001/83/EC)

Fasenra - benralizumab - EMEA/H/C/004433

Applicant: AstraZeneca AB, treatment of severe asthma with an eosinophilic phenotype, New active substance (Article 8(3) of Directive No 2001/83/EC)

Fulvestrant Mylan - fulvestrant - EMEA/H/C/004649

Applicant: Mylan S.A.S, treatment of breast cancer, Generic, Generic of Faslodex, Generic application (Article 10(1) of Directive No

2001/83/EC)

Intrarosa - prasterone - EMEA/H/C/004138
Applicant: Endoceutics Limited, treatment of vulvovaginal atrophy, New active substance (Article 8(3) of Directive No 2001/83/EC)

Jorveza - budesonide - EMEA/H/C/004655, Orphan
Applicant: Dr. Falk Pharma GmbH, treatment of eosinophilic esophagitis (EoE), Known active substance (Article 8(3) of Directive No 2001/83/EC)

Mvasi - bevacizumab - EMEA/H/C/004728
Applicant: Amgen Europe B.V., treatment of metastatic breast cancer, unresectable advanced, metastatic or recurrent non-small cell lung cancer, unresectable advanced metastatic or recurrent non-squamous non-small cell lung cancer, advanced and/or metastatic renal cell cancer, advanced epithelial ovarian, fallopian tube, or primary peritoneal cancer, platinum-sensitive epithelial ovarian, fallopian tube or primary peritoneal cancer, Duplicate, Duplicate of KYOMARC, Similar biological application (Article 10(4) of Directive No 2001/83/EC)

Ocrevus - ocrelizumab - EMEA/H/C/004043
Applicant: Roche Registration Limited, treatment of multiple sclerosis
treatment of multiple sclerosis, New active substance (Article 8(3) of Directive No 2001/83/EC)

Prevymis - letermovir - EMEA/H/C/004536, Orphan
Applicant: Merck Sharp & Dohme Limited, prophylaxis of cytomegalovirus (CMV) reactivation and disease, New active substance (Article 8(3) of Directive No 2001/83/EC)

Qizenday - d-biotin - EMEA/H/C/004153
Applicant: Medday Pharmaceuticals, treatment of progressive multiple sclerosis (primary or secondary), Known active substance (Article 8(3) of Directive No 2001/83/EC)
WPAR

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

Avastin - bevacizumab - EMEA/H/C/000582/II/0098 MAH: Roche Registration Limited, Rapporteur: Sinan B. Sarac Opinion adopted on 23.11.2017. Request for Supplementary Information adopted on 12.10.2017.	Positive Opinion adopted by consensus on 23.11.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Axumin - fluciclovine (18F) - EMEA/H/C/004197/II/0001/G MAH: Blue Earth Diagnostics Ltd, Rapporteur: Harald Enzmann	Weekly start timetable.
Axumin - fluciclovine (18F) - EMEA/H/C/004197/II/0002/G MAH: Blue Earth Diagnostics Ltd, Rapporteur: Harald Enzmann	Weekly start timetable.
Blincyto - blinatumomab - EMEA/H/C/003731/II/0020/G, Orphan MAH: Amgen Europe B.V., Rapporteur: Alexandre Moreau	Weekly start timetable.
Brineura - cerliponase alfa - EMEA/H/C/004065/II/0001/G, Orphan MAH: BioMarin International Limited, Rapporteur: Martina Weise Request for Supplementary Information adopted on 12.10.2017.	Weekly start timetable.
Daptomycin Hospira - daptomycin - EMEA/H/C/004310/II/0003 MAH: Hospira UK Limited, Generic, Generic of Cubicin, Rapporteur: Kolbeinn Gudmundsson Request for Supplementary Information adopted on 19.10.2017.	Weekly start timetable.
DuoTrav - travoprost / timolol - EMEA/H/C/000665/II/0051 MAH: Novartis Europharm Limited, Rapporteur: Concepcion Prieto Yerro Opinion adopted on 23.11.2017.	Positive Opinion adopted by consensus on 23.11.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Elaprase - idursulfase - EMEA/H/C/000700/II/0071/G MAH: Shire Human Genetic Therapies AB, Rapporteur: Greg Markey	Weekly start timetable.

Request for Supplementary Information adopted on 26.10.2017.	
Eliquis - apixaban - EMEA/H/C/002148/II/0049/G MAH: Bristol-Myers Squibb / Pfizer EEIG, Rapporteur: Johann Lodewijk Hillege Opinion adopted on 16.11.2017.	Positive Opinion adopted by consensus on 16.11.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Elocta - efmoroctocog alfa - EMEA/H/C/003964/II/0016/G MAH: Swedish Orphan Biovitrum AB (publ), Rapporteur: Jan Mueller-Berghaus Opinion adopted on 16.11.2017. Request for Supplementary Information adopted on 14.09.2017.	Positive Opinion adopted by consensus on 16.11.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Elonva - corifollitropin alfa - EMEA/H/C/001106/II/0037/G MAH: Merck Sharp & Dohme Limited, Rapporteur: Paula Boudewina van Hennik Request for Supplementary Information adopted on 19.10.2017.	Weekly start timetable.
Envarsus - tacrolimus - EMEA/H/C/002655/II/0008/G MAH: Chiesi Farmaceutici S.p.A., Rapporteur: John Joseph Borg Request for Supplementary Information adopted on 20.07.2017, 16.03.2017.	Weekly start timetable.
Fluenz Tetra - influenza vaccine (live attenuated, nasal) - EMEA/H/C/002617/II/0075/G MAH: AstraZeneca AB, Rapporteur: Bart Van der Schueren	Weekly start timetable.
Fotivda - tivozanib - EMEA/H/C/004131/II/0001 MAH: EUSA Pharma (UK) Limited, Rapporteur: Bruno Sepodes	Weekly start timetable.
Hizentra - human normal immunoglobulin - EMEA/H/C/002127/II/0089 MAH: CSL Behring GmbH, Rapporteur: Jan Mueller-Berghaus Opinion adopted on 30.11.2017. Request for Supplementary Information adopted on 05.10.2017.	Positive Opinion adopted by consensus on 30.11.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Hizentra - human normal immunoglobulin - EMEA/H/C/002127/II/0091 MAH: CSL Behring GmbH, Rapporteur: Jan Mueller-Berghaus	Weekly start timetable.

Kevzara - sarilumab - EMEA/H/C/004254/II/0003/G MAH: sanofi-aventis groupe, Rapporteur: Jan Mueller-Berghaus Opinion adopted on 23.11.2017.	Positive Opinion adopted by consensus on 23.11.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Kevzara - sarilumab - EMEA/H/C/004254/II/0004 MAH: sanofi-aventis groupe, Rapporteur: Jan Mueller-Berghaus	Weekly start timetable.
Kineret - anakinra - EMEA/H/C/000363/II/0058 MAH: Swedish Orphan Biovitrum AB (publ), Rapporteur: Sinan B. Sarac	Weekly start timetable.
Kyprolis - carfilzomib - EMEA/H/C/003790/II/0020, Orphan MAH: Amgen Europe B.V., Rapporteur: Jorge Camarero Jiménez Opinion adopted on 23.11.2017.	Positive Opinion adopted by consensus on 23.11.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Kyprolis - carfilzomib - EMEA/H/C/003790/II/0022/G, Orphan MAH: Amgen Europe B.V., Rapporteur: Jorge Camarero Jiménez	Weekly start timetable.
Mosquirix - plasmodium falciparum and hepatitis B vaccine (recombinant, adjuvanted) - EMEA/H/W/002300/II/0025/G MAH: GlaxoSmithkline Biologicals SA, Rapporteur: Jan Mueller-Berghaus	Weekly start timetable.
Natpar - parathyroid hormone - EMEA/H/C/003861/II/0004/G, Orphan MAH: Shire Pharmaceuticals Ireland Ltd, Rapporteur: Bart Van der Schueren Request for Supplementary Information adopted on 02.11.2017.	Weekly start timetable.
Nimenrix - meningococcal group A, C, W135 and Y conjugate vaccine - EMEA/H/C/002226/II/0069 MAH: Pfizer Limited, Rapporteur: Greg Markey Request for Supplementary Information adopted on 16.11.2017.	Request for Supplementary Information adopted
NovoEight - turoctocog alfa - EMEA/H/C/002719/II/0021/G MAH: Novo Nordisk A/S, Rapporteur: Jan Mueller-Berghaus	Weekly start timetable.
Nucala - mepolizumab - EMEA/H/C/003860/II/0011	Request for Supplementary Information adopted

MAH: GlaxoSmithKline Trading Services Limited, Rapporteur: Nithyanandan Nagercoil Request for Supplementary Information adopted on 23.11.2017.	
NutropinAq - somatropin - EMEA/H/C/000315/II/0068/G MAH: Ipsen Pharma, Rapporteur: Hanne Lomholt Larsen Opinion adopted on 23.11.2017.	Positive Opinion adopted by consensus on 23.11.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Opdivo - nivolumab - EMEA/H/C/003985/II/0037/G MAH: Bristol-Myers Squibb Pharma EEIG, Rapporteur: Jorge Camarero Jiménez Request for Supplementary Information adopted on 14.09.2017.	Weekly start timetable.
Pandemic influenza vaccine H5N1 AstraZeneca - pandemic influenza vaccine (H5N1) (live attenuated, nasal) - EMEA/H/C/003963/II/0009/G MAH: AstraZeneca AB, Rapporteur: Jan Mueller-Berghaus	Weekly start timetable.
Pixuvri - pixantrone - EMEA/H/C/002055/II/0040 MAH: CTI Life Sciences Limited, Rapporteur: Greg Markey	Weekly start timetable.
Praluent - alirocumab - EMEA/H/C/003882/II/0028/G MAH: sanofi-aventis groupe, Rapporteur: Johann Lodewijk Hillege Opinion adopted on 16.11.2017. Request for Supplementary Information adopted on 21.09.2017.	Positive Opinion adopted by consensus on 16.11.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Praluent - alirocumab - EMEA/H/C/003882/II/0030 MAH: sanofi-aventis groupe, Rapporteur: Johann Lodewijk Hillege Opinion adopted on 23.11.2017.	Positive Opinion adopted by consensus on 23.11.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Privigen - human normal immunoglobulin - EMEA/H/C/000831/II/0127 MAH: CSL Behring GmbH, Rapporteur: Jan Mueller-Berghaus	Weekly start timetable.
Scenesse - afamelanotide - EMEA/H/C/002548/II/0017, Orphan MAH: Clinuvel (UK) Limited, Rapporteur: Harald Enzmann Request for Supplementary Information adopted	Request for Supplementary Information adopted

on 30.11.2017.	
Strensiq - asfotase alfa - EMEA/H/C/003794/II/0026/G, Orphan MAH: Alexion Europe SAS, Rapporteur: Greg Markey Opinion adopted on 23.11.2017.	Positive Opinion adopted by consensus on 23.11.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
TachoSil - human thrombin / human fibrinogen - EMEA/H/C/000505/II/0081 MAH: Takeda Austria GmbH, Rapporteur: Jan Mueller-Berghaus Opinion adopted on 16.11.2017. Request for Supplementary Information adopted on 21.09.2017.	Positive Opinion adopted by consensus on 16.11.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Tecentriq - atezolizumab - EMEA/H/C/004143/II/0001 MAH: Roche Registration Limited, Rapporteur: Sinan B. Sarac	Weekly start timetable.
Travatan - travoprost - EMEA/H/C/000390/II/0057 MAH: Novartis Europharm Limited, Rapporteur: Concepcion Prieto Yerro Opinion adopted on 23.11.2017.	Positive Opinion adopted by consensus on 23.11.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Trulicity - dulaglutide - EMEA/H/C/002825/II/0021 MAH: Eli Lilly Nederland B.V., Rapporteur: Greg Markey Request for Supplementary Information adopted on 23.11.2017, 19.10.2017.	Request for Supplementary Information adopted
Vimizim - elosulfase alfa - EMEA/H/C/002779/II/0021/G, Orphan MAH: BioMarin Europe Ltd, Rapporteur: Johann Lodewijk Hillege Opinion adopted on 16.11.2017. Request for Supplementary Information adopted on 21.09.2017.	Positive Opinion adopted by consensus on 16.11.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Xofigo - radium-223 - EMEA/H/C/002653/II/0027 MAH: Bayer AG, Rapporteur: Harald Enzmann Opinion adopted on 30.11.2017.	Positive Opinion adopted by consensus on 30.11.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Zaltrap - aflibercept - EMEA/H/C/002532/II/0040 MAH: sanofi-aventis groupe, Rapporteur: Filip Josephson	Weekly start timetable.
WS1232 **Infanrix hexa-** **EMEA/H/C/000296/WS1232/0232**	Weekly start timetable.

MAH: GlaxoSmithkline Biologicals SA, Lead Rapporteur: Bart Van der Schueren	
WS1233/G **Hexacima-** **EMEA/H/C/002702/WS1233/0070/G** **Hexaxim-** **EMEA/H/W/002495/WS1233/0075/G** **Hexyon-** **EMEA/H/C/002796/WS1233/0074/G** MAH: Sanofi Pasteur SA, Lead Rapporteur: Jan Mueller-Berghaus Request for Supplementary Information adopted on 16.11.2017.	Request for Supplementary Information adopted
WS1254/G **Hirobriz Breezhaler-** **EMEA/H/C/001211/WS1254/0042/G** **Onbrez Breezhaler-** **EMEA/H/C/001114/WS1254/0041/G** **Oslif Breezhaler-** **EMEA/H/C/001210/WS1254/0041/G** **Ultibro Breezhaler-** **EMEA/H/C/002679/WS1254/0017/G** **Ulunar Breezhaler-** **EMEA/H/C/003875/WS1254/0017/G** **Xoterna Breezhaler-** **EMEA/H/C/003755/WS1254/0020/G** MAH: Novartis Europharm Limited, Lead Rapporteur: Hanne Lomholt Larsen Request for Supplementary Information adopted on 19.10.2017.	Weekly start timetable.
WS1262 **Cerezyme-** **EMEA/H/C/000157/WS1262/0106** **Fabrazyme-** **EMEA/H/C/000370/WS1262/0101** **Myozyme-** **EMEA/H/C/000636/WS1262/0067** **Thyrogen-** **EMEA/H/C/000220/WS1262/0093** MAH: Genzyme Europe BV, Lead Rapporteur: Johann Lodewijk Hillege	Weekly start timetable.

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

Afstyla - lonoctocog alfa - **EMEA/H/C/004075/II/0007** MAH: CSL Behring GmbH, Rapporteur: Jan Mueller-Berghaus, “Update of section 4.8 of the SmPC in order to include information on	Weekly start timetable.

inhibitor development in Previously Untreated Patients (PUPs), based on the ongoing Phase III study CSL627_3001 which aims to evaluate the long-term safety and efficacy of rVIII-Single Chain for routine prophylaxis and on-demand treatment of bleeding episodes in children, adolescents and adults with severe hemophilia A (ie, FVIII activity of ≤ 1%). The Package Leaflet is updated accordingly. In addition, the Marketing authorisation holder (MAH) took the opportunity to make editorial changes related to the secondary packaging in section 6.5 and 6.6 of the SmPC, in section 4 of the Labelling and sections 3 and 6 of the Package leaflet. Moreover, the MAH took the opportunity to update the list of local representatives (for Bulgaria) in the Package Leaflet."	
Blincyto - blinatumomab - EMEA/H/C/003731/II/0009, Orphan MAH: Amgen Europe B.V., Rapporteur: Alexandre Moreau, "Update of sections 4.4, 4.8, 5.1 and 5.2 of the SmPC in order to update the safety information with the data from the study 103311. This study is fulfilling the specific obligation for the conditional MA. The SO is removed from annex II. The Package Leaflet is updated accordingly. The MAH takes this opportunity to amend the format of the preparation instructions to improve clarity. The content is not impacted." Request for Supplementary Information adopted on 20.07.2017, 26.01.2017.	See 9.1
Buccolam - midazolam - EMEA/H/C/002267/II/0035 MAH: Shire Services BVBA, Rapporteur: Johann Lodewijk Hillege, "Update of SmPC sections 4.4 and 4.5 to strengthen the warning regarding concomitant administration of benzodiazepines and opioids following a recent review of the MAH's safety databases and literature. The Package Leaflet has been updated accordingly. In addition, the MAH took the opportunity to align the annexes with the latest QRD template and to update the contact details of the MAH in the Package Leaflet." Request for Supplementary Information adopted on 23.11.2017.	Request for Supplementary Information adopted
CellCept - mycophenolate mofetil -	Positive Opinion adopted by consensus on

EMEA/H/C/000082/II/0136 MAH: Roche Registration Limited, Rapporteur: Greg Markey, "Update of sections 4.4 and 4.5 of the SmPC of all pharmaceutical forms, in order to update information regarding potential interactions with antibiotics and drugs interfering with glucuronidation pathway, based on a review of published literature. The Package Leaflet is updated accordingly. In addition, update of section 6.6 of the SmPC and section 3 of the package leaflet to improve the recommendations regarding safe handling of the powder for oral suspension formulation as well as other minor editorial changes." Opinion adopted on 23.11.2017.	23.11.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
DuoTrav - travoprost / timolol - EMEA/H/C/000665/II/0052 MAH: Novartis Europharm Limited, Rapporteur: Concepcion Prieto Yerro, "Update of sections 4.8 of the SmPC in order to add "lid sulcus deepened" and "iris hyperpigmentation" as new adverse drug reactions and to upgrade the frequency of "skin hyperpigmentation (periocular)" from rare to uncommon based on the post-approval review of the safety data. In addition, section 4.8 of SmPC has been updated to align Adverse Drug Reactions table for the travoprost monotherapy. Based on the same safety review, section 4.6 of SmPC has been updated with dose margin estimates. In addition, the MAH took the opportunity to align the Product information with the currently approved travoprost EU SmPC and QRD version 10 and to update the list of local representatives."	Weekly start timetable.
Epivir - lamivudine - EMEA/H/C/000107/II/0104 MAH: ViiV Healthcare UK Limited, Rapporteur: Joseph Emmerich, "Update of section 4.5 of the SmPC of both Epivir tablets and oral solution, and section 4.4 of the SmPC for Epivir Oral solution only, to add information regarding the potential for interaction between lamivudine and sorbitol based on the results of Study 204857. Further, a minor amendment has been implemented throughout the SmPC to update the clinical terminology for 'Pneumocystis carinii pneumonia' to 'Pneumocystis jiroveci	Weekly start timetable.

pneumonia'. In addition, the MAH has taken the opportunity to align the product information with the QRD template version 10, to make minor editorial changes in the annexes and to update the contact details of the local representative in Norway in the Package Leaflet." Request for Supplementary Information adopted on 21.09.2017, 05.05.2017.	
Humira - adalimumab - EMEA/H/C/000481/II/0170 MAH: AbbVie Limited, Rapporteur: Kristina Dunder, "Update of section 4.6 of the SmPC in order to update information on pregnancy and lactation based on results from pregnancy registry (OTIS; study number M03-604) and supported by relevant published literature. The Package Leaflet is updated accordingly." Request for Supplementary Information adopted on 30.11.2017.	Request for Supplementary Information adopted
Ibrance - palbociclib - EMEA/H/C/003853/II/0006 MAH: Pfizer Limited, Rapporteur: Filip Josephson, "Update of sections 4.2, 4.8 and 5.1 in order to reflect the results of the study A5481008 (PALOMA-2) and of the Phase 2 portion of A5481010 single-arm study. The MAH took the opportunity to implement minor editorial changes to the PIL." Request for Supplementary Information adopted on 12.10.2017, 22.06.2017.	
Iclusig - ponatinib - EMEA/H/C/002695/II/0041, Orphan MAH: Incyte Biosciences UK Ltd, Rapporteur: Greg Markey, "Update of section 4.8 of the SmPC in order to update the safety information to include a paragraph in the SmPC section 4.8 on severe cutaneous reaction. The Package Leaflet is updated accordingly." Request for Supplementary Information adopted on 12.10.2017.	Weekly start timetable.
Invirase - saquinavir - EMEA/H/C/000113/II/0122 MAH: Roche Registration Limited, Rapporteur: Milena Stain, "Update of sections 4.2, 4.3, and 4.5 of the SmPC following an update to the Company Core Data Sheet in order to include a cross-reference to a new contraindication	Weekly start timetable.

against switching from rilpivirine to invirase/ritonavir (section 4.2), to include lurasidone in the contraindications section (section 4.3) and to add information on additional interactions regarding lurasidone, rilpivirine, dasatinib, and sunitinib (section 4.5). The existing information regarding the interaction with alfuzosin has been updated to include a warning that co-administration may also cause potentially life-threatening cardiac arrhythmia. The existing information regarding interaction with medicines listed in the section 'neuroleptics' has been moved to the section 'antipsychotics' (section 4.5). The Package Leaflet is updated accordingly. In addition, the Marketing authorisation holder (MAH) took the opportunity the PI to correct formatting and minor typographical errors."	
Isentress - raltegravir - EMEA/H/C/000860/II/0069 MAH: Merck Sharp & Dohme Limited, Rapporteur: Greg Markey, "Update of sections 4.8 and 5.1 of the SmPC based on the final results (i.e. through 96 weeks) from study PN292 (ONCEMRK), the pivotal Phase 3 study evaluating the safety and efficacy of raltegravir 1200 mg QD (2 x 600 mg tablets) versus raltegravir 400 mg BID, each in combination with emtricitabine / tenofovir disoproxil fumarate in treatment-naïve HIV-1 infected adult subjects. In addition, the MAh took the opportunity to implement minor editorial changes in the SmPC."	Weekly start timetable.
Izba - travoprost - EMEA/H/C/002738/II/0008 MAH: Novartis Europharm Limited, Rapporteur: Concepcion Prieto Yerro, "Update of sections 4.4 and 4.8 of the SmPC in order to update the safety information in line with Travoprost 40 µg/mL Eye Drops PI, based on the review of clinical trial and post-marketing data along with literature references. The package leaflet section 4 is updated accordingly." Request for Supplementary Information adopted on 05.10.2017.	Weekly start timetable.
Jevtana - cabazitaxel - EMEA/H/C/002018/II/0038 MAH: sanofi-aventis groupe, Rapporteur:	Weekly start timetable.

Alexandre Moreau, “Submission of the final PK analysis report with data from studies EFC11784, EFC11785, TCD11068, and TCD 11870 to provide information on relationship between allelic variants of genes coding for CY3A4 enzyme and cabazitaxel. No changes to the PI are proposed.”	
Maviret - glecaprevir / pibrentasvir - EMEA/H/C/004430/II/0002 MAH: AbbVie Limited, Rapporteur: Joseph Emmerich, “Update of sections 4.8 and 5.1 of the SmPC in order to add information on clinical efficacy and safety in HCV/HIV-1 co-infected subjects, based on new clinical data from Study M14-730 (EXPEDITION-2), a post-registrational Phase 3 study which evaluated the efficacy and safety of the glecaprevir/pibrentasvir regimen in chronic HCV GT1-GT6/HIV-1 co-infected subjects who were HCV treatment-naïve or treatment-experienced. In addition, the SmPC was revised to make minor grammatical and formatting amendments and to correct errors in section 5.2.”	Weekly start timetable.
Maviret - glecaprevir / pibrentasvir - EMEA/H/C/004430/II/0003 MAH: AbbVie Limited, Rapporteur: Joseph Emmerich, “Update of section 4.5 of the SmPC in order to remove the restriction relating to co-administration with omeprazole, based on new analyses of previously submitted data from the Phase 1 study M14-715 (Open-label study to assess the effect of acid reducing agent on the pharmacokinetics, safety and tolerability of ABT-493/ABT-530 in healthy adult subjects) and on pharmacokinetic as well as efficacy results from Phase 2 and 3 clinical studies for the subjects who were coadministered GLE/PIB and PPIs including omeprazole 40 mg daily. The Package Leaflet is updated accordingly.”	Weekly start timetable.
Mysimba - naltrexone hydrochloride / bupropion hydrochloride - EMEA/H/C/003687/II/0023 MAH: Orexigen Therapeutics Ireland Limited, Rapporteur: Hanne Lomholt Larsen, “Update of sections 4.2, 4.3, 4.4, 4.8 and 5.2 of the SmPC in order to update the dosage recommendation and safety information for patients with moderate renal impairment based on final results from study NaltrexBuprop-1006 - A	Weekly start timetable.

Phase 1, Open-Label, Parallel Study to Evaluate the Pharmacokinetics of a Single Oral Dose of Extended-Release Combination of Naltrexone and Bupropion in Subjects With Normal Renal Function or Varying Degrees of Impaired Renal Function. The Package Leaflet is updated accordingly."	
Natpar - parathyroid hormone - EMEA/H/C/003861/II/0006, Orphan MAH: Shire Pharmaceuticals Ireland Ltd, Rapporteur: Bart Van der Schueren, "Submission of the final report from studies (R7661M-SHP634 and R7673M-SHP634) listed as a category 3 studies in the RMP. Study R7661M-SHP634 is a Comparison of the Effects of Once- versus Twice-Daily Dosing with NPSP558 (Recombinant Human Parathyroid Hormone (1-84)) on Osteoblast Proliferation and Bone Formation in the Male Fischer 344 Rats. Study R7673M-SHP634 is A 13-Week Subcutaneous Injection Study of NPSP558 (Recombinant Human PTH (1-84)) with an 8-Week Recovery Period in Juvenile Rats." Opinion adopted on 30.11.2017.	Positive Opinion adopted by consensus on 30.11.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Nimenrix - meningococcal group A, C, W135 and Y conjugate vaccine - EMEA/H/C/002226/II/0071 MAH: Pfizer Limited, Rapporteur: Greg Markey, "Update of sections 4.2, 4.4, 4.8 and 5.1 of the SmPC in order to update the posology for infants from 6 weeks to less than 12 months of age and to remove the recommendation of a second dose in children above 12 months of age, and to add information regarding antibody persistence as measured by serum bactericidal assays 1 year after 1 or 2 doses of MenACWY-TT in toddlers. The posology update is based on results from Study 087 and antibody persistence update is based on results from Study 104 (assessed in procedure ANX 13.3). Study MenACWY-TT-087 is a phase IIIb, open, multi-country, controlled, randomised study to demonstrate the immunogenicity and safety of GSK Biologicals' meningococcal conjugate vaccine, MenACWY-TT in healthy infants, given on a 3+1 primary and booster (2, 4, 6 and 15-18 months of age), a 1+1 primary and booster (6 and 15-18 months of age) or as a single dose at 15-18 months of age.	

The Package Leaflet is updated accordingly. Annex II is also updated to take into account that the 1 year timeline was fulfilled in ANX 13.3."	
Rapiscan - regadenoson - EMEA/H/C/001176/II/0026 MAH: Rapidscan Pharma Solutions EU Ltd., Rapporteur: Greg Markey, "Update of section 4.5 of the SmPC in order to add further information on molecular transporters based 5 non-clinical studies: Study-OPT-2016-045, Study-OPT-2016-046, Study-OPT-2016-099, Study-OPT-2016-100 and Study-OPT-2016-101." Request for Supplementary Information adopted on 12.10.2017.	Weekly start timetable.
Roteas - edoxaban - EMEA/H/C/004339/II/0003 MAH: Daiichi Sankyo Europe GmbH, Rapporteur: Concepcion Prieto Yerro, "Update of sections 4.2 and 5.1 of the SmPC in line with changes already introduce to Lixiana (EMEA/H/C/002629/II/0012) in order to add information deriving from clinical data for the use of edoxaban as anticoagulant therapy for patients with non-valvular atrial fibrillation undergoing cardioversion. The Package Leaflet is updated accordingly. In addition, the Marketing authorisation holder (MAH) took the opportunity to update section 4.8 of the SmPC as per the requirement of the finalised PSUSA/00010387/201610 procedure to include headache, abdominal pain and dizziness with a common frequency as new adverse drug reactions. The MAH also took the opportunity to bring the PI in line with the latest QRD template version 10.0 and to introduce some editorial changes and minor corrections." Opinion adopted on 30.11.2017.	Positive Opinion adopted by consensus on 30.11.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Simponi - golimumab - EMEA/H/C/000992/II/0078/G MAH: Janssen Biologics B.V., Rapporteur: Kristina Dunder, "Update of sections 4.4 and 4.8 of the SmPC in order to add a warning on agranulocytosis and update neutropenia from uncommon to common based on new safety information in the Company Core Data Sheet (CCDS). The Marketing Authorisation Holder has taken	Weekly start timetable.

the opportunity to include the safety data from the intravenous (IV) psoriatic arthritis (PsA), and IV ankylosing spondylitis (AS) studies that were recently included in the CCDS. The Package Leaflet is updated accordingly."	
Starlix - nateglinide - EMEA/H/C/000335/II/0033 MAH: Novartis Europharm Limited, Rapporteur: Greg Markey, "Update of section 5.2 of the SmPC to add information on the accumulation of M1 metabolite in diabetic patients with end-stage renal disease (ESRD), based on the review of the Core Data Sheet. This information is reflected in section 4.4 of the SmPC. The Package Leaflet and Labelling are updated accordingly. In addition, the MAH took the opportunity to bring the PI in line with QRD template 10, combine the three SmPC into a single SmPC, align sections 1 and 2 of the package leaflet with the SmPC, and correct the name of the local representatives for Latvia and Bulgaria."	Weekly start timetable.
Sutent - sunitinib - EMEA/H/C/000687/II/0067 MAH: Pfizer Limited, Rapporteur: Daniela Melchiorri, "Update of sections 4.5 and 5.2 of the SmPC in order to include information regarding a possible interaction between sunitinib and other medicinal products that are inhibitors of the efflux transporter breast cancer resistance protein (BCRP) following assessment of PAM (REC 052)."	Weekly start timetable.
Tafinlar - dabrafenib - EMEA/H/C/002604/II/0027 MAH: Novartis Europharm Limited, Rapporteur: Filip Josephson, "Submission of the final report from study BRF113683 (BREAK-3) listed as a category 3 study in the RMP. This is a phase III, randomised, two-arm, open label study comparing dabrafenib to dacarbazine (DTIC) in previously untreated patients with BRAF mutation positive advanced (stage III) or metastatic (stage IV) melanoma. This study is aimed to confirm the superior efficacy of dabrafenib compared to DTIC."	Weekly start timetable.
Torisel - temsirolimus - EMEA/H/C/000799/II/0069, Orphan MAH: Pfizer Limited, Rapporteur: Harald	

Enzmann, “Update of section 4.3 of the SmPC in order to specify that the use of temsirolimus in patients with mantle cell lymphoma (MCL) with moderate or severe hepatic impairment is an absolute contraindication, as requested to be clarified during the renewal procedure (EMEA/H/C/000799/R/0065). In addition, the MAH took the opportunity to make minor editorial changes in the Package Leaflet.”	
Translarna - ataluren - EMEA/H/C/002720/II/0039, Orphan MAH: PTC Therapeutics International Limited, Rapporteur: Johann Lodewijk Hillege, “Update of sections 4.2, 4.4, and 5.2 of the SmPC to include new clinical information based on final results from study PTC124-GD-033-HV (Study 033) listed as a category 3 study in the RMP (MEA009); this is a Safety and PK study in patients with moderate to severe hepatic impairment; the Package Leaflet and Labelling are updated accordingly. In addition, the Marketing authorisation holder (MAH) took the opportunity to bring the PI in line with the latest QRD template version 10.0 and to implement some editorial changes.”	
Trumenba - meningococcal group B vaccine (recombinant, adsorbed) - EMEA/H/C/004051/II/0002/G MAH: Pfizer Limited, Rapporteur: Johann Lodewijk Hillege, “Update of sections 4.4 and 4.8 of the SmPC in order to add a warning on syncope and to add this adverse reaction based on post-marketing data. Update of section 4.8 of the SmPC in order to update the safety information regarding booster vaccination based a review of adverse events data reported in the interim clinical study report (B1971033). The package leaflet is updated accordingly. In addition, the MAH took the opportunity to make a clarification on interchangeability of Trumenba in section 4.2 of the SmPC and to update the list of local representatives in the package leaflet.”	Weekly start timetable.
Vectibix - panitumumab - EMEA/H/C/000741/II/0086 MAH: Amgen Europe B.V., Rapporteur: Robert James Hemmings, “Update of section 4.4 and section 4.8 of the SmPC and relevant sections of the PL to reflect a re-analysis of the safety information which pooled data from all the	Weekly start timetable.

indications requiring a change in the overall incidence, severity, and seriousness of some of the currently labelled ADRs."	
Velphoro - mixture of polynuclear iron(III)-oxyhydroxide, sucrose and starches - EMEA/H/C/002705/II/0012 MAH: Vifor Fresenius Medical Care Renal Pharma France, Rapporteur: Johann Lodewijk Hillege, "Update of section 4.4 of the SmPC in order to remove a warning related to allergy to gluten. The Package Leaflet has been updated accordingly. In addition, the Marketing authorisation holder (MAH) took the opportunity to make minor corrections in the SmPC in line with the Company Core Data Sheet (CCDS). Moreover, the MAH took the opportunity to bring the Annex IIIA in line with the latest QRD template version 10." Opinion adopted on 16.11.2017.	Positive Opinion adopted by consensus on 16.11.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Venclyxto - venetoclax - EMEA/H/C/004106/II/0007/G, Orphan MAH: AbbVie Limited, Rapporteur: Filip Josephson, "Update of section 4.5 of the SmPC in order to update the drug-drug interaction between venetoclax and dogixin based on final results from study M16-042; this is study to assess the effect of venetoclax on the pharmacokinetics of digoxin in healthy female subjects. Update of section 4.5 of the SmPC in order to update the drug-drug interaction between venetoclax and ritonavir, based on final results from study M15-719; this is study to assess the effect of ritonavir on the pharmacokinetics of venetoclax in healthy female subjects of non-childbearing potential. Update of section 4.5 of the SmPC in order to update the drug-drug interaction between venetoclax and azithomycin, based on final results from study M16-068; this is study to assess effect of azithromycin on the pharmacokinetics of venetoclax in healthy female subjects. The MAH took the opportunity to update the Product Information with minor editorial and QRD updates."	Weekly start timetable.
Xeljanz - tofacitinib - EMEA/H/C/004214/II/0003	Weekly start timetable.

MAH: Pfizer Limited, Rapporteur: Robert James Hemmings, “Update of section 4.5 and 5.2 of the SmPC to add results of in vitro transporter inhibition studies evaluating tofacitinib for its potential to inhibit organic anion transporter (OAT1, OAT3) and to interact with MRP2 Transporter, in order to fulfil a CHMP recommendation.” Request for Supplementary Information adopted on 14.09.2017.	
Zeffix - lamivudine - EMEA/H/C/000242/II/0069 MAH: Glaxo Group Ltd, Duplicate, Duplicate of Epivir, Rapporteur: Joseph Emmerich, “Update of section 4.5 of the SmPC to add information regarding a potential interaction with sorbitol-containing medicines. The Package Leaflet has been updated accordingly. In addition, the MAH took the opportunity to implement a minor change in the labelling in line with the QRD template version 10.” Request for Supplementary Information adopted on 21.09.2017, 05.05.2017.	Weekly start timetable.
Zykadia - ceritinib - EMEA/H/C/003819/II/0016 MAH: Novartis Europharm Limited, Rapporteur: Jorge Camarero Jiménez, “Update of sections 4.2, 4.4, 4.8 and 5.2 of the SmPC in order to include amendments to the posology in hepatically impaired patients and update the safety information, respectively. The updates are based on the results from the hepatic function Study CLDK378A2110 which evaluated the PK, safety and tolerability of a single oral dose of ceritinib in subjects with varying degrees of impaired hepatic function and results from physiology-based pharmacokinetic (PBPK) modeling at steady-state. Submission of the Report for Study A2110 fulfils MEA 001 for Zykadia.” Request for Supplementary Information adopted on 09.11.2017, 14.09.2017.	
WS1137 **Lyrica-EMEA/H/C/000546/WS1137/0087** **Pregabalin Pfizer-EMEA/H/C/003880/WS1137/0017** MAH: Pfizer Limited, Lead Rapporteur: Johann Lodewijk Hillege, “Update of sections 4.8 and 5.1 of the SmPC in order to reflect final results	Positive Opinion adopted by consensus on 16.11.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

from paediatric study A0081041: “A Double-Blind, Placebo-Controlled, Parallel-Group, Multicenter Study of the Efficacy and Safety of Pregabalin as Adjunctive Therapy in Children 4-16 Years of Age with Partial Onset Seizures”.” Opinion adopted on 16.11.2017. Request for Supplementary Information adopted on 14.09.2017, 08.05.2017.	
WS1156 **Combivir-EMEA/H/C/000190/WS1156/0090** **Kivexa-EMEA/H/C/000581/WS1156/0072** **Triumeq-EMEA/H/C/002754/WS1156/0042** **Trizivir-EMEA/H/C/000338/WS1156/0104** MAH: ViiV Healthcare UK Limited, Lead Rapporteur: Joseph Emmerich, “Update of section 4.5 of the SmPC to add information regarding the potential interaction between lamivudine and sorbitol based on the results of Study 204857. The Package Leaflet has been updated accordingly. Further, a minor amendment has been implemented throughout the SmPC in order to update the clinical terminology of Pneumocystis carinii pneumonia to Pneumocystis jiroveci pneumonia. In addition, the MAH takes the opportunity to make minor editorial changes , to align the annexes with the QRD template version 10 and to update the contact details of the local representative in Norway in the Package Leaflet.” Request for Supplementary Information adopted on 21.09.2017, 05.05.2017.	Weekly start timetable.
WS1210/G **Mekinist-EMEA/H/C/002643/WS1210/0021/G** **Tafinlar-EMEA/H/C/002604/WS1210/0026/G** MAH: Novartis Europharm Limited, Lead Rapporteur: Paula Boudewina van Hennik, “Update of section 5.1 of the Mekinist (trametinib) and Tafinlar (dabrafenib) SmPC to include the 3-years overall survival (OS) results from study MEK115306 (COMBI-d), a phase III, randomised, double-blinded study comparing the combination of dabrafenib and trametinib to dabrafenib and placebo in first-line therapy for subjects with unresectable or metastatic BRAF V600/K mutation-positive cutaneous melanoma.	Request for Supplementary Information adopted

Update of section 5.1 of the Mekinist (trametinib) and Tafinlar (dabrafenib) SmPC to include the 3-years overall survival (OS) results from study MEK116513 (COMBI-v), a phase III, open-label, 2 arm, randomised study comparing dabrafenib and trametinib combination therapy with vemurafenib monotherapy in BRAF V600 mutation-positive metastatic melanoma." Request for Supplementary Information adopted on 30.11.2017, 05.10.2017.	
WS1219 **Brimica Genuair-** **EMEA/H/C/003969/WS1219/0014** **Duaklir Genuair-** **EMEA/H/C/003745/WS1219/0014** MAH: AstraZeneca AB, Lead Rapporteur: Nithyanandan Nagercoil, "Update of section 5.2 of the SmPC in order to update information based on results from study KRP-AB1102F-302 [KRP-AB1102F Phase II Clinical Pharmacology Study - An Investigation into the Pharmacokinetics upon Repeated Administration of KRP-AB1102F to COPD Patients as Subjects]. In addition, the Worksharing applicant (WSA) took the opportunity to update footnotes of the table in section 4.8 as requested during PSUR procedure EMEA/H/C/PSUSA/00010307/201511 and to amend annex II following request from procedure EMEA/H/C/PSA/S/0017." Opinion adopted on 16.11.2017. Request for Supplementary Information adopted on 14.09.2017.	Positive Opinion adopted by consensus on 16.11.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

B.5.3. CHMP-PRAC assessed procedures

Caprelsa - vandetanib - **EMEA/H/C/002315/II/0028** MAH: Genzyme Europe BV, Rapporteur: Alexandre Moreau, PRAC Rapporteur: Ghania Chamouni, "Update of sections 4.1, 4.4 and 5.1 of the SmPC in order to delete the information regarding Rearranged during Transfection (RET) mutation. The application addresses SOB 001 and the MAH proposes to revert from conditional marketing authorisation to standard marketing authorisation. Annex II and Package Leaflet are updated accordingly. The RMP version 12.2 has also been submitted. In addition, the MAH took the opportunity to	

bring the PI in line with the latest QRD template version 10."	
Cerdelga - eliglustat - EMEA/H/C/003724/II/0015/G, Orphan MAH: Genzyme Europe BV, Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Dolores Montero Corominas, "Update of sections 4.2., 4.3. ,4.4, 4.5. and 5.2. of the SmPC based on the final data from studies POP13777 "Pharmacokinetics of Oral Single-Dose eliglustat in Subjects with Hepatic Impairment" (MEA003.3) and POP13778 "Pharmacokinetics of Oral Single-Dose eliglustat in Subjects with Renal Impairment" (MEA004.3). Annex II D - Conditions Or Restrictions With Regard To The Safe And Effective Use Of The Medicinal Product of the Product Information, additional risk minimisation measures has likewise been amended. The Package Leaflet is updated accordingly. The RMP version 5.0 has also been submitted."	
Cimzia - certolizumab pegol - EMEA/H/C/001037/II/0060 MAH: UCB Pharma S.A., Rapporteur: Kristina Dunder, PRAC Rapporteur: Ulla Wändel Liminga, "Update of section 4.6 of the SmPC in order to update the information on pregnancy and lactation based on two pharmacokinetics studies evaluation the transfer of Cimzia into breastmilk and via the placenta (UP0016 and UP0017). The Package Leaflet is updated accordingly. The RMP version 12 has also been submitted." Request for Supplementary Information adopted on 09.11.2017, 14.09.2017, 22.06.2017.	
Defitelio - defibrotide - EMEA/H/C/002393/II/0026, Orphan MAH: Gentium S.r.l., Rapporteur: Nithyanandan Nagercoil, PRAC Rapporteur: Julie Williams, "Update of sections 4.8 and 5.1 of the SmPC in order to update the frequencies of adverse reactions included in the tabulated list of adverse reactions and to update the clinical efficacy and safety information based on the results from study 2006-05 listed as category 3 in the RMP. This is a phase 3, open-label expanded access study designed to provide access to defibrotide as an investigational new drug to patients with severe hepatic veno-occlusive disease. The final study report is being	Request for Supplementary Information adopted

submitted together with the revised risk management plan (version 3.0). The package leaflet is also being updated accordingly. In addition, the MAH took the opportunity to bring the SmPC in line with the latest QRD template (version 10), to update the list of local representatives in the package leaflet and to correct a translation error in the Polish, Finnish, Danish and Latvian languages." Request for Supplementary Information adopted on 30.11.2017, 28.09.2017.	
Dificlir - fidaxomicin - EMEA/H/C/002087/II/0032/G MAH: Astellas Pharma Europe B.V., Rapporteur: Filip Josephson, PRAC Rapporteur: Qun-Ying Yue, "C.I.11.b) Update of sections 4.2, 4.4 and 5.1 of the SmPC in order to update the safety information following final results from the drug utilisation study ANEMONE listed as an additional pharmacovigilance activity in the RMP. 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. C.I.3 Update of sections 4.4 and 5.2 of the SmPC in order to update the safety information based on results from the PROFILE study, an open label study designed to evaluate the pharmacokinetics of fidaxomicin in IBD subjects with CD. The Package Leaflet is updated accordingly. The RMP version 9.0 has also been submitted." Request for Supplementary Information adopted on 30.11.2017.	Request for Supplementary Information adopted
Ibrance - palbociclib - EMEA/H/C/003853/II/0007 MAH: Pfizer Limited, Rapporteur: Filip Josephson, PRAC Rapporteur: Doris Stenver, "Update of sections 4.2, 4.4 and 5.2 of the SmPC to reflect the results of studies A5481013 and A5481014. The mentioned studies provide information of the impact of hepatic impairment (Study A5481013) on the PK of a single oral dose of 75 mg palbociclib and the impact of renal impairment (Study A5481014) on the PK of a single oral dose of 125 mg palbociclib both administered under fed conditions to subjects	

with varying degrees of hepatic function or renal function. The RMP (version 1.4) is proposed to be amended to reflect the completion of these studies." Request for Supplementary Information adopted on 14.09.2017.	
Imraldi - adalimumab - EMEA/H/C/004279/II/0002/G MAH: Samsung Bioepis UK Limited (SBUK), Rapporteur: Outi Mäki-Ikola, PRAC Rapporteur: Ulla Wändel Liminga Request for Supplementary Information adopted on 30.11.2017.	Request for Supplementary Information adopted
INOmax - nitric oxide - EMEA/H/C/000337/II/0051 MAH: Linde Healthcare AB, Rapporteur: Nithyanandan Nagercoil, PRAC Rapporteur: Julie Williams Request for Supplementary Information adopted on 12.10.2017.	
Keytruda - pembrolizumab - EMEA/H/C/003820/II/0037/G MAH: Merck Sharp & Dohme Limited, Rapporteur: Daniela Melchiorri, PRAC Rapporteur: Sabine Straus, "Update of sections 4.4 and 4.8 of the SmPC to add information regarding the risks of encephalitis, sarcoidosis and graft versus host disease (GVHD) that have been reported in patients treated with pembrolizumab. The Package Leaflet and the 'additional risk minimization measures' section (educational material) in the Annex II have been updated accordingly. In addition, the MAH has implemented minor changes in the SmPC section 5.1 and editorial changes in the Package Leaflet. An updated RMP version 13.0 was provided as part of the application."	
Kuvan - sapropterin - EMEA/H/C/000943/II/0052, Orphan MAH: BioMarin International Limited, Rapporteur: Peter Kiely, PRAC Rapporteur: Almath Spooner, "Based on a review of the post-marketing experience and in order to harmonize the safety information with the CCDS, update of section 4.4 of the Kuvan SmPC to add a warning regarding gastritis and update of section 4.8 to add the following adverse events regarding gastrointestinal tract and	Request for Supplementary Information adopted

respiratory irritation: oropharyngeal pain, oesophageal pain, dyspepsia, nausea, gastritis and pharyngitis.
The Package Leaflet is updated accordingly. The RMP version 13.0 has also been submitted."
Request for Supplementary Information adopted on 30.11.2017.

Kyprolis - carfilzomib - EMEA/H/C/003790/II/0017/G, Orphan
MAH: Amgen Europe B.V., Rapporteur: Jorge Camarero Jiménez, PRAC Rapporteur: Nikica Mirošević Skvrce, "C.I.4
Update of sections 4.8 and 5.1 of the SmPC in order to update the efficacy and safety information based on the second interim analysis of the overall survival data from study ENDEAVOR (study 20130398); this is a randomised, multicentre, open-label, phase 3 study of carfilzomib and dexamethasone compared to bortezomib with dexamethasone in patients with relapse multiple myeloma. The Package Leaflet is updated accordingly. The RMP version 9.0 has also been submitted.

C.I.4
Update of section 4.8 of the SmPC in order to revise the frequencies of certain adverse drug reactions based on the pooled data set including ENDEAVOR and 7 recently completed studies.

In addition, the Marketing authorisation holder (MAH) took the opportunity to add editorial changes in sections 4.2, 4.4, 6.3 and 6.6 of the SmPC. Editorial changes have also been included in the package leaflet and labelling."
Request for Supplementary Information adopted on 12.10.2017.

Neulasta - pegfilgrastim - EMEA/H/C/000420/II/0093/G
MAH: Amgen Europe B.V., Rapporteur: Robert James Hemmings, PRAC Rapporteur: Patrick Batty
Request for Supplementary Information adopted on 14.09.2017, 22.06.2017.

Olumiant - baricitinib - EMEA/H/C/004085/II/0003
MAH: Eli Lilly Nederland B.V., Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Patrick Batty, "Update of section 4.4 of the SmPC in order to include results of a vaccination

sub-study of the long term extension study I4V-MC-JADY (I4V-MC-JADY: 'A Phase 3, Multicenter Study to Evaluate the Long-Term Safety and Efficacy of Baricitinib in Patients with Rheumatoid Arthritis'). In addition, the updated RMP version 4.0 has been submitted as part of this application."	
Orkambi - lumacaftor / ivacaftor - EMEA/H/C/003954/II/0017 MAH: Vertex Pharmaceuticals (Europe) Ltd., Rapporteur: Nithyanandan Nagercoil, PRAC Rapporteur: Almath Spooner, "Updates of sections 4.8 and 5.1 of the SmPC in order to reflect the long-term safety and efficacy data from Study VX12 809 105. Study VX12 809 105 was a Phase 3, rollover study to evaluate the safety and efficacy of long term treatment with LUM/IVA in subjects aged 12 years and older with cystic fibrosis, homozygous or heterozygous for the F508del CFTR mutation (MEA 001). A new version of the RMP (ver. 3.5) included in this submission has been updated to include the final data from Study 105. In addition, the MAH took the opportunity to bring the PI in line with the latest QRD template version 10 and to make editorial corrections." Request for Supplementary Information adopted on 12.10.2017, 18.05.2017, 23.02.2017.	
Sivextro - tedizolid phosphate - EMEA/H/C/002846/II/0019 MAH: Merck Sharp & Dohme Limited, Rapporteur: Bruno Sepodes, PRAC Rapporteur: Dolores Montero Corominas, "Update of section 4.8 of the SmPC of the concentrate for solution for infusion formulation, in order to add information from BAY119-2631/16121, a Phase 3 randomized, double-blind, multi-centre study comparing the efficacy and safety of intravenous to oral 6-day tedizolid phosphate and intravenous to oral 10 day linezolid for the treatment of acute bacterial skin and skin structure infections (ABSSSI) and change the reported expected frequency of the adverse reaction "infusion site phlebitis" from "uncommon" to "common". The Package Leaflet is updated accordingly. An updated RMP (version 3.2) has also been submitted, proposing to collect safety information regarding tedizolid phosphate by conducting three investigator initiated studies and deleting the	Positive Opinion adopted by consensus on 30.11.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

original proposed long term safety study. The MAH also took the opportunity to make minor editorial corrections throughout the product information." Opinion adopted on 30.11.2017. Request for Supplementary Information adopted on 28.09.2017, 06.07.2017.	
Soliris - eculizumab - EMEA/H/C/000791/II/0098, Orphan MAH: Alexion Europe SAS, Rapporteur: Jorge Camarero Jiménez, PRAC Rapporteur: Eva A. Segovia, "Update of sections 4.6 and 5.3 of the SmPC in order to update the safety information related to pregnancy, lactation and fertility following the review of data in PSUR 13 and 14. Annex II and the Package Leaflet are updated accordingly. The RMP version 17 has also been submitted with updated information on pregnancy and lactation and fertility." Request for Supplementary Information adopted on 14.09.2017.	
Sylvant - siltuximab - EMEA/H/C/003708/II/0026/G, Orphan MAH: Janssen-Cilag International NV, Rapporteur: Concepcion Prieto Yerro, PRAC Rapporteur: Brigitte Keller-Stanislawski, "Update of sections 4.4, 4.8, 5.1 and 5.2 of the SmPC in order to update the product information following final results from studies CNTO328MCD2001 and CNTO328MCD2002 listed as imposed obligation in the Annex II. The Package Leaflet are updated accordingly. The RMP version 4.0 has also been submitted." Request for Supplementary Information adopted on 30.11.2017.	Request for Supplementary Information adopted
Tecentriq - atezolizumab - EMEA/H/C/004143/II/0002/G MAH: Roche Registration Limited, Rapporteur: Sinan B. Sarac, PRAC Rapporteur: Marcia Sofia Sanches de Castro Lopes Silva, "Grouped variations consisting of: 1) update of sections 4.2, 4.4 and 4.8 of the SmPC in order to add myocarditis as a new adverse reaction based on the results of a cumulative review of cases of suspected myocarditis. As a consequence, the information regarding the posology and special warnings have been updated. Annex II, the Package Leaflet and the RMP (version 2.0) have	

been updated accordingly; 2) update of the RMP to add haemolytic anaemia as a new important identified potential risk”	
Trulicity - dulaglutide - EMEA/H/C/002825/II/0022 MAH: Eli Lilly Nederland B.V., Rapporteur: Greg Markey, PRAC Rapporteur: Carmela Macchiarulo, “Update of sections 4.2, 5.1 and 5.2 of the SmPC for Trulicity following completion of a Phase 3 study (H9X-MCGBDX (GBDX)) comparing the effect of once-weekly Trulicity with insulin glargine on glycaemic control over 52 weeks in patients with type 2 Diabetes Mellitus and moderate or severe chronic kidney disease. In addition, an update to the ATC code and a correction to the "Instructions for use" in Section 6.6 of the SmPC to make it consistent with instructions on "How to store Trulicity" in the Package Insert Leaflet (PL) are proposed. The RMP version 1.11 has also been submitted.” Request for Supplementary Information adopted on 30.11.2017.	Request for Supplementary Information adopted
Xalkori - crizotinib - EMEA/H/C/002489/II/0050 MAH: Pfizer Limited, Rapporteur: Alexandre Moreau, PRAC Rapporteur: Ghania Chamouni, “Update of sections 4.2, 4.3, 4.4, 4.8 and 5.2 of the SmPC in order to update the information about hepatic impairment based on the results of study (A8081012) which evaluated the effect of hepatic impairment on the pharmacokinetics and safety of crizotinib in advanced cancer patients. The package leaflet is updated accordingly. In addition, the final study report of study (A8081012) and an updated RMP version 7.4 are also being submitted.” Request for Supplementary Information adopted on 12.10.2017.	
Zydelig - idelalisib - EMEA/H/C/003843/II/0035/G MAH: Gilead Sciences International Limited, Rapporteur: Filip Josephson, PRAC Rapporteur: Patrick Batty, “Update of section 5.3 of the SmPC in order to revise the carcinogenicity information for idelalisib based on final results from two long term carcinogenicity studies (TX-312-2017, TX-312-2019). The RMP version 2.3	Positive Opinion adopted by consensus on 30.11.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

has also been submitted. In addition, the Marketing authorisation holder (MAH) took the opportunity to bring the PI in line with the latest QRD template version 10.0."
Opinion adopted on 30.11.2017.
Request for Supplementary Information adopted on 28.09.2017.

Zydelig - idelalisib -
EMEA/H/C/003843/II/0038
MAH: Gilead Sciences International Limited, Rapporteur: Filip Josephson, PRAC Rapporteur: Patrick Batty, "Update of sections 4.2, 4.4 and 4.8 of the SmPC in order to reflect information from a recent cumulative safety review of cases of organising pneumonia. The safety review resulted from the Marketing authorisation holder (MAH) MAH ongoing pharmacovigilance and signal detection for Zydelig.

The RMP version 2.6 has also been submitted to extend the deadlines for submission of final CSRs for three studies linked with Annex II conditions. The Package Leaflet and Labelling are updated accordingly."

Zykadia - ceritinib -
EMEA/H/C/003819/II/0015
MAH: Novartis Europharm Limited, Rapporteur: Jorge Camarero Jiménez, PRAC Rapporteur: Ulla Wändel Liminga, "Update of sections 4.2, 4.4, 4.5, 4.8 and 5.2 of the SmPC in order to update the safety informatio based on based on the primary PK and preliminary safety results of the food effect study CLDK378A2112. The Package Leaflet is updated accordingly.

The RMP version 9.0 has also been submitted."
Request for Supplementary Information adopted on 12.10.2017, 22.06.2017.

WS1026
Rasilez-EMEA/H/C/000780/WS1026/0110
Rasilez HCT-
EMEA/H/C/000964/WS1026/0080
MAH: Noden Pharma DAC, Lead Rapporteur: Daniela Melchiorri, Lead PRAC Rapporteur: Carmela Macchiarulo, "Update of section 5.1 of the SmPC in order to reflect the results of study SPP100F2301 (ATMOSPHERE) a multicenter, randomized, doubleblind, parallel group, active-controlled study to evaluate the efficacy and safety of both aliskiren monotherapy and

aliskiren/enalapril combination therapy compared to enalapril monotherapy, on morbidity and mortality in patients with chronic heart failure (NYHA Class II - IV). The RMP (v 13) has also been updated to reflect the study results." Request for Supplementary Information adopted on 09.11.2017, 14.09.2017, 22.06.2017, 21.04.2017, 15.12.2016.	
WS1190/G **Enbrel-** **EMEA/H/C/000262/WS1190/0210/G** **LIFMIOR-** **EMEA/H/C/004167/WS1190/0009/G** MAH: Pfizer Limited, Lead Rapporteur: Robert James Hemmings, Lead PRAC Rapporteur: Patrick Batty Request for Supplementary Information adopted on 30.11.2017, 28.09.2017.	See Agenda 9.1
WS1248/G **Blitzima-** **EMEA/H/C/004723/WS1248/0002/G** **Ritemvia-** **EMEA/H/C/004725/WS1248/0002/G** **Rituzena-** **EMEA/H/C/004724/WS1248/0003/G** MAH: Celltrion Healthcare Hungary Kft., Duplicate, Duplicate of Truxima, Lead Rapporteur: Sol Ruiz, Lead PRAC Rapporteur: Doris Stenver Opinion adopted on 30.11.2017.	Positive Opinion adopted by consensus on 30.11.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1292 **Evotaz-EMEA/H/C/003904/WS1292/0019** **Reyataz-** **EMEA/H/C/000494/WS1292/0114** MAH: Bristol-Myers Squibb Pharma EEIG, Lead Rapporteur: Joseph Emmerich, Lead PRAC Rapporteur: Caroline Laborde, "Update of section 4.3 and 4.5 of the SmPC in order to add a contraindication with lurasidone to reflect this interaction based on literature data. The Package Leaflet is updated accordingly. The RMP of Reyataz/Evotaz versions 14 and 6 respectively have been submitted."	

B.5.4. PRAC assessed procedures

PRAC Led	Positive Opinion adopted by consensus on

Arzerra - ofatumumab - EMEA/H/C/001131/II/0054, Orphan MAH: Novartis Europharm Limited, Rapporteur: Sinan B. Sarac, PRAC Rapporteur: Doris Stenver, PRAC-CHMP liaison: Sinan B. Sarac, "Update of the RMP (version 14.1) in order to amend the study milestones for two studies: - study OMB112517, a phase III, open label, randomized, multicentre trial of ofatumumab maintenance treatment versus no further treatment in subjects with relapsed CLL who have responded to induction therapy: final study report to be submitted by 30 June 2019 - study OMB110913, a phase III, open label, randomized trial of ofatumumab added to fludarabine-cyclophosphamide versus fludarabine-cyclophosphamide combination in subjects with relapsed CLL: final study report to be submitted by 30 June 2019. In addition, changes have been implemented in the safety specifications as previously agreed during the last PSUR outcome (EMEA/H/C/PSUSA/00002202/201610) to upgrade the potential risk of infections and progressive multifocal leukoencephalopathy from potential to an identified risk." Opinion adopted on 30.11.2017.	30.11.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
PRAC Led **Betaferon - interferon beta-1b - EMEA/H/C/000081/II/0118** MAH: Bayer AG, PRAC Rapporteur: Julie Williams, PRAC-CHMP liaison: Greg Markey, "Submission of the final report from study BETAPAEDIC, listed as a category 3 study in the RMP. This was a non-interventional study evaluating safety and tolerability of Betaferon in paediatric patients with multiple sclerosis. The RMP version 3.2 has also been submitted." Request for Supplementary Information adopted on 30.11.2017.	Request for Supplementary Information adopted
PRAC Led **Eliquis - apixaban - EMEA/H/C/002148/II/0048** MAH: Bristol-Myers Squibb / Pfizer EEIG, Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Menno van der Elst, PRAC-CHMP liaison: Johann Lodewijk Hillege, "Submission of the final report from study (B0661073) listed as a category 4 study in the RMP. This is a non-interventional post-authorisation safety study	Positive Opinion adopted by consensus on 30.11.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

(PASS) of the utilisation patterns of apixaban in Denmark. In addition, a revised RMP (version 18.0) is submitted." Opinion adopted on 30.11.2017.	
PRAC Led **Eperzan - albiglutide -** **EMEA/H/C/002735/II/0029/G** MAH: GlaxoSmithKline Trading Services Limited, Rapporteur: TBA, PRAC Rapporteur: Julie Williams, PRAC-CHMP liaison: Greg Markey, "II: C.I.11.b - Update of the RMP to amend Study 201805 (category 3 study): "Observational Study of the Risk of Common Malignant Neoplasms and Malignant Neoplasms of Special Interest (Thyroid and Pancreatic Cancer) in Subjects Prescribed Albiglutide Compared to Those Prescribed Other Antidiabetic Agents", in order to use a different database to study the risk of neoplasms in association with albiglutide exposure II: C.I.11.b – Update of the RMP to add a new category 3 study as an additional pharmacovigilance activity – Study 207351: "Observational Study to Assess Maternal and Fetal Outcomes following exposure to Albiglutide during Pregnancy"" Request for Supplementary Information adopted on 09.11.2017, 22.06.2017, 26.01.2017. Letter from the applicant dated 15.11.2017 requesting a clock stop extension. **For information.**	
PRAC Led **Eylea - aflibercept -** **EMEA/H/C/002392/II/0039** MAH: Bayer AG, Rapporteur: Alexandre Moreau, PRAC Rapporteur: Ghania Chamouni, PRAC-CHMP liaison: Alexandre Moreau, "Submission of the final report from the post authorisation safety study 16526, listed as a category 3 study in the RMP. This is an observational study to evaluate the physician and patient knowledge of safety and safe use information for Aflibercept in Europe as stated in the EU Educational Material of Eylea." Request for Supplementary Information adopted on 30.11.2017.	Request for Supplementary Information adopted
PRAC Led **Nulojix - belatacept -** **EMEA/H/C/002098/II/0047/G**	Request for Supplementary Information adopted

MAH: Bristol-Myers Squibb Pharma EEIG, Rapporteur: Filip Josephson, PRAC Rapporteur: Ulla Wändel Liminga, PRAC-CHMP liaison: Filip Josephson, “Submission of the final report from studies IM103061 and IM103089, listed as a category 3 studies in the RMP. IM103061 is an epidemiological study on pregnancy outcome among belatacept users in the US. IM103089 evaluates data retrospectively to assess the association between belatacept and the risk of PTDL in renal transplant recipients in Europe. An updated RMP, reflecting completion of the two above studies is being submitted as part of this variation (Version 15).” Request for Supplementary Information adopted on 30.11.2017.	
PRAC Led **Sebivo - telbivudine - EMEA/H/C/000713/II/0048** MAH: Novartis Europharm Limited, Rapporteur: Joseph Emmerich, PRAC Rapporteur: Caroline Laborde, PRAC-CHMP liaison: Joseph Emmerich, “Submission of an updated RMP version 11.0 in order to upgrade the risk of lactic acidosis from an important potential to an important identified risk and to include a targeted questionnaire for fatal cases as additional risk minimisation measure as requested by the PRAC as part of the assessment of PSUSA/00002880/201608.” Request for Supplementary Information adopted on 30.11.2017.	Request for Supplementary Information adopted
PRAC Led **Xarelto - rivaroxaban - EMEA/H/C/000944/II/0055** MAH: Bayer AG, Rapporteur: Kristina Dunder, PRAC Rapporteur: Qun-Ying Yue, PRAC-CHMP liaison: Kristina Dunder, “Submission of the final study report of a non-interventional PASS listed as a category 3 study in the RMP (MEA 019): An Observational Post-Authorization Safety Specialist Cohort Event Monitoring Study (SCEM) to Monitor the Safety and Utilization of Rivaroxaban (Xarelto) for the Prevention of Stroke in Patients with AF, Treatment of DVT and PE, and the Prevention of Recurrent DVT and PE in the Secondary Care Setting in England	Positive Opinion adopted by consensus on 30.11.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

and Wales (The ROSE Study), study number 16171." Opinion adopted on 30.11.2017. Request for Supplementary Information adopted on 28.09.2017.	
PRAC Led **WS1164** **Glyxambi-** **EMEA/H/C/003833/WS1164/0008** **Jardiance-** **EMEA/H/C/002677/WS1164/0033** **Synjardy-** **EMEA/H/C/003770/WS1164/0030** MAH: Boehringer Ingelheim International GmbH, Lead Rapporteur: Johann Lodewijk Hillege, Lead PRAC Rapporteur: Dolores Montero Corominas, PRAC-CHMP liaison: Concepcion Prieto Yerro, "C.I.11: Submission of an updated RMP for Jardiance (v12.1), for Synjardy (9.2) and for Glyxambi (v3.0) in order to address the PRAC recommendation concluded in the Article 20 referral for SGLT2 inhibitors on the important potential risk for lower limb amputation. Additionally, the PRAC request to include pancreatitis as important potential risk for empagliflozin-containing medicines following the conclusion adopted by the PRAC after the review of PSUSA/00010077/201603 (canagliflozin) is discussed." Opinion adopted on 30.11.2017. Request for Supplementary Information adopted on 28.09.2017.	Positive Opinion adopted by consensus on 30.11.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
PRAC Led **WS1207** **Bretaris Genuair-** **EMEA/H/C/002706/WS1207/0034** **Eklira Genuair-** **EMEA/H/C/002211/WS1207/0034** MAH: AstraZeneca AB, Lead Rapporteur: Nithyanandan Nagercoil, Lead PRAC Rapporteur: Julie Williams, PRAC-CHMP liaison: Robert James Hemmings, "Submission of the final report from study D6560R00005, (Aclidinium Bromide Drug Utilisation Post-Authorisation Safety Studies (DUS 1) in the United Kingdom, Denmark, and Germany) listed as a category 3 study in the RMP (MEA002). The updated RMP version 6.0 has also been submitted." Opinion adopted on 30.11.2017. Request for Supplementary Information adopted	Positive Opinion adopted by consensus on 30.11.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

on 28.09.2017.	
PRAC Led **WS1229** **Ebymect-EMEA/H/C/004162/WS1229/0025** **Edistride-EMEA/H/C/004161/WS1229/0019** **Forxiga-EMEA/H/C/002322/WS1229/0039** **Xigduo-EMEA/H/C/002672/WS1229/0036** MAH: AstraZeneca AB, Lead Rapporteur: Kristina Dunder, Lead PRAC Rapporteur: Qun-Ying Yue, PRAC-CHMP liaison: Kristina Dunder, "Submission of the final report from study D1690R00013 listed as a category 3 study in the RMP: Incidence of Diabetic Ketoacidosis among Patients with Type 2 Diabetes in the United States. The RMP version 15 (Forxiga/Edistride) and version 10 (Xigduo/Ebymect) have been consequentially updated." Opinion adopted on 30.11.2017.	Positive Opinion adopted by consensus on 30.11.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
PRAC Led **WS1256** **Harvoni-EMEA/H/C/003850/WS1256/0059** **Sovaldi-EMEA/H/C/002798/WS1256/0044** MAH: Gilead Sciences International Limited, Lead Rapporteur: TBA, Lead PRAC Rapporteur: Julie Williams, PRAC-CHMP liaison: Greg Markey, "Submission of the final report from study GS-EU-337-2030, listed as a category 3 study in the RMP. This is an observational, cross-sectional post-authorisation safety study to assess healthcare provider awareness of risks related to sofosbuvir and ledipasvir/sofosbuvir." Opinion adopted on 30.11.2017.	Positive Opinion adopted by consensus on 30.11.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
PRAC Led **WS1259** **Ebymect-EMEA/H/C/004162/WS1259/0024** **Edistride-EMEA/H/C/004161/WS1259/0018** **Forxiga-EMEA/H/C/002322/WS1259/0038** **Xigduo-EMEA/H/C/002672/WS1259/0035** MAH: AstraZeneca AB, Lead Rapporteur: Kristina Dunder, Lead PRAC Rapporteur: Qun-Ying Yue, PRAC-CHMP liaison: Kristina Dunder,	Positive Opinion adopted by consensus on 30.11.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

"Submission of the final report for the Drug Utilisation Study MB102-134 listed as a category 3 study in the RMP: Observational Single-cohort Data Base Study of Dapagliflozin Utilization in Europe. The RMP version 15 (Forxiga/Edistride) and version 10 (Xigduo/Ebymect) have been consequentially updated." Opinion adopted on 30.11.2017.	
PRAC Led **WS1264** **Ariclaim-EMEA/H/C/000552/WS1264/0068** **Cymbalta-EMEA/H/C/000572/WS1264/0072** **Duloxetine Lilly-EMEA/H/C/004000/WS1264/0008** **Xeristar-EMEA/H/C/000573/WS1264/0075** **Yentreve-EMEA/H/C/000545/WS1264/0058** MAH: Eli Lilly Nederland B.V., Duplicate, Duplicate of Yentreve, Lead Rapporteur: Concepcion Prieto Yerro, Lead PRAC Rapporteur: Dolores Montero Corominas, PRAC-CHMP liaison: Concepcion Prieto Yerro, "Submission of the final report from study F1J-MC-B056 listed as a category 3 study in the RMP. This is a non-interventional non-imposed study aimed to investigate the association between duloxetine exposure and suicide-related behaviours and ideation in women with stress urinary inconsistence (SUI). The RMP version 12.3 has also been submitted." Request for Supplementary Information adopted on 30.11.2017.	Request for Supplementary Information adopted
PRAC Led **WS1294** **Actos-EMEA/H/C/000285/WS1294/0078** **Competact-EMEA/H/C/000655/WS1294/0068** **Glubrava-EMEA/H/C/000893/WS1294/0055** **Glustin-EMEA/H/C/000286/WS1294/0077** **Tandemact-EMEA/H/C/000680/WS1294/0056** MAH: Takeda Pharma A/S, Informed Consent of Competact, Lead PRAC Rapporteur: Almath Spooner, PRAC-CHMP liaison: Peter Kiely, "To update the RMPs to version 24.0 for Actos &	

Glustin (pioglitazone), version 22.0 for Tandemact (pioglitazone/ glimepiride), and version 25.0 for Compeact & Glubrava (pioglitazone/metformin). These RMPs have been updated as follows: For all RMPs: Added bone mechanistic (AD4833-402) addendum information per EMA Post-authorization measure (PAM) Report dated 17-Oct-2016: Actos H/C/285 (MEA 68.2), Glustin H/C/286 (MEA 70.2); Competact H/C/655 (MEA 29.2); Glubrava H/C/893 (MEA 18.2); Tandemact H/C/680 (MEA 27.2). The MAH also took the opportunity to make some general edits to these RMPs. For pioglitazone/metformin RMP only: Added lactic acidosis questionnaire per assessor's comment to the final report for Type II Variation EMEA/H/C/000655/WS0991/0062 & EMEA/H/C/000893/WS0991/0047."	

B.5.5. CHMP-CAT assessed procedures

MACI - matrix applied characterised autologous cultured chondrocytes - EMEA/H/C/002522/II/0014/G, ATMP MAH: Vericel Denmark ApS, Rapporteur: Christiane Niederlaender	

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

WS1185/G **Hexacima-** **EMEA/H/C/002702/WS1185/0065/G** **Hexaxim-** **EMEA/H/W/002495/WS1185/0071/G** **Hexyon-** **EMEA/H/C/002796/WS1185/0069/G** MAH: Sanofi Pasteur Europe, Duplicate, Duplicate of Hexacima, Lead Rapporteur: Jan Mueller-Berghaus Opinion adopted on 16.11.2017. Request for Supplementary Information adopted	Positive Opinion adopted by consensus on 16.11.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

on 14.09.2017.	
WS1199/G **ProQuad-** **EMEA/H/C/000622/WS1199/0120/G** **Zostavax-** **EMEA/H/C/000674/WS1199/0114/G** MAH: MSD Vaccins, Lead Rapporteur: Jan Mueller-Berghaus Opinion adopted on 16.11.2017.	Positive Opinion adopted by consensus on 16.11.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1202/G **Efficib-** **EMEA/H/C/000896/WS1202/0084/G** **Janumet-** **EMEA/H/C/000861/WS1202/0084/G** **Januvia-** **EMEA/H/C/000722/WS1202/0058/G** **Ristaben-** **EMEA/H/C/001234/WS1202/0050/G** **Ristfor-** **EMEA/H/C/001235/WS1202/0071/G** **TESAVEL-** **EMEA/H/C/000910/WS1202/0058/G** **Velmetia-** **EMEA/H/C/000862/WS1202/0087/G** **Xelevia-** **EMEA/H/C/000762/WS1202/0062/G** MAH: Merck Sharp & Dohme Limited, Lead Rapporteur: Johann Lodewijk Hillege Opinion adopted on 16.11.2017. Request for Supplementary Information adopted on 14.09.2017.	Positive Opinion adopted by consensus on 16.11.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1245 **Infanrix hexa-** **EMEA/H/C/000296/WS1245/0228** MAH: GlaxoSmithkline Biologicals SA, Lead Rapporteur: Bart Van der Schueren Opinion adopted on 30.11.2017.	Positive Opinion adopted by consensus on 30.11.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1246/G **Epclusa-** **EMEA/H/C/004210/WS1246/0017/G** **Harvoni-** **EMEA/H/C/003850/WS1246/0060/G** **Sovaldi-** **EMEA/H/C/002798/WS1246/0045/G** **Vosevi-** **EMEA/H/C/004350/WS1246/0005/G** MAH: Gilead Sciences International Limited, Lead Rapporteur: Filip Josephson	Positive Opinion adopted by consensus on 30.11.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

Opinion adopted on 30.11.2017.	
WS1250/G **Infanrix hexa-** **EMEA/H/C/000296/WS1250/0230/G** MAH: GlaxoSmithkline Biologicals SA, Lead Rapporteur: Bart Van der Schueren	Weekly start timetable.
WS1255 **Infanrix hexa-** **EMEA/H/C/000296/WS1255/0231** MAH: GlaxoSmithkline Biologicals SA, Lead Rapporteur: Bart Van der Schueren	Weekly start timetable.
WS1257 **Infanrix hexa-** **EMEA/H/C/000296/WS1257/0229** MAH: GlaxoSmithkline Biologicals SA, Lead Rapporteur: Bart Van der Schueren	Weekly start timetable.
WS1263/G **Avamys-** **EMEA/H/C/000770/WS1263/0035/G** **Relvar Ellipta-** **EMEA/H/C/002673/WS1263/0034/G** **Revinty Ellipta-** **EMEA/H/C/002745/WS1263/0030/G** MAH: Glaxo Group Ltd, Lead Rapporteur: Concepcion Prieto Yerro Opinion adopted on 16.11.2017.	Positive Opinion adopted by consensus on 16.11.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1271/G **Ebymect-** **EMEA/H/C/004162/WS1271/0028/G** **Qtern-** **EMEA/H/C/004057/WS1271/0010/G** **Xigduo-** **EMEA/H/C/002672/WS1271/0039/G** MAH: AstraZeneca AB, Lead Rapporteur: Kristina Dunder Opinion adopted on 23.11.2017.	Positive Opinion adopted by consensus on 23.11.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1279 **Helixate NexGen-** **EMEA/H/C/000276/WS1279/0193** **KOGENATE Bayer-** **EMEA/H/C/000275/WS1279/0201** MAH: Bayer AG, Lead Rapporteur: Jan Mueller-Berghaus Opinion adopted on 30.11.2017.	Positive Opinion adopted by consensus on 30.11.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1280 **Blitzima-** **EMEA/H/C/004723/WS1280/0005**	Weekly start timetable.

Ritemvia- **EMEA/H/C/004725/WS1280/0005** **Rituzena-** **EMEA/H/C/004724/WS1280/0006** MAH: Celltrion Healthcare Hungary Kft., Duplicate, Duplicate of Truxima, Lead Rapporteur: Sol Ruiz	
WS1287 **Abseamed-** **EMEA/H/C/000727/WS1287/0066** **Binocrit-** **EMEA/H/C/000725/WS1287/0066** **Epoetin alfa Hexal-** **EMEA/H/C/000726/WS1287/0065** MAH: Sandoz GmbH, Lead Rapporteur: Alexandre Moreau	Weekly start timetable.
WS1290 **Abseamed-** **EMEA/H/C/000727/WS1290/0067** **Binocrit-** **EMEA/H/C/000725/WS1290/0067** **Epoetin alfa Hexal-** **EMEA/H/C/000726/WS1290/0066** MAH: Sandoz GmbH, Lead Rapporteur: Alexandre Moreau	Weekly start timetable.
WS1313/G **AMGEVITA-** **EMEA/H/C/004212/WS1313/0003/G** **SOLYMBIC-** **EMEA/H/C/004373/WS1313/0003/G** MAH: Amgen Europe B.V., Lead Rapporteur: Kristina Dunder“To update section 5.1 of the SmPC in order to add information on nail psoriasis to align the PI to its parent product Humira. The Package Leaflet is updated in accordance. Furthermore, the PI is brought in line with the latest QRD template version 10.0. The MAH has also taken the occasion to correct some editorial mistakes in the PI. To update section 5.1 of the SmPC to reflect the latest results for Enthesitis-related arthritis to align the PI to its parent product Humira. To update section 5.1 of the SmPC in order to update information on the long-term safety, tolerability, and efficacy of adalimumab in subjects with moderate to severe hidradenitis suppurativa to align the PI to its parent product Humira.	

In addition, the PI has been updated to ensure that the paediatric statement is reflected correctly for Amgevita and Solymbic.”	

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

Raplixa - human fibrinogen / human thrombin - EMEA/H/C/002807/II/0028/G MAH: Mallinckrodt Pharmaceuticals Ireland Limited, Rapporteur: Nithyanandan Nagercoil Withdrawal request submitted on 04.12.2017.	The MAH withdrew the procedure on 04.12.2017.

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

doravirine - EMEA/H/C/004747 , treatment of adults infected with HIV-1 without past or present evidence of viral resistance to treatment of adults infected with HIV-1 without past or present evidence of viral resistance to doravirine	
doravirine / lamivudine / tenofovir disoproxil - EMEA/H/C/004746 , treatment of adults infected with HIV-1 without past or present evidence of viral resistance to doravirine, lamivudine, or tenofovir	
galcanezumab - EMEA/H/C/004648 , prophylaxis of migraine	
pegfilgrastim - EMEA/H/C/004915 , treatment of neutropenia	
pegfilgrastim - EMEA/H/C/004556 , Reduction in the duration of neutropenia and the incidence of febrile neutropenia in adult patients	
adalimumab - EMEA/H/C/004475 , treatment of rheumatoid arthritis, psoriatic arthritis and ankylosing spondylitis	
inotersen - EMEA/H/C/004782, Orphan , treatment of transthyretin amyloidosis (hATTR)	**Accelerated review**
tisagenlecleucel - EMEA/H/C/004090,	**Accelerated review**

Orphan, ATMP
, treatment of B cell acute lymphoblastic leukaemia (ALL) and diffuse large B cell lymphoma (DLBCL)

macimorelin - EMEA/H/C/004660
, Diagnosis of Adult growth hormone deficiency (AGHD)

trastuzumab - EMEA/H/C/004916
, treatment of metastatic and early breast cancer and metastatic gastric cancer (MGC)

buprenorphine - EMEA/H/C/004743
, Substitution treatment for opioid drug dependence

influenza vaccine surface antigen, inactivated, prepared in cell cultures - EMEA/H/C/004814,
, prophylaxis of influenza in adults and children from 4 years of age

human fibrinogen / human thrombin - EMEA/H/D/004308
, to support the endogenous clotting process and increase of haemostasis in surgical procedures

daunorubicin / cytarabine - EMEA/H/C/004282, Orphan
, treatment of adults with high-risk acute myeloid leukaemia (AML)

Accelerated review

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

fingolimod - EMEA/H/C/002202/X/0044/G

enoxaparin sodium - EMEA/H/C/004264/X/0026

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

bortezomib - EMEA/H/C/003984/X/0008
List of Questions adopted on 20.07.2017.

caplacizumab - EMEA/H/C/004426, Orphan
, indicated for the treatment of acquired thrombotic thrombocytopenic purpura (aTTP).
List of Questions adopted on 22.06.2017.

emicizumab - EMEA/H/C/004406
, routine prophylaxis to prevent bleeding or reduce the frequency of bleeding episodes in patients with haemophilia A (congenital factor VIII deficiency) with factor VIII inhibitors.
List of Questions adopted on 10.10.2017.

dolutegravir / rilpivirine - EMEA/H/C/004427
, treatment of HIV
List of Questions adopted on 12.10.2017.

pemetrexed - EMEA/H/C/003958
, treatment of malignant pleural mesothelioma and non-small cell lung cancer, Generic, Generic of Alimta
List of Questions adopted on 12.10.2017.

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

mecasermin - EMEA/H/C/000704/S/0050

susoctocog alfa - EMEA/H/C/002792/S/0016

cholic acid - EMEA/H/C/001250/S/0022, Orphan

tocofersolan - EMEA/H/C/000920/S/0027

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

Atosiban SUN - atosiban - EMEA/H/C/002329/R/0012
MAH: Sun Pharmaceutical Industries Europe B.V., Generic, Generic of Tractocile, Rapporteur: John Joseph Borg, PRAC Rapporteur: Amelia Cupelli

Deltyba - delamanid - EMEA/H/C/002552/R/0027, Orphan
MAH: Otsuka Novel Products GmbH, Rapporteur: Greg Markey, Co-Rapporteur: Koenraad Norga, PRAC Rapporteur: Julie Williams

Imatinib medac - imatinib - EMEA/H/C/002692/R/0008
MAH: medac Gesellschaft fur klinische Spezialpraparate mbH, Generic, Generic of Glivec, Rapporteur: John Joseph Borg, PRAC Rapporteur: Eva A. Segovia

Imnovid - pomalidomide -
EMEA/H/C/002682/R/0028, Orphan
MAH: Celgene Europe Limited, Rapporteur: Robert James Hemmings, Co-Rapporteur: Jorge Camarero Jiménez, PRAC Rapporteur: Patrick Batty

Imvanex - modified vaccinia ankara virus -
EMEA/H/C/002596/R/0032
MAH: Bavarian Nordic A/S, Rapporteur: Greg Markey, Co-Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Julie Williams

Inflectra - infliximab -
EMEA/H/C/002778/R/0056
MAH: Hospira UK Limited, Duplicate, Duplicate of Remsima, Rapporteur: Greg Markey, Co-Rapporteur: Outi Mäki-Ikola, PRAC Rapporteur: Patrick Batty

Lojuxta - lomitapide -
EMEA/H/C/002578/R/0029
MAH: Aegerion Pharmaceuticals Limited, Rapporteur: Johann Lodewijk Hillege, Co-Rapporteur: Robert James Hemmings, PRAC Rapporteur: Menno van der Elst

Lonquex - lipegfilgrastim -
EMEA/H/C/002556/R/0039
MAH: Sicor Biotech UAB, Rapporteur: Greg Markey, Co-Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Patrick Batty

Nexium Control - esomeprazole -
EMEA/H/C/002618/R/0021
MAH: Pfizer Consumer Healthcare Limited, Rapporteur: Romaldas Mačiulaitis, Co-Rapporteur: Robert James Hemmings, PRAC Rapporteur: Simona Kudeliene

Ovaleap - follitropin alfa -
EMEA/H/C/002608/R/0023
MAH: Teva B.V., Rapporteur: Paula Boudewina van Hennik, Co-Rapporteur: Agnes Gyurasics, PRAC Rapporteur: Menno van der Elst

Pixuvri - pixantrone -
EMEA/H/C/002055/R/0042
MAH: CTI Life Sciences Limited, Rapporteur: Greg Markey, Co-Rapporteur: Filip Josephson, PRAC Rapporteur: Patrick Batty

Remsima - infliximab -
EMEA/H/C/002576/R/0047

MAH: Celltrion Healthcare Hungary Kft., Rapporteur: Greg Markey, Co-Rapporteur: Outi Mäki-Ikola, PRAC Rapporteur: Patrick Batty

Stivarga - regorafenib - EMEA/H/C/002573/R/0025
MAH: Bayer AG, Rapporteur: Paula Boudewina van Hennik, Co-Rapporteur: Daniela Melchiorri, PRAC Rapporteur: Sabine Straus

Tafinlar - dabrafenib - EMEA/H/C/002604/R/0030
MAH: Novartis Europharm Limited, Rapporteur: Filip Josephson, Co-Rapporteur: Bjorg Bolstad, PRAC Rapporteur: Ulla Wändel Liminga

Voncento - human coagulation factor VIII / human von willebrand factor - EMEA/H/C/002493/R/0032
MAH: CSL Behring GmbH, Rapporteur: Paula Boudewina van Hennik, Co-Rapporteur: Nithyanandan Nagercoil, PRAC Rapporteur: Sabine Straus

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

Tagrisso - osimertinib - EMEA/H/C/004124/II/0019
MAH: AstraZeneca AB, Rapporteur: Jorge Camarero Jiménez, Co-Rapporteur: Bjorg Bolstad, PRAC Rapporteur: Sabine Straus, "Extension of Indication to include first-line treatment of adult patients with locally advanced or metastatic non-small cell lung cancer whose tumours have epidermal growth factor receptor exon 19 deletions or exon 21 (L858R) substitution mutations, based on data from the FLAURA study (D5160C00007); a phase III, double-blind, randomised study to assess the efficacy and safety of AZD9291 versus a standard of care epidermal growth factor receptor-Tyrosine Kinase Inhibitor as first-line treatment in patients with epidermal growth factor receptor mutation-positive, locally-advanced or metastatic non-small-cell lung cancer.
As a consequence, sections 4.1, 4.2, 4.4, 4.8, 5.1, 5.2 of the SmPC have been updated and the Package Leaflet has been updated

accordingly.
In addition, the MAH took the opportunity to implement editorial changes in the SmPC and Package Leaflet.
As part of this application the MAH is requesting an additional year of market protection.
An updated RMP version 8 was submitted as part of the application."

Xarelto - rivaroxaban - EMEA/H/C/000944/II/0058

MAH: Bayer AG, Rapporteur: Kristina Dunder, PRAC Rapporteur: Qun-Ying Yue, "Extension of Indication to include the prevention of stroke, myocardial infarction and cardiovascular death, and for the prevention of acute limb ischaemia and mortality in adult patients with coronary artery disease (CAD) or peripheral artery disease (PAD) for Xarelto 2.5 mg co-administered with acetylsalicylic acid; as a consequence, sections 4.1, 4.2, 4.4, 4.8, and 5.1 of the SmPC are updated. The Package Leaflet and Labelling are updated in accordance.
In addition, section 4.8 of the SmPC is updated for all other dose strengths (10/15/20 mg) of Xarelto with relevant exposure information based on the provided clinical data.
The updated RMP version 11.1 has also been submitted."

Xultophy - insulin degludec / liraglutide - EMEA/H/C/002647/II/0023

MAH: Novo Nordisk A/S, Rapporteur: Kristina Dunder, PRAC Rapporteur: Menno van der Elst, "Update of sections 4.1, 4.2, 4.4, 4.8 and 5.1 of the SmPC in order to update the safety information based on cardiovascular outcomes studies conducted a for each of the monocomponents
of Xultophy: LEADER (Liraglutide Cardiovascular Outcomes Trial) and DEVOTE (Insulin Degludec Cardiovascular Outcomes Trial).

The MAH is also proposing to reorganise parts of section 5.1 to improve the reader friendliness and to remove Xultophy from the list of medicines under additional monitoring.

The Package Leaflet is updated accordingly.

The RMP version 7.0 has also been submitted."

WS1274
Mekinist-

EMEA/H/C/002643/WS1274/0023
Tafinlar-
EMEA/H/C/002604/WS1274/0031

MAH: Novartis Europharm Limited, Lead Rapporteur: Paula Boudewina van Hennik, Lead Co-Rapporteur: Filip Josephson, Lead PRAC Rapporteur: Ulla Wändel Liminga, “Extension of indication to include the combination adjuvant treatment with trametinib and dabrafenib of adult patients with stage III melanoma with a BRAF V600 mutation, following complete resection. As a consequence, sections 4.1, 4.2, 4.4, 4.8, 5.1 of the Mekinist and Tafinlar SmPCs are updated. The Package Leaflet and the Risk Management plan (version 14.0 for Mekinist and version 9.0 for Tafinlar, according to GVP module V revision 2) are updated in accordance. In addition, the Worksharing applicant (WSA) took the opportunity to correct some typos throughout the Mekinist and Tafinlar product information, to include a cross reference to the Mekinist SmPC in section 4.6 of the Tafinlar SmPC regarding fertility, to update the list of local representatives for Bulgaria, Hungary, Estonia, Latvia and Lithuania in the Package Leaflet of both products.”

WS1278
Opdivo-EMEA/H/C/003985/WS1278/0042
Yervoy-EMEA/H/C/002213/WS1278/0053

MAH: Bristol-Myers Squibb Pharma EEIG, Lead Rapporteur: Jorge Camarero Jiménez, Lead Co-Rapporteur: Paula Boudewina van Hennik, Lead PRAC Rapporteur: Brigitte Keller-Stanislawski, “Extension of indication to include the combination treatment with nivolumab and ipilimumab of adult patients with intermediate/poor-risk advanced renal cell carcinoma. As a consequence sections 4.1, 4.2, 4.4, 4.8, 5.1 and 5.2 of the Opdivo and Yervoy SmPCs are updated. The Package Leaflet and the Risk Management Plan (version 19.0 for Yervoy and version 13.0 for Opdivo) are updated in accordance. In addition, the Worksharing applicant (WSA) took the opportunity to correct some typos throughout the Yervoy and Opdivo product information.”

B.6.8. CHMP assessed procedures scope: Pharmaceutical aspects

Bortezomib Accord - bortezomib -

EMEA/H/C/003984/II/0012
MAH: Accord Healthcare Ltd, Generic, Generic of VELCADE, Rapporteur: Milena Stain

Cinryze - C1-esterase inhibitor, human - EMEA/H/C/001207/II/0058/G
MAH: Shire Services BVBA, Rapporteur: Jan Mueller-Berghaus

Cosentyx - secukinumab - EMEA/H/C/003729/II/0031/G
MAH: Novartis Europharm Limited, Rapporteur: Tuomo Lapveteläinen

Erelzi - etanercept - EMEA/H/C/004192/II/0005/G
MAH: Sandoz GmbH, Rapporteur: Johann Lodewijk Hillege

Evicel - human fibrinogen / human thrombin - EMEA/H/C/000898/II/0053
MAH: Omrix Biopharmaceuticals N. V., Rapporteur: Jan Mueller-Berghaus

Flixabi - infliximab - EMEA/H/C/004020/II/0020
MAH: Samsung Bioepis UK Limited, Rapporteur: Jan Mueller-Berghaus

Ilaris - canakinumab - EMEA/H/C/001109/II/0053/G
MAH: Novartis Europharm Limited, Rapporteur: Jan Mueller-Berghaus

Lojuxta - lomitapide - EMEA/H/C/002578/II/0028
MAH: Aegerion Pharmaceuticals Limited, Rapporteur: Johann Lodewijk Hillege

Memantine ratiopharm - memantine - EMEA/H/C/002671/II/0012
MAH: ratiopharm GmbH, Generic, Generic of Ebixa, Rapporteur: Bart Van der Schueren

Nplate - romiplostim - EMEA/H/C/000942/II/0066, Orphan
MAH: Amgen Europe B.V., Rapporteur: Concepcion Prieto Yerro

Nucala - mepolizumab - EMEA/H/C/003860/II/0012
MAH: GlaxoSmithKline Trading Services Limited, Rapporteur: Nithyanandan Nagercoil

Opsumit - macitentan - EMEA/H/C/002697/II/0025/G, Orphan

MAH: Actelion Registration Limited, Rapporteur: Concepcion Prieto Yerro

Oxervate - cenegermin - EMEA/H/C/004209/II/0002, Orphan
MAH: Dompe farmaceutici S.p.A., Rapporteur: Concepcion Prieto Yerro

Plegridy - peginterferon beta-1a - EMEA/H/C/002827/II/0040
MAH: Biogen Idec Ltd, Rapporteur: Johann Lodewijk Hillege

Praluent - alirocumab - EMEA/H/C/003882/II/0032/G
MAH: sanofi-aventis groupe, Rapporteur: Johann Lodewijk Hillege

Respreeza - human alpha1-proteinase inhibitor - EMEA/H/C/002739/II/0020
MAH: CSL Behring GmbH, Rapporteur: Kristina Dunder

Torisel - temsirolimus - EMEA/H/C/000799/II/0070/G, Orphan
MAH: Pfizer Limited, Rapporteur: Harald Enzmann

Trulicity - dulaglutide - EMEA/H/C/002825/II/0023
MAH: Eli Lilly Nederland B.V., Rapporteur: Greg Markey

Vpriv - velaglucerase alfa - EMEA/H/C/001249/II/0035, Orphan
MAH: Shire Pharmaceuticals Ireland Ltd, Rapporteur: Harald Enzmann

WS1237/G
Ambirix- EMEA/H/C/000426/WS1237/0089/G
Fendrix- EMEA/H/C/000550/WS1237/0061/G
Infanrix hexa- EMEA/H/C/000296/WS1237/0233/G
Twinrix Adult- EMEA/H/C/000112/WS1237/0123/G
Twinrix Paediatric- EMEA/H/C/000129/WS1237/0124/G
MAH: GlaxoSmithKline Biologicals, Lead Rapporteur: Bart Van der Schueren

WS1276/G
Incruse- EMEA/H/C/002809/WS1276/0017/G

Rolufta-
EMEA/H/C/004654/WS1276/0003/G
MAH: Glaxo Group Ltd, Lead Rapporteur: Concepcion Prieto Yerro

WS1281/G
Hexacima-
EMEA/H/C/002702/WS1281/0072/G
Hexaxim-
EMEA/H/W/002495/WS1281/0077/G
Hexyon-
EMEA/H/C/002796/WS1281/0076/G
MAH: Sanofi Pasteur SA, Lead Rapporteur: Jan Mueller-Berghaus

WS1311/G
Aflunov-
EMEA/H/C/002094/WS1311/0040/G
Foclivia-
EMEA/H/C/001208/WS1311/0034/G
MAH: Seqirus S.r.l, Lead Rapporteur: Daniela Melchiorri

WS1314
Abasaglar-
EMEA/H/C/002835/WS1314/0017
Humalog-
EMEA/H/C/000088/WS1314/0162
Liprolog-
EMEA/H/C/000393/WS1314/0124
MAH: Eli Lilly Regional Operations GmbH, Lead Rapporteur: Robert James Hemmings

WS1317/G
Helixate NexGen-
EMEA/H/C/000276/WS1317/0194/G
KOGENATE Bayer-
EMEA/H/C/000275/WS1317/0202/G
MAH: Bayer AG, Lead Rapporteur: Jan Mueller-Berghaus

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

Advate - octocog alfa -
EMEA/H/C/000520/II/0088
MAH: Baxter AG, Rapporteur: Jan Mueller-Berghaus, “Submission of the final report from study 061301. This is an interventional, open-label study aimed to evaluate the efficacy and safety of Advate in the treatment of previously treated patients with haemophilia A.”

Advate - octocog alfa -

EMEA/H/C/000520/II/0090
MAH: Baxter AG, Rapporteur: Jan Mueller-Berghaus, “Submission of the final clinical study report from study 060402. This was an interventional, randomised, controlled study aimed to compare the efficacy and safety of continuous infusion versus intermittent bolus infusion in patients with haemophilia A undergoing major orthopaedic surgery.”

CellCept - mycophenolate mofetil - EMEA/H/C/000082/II/0137
MAH: Roche Registration Limited, Rapporteur: Greg Markey, “Update of section 4.4 of the SmPC in order to update the information on concomitant use of tacrolimus with CellCept and to provide recommendations on therapeutic drug monitoring for management of transplant patients, based on reviews of the medical literature and clinical treatment guidelines. In addition, the Marketing authorisation holder (MAH) took the opportunity to correct inconsistencies in the Package Leaflet.”

PRAC Led
Cetrotide - cetrorelix - EMEA/H/C/000233/II/0064
MAH: Merck Serono Europe Limited, Rapporteur: Martina Weise, PRAC Rapporteur: Valerie Strassmann, PRAC-CHMP liaison: Martina Weise, “Submission of an updated RMP version 5.0 in order to update the list of important identified risks by adding Ovarian Hyperstimulation Syndrome (OHSS) and removing injection site reactions (ISRs). In addition, further minor RMP updates were introduced.”

Cubicin - daptomycin - EMEA/H/C/000637/II/0066
MAH: Merck Sharp & Dohme Limited, Rapporteur: Greg Markey, “Update of section 4.8 of the SmPC in order to add the adverse events Leukocytosis, Muscle cramps and Eye irritation with the frequency Uncommon, based on data from two previously submitted adult multicentre, randomized clinical studies investigating the safety and efficacy of IV daptomycin compared with that of vancomycin or a semi-synthetic penicillin, in the treatment of complicated skin and skin structure infections due to Gram-positive bacteria (DAP-SST-98-01)

or in the treatment of adult hospitalized subjects with complicated bacterial skin and soft tissue infections due, at least in part, to Gram-positive bacteria (DAP-SST9901)."

Descovy - emtricitabine / tenofovir alafenamide - EMEA/H/C/004094/II/0025

MAH: Gilead Sciences International Limited, Rapporteur: Robert James Hemmings, "Update of sections 4.8, 5.1 and 5.2 of the SmPC in order to provide week 48 results from study GS-US-311-1717(include study identifier) listed as a category 3 study in the RMP; this is a Phase 3b, Randomized, Double-Blind, Switch Study to Evaluate F/TAF in HIV-1 Infected Subjects who are Virologically Suppressed on Regimens containing ABC/3TC.

In addition, the Marketing authorisation holder (MAH) took the opportunity to make administrative updateds and Minor Linguistic Amendments to the Product Information."

Fluenz Tetra - influenza vaccine (live attenuated, nasal) - EMEA/H/C/002617/II/0076

MAH: AstraZeneca AB, Rapporteur: Bart Van der Schueren, "Update of section 4.6 of the SmPC with regards to pregnancy and breast-feeding information based on the review and summary of pregnancy and lactation data from published literature and MAH pharmacovigilance database. The package leaflet has been updated accordingly."

Glivec - imatinib - EMEA/H/C/000406/II/0109

MAH: Novartis Europharm Limited, Rapporteur: Jorge Camarero Jiménez, "Update of section 4.8 of the SmPC to add the new adverse drug reaction (ADR) 'pseudoporphyria' following a revision of the company's core data sheet (CDS). The package leaflet has been updated accordingly.
In addition, the Marketing authorisation holder (MAH) took the opportunity to update the contact details of the local representatives in Bulgaria, Hungary and Latvija in the Package Leaflet."

Humira - adalimumab - EMEA/H/C/000481/II/0172

MAH: AbbVie Limited, Rapporteur: Kristina Dunder, “Update of sections 5.1 and 5.2 of the SmPC for 40mg/0.8ml and 40mg/0.4 ml Prefilled pen and prefilled syringe in order to add information on non-radiographic axial spondyloarthritis following final results from Humira remission-withdrawal-retreatment study (M13-375) listed in the RMP.”

Imbruvica - ibrutinib - EMEA/H/C/003791/II/0039, Orphan

MAH: Janssen-Cilag International NV, Rapporteur: Filip Josephson, “Submission of the final report from the in vitro rabbit ventricular and atrial wedge study (study 16-088-B-X-IV-CT), listed as a category 3 study in the RMP. This in vitro exploratory safety pharmacology study was designed to further elucidate a mechanism or potential association of ibrutinib’s effects on ECG signaling.”

Infanrix hexa - diphtheria (D), tetanus (T), pertussis (acellular, component) (Pa), hepatitis B (rDNA) (HBV), poliomyelitis (inactivated) (IPV) and Haemophilus influenzae type b (Hib) conjugate vaccine (adsorbed) - EMEA/H/C/000296/II/0235

MAH: GlaxoSmithkline Biologicals SA, Rapporteur: Bart Van der Schueren, “Update of section 4.5 of the SmPC in order to update the interactions section with additional data on the co-administration with Meningococcal serogroup B vaccine (MenB) in order to facilitate the administration of Infanrix hexa and Bexsero to infants and toddlers based on final results from clinical studies V72P12, V72P13 and V72P16.”

Invokana - canagliflozin - EMEA/H/C/002649/II/0033/G

MAH: Janssen-Cilag International NV, Rapporteur: Martina Weise, “Update of section 5.1 of the SmPC in order to update the safety information on 'Canagliflozin as initial combination therapy with metformin' based on final results from the DIA3011 study, which is Randomized, Double-Blind, 5-Arm, Parallel-Group, 26-Week, Multicenter Study to Evaluate the Efficacy, Safety, and Tolerability of Canagliflozin in Combination With Metformin as Initial Combination Therapy in the Treatment of Subjects With Type 2 Diabetes Mellitus With Inadequate Glycemic Control With Diet and

Exercise.

Update of section 5.1 of the SmPC in order to update the safety information on 'Add-on combination therapy with Metformin and Dipeptidyl-peptidase-4 Inhibitor' based on final results from the DIA4004 study, which is a Randomized, Double-blind, Placebo Controlled, 2-arm, Parallel-group, 26-week, Multicenter Study to Evaluate the Efficacy, Safety, and Tolerability of Canagliflozin in the Treatment of Subjects with Type 2 Diabetes Mellitus with Inadequate Glycemic Control on Metformin and Sitagliptin Therapy."

NexoBrid - concentrate of proteolytic enzymes enriched in bromelain - EMEA/H/C/002246/II/0032, Orphan

MAH: MediWound Germany GmbH, Rapporteur: Harald Enzmann, "Submission of the clinical study report MW2008-09-03 "Enzymatic Debridement in Patients with Partial Thickness Burns."
an open label, single-arm study evaluating the safety (primary), PK (NexoBrid transcutaneous absorption) and efficacy (exploratory) of NexoBrid in hospitalized adult with partial thickness (mid and deep dermal) thermal burns of 4-30% total body surface area."

Reagila - cariprazine - EMEA/H/C/002770/II/0002

MAH: Gedeon Richter Plc., Rapporteur: Kristina Dunder, "Update of section 4.8 of the SmPC to add the new ADR Steven-Johnson syndrome with an unknown frequency. The Package Leaflet has been updated accordingly."

Sirturo - bedaquiline - EMEA/H/C/002614/II/0026, Orphan

MAH: Janssen-Cilag International NV, Rapporteur: Filip Josephson, "Submission of the final report from study TMC207TBC3001 listed as a category 3 study in the RMP. This is an interventional, open-label, non-comparative, uncontrolled study without formal efficacy objectives and associated statistical analyses to provide early access to BDQ for subjects with pulmonary infection due to pre-XDR or XDR strains of M. tuberculosis."

Stelara - ustekinumab - EMEA/H/C/000958/II/0060

MAH: Janssen-Cilag International NV, Rapporteur: Greg Markey, “Update of sections 4.4, 4.8 and 5.1 of the SmPC in order to update the efficacy data following completion of extension of study IM-UNITI - A Phase 3, Randomized, Double-blind, Placebo-controlled, Parallel-group, Multicenter Study to Evaluate the Safety and Efficacy of Ustekinumab Maintenance Therapy in Subjects with Moderately to Severely Active Crohns Disease.

In addition, the marketing authorisation holder took the opportunity to introduce editorial changes in the SmPC and PL.”

Sutent - sunitinib - EMEA/H/C/000687/II/0068

MAH: Pfizer Limited, Rapporteur: Daniela Melchiorri, “Update of section 4.8 of the SmPC in order to include available long-term safety data pooled from 9 Pfizer-sponsored sunitinib clinical studies in patients with metastatic renal cell carcinoma (MRCC) from a recently published journal article by Porta et al (2016).”

Sycrest - asenapine - EMEA/H/C/001177/II/0030

MAH: N.V. Organon, Rapporteur: Greg Markey, “Update of sections 4.4 and 4.8 of the SmPC to add safety information regarding falls as a result of postmarketing reports and published literature review. The package leaflet is updated accordingly. In addition, the Marketing authorisation holder (MAH) took the opportunity to update the list of local representatives for Denmark, Norway, Slovenia and Slovakia in the Package Leaflet and to bring the PI in line with the latest QRD template version 10.0.”

Tafinlar - dabrafenib - EMEA/H/C/002604/II/0029

MAH: Novartis Europharm Limited, Rapporteur: Filip Josephson, “Update of section 5.2 of the SmPC in order to update the information on the in vitro evaluation of drug-drug interaction potential (to include that dabrafenib is a human BCRP substrate and a OCT2 inhibitor but that the risk of a drug drug interaction is minimal with substrates of OAT1, OAT3 and OTC2), based on the results of non-clinical studies 2014N220059 and 2015N235499.”

Viekirax - ombitasvir / paritaprevir /

ritonavir - EMEA/H/C/003839/II/0039
MAH: AbbVie Limited, Rapporteur: Filip Josephson, “Update of sections 4.3 and 4.5 of the SmPC to add disopyramide in the list of contraindicated medicines and in the list of medicines which interact with Viekirax. The Package Leaflet is updated accordingly.”

Vokanamet - canagliflozin / metformin - EMEA/H/C/002656/II/0033/G
MAH: Janssen-Cilag International NV, Rapporteur: Martina Weise, “Update of section 5.1 of the SmPC in order to update the safety information on 'Canagliflozin as initial combination therapy with metformin' based on final results from the DIA3011 study, which is Randomized, Double-Blind, 5-Arm, Parallel-Group, 26-Week, Multicenter Study to Evaluate the Efficacy, Safety, and Tolerability of Canagliflozin in Combination With Metformin as Initial Combination Therapy in the Treatment of Subjects With Type 2 Diabetes Mellitus With Inadequate Glycemic Control With Diet and Exercise.

Update of section 5.1 of the SmPC in order to update the safety information on 'Add-on combination therapy with Metformin and Dipeptidyl-peptidase-4 Inhibitor' based on final results from the DIA4004 study, which is a Randomized, Double-blind, Placebo Controlled, 2-arm, Parallel-group, 26-week, Multicenter Study to Evaluate the Efficacy, Safety, and Tolerability of Canagliflozin in the Treatment of Subjects with Type 2 Diabetes Mellitus with Inadequate Glycemic Control on Metformin and Sitagliptin Therapy.”

Votrient - pazopanib - EMEA/H/C/001141/II/0043
MAH: Novartis Europharm Limited, Rapporteur: Sinan B. Sarac, “Update of section 4.8 of the SmPC in order to update the frequency of the adverse drug reaction ‘infection’ from uncommon to common.
In addition, the Marketing authorisation holder (MAH) took the opportunity to correct some discrepancies, as noted by the MHRA, in sections 4.4, 4.5 and 4.8 of the SmPC. The PL is updated accordingly.”

Xeloda - capecitabine - EMEA/H/C/000316/II/0074

MAH: Roche Registration Limited, Rapporteur: Harald Enzmann, “Update of section 4.4 of the SmPC with regards to DPD deficiency genotyping, following a request from the PRAC after assessment of LEG-33.1.”

Zaltrap - aflibercept - EMEA/H/C/002532/II/0044

MAH: sanofi-aventis groupe, Rapporteur: Filip Josephson, “Submission of the final report from study EFC11338 / AFLAME, “A Multinational, Randomized, Double-Blind Study of Aflibercept Versus Placebo with Irinotecan/5-FU Combination (FOLFIRI) in Patients with Metastatic Colorectal Cancer (MCRC) After Failure of an Oxaliplatin Based Regimen"”

WS1258
Clopidogrel Zentiva- EMEA/H/C/000975/WS1258/0059
Clopidogrel/Acetylsalicylic acid Zentiva- EMEA/H/C/001144/WS1258/0050
DuoPlavin- EMEA/H/C/001143/WS1258/0049
Iscover- EMEA/H/C/000175/WS1258/0132
Plavix-EMEA/H/C/000174/WS1258/0129

MAH: Sanofi Clir SNC, Lead Rapporteur: Bruno Sepodes, “Update of section 4.8 of the SmPC in order to add the undesirable effect ‘ageusia’. The labelling is updated accordingly.
In addition, the Worksharing applicant (WSA) took the opportunity to introduce a clarification in section 4.2 of the SmPC; update the German local representative in the Package Leaflet; and bring the PI in line with the latest QRD template version 10.”

WS1273/G
Effentora- EMEA/H/C/000833/WS1273/0047/G

MAH: Teva B.V., Lead Rapporteur: Martina Weise, “Update of sections 4.2 and 4.4 of the SmPC in order to add a warning on Hyperalgesia following an internal cumulative review. The Package Leaflet is updated accordingly.

Update of sections 4.4 and 4.45 of the SmPC in order to add a warning on the interaction of fentanyl with benzodiazepines or other CNS depressants including alcohol following an internal cumulative review. The Package Leaflet

is updated accordingly.

In addition, the MAH took the opportunity to introduce editorial and format changes in the SmPC and PL."

WS1302
Exviera-EMEA/H/C/003837/WS1302/0032
Viekirax-
EMEA/H/C/003839/WS1302/0037
MAH: AbbVie Limited, Lead Rapporteur: Filip Josephson, "Submission of the final report from study (M13-102) listed as a category 3 study in the RMP. This is a phase 3, long-term follow-up study to assess resistance and durability of response to direct-acting antiviral agent (DAA) therapy in subjects who participated in phase 2 or 3 clinical studies for the treatment of chronic hepatitis C virus (HCV) infection."

WS1308/G
Exviera-
EMEA/H/C/003837/WS1308/0033/G
Viekirax-
EMEA/H/C/003839/WS1308/0038/G
MAH: AbbVie Limited, Lead Rapporteur: Filip Josephson, "Update of section 4.8 of the SmPC to add the adverse reactions anaphylactic reactions and erythema multiforme with unknown frequency following a safety review. The package leaflet is updated accordingly."

WS1310
Descovy-
EMEA/H/C/004094/WS1310/0026
Genvoya-
EMEA/H/C/004042/WS1310/0040
Odefsey-
EMEA/H/C/004156/WS1310/0026
Vemlidy-
EMEA/H/C/004169/WS1310/0008
MAH: Gilead Sciences International Limited, Lead Rapporteur: Robert James Hemmings, "Update of section 4.5 of the Descovy, Genvoya, Odefsey and Vemlidy SmPCs in order to include some information on the drug-drug interaction with sofosbuvir/velpatasvir/voxilaprevir fixed dose combination based on the results of study GS-US0367-1657, listed as a category 3 in the Vemlidy RMP, in order to fulfil MEA 006 for Vemlidy. Study GS-US0367 is a phase I multiple dose study to evaluate the drug-drug interaction potential between

sofosbuvir/velpatasvir/voxilaprevir fixed dose combination and HIV anti-retrovirals in healthy subjects. In addition, the Worksharing applicant (WSA) took the opportunity to make some small corrections to section 4.5 of the SmPC for Descovy, Genvoya, Odefsey and Vemlidy and to make corrections to the DE, ES, HU, IS, IT, LV, NO, PT, SL and SV translations for Vemlidy."

WS1316
Glyxambi-
EMEA/H/C/003833/WS1316/0011
Jardiance-
EMEA/H/C/002677/WS1316/0037
Synjardy-
EMEA/H/C/003770/WS1316/0032
MAH: Boehringer Ingelheim International GmbH, Lead Rapporteur: Johann Lodewijk Hillege, "Update of section 5.1 of the SmPC for Jardiance, Synjardy and Glyxambi in order to add clinically relevant information on hear failure and microvascular endpoints based on the results from trial 1245.25 (EMPA-REG OUTCOME study).

The Package Leaflet is updated accordingly.

In addition, the Worksharing applicant (WSA) took the opportunity to update section 4.4 of the SmPC to align the statement regarding diabetic ketoacidosis for all SGLT-2 Inhibitors."

B.6.10. CHMP-PRAC assessed procedures

Amyvid - florbetapir (18F) -
EMEA/H/C/002422/II/0029
MAH: Eli Lilly Nederland B.V., Rapporteur: Harald Enzmann, PRAC Rapporteur: Valerie Strassmann, "Submission of the final report from study I6E-MC-AVBF listed as a category 3 study in the RMP. This is a non-interventional category 3 PASS: European Drug Usage Survey for Amyvid to assess the usage pattern of Amyvid in the EU.
Section 4.4 of SmPC has been reformatted as result of this study.

The RMP version 3.1 has also been submitted."

Benlysta - belimumab -
EMEA/H/C/002015/II/0052
MAH: Glaxo Group Ltd, Rapporteur: Kristina Dunder, PRAC Rapporteur: Ulla Wändel Liminga,

“Submission of the final report from study HGS1006-C1074 (BEL112234) "A Multi-Center, Continuation Trial of Belimumab (HGS1006, LymphoStat-B™), a Fully Human Monoclonal Anti-BLyS Antibody, in Subjects with Systemic Lupus Erythematosus (SLE) who Completed the Phase 3 Protocol HGS1006-C1056 or HGS1006-C1057”. The study is listed as a category 3 study in the RMP (MEA012). The RMP version 26.0 is updated accordingly. In addition the MAH has taken the occasion to update the RMP for the due date for final study report and introduction of protocol changes (reduced study sample size) already discussed and agreed in recent procedure EMEA/H/C/002015/MEA/006.4 and EMEA/H/C/002015/MEA/006.5 for study BEL116027.”	
Eliquis - apixaban - EMEA/H/C/002148/II/0050 MAH: Bristol-Myers Squibb / Pfizer EEIG, Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Menno van der Elst, “Update of sections 4.2 and 5.1 of the SmPC in order to update the posology, method of administration and efficacy and safety information based on final results from study (EMANATE – b0661025/CV185267) listed as a PAES in the RMP; this is a phase 4 study to assess the effectiveness of apixaban compared with usual care anticoagulant in subjects with non-valvular atrial fibrillation (NVAF) undergoing cardioversion; the Package Leaflet is updated accordingly. The RMP version 19 has also been submitted. In addition, the Marketing authorisation holder took the opportunity to update the address of the MAH in the product information.”	
Giotrif - afatinib - EMEA/H/C/002280/II/0025 MAH: Boehringer Ingelheim International GmbH, Rapporteur: Filip Josephson, PRAC Rapporteur: Ulla Wändel Liminga, “Submission of the final report from study (1200.217) listed as a category 3 study in the RMP. This is a phase IV study to assess the efficacy and safety of afatinib as second-line therapy for patients with locally advanced or metastatic non-small cell lung cancer harbouring an EGFR mutation who have failed first-line treatment with platinum-based chemotherapy. In addition, an	

updated RMP (version 6.0) has also been submitted."

Imnovid - pomalidomide - EMEA/H/C/002682/II/0027, Orphan

MAH: Celgene Europe Limited, Rapporteur: Robert James Hemmings, PRAC Rapporteur: Patrick Batty, "Update of sections 4.2, 4.4, and 4.8 of the SmPC in order to add new ADRs SJS, TEN and DRESS following a review of reports on severe skin reactions. The Package Leaflet is updated accordingly. The RMP version 12.0 has also been submitted."

Invokana - canagliflozin - EMEA/H/C/002649/II/0034

MAH: Janssen-Cilag International NV, Rapporteur: Martina Weise, PRAC Rapporteur: Valerie Strassmann, "Update of sections 4.1, 4.4, 4.8 and 5.1 of the SmPC in order to update the safety and efficacy information on cardiovascular events following final results from CANVAS Program (DIA3008 and DIA4003); the Package Leaflet is updated accordingly.
Study DIA3008 is phase 3 Randomized, Multicenter, Double-Blind, Parallel, Placebo-Controlled Study of the Effects of JNJ-28431754 on Cardiovascular Outcomes in Adult Subjects With Type 2 Diabetes Mellitus

Study DIA4004 is phase 4 Randomized, Multicenter, Double-Blind, Parallel, Placebo-Controlled Study of the Effects of Canagliflozin on Renal Endpoints in Adult Subjects With Type 2 Diabetes Mellitus

The RMP version 7.2 has also been submitted."

Keppra - levetiracetam - EMEA/H/C/000277/II/0169/G

MAH: UCB Pharma S.A., Rapporteur: Koenraad Norga, PRAC Rapporteur: Laurence de Fays, "1) C.I.4 (type II): Update of section 4.8 of the SmPC to add the ADR Gait Disturbance, to address the CHMP recommendations from P46/085;
2) C.I.4 (type II): Update of section 4.2 of the SmPC to add Dysgeusia as a potential experience post administration and to section 4.5 of the SmPC to remove drug interaction with Methotrexate, in accordance with the latest Levetiracetam Company Core Data Sheet);

3) C.I.4 (type II): Update of Section 4.6 to add information on 'Women of childbearing potential' and to update the Pregnancy section, to address PRAC recommendations from LEG-084.1;
The Package Leaflet is updated accordingly.
An updated to the Risk Management Plan (version 8) is included to address PRAC recommendations from LEG 84.1."

Maviret - glecaprevir / pibrentasvir - EMEA/H/C/004430/II/0004

MAH: AbbVie Limited, Rapporteur: Joseph Emmerich, PRAC Rapporteur: Ana Sofia Diniz Martins, "Update of sections 4.2, 4.4, 4.5, 4.8 and 5.1 of the SmPC in order to update information on the use of Maviret in liver or kidney transplant patients, based on new clinical data from study M13-596 (MAGELLAN-2), a post-registrational Phase 3 study listed as a category 3 study in the RMP, which evaluated the efficacy and safety of the glecaprevir/pibrentasvir regimen in adult subjects with chronic hepatitis C virus genotypes 1-6 infection, who have received a liver or renal transplant. The Package Leaflet is updated accordingly. The RMP version 3.0 has also been submitted."

Odomzo - sonidegib - EMEA/H/C/002839/II/0016

MAH: Sun Pharmaceutical Industries Europe B.V., Rapporteur: Paula Boudewina van Hennik, PRAC Rapporteur: Patrick Batty, "Update of Annex II to delete the condition "Post-authorisation efficacy study (PAES): The MAH should submit the final CSR for Study CLDE225A2201, including an updated analysis of outcomes by aggressive vs non-aggressive histological subtypes."
Consequentially, the updated RMP version 7.0 was provided in order to reflect the changes following the fulfilment of Annex II condition."

Ofev - nintedanib - EMEA/H/C/003821/II/0018/G, Orphan

MAH: Boehringer Ingelheim International GmbH, Rapporteur: Jayne Crowe, PRAC Rapporteur: Nikica Mirošević Skvrce, "Update of section 4.4 in order to remove the current warning on co-administration with pirfenidone and update of section 5.1 to include the results of study 1199.222, a phase IV, 12 week, open

label, randomised, parallel group study to evaluate the safety, tolerability and PK of oral nintedanib in combination with oral pirfenidone in comparison with nintedanib alone in patients with IPF.
Update of section 5.2 of the SmPC in order to include the results of study 1199.229, a phase IV, open label, multi-dose, 2 groups study to investigate the DDI between nintedanib anfd pirfenidone in patients with IPF, a category 3 study in the RMP.
The RMP version 5.0 has also been submitted.
In addition, the Marketing authorisation holder (MAH) took the opportunity to implement some corrections to the French and Swedish translations."

Raxone - idebenone - EMEA/H/C/003834/II/0008, Orphan
MAH: Santhera Pharmaceuticals (Deutschland) GmbH, Rapporteur: John Joseph Borg, PRAC Rapporteur: Carmela Macchiarulo, "Update of SmPC section 4.5 to include that CYP3A4 substrates known to have a narrow therapeutic index should be administered with caution in patients receiving idebenone, based on the final study report for study SNT-I-017: "An open-label study to assess the potential for pre-systemic inhibition of cytochrome P450 3A4 (CYP3A) by idebenone in healthy male subjects using midazolam as a substrate". The Package Leafelt was updated accordingly. An updated RMP version 1.5 was submitted as part of the application. The provision of the study report addresses the post-authorisation measure MEA 005.1."

ReFacto AF - moroctocog alfa - EMEA/H/C/000232/II/0143
MAH: Pfizer Limited, Rapporteur: Hanne Lomholt Larsen, PRAC Rapporteur: Doris Stenver, "Update of sections 4.2, 4.8, 5.1 and 5.2 of the SmPC in order to update the existing safety, efficacy and pharmacokinetic information based on the final results from studies B1831005 and B1831006 listed as category 3 in the RMP (MEA 111 and 113).
Study B1831005 is a non-randomized, open label study to evaluate the safety, efficacy, and pharmacokinetics (PK) of ReFacto AF in previously treated children less than 12 years of age with severe hemophilia A

(FVIII:C<1%)(already submitted in P46-143).
Study B1831006 is an open-label study on the safety and efficacy of ReFacto AF in previously untreated patients (PUPs) in usual care settings (already submitted in P46-145).
In addition, the PI is brought in line with the latest QRD template (version 10). An editorial change has been made to the Package Leaflets (CZ local representative address).
The updated RMP version 12.0 (new template, revision 2) has been submitted in order to add information regarding the above mentioned studies and from study B1831083 an open-label, single-arm, post-authorization pragmatic clinical trial on the safety and efficacy of Xyntha in subjects with Hemophilia A in usual care settings in China, listed as category 3 in the RMP and already submitted as P46-144."

Remicade - infliximab - EMEA/H/C/000240/II/0209

MAH: Janssen Biologics B.V., Rapporteur: Kristina Dunder, PRAC Rapporteur: Ulla Wändel Liminga, "Update of the current warning on colon cancer and dysplasia of Section 4.4 of the SmPC based on final report of the OPUS Registry (Prospective, Observational, Non-Interventional, Post-marketing Safety Surveillance Program in Subjects with UC; P04808) as per MEA 121.
In addition, the MAH is taking the opportunity to add a warning on screening tests for tuberculosis to align it with current medical practice, add a reminder on the patient alert card in package leaflet and include some editorial changes in line with the QRD template."

RoActemra - tocilizumab - EMEA/H/C/000955/II/0074/G

MAH: Roche Registration Limited, Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Brigitte Keller-Stanislawski

Stribild - elvitegravir / cobicistat / emtricitabine / tenofovir disoproxil - EMEA/H/C/002574/II/0087

MAH: Gilead Sciences International Limited, Rapporteur: Robert James Hemmings, PRAC Rapporteur: Julie Williams, "Submission of the final study report for Study GS-EU-236-0141, listed as a category 3 study in the Risk

Management Plan, in order to fulfil a post-authorisation measure (PAM) MEA 006 for Stribild; This study is an Observational Drug Utilization Study of Stribild in Adults with HIV-1 Infection.

With this application and as agreed with the EMA, Gilead is also taking this opportunity to address the outstanding questions from MEA 002.3."

Symtuza - darunavir / cobicistat / emtricitabine / tenofovir alafenamide - EMEA/H/C/004391/II/0002/G
MAH: Janssen-Cilag International N.V., Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Julie Williams, "C.I.13-II to submit the results of the study GS-US-311-1089 "A Phase 3, Randomized, Double-Blind, Switch Study to Evaluate F/TAF in HIV 1 Positive Subjects who are Virologically Suppressed on Regimens containing FTC/TDF". The RMP has been updated to reflect the completion of the study.

C.I.11.z- II to update the RMP to remove pancreatitis, convulsion, and cardiac conduction abnormalities as risks in the RMP in alignment with the RMP for Prezista and Rezolsta."

Symtuza - darunavir / cobicistat / emtricitabine / tenofovir alafenamide - EMEA/H/C/004391/II/0003/G
MAH: Janssen-Cilag International N.V., Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Julie Williams, "Update of sections 4.4, 4.8, 5.1 and 5.2 of the SmPC in order to reflect the week-48 results from 2 studies (TMC114FD2HTX3001 and TMC114IFD3013) listed as category 3 studies in the RMP; these are phase 3 studies to evaluate the efficacy and safety of D/C/F/TAF once daily fixed-dose combination regimen versus a regimen consisting of DRV/COBI FDC co-administered with FTC/TDF FDC in ARV treatment-naïve HIV-1 infected subjects (study TMC114FD2HTX3001) and to evaluate switching to a D/C/F/TAF once-daily single-tablet regimen versus continuing the current regimen consisting of a boosted protease inhibitor combined with FTC/TDF in virologically-suppressed, HIV-1 infected subjects (study TMC114IFD3013). The RMP version 2.0 has also been submitted. In

addition, the MAH took the opportunity to update the list of local representatives in the Package Leaflet and to make minor editorial revision in the product information."

Tyverb - lapatinib - EMEA/H/C/000795/II/0050/G

MAH: Novartis Europharm Limited, Rapporteur: Filip Josephson, PRAC Rapporteur: Ulla Wändel Liminga, "1. Type II- C.I.13: Submission of the final non-clinical study report 09DMR047 listed as a category 3 study in the RMP. This is a non-clinical mechanistic study related to lapatinib metabolite identification in dog plasma, bile and liver. An updated RMP (version 33) is included to reflect the completion of a dog study and integration of the results.
2. Type IB- C.I.11.Z: Change to the final due date of study EGF117165 to evaluate biomarkers of drug resistance in patients with HER2+ metastatic breast cancer whilst on treatment with trastuzumab in combination with either lapatinib or chemotherapy (category 1, ANX034.2) from Jun-2018 to Jun-2019 in the RMP and Annex II.
In addition, the MAH took the opportunity to implement the recent PRAC PSUR recommendation into the RMP version 33, including the removal of two identified risks (rash, diarrhoea) and update of missing information wording (hepatic impairment and renal impairment)."

Vaxelis - diphtheria, tetanus, pertussis (acellular, component), hepatitis B (rDNA), poliomyelitis (inact.) and Haemophilus type B conjugate vaccine (adsorbed) - EMEA/H/C/003982/II/0021

MAH: MCM Vaccine B.V., Rapporteur: Bart Van der Schueren, PRAC Rapporteur: Brigitte Keller-Stanislawski, "Update of section 5.1 of the SmPC in order to update the efficacy section on immune persistence based on the final results from study PRI03C - Long-term Persistence of Hepatitis B and Pertussis Antibody Responses in Healthy 4- to 5-year-old Children Previously Vaccinated with a 2 dose or 3 dose Infants Series and Toddler dose of Vaxelis or INFANRIX hexa listed as P46 study in the PIP.

The RMP version 2.2 has also been submitted.

In addition, the MAH took the opportunity to

introduce editorial changes in Annex IIIa”

Vokanamet - canagliflozin / metformin - EMEA/H/C/002656/II/0034

MAH: Janssen-Cilag International NV, Rapporteur: Martina Weise, PRAC Rapporteur: Menno van der Elst, “Update of sections 4.1, 4.4, 4.8 and 5.1 of the SmPC in order to update the safety and efficacy information on cardiovascular events following final results from CANVAS Program (DIA3008 and DIA4003); the Package Leaflet is updated accordingly.
Study DIA3008 is phase 3 Randomized, Multicenter, Double-Blind, Parallel, Placebo-Controlled Study of the Effects of JNJ-28431754 on Cardiovascular Outcomes in Adult Subjects With Type 2 Diabetes Mellitus

Study DIA4004 is phase 4 Randomized, Multicenter, Double-Blind, Parallel, Placebo-Controlled Study of the Effects of Canagliflozin on Renal Endpoints in Adult Subjects With Type 2 Diabetes Mellitus

The RMP version 7.2 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 andto bring the PI in line with the latest QRD template version 10.”

B.6.11. PRAC assessed procedures

PRAC Led

Advate - octocog alfa - EMEA/H/C/000520/II/0089

MAH: Baxter AG, PRAC Rapporteur: Brigitte Keller-Stanislawski, PRAC-CHMP liaison: Jan Mueller-Berghaus, “Submission of the final report from study 061501. This was a retrospective chart review aimed to evaluate safety and tolerability of Advate among previously untreated patients with moderate to severe Haemophilia A.”

PRAC Led

Baraclude - entecavir - EMEA/H/C/000623/II/0053

MAH: Bristol-Myers Squibb Pharma EEIG, Rapporteur: Filip Josephson, PRAC Rapporteur: Qun-Ying Yue, PRAC-CHMP liaison: Filip

Josephson, “Submission of the final study report for Study AI463080, a long-term outcomes study (10 years), to assess the rates of malignant neoplasm (all, non-HCC, and HCC), liver-related events of HBV disease progression, and mortality. Along with this type II variation a consequential updated Risk Management Plan (Version 14) has been submitted.”

PRAC Led

Fiasp - insulin aspart -

EMEA/H/C/004046/II/0003/G

MAH: Novo Nordisk A/S, Rapporteur: Kristina Dunder, PRAC Rapporteur: Julie Williams, PRAC-CHMP liaison: Greg Markey, “Update of the RMP to upgrade the risk of mix-up between basal and bolus insulin from a potential to an important identified risk (RMP version 2.0). In addition, to update the secondary packaging material (carton, Label, IFU) design and change colour of selected plastic components from yellow to red.
Also the MAH submitted as part of this variation a proposal for communication to Health Care Professionals and Patients (indirectly) regarding similarity of Fiasp and Tresiba products that are currently on the market.”

PRAC Led

Mycamine - micafungin -

EMEA/H/C/000734/II/0035

MAH: Astellas Pharma Europe B.V., Rapporteur: Harald Enzmann, PRAC Rapporteur: Martin Huber, PRAC-CHMP liaison: Harald Enzmann, “Submission of the final survey report regarding Educational tools in the RMP and Educational tools as a LEG (39) and updated RMP version 18.0.”

PRAC Led

Orencia - abatacept -

EMEA/H/C/000701/II/0116/G

MAH: Bristol-Myers Squibb Pharma EEIG, Rapporteur: Outi Mäki-Ikola, PRAC Rapporteur: Kirsti Villikka, PRAC-CHMP liaison: Outi Mäki-Ikola, “Grouping of two Type II variations, as follows:

C.I.13: Submission of the final report from study IM101537 listed as a category 3 study in the RMP. This is a non-interventional HCP/patient cross-sectional survey and retrospective chart review Post Authorisation

Safety Study to evaluate the effectiveness of the Patient Alert Card for both IV and SC abatacept in a sample of EU countries.

C.I.11: Submission of an updated RMP (version 24) in order to reflect the early closure of another RMP category 3 study: Study IM101212, which closed in September 2017 and for which no further data will be available. A number of other administrative updates to the RMP are being carried out in the context of this procedure."

PRAC Led

ReFacto AF - moroctocog alfa - EMEA/H/C/000232/II/0142

MAH: Pfizer Limited, PRAC Rapporteur: Doris Stenver, "Submission of the final study report from study B1831016, listed as a category 3 in the RMP (MEA 108.3). This is a non-interventional open-label study conducted at haemophilia treatment centres in Germany and Austria to generate information regarding the safety and effectiveness of treatment with ReFacto AF under routine clinical conditions."

PRAC Led

Scenesse - afamelanotide - EMEA/H/C/002548/II/0018, Orphan

MAH: Clinuvel (UK) Limited, Rapporteur: TBA, PRAC Rapporteur: Valerie Strassmann, PRAC-CHMP liaison: Harald Enzmann, "Submission of an updated RMP version 8.0 which aims to address the comments made in procedure IB/14 and including:

- Updates from pre-approval information to post-marketing information
- Update of number of patients treated in clinical trials, special access schemes and commercial distribution
- Change in development of the custom-made device
- Postponement of pharmacokinetic study CUV052 (no timeframe yet)
- Update on timelines for safety extension study CUV037 from Q12013 to Q12018
- Update on timelines for on-going and planned PV studies
- key elements of educational and training programme (annex 10)
- Correction: replacement of pigmentary lesions by pigmentary expressions

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- General update of safety information”

PRAC Led

Synflorix - pneumococcal polysaccharide conjugate vaccine (adsorbed) - EMEA/H/C/000973/II/0124/G

MAH: GlaxoSmithkline Biologicals SA, Rapporteur: Kristina Dunder, PRAC Rapporteur: Qun-Ying Yue, PRAC-CHMP liaison: Kristina Dunder, “Submission of the final study reports from two 5-year Invasive Pneumococcal Disease (IPD) post-marketing surveillance (PMS) studies “Monitoring the Population Effectiveness of Pneumococcal Conjugate Vaccination in the Finnish National Vaccination Programme” (MEA 019) and “Epidemiology of invasive pneumococcal disease in the Netherlands” (MEA 020), addressing the potential risks of "possible serotype replacement of disease isolates" and "possible breakthrough infections/vaccine failure". The MAH also submitted data from IPD surveillance in 5 other European countries (Austria, Bulgaria, Cyprus, Iceland and Sweden) and 6-year update results from a 5-year PMS in Kenya (Pneumococcal Conjugate Vaccine Impact Study (PCVIS), MEA 021). Submission of an updated RMP version 17 to reflect data from the PMS studies, close MEA 019 and MEA 020, extend MEA 021 and implement the latest RMP template (revision 2). No changes to the Product Information are proposed with this submission.”

PRAC Led

Tecfidera - dimethyl fumarate - EMEA/H/C/002601/II/0049

MAH: Biogen Idec Ltd, Rapporteur: Martina Weise, PRAC Rapporteur: Martin Huber, PRAC-CHMP liaison: Martina Weise, “Submission of the final report from study 109MS409 listed as a category 3 study in the RMP. This is an observation study aimed to estimate the proportion of dimethyl fumarate use that is prescribed “on-label” versus “off-label” in Germany.”

PRAC Led

WS1283

Relvar Ellipta- EMEA/H/C/002673/WS1283/0035

Revinty Ellipta- EMEA/H/C/002745/WS1283/0031

MAH: Glaxo Group Ltd, Lead Rapporteur: Concepcion Prieto Yerro, Lead PRAC Rapporteur: Dolores Montero Corominas, PRAC-CHMP liaison: Concepcion Prieto Yerro, “Submission of the final report from study 205052 (PRJ2214). This is a drug utilization study to identify the extent of any off-label prescribing fluticasone furoate/vilanterol FF/VI in any dose in children less than 12 years of age; and prescribing of FF/VI 200/25 mcg in patients with a diagnosis of chronic obstructive pulmonary disease COPD, considering the presence of a concurrent diagnosis of asthma. The RMP version 9.1 has been updated accordingly.”

PRAC Led

WS1293

Exelon-EMEA/H/C/000169/WS1293/0115

Prometax-EMEA/H/C/000255/WS1293/0115

MAH: Novartis Europharm Limited, Lead Rapporteur: Alexandre Moreau, Lead PRAC Rapporteur: Ghania Chamouni, PRAC-CHMP liaison: Alexandre Moreau, “To update the Rivastigmine RMP with:

- Milestone changes for Drug Utilization Study (DUS) (ENA713D2409) - Based on PRAC Assessment Report (AR) (EMEA/H/C/000169/MEA 034.2 & EMEA/H/C/000255/MEA 035.1) Protocol Amendment version 2, as finalized.

- Removal of important identified risk "pancreatitis" - Based on PRAC PSUR 23 (01-Feb-2014 to 31-Jan-2015) AR (EMEA/H/C/PSUSA/00002654/201501)

- Discontinue the use of the targeted checklist to document cases of medication error/misuse - Based on PRAC PSUR 24 (01-Feb-2015 to 31-Jan-2016) AR (EMEA/H/C/PSUSA/00002654/201601)

- Change of 6 monthly report on “the effectiveness of risk minimization measures for multiple patch use" to annual report - Based on the AR of the fourth 6 monthly report on “the effectiveness of risk minimization measures for multiple patch use.”

The following activities which occurred after the DLP of 31-Jan-16 are also included in the Rivastigmine RMP update: information about the

submission of an interim analysis report for DUS ENA713D2409 dated 10-Mar-2016 to PRAC, information about distribution of a health care professional (HCP) letter in Japan, information about the request from the Brazilian health authority to include a statement in local Exelon patch leaflet to minimize the potential risk of skin irritation, information that the Exelon/Prometax CDS was amended on 04 -Mar -2016 to include "nightmares" as an ADR. Furthermore e.g. updates of RMP Parts and RMP Annexes to align with the status provided in PSUR 24 (DLP 31-Jan-16) are included."

PRAC Led

WS1294

Actos-EMEA/H/C/000285/WS1294/0078

Competact-

EMEA/H/C/000655/WS1294/0068

Glubrava-

EMEA/H/C/000893/WS1294/0055

Glustin-EMEA/H/C/000286/WS1294/0077

Tandemact-

EMEA/H/C/000680/WS1294/0056

MAH: Takeda Pharma A/S, Informed Consent of Competact, Lead Rapporteur: TBA, Lead PRAC Rapporteur: Almath Spooner, PRAC-CHMP liaison: Peter Kiely, "To update the RMPs to version 24.0 for Actos & Glustin (pioglitazone), version 22.0 for Tandemact (pioglitazone/ glimepiride), and version 25.0 for Compeact & Glubrava (pioglitazone/metformin).

These RMPs have been updated as follows: For all RMPs:
Added bone mechanistic (AD4833-402) addendum information per EMA Post-authorization measure (PAM) Report dated 17-Oct-2016: Actos H/C/285 (MEA 68.2), Glustin H/C/286 (MEA 70.2); Competact H/C/655 (MEA 29.2); Glubrava H/C/893 (MEA 18.2); Tandemact H/C/680 (MEA 27.2).
The MAH also took the opportunity to make some general edits to these RMPs. For pioglitazone/metformin RMP only:
Added lactic acidosis questionnaire per assessor's comment to the final report for Type II Variation EMEA/H/C/000655/WS0991/0062 & EMEA/H/C/000893/WS0991/0047."

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

WS1269
Rotarix-EMEA/H/C/000639/WS1269/0103
MAH: GlaxoSmithKline Biologicals S.A., Lead Rapporteur: Bart Van der Schueren

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WS1272
Epclusa-
EMEA/H/C/004210/WS1272/0020
Vosevi-EMEA/H/C/004350/WS1272/0007
MAH: Gilead Sciences International Limited, Lead Rapporteur: Filip Josephson

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WS1305
Descovy-
EMEA/H/C/004094/WS1305/0024
Genvoya-
EMEA/H/C/004042/WS1305/0039
Odefsey-
EMEA/H/C/004156/WS1305/0023
Vemlidy-
EMEA/H/C/004169/WS1305/0007
MAH: Gilead Sciences International Limited, Lead Rapporteur: Robert James Hemmings

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WS1313/G
Amgevita-
EMEA/H/C/004212/WS1313/0003/G
Solymbic-
EMEA/H/C/004373/WS1313/0003/G
MAH: Amgen Europe B.V., Lead Rapporteur: Kristina Dunder

WS1319/G
Helixate NexGen-
EMEA/H/C/000276/WS1319/0195/G
Kogenate Bayer-
EMEA/H/C/000275/WS1319/0203/G
MAH: Bayer AG, Lead Rapporteur: Jan Mueller-Berghaus

WS1321
Exelon-EMEA/H/C/000169/WS1321/0116
Prometax-
EMEA/H/C/000255/WS1321/0116

MAH: Novartis Europharm Limited, Lead Rapporteur: Alexandre Moreau

WS1323
Aflunov-
EMEA/H/C/002094/WS1323/0041
Foclivia-
EMEA/H/C/001208/WS1323/0035
MAH: Seqirus S.r.l, Lead Rapporteur: Daniela Melchiorri

WS1325
M-M-Rvaxpro-
EMEA/H/C/000604/WS1325/0086
ProQuad-
EMEA/H/C/000622/WS1325/0122
MAH: MSD Vaccins, Lead Rapporteur: Jan Mueller-Berghaus

WS1331
Ariclaim-
EMEA/H/C/000552/WS1331/0070
Cymbalta-
EMEA/H/C/000572/WS1331/0074
Duloxetine Lilly-
EMEA/H/C/004000/WS1331/0010
Xeristar-
EMEA/H/C/000573/WS1331/0077
MAH: Eli Lilly Nederland B.V., Duplicate, Duplicate of Ariclaim, Yentreve, Lead Rapporteur: Concepcion Prieto Yerro

Hexacima-
EMEA/H/C/002702/WS1286/0075
Hexaxim-
EMEA/H/W/002495/WS1286/0080
Hexyon-
EMEA/H/C/002796/WS1286/0079
MAH: Sanofi Pasteur SA, Lead Rapporteur: Jan Mueller-Berghaus

Hexacima-
EMEA/H/C/002702/WS1306/0074
Hexaxim-
EMEA/H/W/002495/WS1306/0079
Hexyon-
EMEA/H/C/002796/WS1306/0078
MAH: Sanofi Pasteur SA, Lead Rapporteur: Jan Mueller-Berghaus

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.

Qualification of Biomarkers:

HTA:

G.2. PRIME

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

G.2.1. List of procedures concluding at 11-14 December 2017 CHMP plenary:

G.2.2. List of procedures starting in December 2017 for January 2018 CHMP adoption of outcomes

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