17 July 2017
EMA/CHMP/451869/2017
Inspections, Human Medicines Pharmacovigilance and Committees Division

Committee for medicinal products for human use (CHMP)

Draft agenda for the meeting on 17-20 July 2017

Chair: Tomas Salmonson – Vice-Chair: Harald Enzmann

17 July 2017, 13:00 – 21:00, room 2A

18 July 2017, 08:30 – 21:00, room 2A

19 July 2017, 08:30 – 20:00, room 2A

20 July 2017, 08:30 – 16:00, room 2A

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 17-20 July 2017. See July 2017 CHMP minutes (to be published post August 2017 CHMP meeting).

1.2. Adoption of agenda

CHMP agenda for 17-20 July 2017

1.3. Adoption of the minutes

CHMP minutes for 19-22 June 2017

2. Oral Explanations

2.1. Pre-authorisation procedure oral explanations

2.1.1. etirinotecan pegol - EMEA/H/C/003874

treatment of breast cancer with brain metastases

Scope: Oral explanation/Opinion

Action: Oral explanation to be held on 19 July 2017 at time 09:00

Revised list of experts for the ad hoc expert group meeting adopted via written procedure on 11 July 2017.

Oral explanation 16.05.2017, List of Outstanding Issues adopted on 23.03.2017. List of Questions adopted on 10.11.2016.

See 3.1

2.1.2. midostaurin - Orphan - EMEA/H/C/004095

Novartis Europharm Ltd; treatment of mastocytosis and treatment of acute myeloid leukaemia

Scope: Oral explanation

Action: Oral explanation to be held on 19 July 2017 at time 11:00

List of Outstanding Issues adopted on 22.06.2017, 21.04.2017. List of Questions adopted on 15.12.2016.

See 3.1

2.1.3. padeliporfin - EMEA/H/C/004182

treatment of prostate cancer

Scope: Oral explanation

Action: Oral explanation to be held on 18 July 2017 at time 11:00

Oral explanation 20.04.2017. List of Outstanding Issues adopted on 21.04.2017, 15.12.2016. List of Questions adopted on 26.05.2016.

2.2. Re-examination procedure oral explanations

2.3. Post-authorisation procedure oral explanations

2.3.1. Opdivo - nivolumab - EMEA/H/C/003985/II/0029

Bristol-Myers Squibb Pharma EEIG

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

Scope: Oral explanation

Action: Oral explanation to be held on 19 July 2017 at time 14:00

“Extension of Indication to include the treatment of hepatocellular carcinoma after prior sorafenib therapy in adults for Opdivo.
As a consequence, sections 4.1, 4.2, 4.4, 4.8 and 5.1 of the SmPC are updated. The Package Leaflet is updated in accordance.
Moreover, the updated RMP version 8.0 has been submitted.”

Request for Supplementary Information adopted on 22.06.2017, 23.03.2017.

See 5.1

2.4. Referral procedure oral explanations

2.4.1. Gadolinium-containing contrast agents (GdCA): gadoversetamide – OPTIMARK (CAP)
Gadobenate dimeglumine; gadobutrol; gadodiamide; gadopentetic acid dimeglumine, gadoteric acid (intra articular formulation); gadoteric acid (intrvenous and intravascular formulations); gadoteridol; gadoxetic acid disodium (NAP)

Applicant(s): Mallinckrodt Deutschland GmbH (Optimark); various

Rapporteurs for the Article 31 referral: PRAC Rapporteur: Ulla Wändel Liminga; PRAC Co-rapporteur: Valerie Strassmann

Rapporteurs for Optimark: CHMP Rapporteur: Patrick Salmon, CHMP Co-rapporteur: Johann Lodewijk Hillege

Scope: Oral explanation

Action: Oral explanation to be held on 18 July 2017 at time 14:00

Re-examination procedure under Article 32 of Directive 2001/83/EC of the review of the benefit-risk balance of GdCA following notification by the European Commission of a referral under Article 31 of Directive 2001/83/EC, based on pharmacovigilance data

See 10.6

3. Initial applications

3.1. Initial applications; Opinions

3.1.1. avelumab - Orphan - EMEA/H/C/004338

Merck Serono Europe Limited; treatment of Merkel cell carcinoma (MCC)

Scope: Opinion

Action: For adoption

List of Outstanding Issues adopted on 22.06.2017. List of Questions adopted on 23.02.2017.

3.1.2. dupilumab - EMEA/H/C/004390

treatment of moderate-to-severe atopic dermatitis

Scope: Opinion

Action: For adoption

List of Questions adopted on 23.03.2017.

3.1.3. entecavir - EMEA/H/C/004458

treatment of chronic hepatitis B virus infection

Scope: Opinion

Action: For adoption

List of Outstanding Issues adopted on 22.06.2017. List of Questions adopted on 15.12.2016.

3.1.4. entecavir - EMEA/H/C/004377

treatment of chronic hepatitis B virus infection

Scope: Opinion

Action: For adoption

List of Outstanding Issues adopted on 18.05.2017. List of Questions adopted on 15.12.2016.

3.1.5. iloperidone - EMEA/H/C/004149

treatment of schizophrenia

Scope: Opinion

Action: For adoption

Oral explanation 17.05.2016, List of Outstanding Issues adopted on 18.05.2017, 23.02.2017. List of Questions adopted on 28.04.2016.

3.1.6. lacosamide - EMEA/H/C/004443

treatment of epilepsy

Scope: Opinion

Action: For adoption

List of Outstanding Issues adopted on 22.06.2017, List of Questions adopted on 26.01.2017.

3.1.7. lutetium (177lu) oxodotreotide - Orphan - EMEA/H/C/004123

Advanced Accelerator Applications; treatment of gastro-entero-pancreatic neuroendocrine tumours

Scope: Opinion

Action: For adoption

List of Outstanding Issues adopted on 18.05.2017. List of Questions adopted on 15.09.2016.

3.1.8. etirinotecan pegol - EMEA/H/C/003874

treatment of breast cancer with brain metastases

Scope: Opinion/Oral explanation

Action: For adoption

Revised list of experts for the ad hoc expert group meeting adopted via written procedure on 11 July 2017.

Oral explanation 16.05.2017, List of Outstanding Issues adopted on 18.05.2017, 23.03.2017. List of Questions adopted on 10.11.2016.

See 2.1

3.1.9. midostaurin - Orphan - EMEA/H/C/004095

Novartis Europharm Ltd; treatment of mastocytosis and treatment of acute myeloid leukaemia

Scope: Oral explanation/Opinion

Action: For adoption

List of Outstanding Issues adopted on 22.06.2017, 21.04.2017. List of Questions adopted on 15.12.2016.

See 2.1

3.1.10. darunavir / cobicistat / emtricitabine / tenofovir alafenamide - EMEA/H/C/004391

treatment of human immunodeficiency virus type 1 (HIV-1)

Scope: Opinion

Action: For adoption

List of Outstanding Issues adopted on 18.05.2017. List of Questions adopted on 26.01.2017.

3.1.11. atezolizumab - EMEA/H/C/004143

treatment of locally advanced or metastatic urothelial carcinoma,
treatment of non-small cell lung carcinoma (NSCLC)

Scope: Opinion

Action: For adoption

List of Outstanding Issues adopted on 21.04.2017. List of Questions adopted on 15.09.2016.

3.1.12. ciclosporin - Orphan - EMEA/H/C/004411

Accelerated assessment

Santen Oy; treatment of severe vernal keratoconjunctivitis (VKC)

Scope: Opinion

Action: For adoption

List of Outstanding Issues adopted on 22.06.2017, List of Questions adopted on 19.04.2017.

3.1.13. telotristat ethyl - Orphan - EMEA/H/C/003937

Ipsen Pharma; treatment of carcinoid syndrome

Scope: Opinion

Action: For adoption

List of Outstanding Issues adopted on 21.04.2017. List of Questions adopted on 10.11.2016.

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

3.2.1. abaloparatide - EMEA/H/C/004157

treatment of osteoporosis

Scope: Day 180 list of outstanding issue

Action: For adoption

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

3.2.2. carmustine - EMEA/H/C/004326

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

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 13.10.2016.

3.2.3. adalimumab - EMEA/H/C/004319

treatment of rheumatoid arthritis, axial spondyloarthritis, psoriasis, hidradenitis suppurativa (HS), Crohn's disease, ulcerative colitis and uveitis

Scope: Day 180 list of outstanding issue

Action: For adoption

3.2.4. List of Questions adopted on 23.03.2017. dupilumab - EMEA/H/C/004390

treatment of moderate-to-severe atopic dermatitis

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 23.03.2017.

3.2.5. fluticasone furoate / umeclidinium / vilanterol - EMEA/H/C/004781

treatment of adult patients with chronic obstructive pulmonary disease (COPD)

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 21.04.2017.

3.2.6. guselkumab - EMEA/H/C/004271

treatment of plaque psoriasis

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 21.04.2017.

3.2.7. neratinib - EMEA/H/C/004030

extended adjuvant treatment of adult patients with early-stage HER2-overexpressed/amplified breast cancer who have received prior adjuvant trastuzumab based therapy

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 15.12.2016.

3.2.8. naloxone - EMEA/H/C/004325

intended for emergency use for known or suspected opioid overdose as manifested by respiratory and/or central nervous system depression

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 23.03.2017.

3.2.9. trastuzumab - EMEA/H/C/004323

treatment of breast cancer and metastatic gastric cancer

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 26.01.2017.

3.2.10. ritonavir - EMEA/H/C/004549

treatment of HIV-1

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 21.04.2017.

3.2.11. tacrolimus - EMEA/H/C/004435

prophylaxis of transplant rejection and treatment of allograft rejection

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 21.04.2017.

3.2.12. fluticasone furoate / umeclidinium / vilanterol - EMEA/H/C/004363

treatment of adult patients with chronic obstructive pulmonary disease (COPD)

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 21.04.2017.

3.2.13. buprenorphine / naloxone - EMEA/H/C/004407

treatment for opioid drug dependence, treatment for opioid drug dependence

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 23.02.2017.

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

3.3.1. tildrakizumab - EMEA/H/C/004514

treatment of adults with moderate-to-severe plaque psoriasis

Scope: Day 120 list of questions

Action: For adoption

3.3.2. trastuzumab - EMEA/H/C/004361

treatment of metastatic breast cancer, early breast cancer, metastatic gastric cancer

Scope: Day 120 list of questions

Action: For adoption

3.3.3. letermovir - Orphan - EMEA/H/C/004536

Accelerated assessment

Merck Sharp & Dohme Limited; prophylaxis of cytomegalovirus (CMV) reactivation and disease

Scope: Day 120 list of questions

Action: For adoption

3.3.4. naldemedine - EMEA/H/C/004256

treatment of opioid-induced constipation (OIC) in adult patients

Scope: Day 120 list of questions

Action: For adoption

3.3.5. brexpiprazole - EMEA/H/C/003841

treatment of schizophrenia

Scope: Day 120 list of questions

Action: For adoption

3.3.6. sufentanil - EMEA/H/C/004335

management of acute moderate to severe pain

Scope: Day 120 list of questions

Action: For adoption

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

3.4.1. dapivirine - EMEA/H/W/002168, Article 58

reducing the risk of HIV-1 infection via vaginal intercourse in sexually active HIV-uninfected women

Scope: Start of the procedure

Action: For information

3.4.2. pegfilgrastim - EMEA/H/C/004262

treatment of neutropenia

Scope: Letter from the applicant dated 12 July 2017 requesting an additional extension of clock stop to respond to the List of Questions adopted on 13 October 2016.

Action: For adoption

List of Questions adopted on 13.10.2016.

3.4.3. andexanet alfa - EMEA/H/C/004108

treatment of direct or indirect factor Xa(FXa) inhibitor when reversal of anticoagulation is needed

Scope: Letter from the applicant dated 7 July 2017 requesting an extension of clock stop to respond to the List of Questions adopted on 15 December 2016

Action: For adoption

List of Questions adopted on 15.12.2016.

3.4.4. trastuzumab - EMEA/H/C/004346

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

Scope: Letter from the applicant dated 12 July 2017 requesting an additional extension of clock stop to respond to the List of Outstanding Issues adopted on 18.05.2017.

Action: For adoption

List of Outstanding Issues adopted on 18.05.2017. List of Questions adopted on 15.12.2016.

3.4.5. rotigotine - EMEA/H/C/004286

treatment of idiopathic Restless Legs Syndrome and Parkinson's disease

Scope: Letter from the applicant dated 11 July 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

3.5.1. Masipro - masitinib - Orphan - EMEA/H/C/004159

AB Science; treatment of mastocytosis

Scope: draft SAG list of questions

Letter from the applicant dated 31 May 2017 requesting a re-examination of the Opinion adopted on 18 May 2017 and consultation of a Scientific Advisory Group

Action: For discussion

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

Opinion adopted on 18.05.2017. Oral explanation was held on 20.04.2017. List of Outstanding Issues adopted on 23.02.2017. List of Questions adopted on 15.09.2016.

3.6. Initial applications in the decision-making phase

3.7. Withdrawals of initial marketing authorisation application

3.7.1. fluocinolone acetonide - Orphan - EMEA/H/C/004540

Campharm Limited; treatment and prevention of recurrences of non-infectious uveitis

Scope: Withdrawal of initial marketing authorisation application

Action: For information

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

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

4.1.1. Prolia - denosumab - EMEA/H/C/001120/X/0059/G

Amgen Europe B.V.

Rapporteur: Kristina Dunder, PRAC Rapporteur: Ulla Wändel Liminga

Scope: "Extension application."

Action: For adoption

List of Outstanding Issues adopted on 18.05.2017. List of Questions adopted on 26.01.2017.

4.1.2. Samsca - tolvaptan - EMEA/H/C/000980/X/0024

Otsuka Pharmaceutical Europe Ltd

Rapporteur: Greg Markey, Co-Rapporteur: Daniela Melchiorri, PRAC Rapporteur: Julie Williams

Scope: "Extension application to add a new strength of 7.5 mg tablets."

Action: For adoption

List of Questions adopted on 21.04.2017.

4.1.3. Signifor - pasireotide - Orphan - EMEA/H/C/002052/X/0030/G

Novartis Europharm Ltd

Rapporteur: Kristina Dunder, PRAC Rapporteur: Qun-Ying Yue

Scope: “Extension application to introduce two new strengths of the ‘powder and solvent for suspension for injection pharmaceutical form’ (10 mg and 30 mg) grouped with a type II variation (C.I.6.a) to extend the indication to include ‘Treatment of adult patients with Cushing’s disease for whom surgery is not an option or for whom surgery has failed’ to the intramuscular injection formulations.”

Action: For adoption

List of Questions adopted on 21.04.2017.

4.1.4. Xgeva - denosumab - EMEA/H/C/002173/X/0048/G

Amgen Europe B.V.

Rapporteur: Kristina Dunder, PRAC Rapporteur: Ulla Wändel Liminga

Scope: “Extension application.”

Action: For adoption

List of Outstanding Issues adopted on 18.05.2017. List of Questions adopted on 26.01.2017.

4.1.5. Xtandi - enzalutamide - EMEA/H/C/002639/X/0029

Astellas Pharma Europe B.V.

Rapporteur: Jorge Camarero Jiménez, PRAC Rapporteur: Eva A. Segovia

Scope: “To add new pharmaceutical form and strenghts (film-coated tablets 40 mg and 80 mg) to the currently approved presentations for Xtandi.”

Action: For adoption

List of Questions adopted on 21.07.2016.

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. Tasigna - nilotinib - Orphan - EMEA/H/C/000798/X/0088/G

Novartis Europharm Ltd

Rapporteur: Sinan B. Sarac, Co-Rapporteur: Harald Enzmann, PRAC Rapporteur: Doris Stenver

Scope: “Extension of Indication to include treatment of paediatric patients with newly diagnosed Philadelphia chromosome-positive chronic myelogenous leukemia in chronic phase (Ph+ CML-CP), or with Ph+ CML-CP resistant or intolerant to prior therapy including imatinib, based on results from two clinical studies in paediatric patients conducted in accordance with the approved Tasigna Paediatric Investigation Plan (PIP), the Phase I PK study CAMN107A2120 and the Phase II safety and efficacy study CAMN107A2203. An updated RMP version 18.0 was provided as part of the application.
Extension application to add a new strength of 50mg hard capsules.

In addition, the applicant proposes to merge the SmPCs for the 50 mg and 200 mg strengths."

Action: For adoption

List of Questions adopted on 23.03.2017.

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. Bortezomib Accord - bortezomib - EMEA/H/C/003984/X/0008

Accord Healthcare Ltd

Rapporteur: Milena Stain, PRAC Rapporteur: Carmela Macchiarulo

Scope: "Extension application to add a new strength of powder for solution for injection (1 mg) to the currently approved strength (3.5 mg) of Bortezomib Accord."

Action: For adoption

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

AstraZeneca AB

Rapporteur: Concepcion Prieto Yerro, Co-Rapporteur: David Lyons, 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

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

AstraZeneca AB

Rapporteur: Concepcion Prieto Yerro, Co-Rapporteur: David Lyons, 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

4.3.4. Humira - adalimumab - EMEA/H/C/000481/X/0164/G

AbbVie Limited.

Rapporteur: Kristina Dunder

Scope: "Extension application to add a new strength/potency of 20 mg for adalimumab solution for injection in pre-filled syringe, grouped with a type II variation (C.I.4.z) to update of sections 4.2 of the SmPC in order to introduce new fixed dose regimen (posology)

for the paediatric indications of JIA and Ps. The Package Leaflet and Labelling are updated accordingly.

In addition, the marketing authorisation holder took the opportunity to:
- introduce editorial changes to align wording and layout of the Product Information
- to amend the statement relating to anti-adalimumab antibody development in JIA patients, which will reside in section 5.1 of the Humira SmPCs (20 mg and 40 mg presentations)."

Action: For adoption

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

AstraZeneca AB

Rapporteur: Concepcion Prieto Yerro, Co-Rapporteur: David Lyons, 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

4.3.6. Oncaspar - pegaspargase - EMEA/H/C/003789/X/0008

Baxalta Innovations GmbH

Rapporteur: Alexandre Moreau, PRAC Rapporteur: Patrick Batty

Scope: "Extension application to add a new pharmaceutical form, powder for solution for injection/infusion (750 U/ml)."

Action: For adoption

4.3.7. Orkambi - lumacaftor / ivacaftor - EMEA/H/C/003954/X/0020

Vertex Pharmaceuticals (Europe) Ltd.

Rapporteur: Nithyanandan Nagercoil, Co-Rapporteur: Daniela Melchiorri, PRAC Rapporteur: Almath Spooner

Scope: "Extension application to add a new strength of film-coated tablets (100 mg Lumacaftor / 125 mg Ivacaftor) for paediatric use (6 to 11 years). The RMP (version 3.1) is updated accordingly.
CHMP assessment report on similarity
CHMP assessment report on significant clinical benefit (additional year of market exclusivity)"

Action: For adoption

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

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

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. Adcetris - brentuximab vedotin - Orphan - EMEA/H/C/002455/II/0048

Takeda Pharma A/S

Rapporteur: Paula Boudewina van Hennik, Co-Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Sabine Straus

Scope: "Extension of indication to include the new indication "ADCETRIS is indicated for the treatment of adult patients with CD30+ cutaneous T-cell lymphoma (CTCL) who require systemic therapy", based on data from study C25001 (the 'ALCANZA' study): "A Phase 3 Trial of brentuximab vedotin(SGN-35) Versus Physician's Choice (Methotrexate or Bexarotene) in Patients With CD30-Positive Cutaneous T-Cell Lymphoma". As a consequence, sections 4.1, 4.2, 4.4, 4.8, 5.1 and 5.2 of the SmPC are updated. The Package Leaflet is updated in accordance. An updated RMP (version 10) has also been submitted."

Action: For adoption

5.1.2. Aranesp - darbepoetin alfa - EMEA/H/C/000332/II/0142

Amgen Europe B.V.

Rapporteur: Martina Weise, Co-Rapporteur: Koenraad Norga, PRAC Rapporteur: Valerie Strassmann

Scope: "Extension of Indication to include treatment of anaemia in adult patients with low transfusion demand in low or intermediate-1-risk myelodysplastic syndromes for Aranesp; as a consequence, sections 4.1, 4.2,4.8, 5.1 and 5.2 of the SmPC are updated in order to update the safety and efficacy information. The Package Leaflet is updated in accordance. Updated RMP version 8.0 has been submitted.

In addition, the Marketing authorisation holder (MAH) took the opportunity to introduce QRD editorial changes in the SmPC, Annex IIIA and Annex IIIB."

Action: For adoption

5.1.3. Bydureon - exenatide - EMEA/H/C/002020/II/0041

AstraZeneca AB

Rapporteur: Kristina Dunder, PRAC Rapporteur: Qun-Ying Yue

Scope: "Update of section 4.1 of the SmPC in order to align with more recently approved glucose-lowering agents and with "Reflection paper on the wording of indication for medicinal products for treatment of type 2 diabetes" and update of section 5.1 based on the study D5553C00003 (Duration 8 study) which evaluated concomitant add-on treatment with the combination of exenatide once weekly 2 mg and dapagliflozin 10 mg once daily in patients with type 2 diabetes mellitus who have inadequate glycaemic control on metformin. The Package Leaflet is updated accordingly. In addition, the Marketing authorisation holder (MAH) took the opportunity to make minor editorial changes in the SmPC and Package Leaflet. Furthermore, the updated RMP version 24 has been submitted."

Action: For adoption

Request for Supplementary Information adopted on 23.03.2017.

5.1.4. Gazyvaro - obinutuzumab - Orphan - EMEA/H/C/002799/II/0016

Roche Registration Limited

Rapporteur: Sinan B. Sarac, Co-Rapporteur: Alexandre Moreau, PRAC Rapporteur: Patrick Batty

Scope: "Extension of Indication to include a new indication for Gazyvaro in combination with chemotherapy, followed by Gazyvaro maintenance therapy in patients achieving a response, for the treatment of patients with previously untreated advanced follicular lymphoma.
As a consequence, sections 4.1, 4.2, 4.4, 4.5, 4.8 and 5.1 of the SmPC are updated. The Package Leaflet and the RMP are updated in accordance. In addition, the due date for provision of the final clinical study report of study BO21223/GALLIUM listed in the Gazyvaro RMP as Category 3 has been updated.

Furthermore, the PI is brought in line with the missing information of QRD template version 9.1 regarding annex II C. In addition, clarification or editorial changes to the SmPC are proposed for accuracy and clarity." 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 18.05.2017, 26.01.2017.

5.1.5. Genvoya - elvitegravir / cobicistat / emtricitabine / tenofovir alafenamide - EMEA/H/C/004042/II/0026

Gilead Sciences International Ltd

Rapporteur: Robert James Hemmings, PRAC Rapporteur: Amelia Cupelli

Scope: "Extension of Indication to include paediatric patients from 6 of age to less than 12 years of age, with body weight of at least 25kg, infected with human immunodeficiency virus-1 (HIV-1) without any known mutations associated with resistance to the integrase inhibitor class, emtricitabine or tenofovir, for Genvoya.

As a consequence, sections 4.2, 4.8, 5.1 and 5.2 of the SmPC are updated based on the analysis of the paediatric study GS-US-292-0106 (Cohort 2) "A Phase 2/3, Open-Label Study of the Pharmacokinetics, Safety, and Antiviral Activity of the Elvitegravir/Cobicistat/Emtricitabine/Tenofovir Alafenamide (E/C/F/TAF) Single Tablet Regimen (STR) in HIV-1 Infected Antiretroviral Treatment Naive Adolescents and Virologically Suppressed Children".

The Package Leaflet and the Risk Management Plan (v. 3) are updated in accordance."

Action: For adoption

Request for Supplementary Information adopted on 23.03.2017.

5.1.6. Humira - adalimumab - EMEA/H/C/000481/II/0163

AbbVie Limited

Rapporteur: Kristina Dunder, PRAC Rapporteur: Ulla Wändel Liminga

Scope: "Extension of Indication to include new indication for treatment of chronic non-infectious uveitis in paediatric patients for Humira. As a consequence, sections 4.1, 4.2, 4.8 and 5.1 of the SmPC are updated. The Package Leaflet and the RMP are updated in accordance.
In addition, the Marketing authorisation holder (MAH) took the opportunity to implement an alternative format statement for blind/partially sighted patients into the Package Leaflet as it was introduced with procedure EMEA/H/C/000481/N/0155.
Furthermore, the MAH has made some editorial changes to the Package leaflet."

Action: For adoption

Request for Supplementary Information adopted on 23.03.2017.

5.1.7. Keytruda - pembrolizumab - EMEA/H/C/003820/II/0023/G

Merck Sharp & Dohme Limited

Rapporteur: Daniela Melchiorri, Co-Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Sabine Straus

Scope: "Extension of Indication to add treatment of urothelial carcinoma in patients previously treated with chemotherapy based on the results from study KEYNOTE-045; a phase 3, randomized, active-controlled, multi-site, open-label trial evaluating pembrolizumab administered at 200 mg Q3W versus investigators' choice of paclitaxel, docetaxel, or vinflunine in patients previously treated with chemotherapy.

Extension of Indication to add treatment of urothelial carcinoma in patients ineligible for cisplatin (not previously treated) based on the results from study KEYNOTE-52; a phase 2, single-arm, multisite, open-label trial of pembrolizumab at 200 mg Q3W in the treatment of patients with locally advanced or metastatic urothelial carcinoma who are not eligible for cisplatin-containing chemotherapy.

As a consequence, sections 4.1, 4.2, 4.4, 4.8 and 5.1 of the SmPC have been updated and the Package Leaflet has been updated accordingly.

Further, the MAH is proposing a change to section 4.3 of the SmPC to add that only patients

with severe hypersensitivity should be excluded from therapy, and a change to section 4.4 of the SmPC adding possible hypersensitivity and anaphylaxis as part of infusion reactions.

The application included an updated RMP version 7.0."

Action: For adoption

Request for Supplementary Information adopted on 18.05.2017.

5.1.8. Kineret - anakinra - EMEA/H/C/000363/II/0056

Swedish Orphan Biovitrum AB (publ)

Rapporteur: Sinan B. Sarac, Co-Rapporteur: Fátima Ventura, PRAC Rapporteur: Torbjorn Callreus

Scope: "Extension of indication to include a new indication for Kineret 100 mg/0.67 ml solution for injection in pre-filled syringe for the treatment of active Still's disease, including Systemic Juvenile Idiopathic Arthritis and Adult-Onset Still's Disease. As a consequence, sections 4.1, 4.2, 4.4, 4.8, 4.9, 5.1 and 5.2 of the SmPC are updated. The Package Leaflet and the RMP (version 4.0) are updated accordingly.
In addition, the marketing authorisation holder took the opportunity to make some editorial changes in the SmPC and Package leaflet."

Action: For adoption

5.1.9. Olumiant - baricitinib - EMEA/H/C/004085/II/0001

Eli Lilly Nederland B.V.; treatment of moderate to severe active rheumatoid arthritis (RA)

Scope: "Update of section 4.4 of the SmPC in order to add a warning on venous thromboembolism based on analyses of the occurrence of venous thromboembolic events in clinical trials with baricitinib. The Package Leaflet is updated accordingly. The RMP version 2.0 has been submitted, as part of this application."

Action: For adoption

5.1.10. Opdivo - nivolumab - EMEA/H/C/003985/II/0029

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 the treatment of hepatocellular carcinoma after prior sorafenib therapy in adults for Opdivo.
As a consequence, sections 4.1, 4.2, 4.4, 4.8 and 5.1 of the SmPC are updated. The Package Leaflet is updated in accordance.
Moreover, the updated RMP version 8.0 has been submitted."

Action: For adoption

Request for Supplementary Information adopted on 22.06.2017, 23.03.2017.

See 2.3

5.1.11. Pegasys - peginterferon alfa-2a - EMEA/H/C/000395/II/0091

Roche Registration Limited

Rapporteur: Filip Josephson, PRAC Rapporteur: Qun-Ying Yue

Scope: "Extension of Indication to include paediatric patients from 3 to less than 18 years of age with Chronic Hepatitis B in the immune-active phase for Pegasys.
As a consequence, sections 4.1, 4.2, 4.4, 4.8, 5.1 and 5.2 of the SmPC are updated in order to add efficacy and safety information from study YV25718. The Package Leaflet is updated in accordance.
An updated EU RMP (version 8.0) is included in this application."

Action: For adoption

Request for Supplementary Information adopted on 23.02.2017.

5.1.12. Rapamune - sirolimus - EMEA/H/C/000273/II/0164

Pfizer Limited

Rapporteur: Kristina Dunder, PRAC Rapporteur: Ulla Wändel Liminga

Scope: "Extension of indication to include the treatment of patients with lymphangioleiomyomatosis. As a consequence section 4.1, 4.2, 4.8, 5.1 and 5.2 of the SmPC are updated. The Package Leaflet and the RMP (version 6.0) are updated in accordance. In addition the MAH took the opportunity to make very minor formatting changes in the Labelling."

Action: For adoption

5.1.13. RoActemra - tocilizumab - EMEA/H/C/000955/II/0066

Roche Registration Limited

Rapporteur: Jan Mueller-Berghaus, Co-Rapporteur: Agnes Gyurasics, PRAC Rapporteur: Brigitte Keller-Stanislawski

Scope: "Extension of indication to include an indication in adult patients for the treatment of giant cell arteritis for the subcutaneous formulation of RoActemra. 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 information relevant to this indication. The Package Leaflet is updated in accordance."

Action: For adoption

Request for Supplementary Information adopted on 22.06.2017, 23.03.2017.

5.1.14. Sovaldi - sofosbuvir - EMEA/H/C/002798/II/0036

Gilead Sciences International Ltd

Rapporteur: Filip Josephson, Co-Rapporteur: Alar Irs, PRAC Rapporteur: Julie Williams

Scope: "Extension of indication to add treatment of chronic hepatitis C in adolescents aged 12 to <18 years.
As a consequence, sections 4.1, 4.2, 4.4, 4.8, 5.1 and 5.2 of the SmPC are updated in order

to add information on posology, warnings, safety, efficacy and pharmacokinetics.
The Package Leaflet and Risk Management Plan (RMP version 5.0) are updated in accordance.
In addition, the Marketing authorisation holder (MAH) took the opportunity to update the list of local representatives in the Package Leaflet.
Furthermore, the Product Information is brought in line with the latest QRD template version 10."

Action: For adoption

Request for Supplementary Information adopted on 18.05.2017, 23.02.2017.

5.1.15. Sutent - sunitinib - EMEA/H/C/000687/II/0065

Pfizer Limited

Rapporteur: Daniela Melchiorri, Co-Rapporteur: Sinan B. Sarac, PRAC Rapporteur: Carmela Macchiarulo

Scope: "Extension of Indication to include adjuvant treatment of patients at high risk of recurrent renal cell carcinoma (RCC) following nephrectomy for Sutent; as a consequence, sections 4.1, 4.2, 4.4, 4.8 and 5.1 of the SmPC are updated based on the study A6181109 ("a randomized double-blind phase 3 study of adjuvant sunitinib vs. placebo in subjects at high risk of recurrent RCC"). The Package Leaflet is updated accordingly.
In addition, the Marketing authorisation holder (MAH) took the opportunity to make minor editorial changes to the SmPC and Package Leaflet and in addition, to fulfil PAM (FU2 22.5).
Furthermore, the PI is brought in line with the latest QRD template version 10. Moreover, updated RMP version 16 has been submitted."

Action: For adoption

5.1.16. Vimpat - lacosamide - EMEA/H/C/000863/II/0065/G

UCB Pharma S.A.

Rapporteur: Filip Josephson, Co-Rapporteur: Luca Pani, PRAC Rapporteur: Qun-Ying Yue

Scope: "This is a group of variations including extension of Indication to include monotherapy and adjunctive therapy in the treatment of partial-onset seizures with or without secondary generalisation in children from 4 to less than 16 years old with epilepsy. For the treatment initiation pack it is proposed to extend only adjunctive treatment to adolescents weighting more than 50 kg (not suitable for monotherapy and children and adolescents weighting less than 50 kg). As a consequence, sections 4.1, 4.2, 4.5, 4.8, 5.1 and 5.2 of the SmPC are updated. The Package Leaflet is updated accordingly.
In addition, the Marketing authorisation holder (MAH) took the opportunity to bring Annex IIIA in line with the latest QRD template version 10 and to introduce combined SmPC for film coated tablets. Moreover, updated RMP version 12 has been submitted.
Furthermore, only for syrup presentation, in addition sections 6.3 and 6.5 of the SmPC are updated due to extension of shelf life of finished product after first opening from 4 weeks to 6 months and addition of a 10 mL dosing syringe for syrup, as additional dosing device for paediatric population."

Action: For adoption

Request for Supplementary Information adopted on 18.05.2017, 10.11.2016.

5.1.17. Xgeva - denosumab - EMEA/H/C/002173/II/0055

Amgen Europe B.V.

Rapporteur: Kristina Dunder, Co-Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Ulla Wändel Liminga

Scope: "Extension of Indication to include "Prevention of skeletal related events (pathological fracture, radiation to bone, spinal cord compression or surgery to bone) in adults with multiple myeloma and in adults with bone metastases from solid tumours" for Xgeva; 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."

Action: For adoption

5.1.18. Zytiga - abiraterone acetate - EMEA/H/C/002321/II/0047

Janssen-Cilag International NV

Rapporteur: Jorge Camarero Jiménez, Co-Rapporteur: Robert James Hemmings, PRAC Rapporteur: Eva A. Segovia

Scope: "Extension of indication to include the treatment of newly diagnosed high risk metastatic hormone sensitive prostate cancer and in combination with androgen deprivation therapy. As a consequence, sections 4.1, 4.2, 4.4, 4.8 and 5.1 of the SmPC are updated. The package leaflet and the RMP (version 14.0) are updated accordingly. In addition, the MAH took the opportunity to update the list of local representatives in the Package Leaflet"

Action: For adoption

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

5.2.1. Fluenz Tetra - influenza vaccine (live attenuated, nasal) - EMEA/H/C/02617/II/0072

AstraZeneca UK Ltd

Rapporteur: Bart Van der Schueren

Scope: "To replace the strain of a seasonal vaccine against human influenza in line with the EU recommendations for the seasonal influenza vaccine composition for the season 2017/2018."

Report from BWP

Action: For information

Request for Supplementary Information adopted on 22.06.2017

5.2.2. Orkambi - lumacaftor / ivacaftor - EMEA/H/C/003954/II/17

Vertex Pharmaceuticals (Europe) Ltd.

Rapporteur: Nithyanandan Nagercoil, Co-Rapporteur: Daniela Melchiorri, PRAC Rapporteur: Almath Spooner

Scope: Letter from the the applicant dated 3 July 2017 requesting an extension of clock stop to respond to the second Request for Supplementary Information adopted on 18.05.2017.

Action: For adoption

Request for Supplementary Information adopted on 18.05.2017, 23.02.2017

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

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

6.2. Update of Ancillary medicinal substances in medical devices

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)

8. Pre-submission issues

8.1. Pre-submission issue

8.1.1. durvalumab - H0004771

treatment of patients with locally advanced, unresectable NSCLC whose disease has not progressed following platinum-based chemoradiation therapy.

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

Action: For adoption

8.2. Priority Medicines (PRIME)

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

8.2.1. List of applications received

Action: For information

8.2.2. Recommendation for PRIME eligibility

Action: For adoption

9. Post-authorisation issues

9.1. Post-authorisation issues

9.1.1. CHMP request for PRAC advice on Fluoropyrimidines (Capecitabine-Xeloda and 5-FU), EMEA/H/C/0316/LEG-033.1

Xeloda, Rapporteur: Harald Enzmann, PRAC Rapporteur: Martin Huber

Scope: PRAC Advice to CHMP following formal advice from PGWP regarding the proposed SmPC changes for upfront testing of patients for DPYD variants and dose reduction based on patient's genotype to reduce the toxicity of capecitabine and 5-FU

Action: For adoption

9.1.2. Vedrop - tocofersolan – Orphan - EMEA/H/C/000920/II/0022

Orphan Europe SARL

Rapporteur: Greg Markey, PRAC Rapporteur: Patrick Batty

Scope: "Submission of the final report for the registry of pediatric patients treated with Vedrop (tocofersolan) in Europe for vitamin E deficiency due to digestive malab-sorption in congenital or hereditary chronic cholestasis. Consequentially, the remaining specific obligation is fulfilled and Annexes I, II and IIIB are updated accordingly."

Action: For adoption

10. Referral procedures

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

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

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

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

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

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

10.6.1. Gadolinium-containing contrast agents (GdCA): gadoversetamide – OPTIMARK (CAP) Gadobenate dimeglumine; gadobutrol; gadodiamide; gadopentetic acid dimeglumine, gadoteric acid (intra articular formulation); gadoteric acid (intrvenous and intravascular formulations); gadoteridol; gadoxetic acid disodium (NAP)

Applicant(s): Mallinckrodt Deutschland GmbH (Optimark); various

Rapporteurs for the Article 31 referral: PRAC Rapporteur: Ulla Wändel Liminga; PRAC Co-rapporteur: Valerie Strassmann

Rapporteurs for Optimark: CHMP Rapporteur: Patrick Salmon, CHMP Co-rapporteur: Johann Lodewijk Hillege

Scope: Oral explanation/Opinion

Action: For adoption

Re-examination procedure under Article 32 of Directive 2001/83/EC of the review of the benefit-risk balance of GdCA following notification by the European Commission of a referral under Article 31 of Directive 2001/83/EC, based on pharmacovigilance data

See 2.4

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

10.7.1. Alcover 750 mg, 1250 mg, 1750 mg Granulat im Beutel – Sodium oxybate – EMEA/H/A-29(4)/1451

D&A Pharma

Rapporteur: Andrea Laslop, Co-Rapporteur: Fatima Ventura,

Scope: Appointment of re-examination rapporteurs, draft timetable, SAG involvement

Letter from the applicant dated 30 June 2017 requesting a re-examination of the Opinion adopted on 22 June 2017 and consultation of a Scientific Advisory Group

Action: For adoption

Decentralised Procedure number: AT/H/0552/01-03/DC, notification by the Austrian Agency dated 22 December 2016 notifying of the start of a referral under Article 29(4) of Directive 2001/83/EC.

Opinion adopted on 22 June 2017, List of Outstanding issues adopted on 21.04.2017. List of Questions adopted on 26.01.2017.

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

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

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

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

11. Pharmacovigilance issue

11.1. Early Notification System

July 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

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

12.2. GCP inspections

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

12.3. Pharmacovigilance inspections

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

12.4. GLP inspections

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

13. Innovation Task Force

13.1. Minutes of Innovation Task Force

Action: For information

13.2. Innovation Task Force briefing meetings

Disclosure of information related to briefing meetings taking place with applicants cannot be released at present time as deemed to contain commercially confidential information

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

13.4. Nanomedicines activities

14. Organisational, regulatory and methodological matters

14.1. Mandate and organisation of the CHMP

14.1.1. Time schedule of CHMP August written procedure

Action: For information

14.1.2. Pilot phase for abolition of signatures for divergent positions for referral procedures

The proposal is to start new process from September onwards

Action: For information

14.1.3. Update to the CHMP templates on initial Marketing Authorisation

Update to the Rapporteurs' D80 AR overview guidance document to add guidance specific to biosimilars (including a revised Benefit/Risk balance section). When adopted, the changes will be implemented in all relevant templates on initial MA

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 3-6 July 2017

Action: For information

List of Union Reference Dates and frequency of submission of Periodic Safety Update Reports (EURD list) for July 2017

Action: For adoption

14.2.2. Committee for Advanced Therapies (CAT)

CAT draft minutes of meeting held on 12-14 July 2017

Action: For information

Revision of Procedural advice on the evaluation of Advanced Therapy Medicinal Product in accordance with Article 8 of Regulation (EC) NO 1394/2007

Action: For discussion

14.2.3. Paediatric Committee (PDCO)

PIPs reaching D30 at July 2017 PDCO

Action: For information

Report from the PDCO meeting held on 18-21 July 2017

Action: For information

Advice from the CHMP and PDCO task force on how to address issues related to therapeutic equivalence for orally inhaled products for children

Action: For adoption

EMA/FDA/Health Canada workshop on paediatric pulmonary hypertension (PAH) – meeting highlights

Action: For information

14.2.4. Committee for Orphan Medicinal Products (COMP)

Report from the COMP meeting held on11-13 July 2017
Action: For information

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

Report from the Coordination Group for Mutual Recognition and Decentralised Procedures – Human (CMDh) on the meeting held on 17-19 July 2017.

Action: For information

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 3-6 July 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.

14.3.2. Quality Working Party (QWP)

Chair: Keith Pugh

Election of QWP Vice Chair, the term of the current Chair ending in July 2017.

Action: For adoption

Guideline on Manufacture of the Finished Dosage Form (EMA/CHMP/QWP/BWP/245074)

Action: For adoption

14.3.3. Radiopharmaceutical Drafting Group (RadDG)

Chair: Anabel Cortes

Guideline on core SmPC and Package Leaflet for (68Ge-68Ga) generator (EMA/313282/2017)

Action: For adoption

Overview of comments 'Guideline on core SmPC and Package Leaflet for (68Ge68Ga) generator (EMA/313283/2017)

Action: For information

14.4. Cooperation within the EU regulatory network

14.5. Cooperation with International Regulators

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

14.7. CHMP work plan

14.7.1. CHMP 2017 Work Plan: mid-year update

Action: For information

14.8. Planning and reporting

14.9. Others

15. Any other business

15.1. AOB topic

15.1.1. Working group on Committees' operational preparedness for human medicines

Scope: CHMP representatives to this Cross-Committee working group

Action: For adoption

15.1.2. Revision of the 'Guideline on strategies to identify and mitigate risks for first-in-human clinical trials with investigational medicinal products'

CHMP Rapporteur: Harald Enzmann

Scope: Final revised guideline to be published

Action: For adoption

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)*

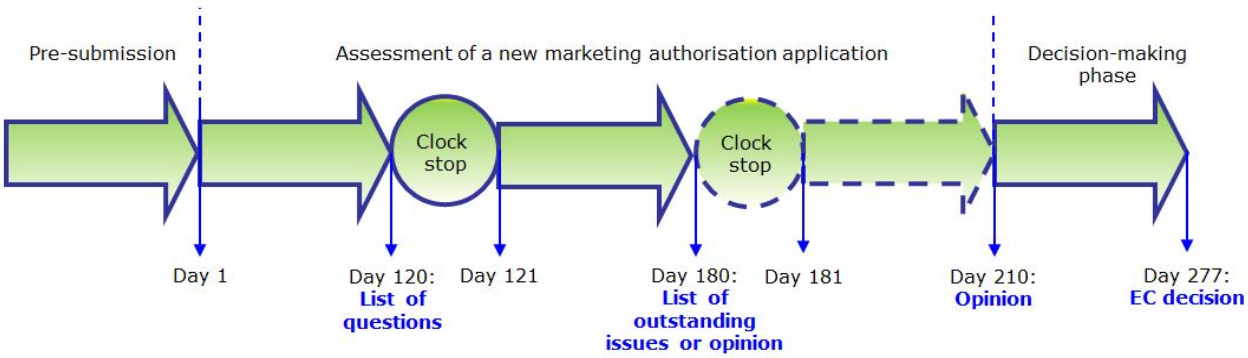
The items listed in this section are those for which marketing authorisation holders (MAHs) or applicants have been invited to the CHMP plenary meeting to address questions raised by the Committee. Oral explanations normally relate to on-going applications (section 3, 4 and 5) or referral procedures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

17 July 2017
EMA/CHMP/451868/2017

Annex to July 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
July 2017: **For adoption**

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

Final Outcome of Rapporteurship allocation for
July 2017: **For adoption**

A.3. PRE-SUBMISSION ISSUES FOR INFORMATION

Disclosure of information related to pre-submission of initial applications cannot be released at 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

Elaprase - idursulfase -
EMEA/H/C/000700/S/0070
MAH: Shire Human Genetic Therapies AB,
Rapporteur: Greg Markey, PRAC Rapporteur:
Patrick Batty

B.2. RENEWALS OF MARKETING AUTHORISATIONS OUTCOMES

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

Betmiga - mirabegron -
EMEA/H/C/002388/R/0026
MAH: Astellas Pharma Europe B.V., Rapporteur:
Concepcion Prieto Yerro, Co-Rapporteur:
Nithyanandan Nagercoil, PRAC Rapporteur:
Dolores Montero Corominas
Request for Supplementary Information adopted
on 18.05.2017.

Ibandronic acid Accord - ibandronic acid -
EMEA/H/C/002638/R/0013
MAH: Accord Healthcare Limited, Generic,
Generic of Bondronat, Rapporteur: Alar Irs, PRAC
Rapporteur: Doris Stenver

Request for Supplementary Information adopted on 18.05.2017.

B.2.2. Renewals of Marketing Authorisations for unlimited validity

Amyvid - florbetapir (18F) - EMEA/H/C/002422/R/0026
MAH: Eli Lilly Nederland B.V., Rapporteur: Harald Enzmann, Co-Rapporteur: Concepcion Prieto Yerro, PRAC Rapporteur: Valerie Strassmann

Bexsero - meningococcal group B vaccine (rDNA, component, adsorbed) - EMEA/H/C/002333/R/0053
MAH: GSK Vaccines S.r.l, Rapporteur: Kristina Dunder, Co-Rapporteur: Svein Rune Andersen, PRAC Rapporteur: Qun-Ying Yue

Imatinib Teva - imatinib - EMEA/H/C/002585/R/0028
MAH: Teva B.V., Generic, Generic of Glivec, Rapporteur: Jorge Camarero Jiménez, PRAC Rapporteur: Eva A. Segovia

Lyxumia - lixisenatide - EMEA/H/C/002445/R/0023
MAH: sanofi-aventis groupe, Rapporteur: Kristina Dunder, Co-Rapporteur: Bart Van der Schueren, PRAC Rapporteur: Qun-Ying Yue

Ryzodeg - insulin degludec / insulin aspart - EMEA/H/C/002499/R/0024
MAH: Novo Nordisk A/S, Rapporteur: Kristina Dunder, Co-Rapporteur: Hanne Lomholt Larsen, PRAC Rapporteur: Qun-Ying Yue

Tresiba - insulin degludec - EMEA/H/C/002498/R/0027
MAH: Novo Nordisk A/S, Rapporteur: Kristina Dunder, Co-Rapporteur: Hanne Lomholt Larsen, PRAC Rapporteur: Qun-Ying Yue

Zaltrap - aflibercept - EMEA/H/C/002532/R/0037
MAH: sanofi-aventis groupe, Rapporteur: Filip Josephson, Co-Rapporteur: Daniela Melchiorri, PRAC Rapporteur: Ulla Wändel Liminga

B.2.3. Renewals of Conditional Marketing Authorisations

Adcetris - brentuximab vedotin - EMEA/H/C/002455/R/0051, Orphan
MAH: Takeda Pharma A/S, Rapporteur: Paula

Boudewina van Hennik, PRAC Rapporteur: Sabine Straus

Blincyto - blinatumomab - EMEA/H/C/003731/R/0013, Orphan
MAH: Amgen Europe B.V., Rapporteur: Alexandre Moreau, Co-Rapporteur: Daniela Melchiorri, PRAC Rapporteur: Eva Jirsová

Lartruvo - olaratumab - EMEA/H/C/004216/R/0004, Orphan
MAH: Eli Lilly Nederland B.V., Rapporteur: Jorge Camarero Jiménez, Co-Rapporteur: Daniela Melchiorri, PRAC Rapporteur: Sabine Straus

Ninlaro - ixazomib - EMEA/H/C/003844/R/0003, Orphan
MAH: Takeda Pharma A/S, Rapporteur: Greg Markey, Co-Rapporteur: Daniela Melchiorri, PRAC Rapporteur: Ulla Wändel Liminga

Venclyxto - venetoclax - EMEA/H/C/004106/R/0005, Orphan
MAH: AbbVie Limited, Rapporteur: Filip Josephson, PRAC Rapporteur: Patrick Batty

B.3. POST-AUTHORISATION PHARMACOVIGILANCE OUTCOMES

Signal detection

PRAC recommendations on signals adopted at the PRAC meeting held on 3-6 July 2017 PRAC:

Aranesp - Darbepoetin alfa - EMEA/H/C/000332; MAH: Amgen Europe B.V.; Rapporteur: Martina Weise, Co-Rapporteur: Koenraad Norga, PRAC Rapporteur: Valerie Strassmann,

Abseamed - Epoetin alfa - EMEA/H/C/000727; MAH: Medice Arzneimittel Pütter GmbH & Co. KG; **&**
Binocrit - Epoetin alfa - EMEA/H/C/000727; MAH: Sandoz GmbH; **&**
Epoetin alfa Hexal - Epoetin alfa - EMEA/H/C/000726; MAH: Hexal AG; Rapporteur: Alexandre Moreau, Co-Rapporteur: Martina Weise, PRAC Rapporteur: Ghania Chamouni,

NeoRecormon – Epoetin beta - EMEA/H/C/000116; MAH: Roche Registration Limited; Rapporteur: Martina Weise, Co-Rapporteur: Alexandre Moreau,

PRAC Rapporteur: Valerie Strassmann,

Biopoin – Epoetin theta - EMEA/H/C/001036; MAH: TEVA GmbH; **Eporatio – Epoetin theta - EMEA/H/C/001033;** ratiopharm GmbH; Rapporteur: Alexandre Moreau, Co-Rapporteur: Martina Weise, PRAC Rapporteur: Ghania Chamouni,

Retacrit – Epoetin zeta - EMEA/H/C/000872; MAH: Hospira UK Ltd; **& Silapo – Epoetin zeta - EMEA/H/C/000760;** MAH: STADA Arzneimittel AG; Rapporteur: Martina Weise, Co-Rapporteur: Nithyanandan Nagercoil, PRAC Rapporteur: Valerie Strassmann,

Mircera – Methoxy polyethylene glycol-epoetin beta - EMEA/H/C/000739; MAH: Roche Registration Limited; Rapporteur: Alexandre Moreau, Co-Rapporteur: Martina Weise, PRAC Rapporteur: Ghania Chamouni,

Signal of severe cutaneous adverse reactions (SCARs) including Stevens-Johnson syndrome (SJS) and toxic epidermal necrolysis (TEN) – **PRAC recommendation** on a variation, DHPC letter and Communication plan (DHPC and Communication plan are being finalised via written procedure): **For adoption**

Faslodex - Fulvestrant – EMEA/H/C/000540; MAH: AstraZeneca UK Ltd; Rapporteur: Filip Josephson, Co-Rapporteur: Tuomo Lapveteläinen, PRAC Rapporteur: Ulla Wändel Liminga, Signal of anaphylactic reactions: **For adoption**

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

EMEA/H/C/PSUSA/00000425/201611 (bosentan)
CAPS:
Stayveer (EMEA/H/C/002644) (bosentan), MAH: Marklas Nederlands BV, Rapporteur: Alexandre Moreau
Tracleer (EMEA/H/C/000401) (bosentan), MAH: Actelion Registration Ltd., Rapporteur: Alexandre Moreau
NAPS:

Klimurtan - ELPEN PHARMACEUTICAL CO. INC. , PRAC Rapporteur: Caroline Laborde, “20 November 2015 - 19 November 2016”

EMEA/H/C/PSUSA/00001730/201611
(indacaterol)
CAPS:
Hirobriz Breezhaler (EMEA/H/C/001211) (indacaterol), MAH: Novartis Europharm Ltd, Rapporteur: Hanne Lomholt Larsen
Onbrez Breezhaler (EMEA/H/C/001114) (indacaterol), MAH: Novartis Europharm Ltd, Rapporteur: Hanne Lomholt Larsen
Oslif Breezhaler (EMEA/H/C/001210) (indacaterol), MAH: Novartis Europharm Ltd, Rapporteur: Hanne Lomholt Larsen, PRAC Rapporteur: Torbjorn Callreus, “1-Dec-2013 to 30-Nov-2016”

EMEA/H/C/PSUSA/00001838/201612
(lenalidomide)
CAPS:
Revlimid (EMEA/H/C/000717) (lenalidomide), MAH: Celgene Europe Limited, Rapporteur: Alexandre Moreau, PRAC Rapporteur: Ghania Chamouni, “27/12/2015 - 26/12/2016”

EMEA/H/C/PSUSA/00002451/201612
(plerixafor)
CAPS:
Mozobil (EMEA/H/C/001030) (plerixafor), MAH: Genzyme Europe BV, Rapporteur: Paula Boudewina van Hennik, PRAC Rapporteur: Sabine Straus, “16-Dec-2013 – 15-Dec-2016”

EMEA/H/C/PSUSA/00010028/201612
(concentrate of proteolytic enzymes enriched in bromelain)
CAPS:
NexoBrid (EMEA/H/C/002246) (concentrate of proteolytic enzymes enriched in bromelain), MAH: MediWound Germany GmbH, Rapporteur: Harald Enzmann, PRAC Rapporteur: Valerie Strassmann, “Update of section 4.4 and 4.8 of the SmPC to add serious allergic reactions including anaphylactic reaction with a frequency not known and to add a warning on hypersensitivity reactions. The Package leaflet is updated accordingly.”

EMEA/H/C/PSUSA/00010128/201612
(ponatinib)
CAPS:

Iclusig (EMEA/H/C/002695) (ponatinib), MAH: Incyte Biosciences UK Ltd, Rapporteur: Greg Markey, PRAC Rapporteur: Patrick Batty, “14 June 2016 – 13 December 2016”

EMEA/H/C/PSUSA/00010264/201612
(umeclidinium bromide / vilanterol)
CAPS:
Anoro (EMEA/H/C/002751) (umeclidinium bromide / vilanterol), MAH: Glaxo Group Ltd, Rapporteur: Nithyanandan Nagercoil
Laventair (EMEA/H/C/003754) (umeclidinium / vilanterol), MAH: Glaxo Group Ltd, Rapporteur: Nithyanandan Nagercoil, PRAC Rapporteur: Carmela Macchiarulo, “18 June 2016 – 17 December 2016”

EMEA/H/C/PSUSA/00010322/201612
(olaparib)
CAPS:
Lynparza (EMEA/H/C/003726) (olaparib), MAH: AstraZeneca AB, Rapporteur: Alexandre Moreau, PRAC Rapporteur: Carmela Macchiarulo, “16 June 2016 to 15 December 2016”

EMEA/H/C/PSUSA/00010379/201701
(nivolumab)
CAPS:
Opdivo (EMEA/H/C/003985) (nivolumab), MAH: Bristol-Myers Squibb Pharma EEIG, Rapporteur: Jorge Camarero Jiménez, PRAC Rapporteur: Brigitte Keller-Stanislawski, “04 July 2016 - 03 January 2017”

EMEA/H/C/PSUSA/00010391/201612
(lutetium isotope of mass 177)
CAPS:
EndolucinBeta (EMEA/H/C/003999) (lutetium (177lu) chloride), MAH: ITG Isotope Technologies Garching GmbH, Rapporteur: Patrick Salmon
LuMark (EMEA/H/C/002749) (lutetium, isotope of mass 177), MAH: I.D.B. Holland B.V., Rapporteur: Nithyanandan Nagercoil, PRAC Rapporteur: Almath Spooner, “06.07.2016 to 19.12.2016”

EMEA/H/C/PSUSA/00010460/201612
(blinatumomab)
CAPS:
Blincyto (EMEA/H/C/003731) (blinatumomab), MAH: Amgen Europe B.V., Rapporteur: Alexandre Moreau, PRAC Rapporteur: Eva

Jirsová, “24 May 2016 - 02 December 2016”

B.4. EPARs / WPARs

Efavirenz/Emtricitabine/Tenofovir disoproxil Mylan - efavirenz / emtricitabine / tenofovir disoproxil - EMEA/H/C/004240
Applicant: Mylan S.A.S, treatment of HIV-1 infection, Generic, Generic of Atripla, Generic application (Article 10(1) of Directive No 2001/83/EC)

Fotivda - tivozanib - EMEA/H/C/004131
Applicant: EUSA PHARMA, treatment of adult patients with advanced renal cell carcinoma (RCC), New active substance (Article 8(3) of Directive No 2001/83/EC)

Imraldi - adalimumab - EMEA/H/C/004279
Applicant: Samsung Bioepis UK Limited, treatment of rheumatoid arthritis, psoriatic arthritis and ankylosing spondylitis, Similar biological application (Article 10(4) of Directive No 2001/83/EC)

Kisqali - ribociclib - EMEA/H/C/004213
Applicant: Novartis Europharm Ltd, treatment of breast cancer, New active substance (Article 8(3) of Directive No 2001/83/EC)

Mavenclad - cladribine - EMEA/H/C/004230
Applicant: Merck Serono Europe Limited, treatment of highly active relapsing-remitting multiple sclerosis (MS), Known active substance (Article 8(3) of Directive No 2001/83/EC)

Maviret - glecaprevir / pibrentasvir - EMEA/H/C/004430
Applicant: AbbVie Limited, indicated for the treatment of chronic hepatitis C virus (HCV) infection in adults, New active substance (Article 8(3) of Directive No 2001/83/EC)

Nitisinone MendeliKABS - nitisinone - EMEA/H/C/004281
Applicant: MendeliKABS Europe Ltd, treatment of hepatorenal tyrosinemia type 1, Generic, Generic of Orfadin, Generic application (Article 10(1) of Directive No 2001/83/EC)

Vosevi - sofosbuvir / velpatasvir / voxilaprevir - EMEA/H/C/004350
Applicant: Gilead Sciences International Ltd, Treatment of chronic hepatitis C virus in adults

(HCV) infection in adults, New active substance (Article 8(3) of Directive No 2001/83/EC)	

B.5. TYPE II VARIATION, WORKSHARING PROCEDURE OUTCOMES

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

B.5.1. CHMP assessed procedures scope: Pharmaceutical aspects

Advate - octocog alfa - EMEA/H/C/000520/II/0082/G MAH: Baxter AG, Rapporteur: Jan Mueller-Berghaus Request for Supplementary Information adopted on 23.02.2017.	Weekly start timetable.
Advate - octocog alfa - EMEA/H/C/000520/II/0085 MAH: Baxter AG, Rapporteur: Jan Mueller-Berghaus	Weekly start timetable.
Afstyla - lonoctocog alfa - EMEA/H/C/004075/II/0001 MAH: CSL Behring GmbH, Rapporteur: Jan Mueller-Berghaus Request for Supplementary Information adopted on 21.04.2017.	Weekly start timetable.
Alprolix - eftrenonacog alfa - EMEA/H/C/004142/II/0006/G, Orphan MAH: Swedish Orphan Biovitrum AB (publ), Rapporteur: Andrea Laslop Request for Supplementary Information adopted on 09.06.2017.	Weekly start timetable.
Ciambra - pemetrexed - EMEA/H/C/003788/II/0002/G MAH: Menarini International Operations Luxembourg S.A., Generic, Generic of Alimta, Rapporteur: Juris Pokrotnieks Opinion adopted on 06.07.2017. Request for Supplementary Information adopted on 06.04.2017.	Positive Opinion adopted by consensus on 06.07.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Cosentyx - secukinumab - EMEA/H/C/003729/II/0024 MAH: Novartis Europharm Ltd, Rapporteur: Tuomo Lapveteläinen Opinion adopted on 06.07.2017.	Positive Opinion adopted by consensus on 06.07.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Elonva - corifollitropin alfa - EMEA/H/C/001106/II/0036/G	Weekly start timetable.

MAH: Merck Sharp & Dohme Limited, Rapporteur: Paula Boudewina van Hennik Request for Supplementary Information adopted on 09.06.2017.	
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 16.03.2017.	Weekly start timetable.
Fabrazyme - agalsidase beta - EMEA/H/C/000370/II/0099/G MAH: Genzyme Europe BV, Rapporteur: Johann Lodewijk Hillege Opinion adopted on 06.07.2017.	Positive Opinion adopted by consensus on 06.07.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Fluenz Tetra - influenza vaccine (live attenuated, nasal) - EMEA/H/C/002617/II/0072 MAH: AstraZeneca AB, Rapporteur: Bart Van der Schueren, “To replace the strain of a seasonal vaccine against human influenza in line with the EU recommendations for the seasonal influenza vaccine composition for the season 2017/2018.” Request for Supplementary Information adopted on 22.06.2017.	See 5.2 in main agenda
Imatinib Actavis - imatinib - EMEA/H/C/002594/II/0013 MAH: Actavis Group PTC ehf, Generic, Generic of Glivec, Rapporteur: Hrefna Gudmundsdottir Request for Supplementary Information adopted on 06.07.2017.	Weekly start timetable.
Inflectra - infliximab - EMEA/H/C/002778/II/0050/G MAH: Hospira UK Limited, Duplicate, Duplicate of Remsima, Rapporteur: Greg Markey Opinion adopted on 29.06.2017. Request for Supplementary Information adopted on 05.05.2017.	Positive Opinion adopted by consensus on 29.06.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Jakavi - ruxolitinib - EMEA/H/C/002464/II/0034 MAH: Novartis Europharm Ltd, Rapporteur: Filip Josephson	Weekly start timetable.
Kadcyla - trastuzumab emtansine - EMEA/H/C/002389/II/0034 MAH: Roche Registration Limited, Rapporteur: Sinan B. Sarac Request for Supplementary Information adopted	Weekly start timetable.

on 29.06.2017.	
Keytruda - pembrolizumab - EMEA/H/C/003820/II/0026/G MAH: Merck Sharp & Dohme Limited, Rapporteur: Daniela Melchiorri Request for Supplementary Information adopted on 01.06.2017, 21.04.2017.	Weekly start timetable.
Keytruda - pembrolizumab - EMEA/H/C/003820/II/0030 MAH: Merck Sharp & Dohme Limited, Rapporteur: Daniela Melchiorri Request for Supplementary Information adopted on 06.07.2017.	Weekly start timetable.
Keytruda - pembrolizumab - EMEA/H/C/003820/II/0031/G MAH: Merck Sharp & Dohme Limited, Rapporteur: Daniela Melchiorri	Weekly start timetable.
Memantine ratiopharm - memantine - EMEA/H/C/002671/II/0008 MAH: ratiopharm GmbH, Generic, Generic of Ebixa, Rapporteur: Bart Van der Schueren Request for Supplementary Information adopted on 01.06.2017.	Weekly start timetable.
Myozyme - alglucosidase alfa - EMEA/H/C/000636/II/0063/G MAH: Genzyme Europe BV, Rapporteur: Alexandre Moreau	Weekly start timetable.
Pemetrexed Fresenius Kabi - pemetrexed - EMEA/H/C/003895/II/0002 MAH: Fresenius Kabi Oncology PLC, Generic, Generic of Alimta, Rapporteur: Bjorg Bolstad Opinion adopted on 06.07.2017.	Positive Opinion adopted by consensus on 06.07.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Perjeta - pertuzumab - EMEA/H/C/002547/II/0030 MAH: Roche Registration Limited, Rapporteur: Sinan B. Sarac Request for Supplementary Information adopted on 29.06.2017.	Weekly start timetable.
Plavix - clopidogrel - EMEA/H/C/000174/II/0127/G MAH: Sanofi Clir SNC, Rapporteur: Bruno Sepodes	Weekly start timetable.
Praluent - alirocumab - EMEA/H/C/003882/II/0021/G MAH: sanofi-aventis groupe, Rapporteur: Johann Lodewijk Hillege	Weekly start timetable.

Request for Supplementary Information adopted on 18.05.2017.	
Praluent - alirocumab - EMEA/H/C/003882/II/0024/G MAH: sanofi-aventis groupe, Rapporteur: Johann Lodewijk Hillege	Weekly start timetable.
Privigen - human normal immunoglobulin - EMEA/H/C/000831/II/0118 MAH: CSL Behring GmbH, Rapporteur: Jan Mueller-Berghaus Opinion adopted on 06.07.2017.	Positive Opinion adopted by consensus on 06.07.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Ratiograstim - filgrastim - EMEA/H/C/000825/II/0053/G MAH: ratiopharm GmbH, Rapporteur: Outi Mäki-Ikola Opinion adopted on 06.07.2017. Request for Supplementary Information adopted on 16.03.2017.	Positive Opinion adopted by consensus on 06.07.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Remicade - infliximab - EMEA/H/C/000240/II/0205 MAH: Janssen Biologics B.V., Rapporteur: Kristina Dunder	Weekly start timetable.
Remsima - infliximab - EMEA/H/C/002576/II/0042/G MAH: Celltrion Healthcare Hungary Kft., Rapporteur: Greg Markey Opinion adopted on 29.06.2017. Request for Supplementary Information adopted on 05.05.2017.	Positive Opinion adopted by consensus on 29.06.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Surgiflo Haemostatic Matrix Kit -Ferrosan - human thrombin - EMEA/H/D/002301/II/0016 MAH: Presafe Denmark A/S, Rapporteur: Jan Mueller-Berghaus	Weekly start timetable.
Tevagrastim - filgrastim - EMEA/H/C/000827/II/0063/G MAH: TEVA GmbH, Duplicate, Duplicate of Biograstim, Rapporteur: Outi Mäki-Ikola Opinion adopted on 06.07.2017. Request for Supplementary Information adopted on 16.03.2017.	Positive Opinion adopted by consensus on 06.07.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Vaxelis - diphtheria, tetanus, pertussis (acellular, component), hepatitis B (rDNA), poliomyelitis (inact.) and Haemophilus type B conjugate vaccine (adsorbed) - EMEA/H/C/003982/II/0012/G	Weekly start timetable.

MAH: MCM Vaccine B.V., Rapporteur: Bart Van der Schueren	
Vihuma - simoctocog alfa - EMEA/H/C/004459/II/0001/G MAH: Octapharma AB, Rapporteur: Jan Mueller-Berghaus Request for Supplementary Information adopted on 15.06.2017.	Weekly start timetable.
Xultophy - insulin degludec / liraglutide - EMEA/H/C/002647/II/0019 MAH: Novo Nordisk A/S, Rapporteur: Kristina Dunder Request for Supplementary Information adopted on 01.06.2017.	Weekly start timetable.
Yervoy - ipilimumab - EMEA/H/C/002213/II/0048/G MAH: Bristol-Myers Squibb Pharma EEIG, Rapporteur: Paula Boudewina van Hennik Opinion adopted on 06.07.2017.	Positive Opinion adopted by consensus on 06.07.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1143 Aflunov-EMEA/H/C/002094/WS1143/0033 Foclivia-EMEA/H/C/001208/WS1143/0028 MAH: Seqirus S.r.l, Lead Rapporteur: Daniela Melchiorri Request for Supplementary Information adopted on 15.06.2017, 05.05.2017.	Weekly start timetable.
WS1159 Neulasta-EMEA/H/C/000420/WS1159/0095 Ristempa-EMEA/H/C/003910/WS1159/0011 MAH: Amgen Europe B.V., Lead Rapporteur: Robert James Hemmings Opinion adopted on 06.07.2017.	Positive Opinion adopted by consensus on 06.07.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

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

Arzerra - ofatumumab - EMEA/H/C/001131/II/0050, Orphan MAH: Novartis Europharm Ltd, Rapporteur: Sinan B. Sarac, "Update of sections 4.2, 4.4 and 4.8 of the SmPC in order to add a recommendation to permanently discontinue Arzerra in case of anaphylactic reaction and revise the adverse drug reaction profile based on safety pool data analysis and updated Company Core	Weekly start timetable.

Data Sheet. The Package Leaflet is updated accordingly." Request for Supplementary Information adopted on 15.06.2017.	
Arzerra - ofatumumab - EMEA/H/C/001131/II/0051, Orphan MAH: Novartis Europharm Ltd, Rapporteur: Sinan B. Sarac, "Update to Section 4.6 Fertility, pregnancy and lactation and section 5.3 Preclinical safety data following implementation of the new Novartis Core Data Sheet (CDS) template. Update to the Section 4.5 Interaction with other medicinal products and other forms of interaction to reflect the results of a clinical study OMB113603 investigating the potential pharmacokinetic interactions between ofatumumab and bendamustine. Update to the Section 5 Pharmacological properties to update the information on immunogenicity for precision, addition of a table summarizing main pharmacokinectic (PK) parameters for brevity and to facilitate understanding, simplification of the PK section. Editorial changes for Arzerra 100 mg and Arzerra 1000 mg concentrate for solution for infusion consisting of updates to Sections 2 Qualitative and quantitative composition, 6.5 Nature and contents of container, and 6.6 Special precautions for disposal and other handling. Editorial changes to clarify the doses for various indications have been added to Section 4.2 Posology and method of administration. In addition, the Marketing authorisation holder (MAH) took the opportunity to bring the PI in line with the latest QRD template version 10." Request for Supplementary Information adopted on 15.06.2017.	Weekly start timetable.
Bexsero - meningococcal group B vaccine (rDNA, component, adsorbed) - EMEA/H/C/002333/II/0054 MAH: GSK Vaccines S.r.l, Rapporteur: Kristina Dunder, "Update of section 4.8 of the SmPC in order to add the adverse reactions "injection site reactions (including extensive swelling of the vaccinated limb)" and "injection site nodule which may persist for more than one month" with a	Positive Opinion adopted by consensus on 06.07.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

frequency not known. The Package Leaflet is updated accordingly. In addition, the MAH took the opportunity to bring the product information in line with the latest QRD template version 10.0." Opinion adopted on 06.07.2017.	
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 26.01.2017.	
Cosentyx - secukinumab - EMEA/H/C/003729/II/0020 MAH: Novartis Europharm Ltd, Rapporteur: Tuomo Lapveteläinen, "Update of section 4.5 of the SmPC in order to revise general information on CYP450/CYP3A4 as a result of data provided by study A2110 demonstrating that enzyme activity in moderate to severe psoriasis patients at baseline is similar to the activity observed in healthy volunteers." Request for Supplementary Information adopted on 01.06.2017.	Weekly start timetable.
Cyanokit - hydroxocobalamin - EMEA/H/C/000806/II/0031 MAH: SERB SA, Rapporteur: Alexandre Moreau, "Update of sections 4.4 and 4.8 of the SmPC in order to add a warning on renal disorders and to update the safety information on skin and subcutaneous tissue disorders, renal and urinary disorders following a safety signal on renal disorders. The package leaflet is updated accordingly." Request for Supplementary Information adopted on 18.05.2017.	Weekly start timetable.
Dacogen - decitabine - EMEA/H/C/002221/II/0031, Orphan MAH: Janssen-Cilag International NV, Rapporteur: Alexandre Moreau, "Update of	

section 6.6. of the SmPC in order to update the reconstitution procedure based on new quality data, as to obtain a final concentration of 0.15 to 1.0 mg/ml prior administration."	
Edurant - rilpivirine - EMEA/H/C/002264/II/0025 MAH: Janssen-Cilag International NV, Rapporteur: Johann Lodewijk Hillege, "Update of section 4.5 of the SmPC in order to include Pharmacokinetics data of drug-drug interactions between simeprevir and rilpivirine, based on final result from study TMC435-TiDP16-C114; this is a Phase I, 2-panel, open-label, randomized, cross-over study in healthy subjects to investigate the potential drug-drug interaction between simeprevir and RPV. In addition, the drug-drug interaction information for telaprevir is removed from the SmPC as this product is no longer available since its marketing authorization was not renewed. The Marketing authorisation holder (MAH) took the opportunity to bring the PI in line with the latest QRD template version 10." Opinion adopted on 06.07.2017. Request for Supplementary Information adopted on 11.05.2017.	Positive Opinion adopted by consensus on 06.07.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Elonva - corifollitropin alfa - EMEA/H/C/001106/II/0034 MAH: Merck Sharp & Dohme Limited, Rapporteur: Paula Boudewina van Hennik, "Update of section 4.5 of the SmPC to add information pertaining to potential hCG cross-reactivity resulting in a false positive pregnancy test. In addition, the MAH is taking the opportunity to implement changes in the annexes in line with the QRD templates (versions 9.1 and 10) and to propose combined versions of the SmPCs and Package Leaflets for the different strengths." Request for Supplementary Information adopted on 18.05.2017.	Weekly start timetable.
EMEND - aprepitant - EMEA/H/C/000527/II/0055 MAH: Merck Sharp & Dohme Limited, Rapporteur: Filip Josephson, "Update of sections 4.2 of the SmPC in order to replace the nomogram for the paediatric formulation provided in ml/kg with purely weight-based	

dosing instructions (in mg/kg) This is based on data that were already submitted as part of the paediatric application X/49. The Package Leaflet is 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."	
Empliciti - elotuzumab - EMEA/H/C/003967/II/0006 MAH: Bristol-Myers Squibb Pharma EEIG, Rapporteur: Paula Boudewina van Hennik, "Update of section 5.2 of the SmPC to update the volume of distribution and elimination of elotuzumab based on an updated analysis of study HuLuc63-1701."	Weekly start timetable.
Esbriet - pirfenidone - EMEA/H/C/002154/II/0043, Orphan MAH: Roche Registration Limited, Rapporteur: Greg Markey, "Update of sections 4.2 and 5.2 of the SmPC in order to update the existing safety information with revised recommendations for patients with moderate renal impairment based on the totality of data from clinical studies; the Package Leaflet is updated accordingly." Request for Supplementary Information adopted on 29.06.2017.	Weekly start timetable.
Forsteo - teriparatide - EMEA/H/C/000425/II/0046 MAH: Eli Lilly Nederland B.V., Rapporteur: Greg Markey, "Update of section 5.1 of the SmPC of the SmPC based on the results of study B3D-EW-GHDW (VERO), a phase 4 multi-centre, prospective, randomized, parallel, double-blind, double-dummy, active controlled study comparing the effect of teriparatide for injection versus risedronate on the incidence of fractures and low bone mass. In addition, the Marketing authorisation holder (MAH) took the opportunity to correct the formatting throughout the Product Information and to bring Annex II in line with the latest QRD template version 10."	Weekly start timetable.
Galafold - migalastat - EMEA/H/C/004059/II/0010, Orphan MAH: Amicus Therapeutics UK Ltd, Rapporteur: Johann Lodewijk Hillege, "Submission of the final report from study AT1001-041: A phase 3 open label extension study to assess the safety and efficacy of 150 mg migalastat HCl QOD in subjects with Fabry disease who have completed	Weekly start timetable.

Studies AT1001-011, AT1001- 012 or FAB-CL-205, listed as a category 3 study in the RMP." Request for Supplementary Information adopted on 13.07.2017.	
Giotrif - afatinib - EMEA/H/C/002280/II/0023 MAH: Boehringer Ingelheim International GmbH, Rapporteur: Filip Josephson, "Update of section 4.8 of the SmPC in order to add the adverse reaction nail disorders with a frequency common based on the results of study 1200.131 and supportive evidence from EGFR TKJ comparator studies. The package leaflet is updated accordingly."	Weekly start timetable.
Harvoni - ledipasvir / sofosbuvir - EMEA/H/C/003850/II/0053 MAH: Gilead Sciences International Ltd, Rapporteur: Filip Josephson, "Update of section 4.5 of the SmPC in order to revise information related to the Cytochrome P450 3A (CYP3A) mediated drug-drug interaction potential of ledipasvir based on final results from study GS-US-337-1887, listed as a category 3 study in the RMP"	Weekly start timetable.
Hetlioz - tasimelteon - EMEA/H/C/003870/II/0008, Orphan MAH: Vanda Pharmaceuticals Ltd., Rapporteur: Greg Markey, "Update of sections 4.4, 4.5 of the SmPC, based on in vitro studies and pooled analyses from clinical studies, with cautionary statements regarding the coadministration of Hetlioz with strong CYP1A2 inhibitors and removal, from the Risk Management Plan (RMP), of the commitment to conduct a human CYP2C19 Drug-Drug Interaction Study to evaluate the single-dose pharmacokinetics of tasimelteon 20 mg alone and in combination with a CYP2C19 inhibitor, omeprazole, at steady-state. Section 5.2 is also updated with data of the pooled analyses of the clinical studies and the in vitro studies." Opinion adopted on 06.07.2017. Request for Supplementary Information adopted on 06.04.2017.	Positive Opinion adopted by consensus on 06.07.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Humira - adalimumab - EMEA/H/C/000481/II/0168 MAH: AbbVie Limited, Rapporteur: Kristina Dunder, "Update of section 5.1 of the SmPC in	Weekly start timetable.

order to update information on the long-term safety, tolerability, and efficacy of adalimumab in subjects with moderate to severe hidradenitis suppurativa after finalization of phase III open-label extension studyM12-555." Request for Supplementary Information adopted on 15.06.2017.	
Humira - adalimumab - EMEA/H/C/000481/II/0169 MAH: AbbVie Limited, Rapporteur: Kristina Dunder, "Update of section 5.1 of the SmPC in order to update the clinical data section based on interim data from the OLE Study M11-327 in non-infectious uveitis (A Multicenter Open-Label Study of the Long-term Safety and Efficacy of the Human Anti-TNF Monoclonal Antibody Adalimumab in Subjects with Non-infectious Intermediate, Posterior, or Panuveitis)"	Weekly start timetable.
Imnovid - pomalidomide - EMEA/H/C/002682/II/0025, Orphan MAH: Celgene Europe Limited, Rapporteur: Robert James Hemmings, "Submission of a biomarker analysis report following a recommendation from CHMP in MAA procedure (EMEA/H/C/2682/0000) to present the biomarker analysis report based on clinical studies CC-4047-MM-008 and CC-4047-MM-010." Request for Supplementary Information adopted on 06.07.2017.	Weekly start timetable.
Intelence - etravirine - EMEA/H/C/000900/II/0050 MAH: Janssen-Cilag International NV, Rapporteur: Joseph Emmerich, "Update of the Product Information (PI) of Etravirine (Intelence®). The PI for Intelence has been updated to include additions to the drug-drug interaction (DDI) information of Etravirine with hepatitis C virus (HCV) direct-acting antivirals (DAAs) and human immunodeficiency virus (HIV) protease inhibitors (PIs) (SmPC sections 4.3 and 4.5 and Patient Information Leafleft (PIL) section 2) and to include a change in the recommended treatment for Intelence overdose (SmPC section 4.9). Specifically, the product information of Intelence is updated with information about the drug-drug interactions of Etravirine with Elbasvir/Grazoprevir (Zepatier), Daclastavir and Simeprevir.	Weekly start timetable.

In addition, the Marketing Authorisation Holder (MAH) took the opportunity to update the PI of Intelence with: Quality Review of Documents (QRD) template v9.1 A combined SmPC for the 3 different strengths of Intelence (25-, 100- and 200 mg tablets) is provided within this submission Although as a result of this update the content of the PI is not impacted, significant text formatting is introduced resulting in a rather complex track changes updated PI. Thus, for clarity purposes, the MAH refers the Agency to the submitted track version PI version within this procedure rather than listing all changes in current/proposed section of the Application form. Quality Review of Documents (QRD) template v10.0 For consistency with section 4.5 of the currently approved Intelence SmPC, guidance related to co-administration with anti-HIV medicines efavirenz, nevirapine, rilpivirine, indinavir, nelfinavir has been added to section 2 of the PIL. The address of the Netherlands Local Operating Company (section 6, PIL) and Minor editorial changes.”	
Invega - paliperidone - EMEA/H/C/000746/II/0056/G MAH: Janssen-Cilag International NV, Rapporteur: Kristina Dunder, “Update of section 4.2 and 4.9 of the SmPC in order to add 3 mg every other day dosing for patients with moderate and severe renal impairment and to delete the recommendation for gastric lavage in accordance with current best practices for management of overdose respectively. Furthermore, the MAH is proposing deletion of INVEGA 1.5 mg strength (all presentations) which has never been marketed in the EU. In addition, the details of the local representatives for Latvia, the Netherlands, Estonia and Lithuania are updated in the PL. The Company also proposes to combine the SmPCs for the different INVEGA strengths (1.5mg, 3mg, 6mg, 9mg, 12mg) in the frame of the alignment of the package leaflet to QRD 10.0.”	
Lenvima - lenvatinib - EMEA/H/C/003727/II/0008, Orphan MAH: Eisai Europe Ltd., Rapporteur: Bart Van der	Weekly start timetable.

Schueren, PRAC Rapporteur: Ulla Wändel Liminga, “Submission of Clinical Study Report for Study E78080-J081-208” Request for Supplementary Information adopted on 06.04.2017.	
Norvir - ritonavir - EMEA/H/C/000127/II/0146 MAH: AbbVie Limited, Rapporteur: Johann Lodewijk Hillege, “Update of section 4.6 of the SmPC in order to update the safety information on pregnancy and lactation based on the company’s core data sheet information.”	Weekly start timetable.
Norvir - ritonavir - EMEA/H/C/000127/II/0147 MAH: AbbVie Limited, Rapporteur: Johann Lodewijk Hillege, “Update of section 4.3 and 4.5 of the SmPC in order to add a contraindication regarding the interaction between ritonavir and venetoclax based on the company’s core data sheet. The Package Leaflet is updated accordingly to also include some minor editorial updates.”	Weekly start timetable.
Ofev - nintedanib - EMEA/H/C/003821/II/0016, Orphan MAH: Boehringer Ingelheim International GmbH, Rapporteur: David Lyons, “Update of section 4.4 of the SmPC to amend the current warning on the hepatic function to include low body weight, Asian origin, female sex and age as factors of increased risk of liver enzymes elevations, update of section 4.8 of the SmPC to revised the frequency of the ADR ‘drug-induced liver injury’ (DILI) from ‘not known’ to ‘uncommon’ and update of section 5.2 of the SmPC to amend the current information related to the mean exposure to nintedanib by race, based on a review of clinical trials and post-marketing data on DILI and on the exposure safety relationship between nintedanib plasma exposure and liver enzyme elevations. The Package Leaflet is updated accordingly.”	Weekly start timetable.
Pradaxa - dabigatran etexilate - EMEA/H/C/000829/II/0103 MAH: Boehringer Ingelheim International GmbH, Rapporteur: Hanne Lomholt Larsen, “Update of sections 4.2, 4.4 and 5.1 to reflect the final study results of the phase IV study 1160.204 (The RE-CIRCUIT Trial), “ A Randomised Evaluation of dabigatran etexilate Compared to warfar/n inpulmonaRy vein ablation: assessment of an	

uninterrupted periproCedUralant/coagulation sTrategy””	
Pyramax - pyronaridine / artesunate - EMEA/H/W/002319/II/0015 MAH: Shin Poong Pharmaceutical Co., Ltd., Rapporteur: Joseph Emmerich, “Update of sections 4.8 and 5.1 of the SmPC in order to reflect the main efficacy and safety results from the final report from study SP-C-013-11 listed as a category 3 study in the RMP. This is a phase IIIb/IV comparative, randomised, multi-centre, open label, parallel 3-arm clinical study to assess the safety and efficacy of repeated administration of pyronaridine-artesunate, dihydroartemisinin-piperaquine or artemether-lumefantrine or artesunate-amodiaquine over a 2-year period in children and adult patients with acute uncomplicated Plasmodium sp. malaria. In addition, the SOH took the opportunity to make some editorial changes to the product information.” Request for Supplementary Information adopted on 11.05.2017.	Weekly start timetable.
Selincro - nalmefene - EMEA/H/C/002583/II/0020/G MAH: H. Lundbeck A/S, Rapporteur: Harald Enzmann, “Update of section 4.7 of the SmPC to add new information regarding effects on ability to drive and use machines, based on clinical study and post-marketing data. Update of section 4.8 of the SmPC in order to add the adverse drug reaction “diarrhoea” with frequency “common”, based on clinical study and post-marketing data. 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.” Opinion adopted on 06.07.2017. Request for Supplementary Information adopted on 01.06.2017.	Positive Opinion adopted by consensus on 06.07.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Sirturo - bedaquiline - EMEA/H/C/002614/II/0021, Orphan MAH: Janssen-Cilag International NV, Rapporteur: Filip Josephson, “Update of section 4.4 of the SmPC in order to add delamanid as an example of a drug that prolongs the QT interval following the review of the global safety database	Positive Opinion adopted by consensus on 29.06.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

for all serious cases received from 28 December 2012 to 30 September 2016. 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." Opinion adopted on 29.06.2017. Request for Supplementary Information adopted on 18.05.2017, 06.04.2017.	
Tafinlar - dabrafenib - EMEA/H/C/002604/II/0024 MAH: Novartis Europharm Ltd, Rapporteur: Filip Josephson, "Update of section 4.5 of the SmPC in order to include some warning on a drug-drug interaction between dabrafenib and rifampicin (a CYP3A4/CYP2C8 inducer) and between dabrafenib and rabeprazole (a pH elevating agent), based on the final results of study 200072, a phase I open-label fixed sequence study to evaluate the effects of potent CYP3A4 inducer (rifampicin) and of a pH elevating agent (rabeprazole) on the repeat dose pharmacokinetics of dabrafenib in subjects with BRAFV60 mutation positive tumours, to fulfil MEA 005." Opinion adopted on 06.07.2017.	Positive Opinion adopted by consensus on 06.07.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Tagrisso - osimertinib - EMEA/H/C/004124/II/0014/G MAH: AstraZeneca AB, Rapporteur: Jorge Camarero Jiménez, "Update of section 5.3 of the SmPC to include information regarding CNS distribution based on non-clinical data." Opinion adopted on 06.07.2017.	Positive Opinion adopted by consensus on 06.07.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Tagrisso - osimertinib - EMEA/H/C/004124/II/0015 MAH: AstraZeneca AB, Rapporteur: Jorge Camarero Jiménez, "Update of section 5.2 of the SmPC to include data from studies performed to investigate human plasma protein binding (Study No. BS001265-53-AZD9291), the assessment of non-specific incubational binding in transporter inhibition assays (Study No. BS000760-92) and the implications on transporter DDI risk assessment. In addition, the MAH took the opportunity to implement minor updates and editorial changes in the SmPC and to update the address of the MAH and manufacturer in SmPC section 7, the labelling and the Package Leaflet."	Positive Opinion adopted by consensus on 06.07.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

Opinion adopted on 06.07.2017.	
Tarceva - erlotinib - EMEA/H/C/000618/II/0052 MAH: Roche Registration Limited, Rapporteur: Sinan B. Sarac, “Update of section 4.4 of the SmPC in order to include recommendations on Epidermal Growth Factor Receptor (EGFR) mutation status testing, to be in line with current technical and scientific progress. In addition, the Marketing authorisation holder (MAH) took the opportunity to make minor editorial changes and to bring the PI in line with the latest QRD template version 10. Moreover, the MAH took the opportunity to make minor correction of section 4.2 of the SmPC. Furthermore, the Annex II has been corrected, as requested by the EMA, to include Educational Material as an additional risk minimisation measure, which has been already in place in the RMP.”	
Torisel - temsirolimus - EMEA/H/C/000799/II/0066, Orphan MAH: Pfizer Limited, Rapporteur: Harald Enzmann, “Submission of the further analysis of a possible association of corticosteroid (pre-)treatment and frequency and severity of hypersensitivity/infusion reactions in study 3066K1-4438-WW (B1771007), as requested by the CHMP during procedures EMEA/H/C/799/MEA 023.1 and EMEA/H/C/799/MEA 024.1.” Opinion adopted on 06.07.2017. Request for Supplementary Information adopted on 16.03.2017.	Positive Opinion adopted by consensus on 06.07.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Victrelis - boceprevir - EMEA/H/C/002332/II/0042 MAH: Merck Sharp & Dohme Limited, Rapporteur: Joseph Emmerich, “Update of section 4.4 of the SmPC to add a warning regarding HBV reactivation observed in patients with HCV/HBV co-infection treated with some direct-acting antivirals not given in combination with peginterferon alfa and ribavirin. The Package Leaflet is updated accordingly. In addition, the MAH took the opportunity to implement minor editorial updates in the Product Information.” Opinion adopted on 29.06.2017.	Positive Opinion adopted by consensus on 29.06.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Xarelto - rivaroxaban - EMEA/H/C/000944/II/0050	

MAH: Bayer AG, Rapporteur: Kristina Dunder, "Update of the Summary of Product Characteristics (SmPC) to add a new posology in the patients with non valvular atrial fibrillation and information on safety and efficacy in patients who undergo PCI (percutaneous coronary intervention) with stent placement based on the final results of study 16523 (PIONEER AF-PCI): An Open-label, Randomized, Controlled, Multicenter Study Exploring Two Treatment Strategies of Rivaroxaban and a Dose- Adjusted Oral Vitamin K Antagonist Treatment Strategy in Subjects With Atrial Fibrillation Who Undergo Percutaneous Coronary Intervention. Sections 4.2, 4.4 and 5.1 of the SmPC are updated. Package Leaflet is updated accordingly.
In addition, the Marketing authorisation holder took the opportunity to update the telephone number of local representatives for UK in the Package Leaflet."
Request for Supplementary Information adopted on 22.06.2017, 21.04.2017.

Xeplion - paliperidone - EMEA/H/C/002105/II/0035

MAH: Janssen-Cilag International NV, Rapporteur: Kristina Dunder, "Update of section 4.2 of the SmPC in order to add a dosage conversion table to provide guidance for healthcare professionals when switching patients from paliperidone ER tablets to paliperidone palmitate long acting injection (PP1M). The Package Leaflet is updated accordingly."

WS1092
Ebymect-EMEA/H/C/004162/WS1092/0017
Edistride-EMEA/H/C/004161/WS1092/0013
Forxiga-EMEA/H/C/002322/WS1092/0032
Xigduo-EMEA/H/C/002672/WS1092/0028

MAH: AstraZeneca AB, Lead Rapporteur: Kristina Dunder, "Update of sections 4.4 and 5.1 of the SmPC in order to reflect the results of the Phase 3 study D5553C00003: 28-week safety and efficacy, randomised, double-blind comparison of simultaneous administration of exenatide once weekly 2 mg an dapagliflozin once daily 10 mg to exenatide once weekly 2 mg alone and dapagliflozin once daily 10 mg alone in patients with type 2 diabetes with inadequate glycaemic

control on metformin. The Package Leaflet is updated accordingly. In addition, the Worksharing applicant (WSA) took the opportunity to update the list of local representatives in the Package Leaflets for Ebymect and Edistride and to introduce minor editorial changes throughout the Product Informations." Request for Supplementary Information adopted on 23.03.2017.	
WS1136 **Descovy-EMEA/H/C/004094/WS1136/0017** **Genvoya-EMEA/H/C/004042/WS1136/0031** **Odefsey-EMEA/H/C/004156/WS1136/0013** MAH: Gilead Sciences International Ltd, Lead Rapporteur: Robert James Hemmings, "Update of sections 4.4, 4.8. 5.1 and 5.2 of the SmPC in order to provide 48 weeks data from Study GS-US-292-1249; this is a Phase 3b open-label study of the efficacy and safety of elvitegravir/cobicistat/emtricitabine/tenofovir alafenamide single-tablet regimen in HIV-1/Hepatitis B co-infected adults. The Package Leaflet is updated accordingly. In addition, the Worksharing applicant (WSA) took the opportunity to make minor administrative changes in the SmPC and the Package Leaflet." Opinion adopted on 06.07.2017. Request for Supplementary Information adopted on 05.05.2017.	Positive Opinion adopted by consensus on 06.07.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1144/G **Afinitor-EMEA/H/C/001038/WS1144/0052/G** **Votubia-EMEA/H/C/002311/WS1144/0042/G** MAH: Novartis Europharm Ltd, Lead Rapporteur: Harald Enzmann, "Update of sections 4.4 and 5.1 of the SmPC in order to include new safety information on stomatitis and its management based on final results from study CRAD001JUS226: a phase II, single arm study of the use of steroid-based mouthwash to prevent stomatitis in postmenopausal women with advanced or metastatic hormone receptor	Positive Opinion adopted by consensus on 29.06.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

positive breast cancer being treated with everolimus plus exemestane Update of section 4.6 of the SmPC in order to add new information on breast-feeding based on pre-clinical data. The Package Leaflets were updated accordingly. In addition, the Worksharing applicant (WSA) took the opportunity to bring the Afinitor PI in line with the latest QRD template version 10." Opinion adopted on 29.06.2017. Request for Supplementary Information adopted on 27.04.2017.	
WS1167 **Ebymect-EMEA/H/C/004162/WS1167/0021** **Edistride-EMEA/H/C/004161/WS1167/0016** **Forxiga-EMEA/H/C/002322/WS1167/0036** **Xigduo-EMEA/H/C/002672/WS1167/0032** MAH: AstraZeneca AB, Lead Rapporteur: Kristina Dunder, "Update of sections 4.8 and 5.1 of the SmPC in order to add information regarding two initial combination studies (MB102021 and MB102034) in treatment-naïve patients of dapagliflozin 5 mg + metformin and dapagliflozin 10 mg + metformin, respectively, compared to each component separately. In addition, the Worksharing applicant (WSA) took the opportunity to bring the PI in line with the latest QRD template version 10."	Weekly start timetable.
WS1178 **Aluvia-EMEA/H/W/000764/WS1178/0102** **Kaletra-EMEA/H/C/000368/WS1178/0164** MAH: AbbVie Limited., Lead Rapporteur: Joseph Emmerich, "Update of sections 4.3 and 4.5 of the SmPC in order to add new contraindications regarding the interaction of lopinavir/ritonavir with venetoclax, elbasvir/grazoprevir and to add new information on the interaction with ombitasvir/paritaprevir/ritonavir based on the company's core data sheet. The package Leaflet is updated accordingly. In addition, the MAH/SOH is taking the opportunity to update section 4.5 of the SmPC to reflect information already contained in section 4.3 for the following drug-drug interactions: astemizole, terfenadine, pimozide, ergot	Weekly start timetable.

alkaloids and cisapride.”	
WS1179 **Invega-EMEA/H/C/000746/WS1179/0055** **Trevicta-EMEA/H/C/004066/WS1179/0010** **Xeplion-EMEA/H/C/002105/WS1179/0034** MAH: Janssen-Cilag International NV, Lead Rapporteur: Kristina Dunder, “Update of section 4.6 (Fertility, pregnancy and lactation) of the SmPC in order to add new information concerning a retrospective observational cohort study with risperidone and risk of congenital malformations. Nationally approved products are also affected by this variation.”	Weekly start timetable.
WS1189 **ANORO-EMEA/H/C/002751/WS1189/0017** **Laventair-EMEA/H/C/003754/WS1189/0019** MAH: Glaxo Group Ltd, Lead Rapporteur: Nithyanandan Nagercoil, “Update of section 4.8 of the SmPC and relevant section of the PL to add “dysphonia” with rare frequency.”	Weekly start timetable.
WS1191 **Incruse-EMEA/H/C/002809/WS1191/0016** **Rolufta-EMEA/H/C/004654/WS1191/0001** MAH: Glaxo Group Ltd, Lead Rapporteur: Concepcion Prieto Yerro, “Update of section 4.8 of the SmPC and relevant section of the PL to add "Eye pain" with a rare frequency.”	Weekly start timetable.

B.5.3. CHMP-PRAC assessed procedures

Adcetris - brentuximab vedotin - EMEA/H/C/002455/II/0043, Orphan MAH: Takeda Pharma A/S, Rapporteur: Paula Boudewina van Hennik, PRAC Rapporteur: Sabine Straus, “Update of sections 4.8 and 5.1 of the SmPC in order to add data from study C25007. The RMP (version 8.0) was updated accordingly. The submission of the clinical study report fulfils SOB 011 of the conditional marketing authorisation for Adcetris.” Request for Supplementary Information adopted on 18.05.2017, 23.02.2017.	
Aranesp - darbepoetin alfa - EMEA/H/C/000332/II/0141 MAH: Amgen Europe B.V., Rapporteur: Martina	

Weise, PRAC Rapporteur: Valerie Strassmann, "Update of sections 4.4 and 4.8 of the SmPC in order to add a warning on severe cutaneous conditions including Erythema multiforme and Stevens-Johnson Syndrome (SJS) and Toxic Epidermal Necrolysis (TEN) following a request for cumulative review triggered by EMA signal adopted by PRAC on 09 February 2017. The Package Leaflet is updated accordingly. The RMP version 7 has also been submitted."	
Cervarix - human papillomavirus vaccine [types 16, 18] (recombinant, adjuvanted, adsorbed) - EMEA/H/C/000721/II/0085 MAH: GlaxoSmithkline Biologicals SA, Rapporteur: Bart Van der Schueren, Co-Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Jean-Michel Dogné, "Submission of Study EPI-HPV-069, a meta-analysis assessing the risk of three autoimmune diseases following vaccination with Cervarix: autoimmune thyroiditis (AIT), Guillain-Barre Syndrome (GBS) and Inflammatory Bowel Disease (IBD). The EPI-HPV-069 study is a post-licensure commitment to the EMA (PASS register number EUPAS13332). As part of this submission, an updated RMP (version 18) is provided, including changes related to the EPI-HPV-069 meta-analysis submitted and minor updates related to other studies." Request for Supplementary Information adopted on 18.05.2017, 15.12.2016. Clockstop extension of 3 months requested to respond to RSI, responses expected 15.08.2017. For adoption.	
Dificlir - fidaxomicin - EMEA/H/C/002087/II/0028 MAH: Astellas Pharma Europe B.V., Rapporteur: Filip Josephson, PRAC Rapporteur: Qun-Ying Yue, "C.I.11: Submission of an updated RMP version 7 in order to remove the post-authorization measure (PAM) MEA003 (concerning clinical study 2819-CL-2001 in patients with Clostridium difficile Infection who will receive a second course offidaxomicin) due to the non-feasibility of the study." Request for Supplementary Information adopted on 05.05.2017.	Weekly start timetable.
Edurant - rilpivirine -	Positive Opinion adopted by consensus on

EMEA/H/C/002264/II/0024 MAH: Janssen-Cilag International NV, Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Menno van der Elst, "Update of sections 4.2, 4.4, 4.6, 5.1 and 5.2 of the SmPC in order to include information: use of rilpivirine in combination with a background regimen for the treatment of HIV-1 infection during pregnancy and postpartum, without dose adjustment following final results from study TMC114HIV3015 listed as a category 3 study in the RMP. This is a single arm, open-label trial to assess the pharmacokinetics of darunavir/ritonavir, etravirine, and rilpivirine in HIV-1-infected pregnant women. The Package Leaflet is updated accordingly. The RMP version 7.0 has also been submitted. In addition, the Marketing authorisation holder (MAH) took the opportunity to introduce the latest renewal date in section 9 of the SmPC and the physical address of the Netherlands Local Operating Company in the PIL section 6." Opinion adopted on 06.07.2017. Request for Supplementary Information adopted on 09.06.2017, 06.04.2017.	06.07.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Jinarc - tolvaptan - EMEA/H/C/002788/II/0006 MAH: Otsuka Pharmaceutical Europe Ltd, Rapporteur: Greg Markey, PRAC Rapporteur: Julie Williams, "Update of section 5.1 of the SmPC based on final results from study 156-08-271 (TEMPO 4:4) listed as a PAES in Annex II. This study is a Multicenter, Open-label, Extension Study (Extension of Trial 156-04-251) to Evaluate the Long-term Safety and Efficacy of Oral Tolvaptan Tablet Regimens in Patients With Autosomal Dominant Polycystic. It provides data for Jinarc treatment of autosomal dominant polcystic kidney disease (ADPKD) over 5 years. Reference to submission of this study is being deleted from Annex II. In addition, the Marketing authorisation holder (MAH) took the opportunity to add the current ATC code applicable for tolvaptan as it has been assigned by by WHO. The RMP version 13.1 has also been submitted to reflect the completion of the 156-08-271 study." Opinion adopted on 06.07.2017. Request for Supplementary Information adopted	Positive Opinion adopted by consensus on 06.07.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

on 05.05.2017.	
Keytruda - pembrolizumab - EMEA/H/C/003820/II/0028 MAH: Merck Sharp & Dohme Limited, Rapporteur: Daniela Melchiorri, PRAC Rapporteur: Sabine Straus, "Update sections 4.4 and 4.8 of the SmPC to include the risk of myocarditis that has been reported in patients treated with pembrolizumab. The Package Leaflet has been updated accordingly. An updated RMP version 10.0 was provided as part of the application." Request for Supplementary Information adopted on 22.06.2017.	
Keytruda - pembrolizumab - EMEA/H/C/003820/II/0029 MAH: Merck Sharp & Dohme Limited, Rapporteur: Daniela Melchiorri, PRAC Rapporteur: Sabine Straus, "Submission of the final study report for non-clinical study "Anti-Murine PD-1 Antibody (muDX400 L-005571333): Exploratory Multiple-Dose Subcutaneous Immunotoxicity Study in Mice with Hepatitis B Vaccine (L-005552770)". An updated RMP version 11.0 was agreed during the procedure." Opinion adopted on 06.07.2017.	Positive Opinion adopted by consensus on 06.07.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Latuda - lurasidone - EMEA/H/C/002713/II/0016 MAH: Sunovion Pharmaceuticals Europe Ltd, Rapporteur: Filip Josephson, PRAC Rapporteur: Qun-Ying Yue, "Submission of the final CSR for Study D1001057, an extension of study of SM-13496 evaluating the long-term safety and efficacy of lurasidone (40 mg/day or 80 mg/day) in Pan-Asian (Japanese, Korean, Taiwanese and Malaysian) subjects with schizophrenia. The RMP is being updated (ver. 5.0) with information relative to this study and also information relative to Study D1050301, which has already been assessed in P46/006." Opinion adopted on 06.07.2017.	Positive Opinion adopted by consensus on 06.07.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Lemtrada - alemtuzumab - EMEA/H/C/003718/II/0017 MAH: Genzyme Therapeutics Ltd, Duplicate, Duplicate of Lemtrada (WD), Rapporteur: Hanne Lomholt Larsen, PRAC Rapporteur: Torbjorn Callreus, "Update of sections 4.2, 4.4, 4.8 and 5.1 of the SmPC in order to update the safety and	

long term use information in the posology following final results from study CAMMS03409 - An Extension Protocol For Multiple Sclerosis Patients Who Participated in Genzyme-Sponsored Studies of Alemtuzumab (ongoing at the time of the initial MAA) to evaluate the long term safety and efficacy of alemtuzumab in MS patients who received alemtuzumab during prior company-sponsored studies. The RMP version 3.0 has also been submitted. The PL has been 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 introduce editorial corrections in the PI." Request for Supplementary Information adopted on 21.04.2017.	
Mozobil - plerixafor - EMEA/H/C/001030/II/0032, Orphan MAH: Genzyme Europe BV, Rapporteur: Paula Boudewina van Hennik, PRAC Rapporteur: Sabine Straus, "Update of sections 4.2 and 5.2 of the SmPC in order to reflect the results of the completed study MSC12830 (MOZ11809), entitled "A Phase 4, Multicenter, Randomized, Comparator Trial Evaluating the Standard Weight-Based Dose (0.24 mg/kg) Compared to a Fixed Dose (20 mg) of Plerixafor Injection in Combination with G-CSF to Mobilize and Collect ≥5 x 106 CD34+ cells/kg in ≤4 Days and to Evaluate the Difference in Total Systemic Exposure in Patients with Non-Hodgkin's Lymphoma Weighing ≤70 kg" listed as a category 3 study in the RMP. The Package Leaflet is updated accordingly. The RMP version 9.0 has also been submitted. In addition, the Marketing authorisation holder (MAH) took the opportunity to update the list of local representatives in the Package Leaflet." Request for Supplementary Information adopted on 06.07.2017.	Weekly start timetable.
Ninlaro - ixazomib - EMEA/H/C/003844/II/0002, Orphan MAH: Takeda Pharma A/S, Rapporteur: Greg Markey, PRAC Rapporteur: Ulla Wändel Liminga, "Update of sections 4.8 and 5.1 of the SmPC to reflect the final overall survival analysis of C16010 China continuation study, a phase III	Positive Opinion adopted by consensus on 06.07.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

study comparing ixazomib plus lenalidomide and dexamethasone versus placebo plus lenalidomide in patients with relapsed and/or refractory multiple myeloma, in order to fulfil SOB (Specific Obligation) 002. Annex II.E and the RMP (version 2.0) are updated accordingly. In addition the Marketing Authorisation Holder (MAH) took the opportunity to make a small correction in sections 4.7 and 9 of the SmPC and to the German translations." Opinion adopted on 06.07.2017. Request for Supplementary Information adopted on 05.05.2017.	
Odomzo - sonidegib - EMEA/H/C/002839/II/0011 MAH: Sun Pharmaceutical Industries Europe B.V., Rapporteur: Paula Boudewina van Hennik, PRAC Rapporteur: Patrick Batty, "Submission of the final results from studies CLDE225C2301 and CLDE225X2104. Study CLDE225C2301 is a phase II, multi-center, open-label, single-arm study of the efficacy and safety of oral LDE225 in patients with Hhpathway activated relapsed medulloblastoma. Study CLDE225X2104 is a Phase I/II study of LDE225 in pediatric patients with recurrent or refractory medulloblastoma or other tumors potentially dependent on the Hedgehog-signaling pathway and adult patients with recurrent or refractory medulloblastoma. The RMP has been updated accordingly. The product information remains unchanged." Opinion adopted on 06.07.2017. Request for Supplementary Information adopted on 09.06.2017.	Positive Opinion adopted by consensus on 06.07.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Olumiant - baricitinib - EMEA/H/C/004085/II/0001 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 add a warning on venous thromboembolism based on analyses of the occurrence of venous thromboembolic events in clinical trials with baricitinib. The Package Leaflet is updated accordingly. The RMP version 2.0 has been submitted, as part of this application."	
Olysio - simeprevir - EMEA/H/C/002777/II/0031	

MAH: Janssen-Cilag International NV, Rapporteur: Jorge Camarero Jiménez, PRAC Rapporteur: Julie Williams, “Update of section 5.1 of the SmPC in order to update the efficacy information following results from study HPC3002 A Prospective 3-year Follow-up Study in Subjects Previously Treated in a Phase IIb or Phase III Study with a TMC435-containing Regimen for the Treatment of Hepatitis C Virus (HCV) Infection listed as a category 3 study in the RMP and in fulfilment of MEA005. The RMP version 4.0 has also been submitted which includes updates of changes already agreed in procedures EMEA/H/C/002777/II/0021,EMEA/H/C/002777/II/0027 and EMEA/H/A-20/1438/C/2777/0019.” Request for Supplementary Information adopted on 18.05.2017.

Perjeta - pertuzumab - EMEA/H/C/002547/II/0029

MAH: Roche Registration Limited, Rapporteur: Sinan B. Sarac, PRAC Rapporteur: Doris Stenver, “Update of sections 4.2, 4.4, 4.8, 5.1 of the SmPC, annex II and relevant sections of the PL in order to update information on cardiac safety and reflect the results from study BERNICE (WO29217) listed as a specific obligation in the Annex II; BERNICE is an ongoing Multicenter, Multinational, Phase II Study to Evaluate Perjeta in Combination with Herceptin and Standard Neoadjuvant Anthracycline-Based Chemotherapy in Patients with HER2- Positive, Locally Advanced, Inflammatory, or Early-Stage Breast Cancer.
The RMP v.9 has also been submitted.” Request for Supplementary Information adopted on 22.06.2017.

Praxbind - idarucizumab - EMEA/H/C/003986/II/0007

MAH: Boehringer Ingelheim International GmbH, Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Menno van der Elst, “Update of sections 4.2, 4.8, 5.1 and 5.2 of the SmPC in order to reflect the final results from a study 1321.3 titled “A Phase III, case series clinical study of the reversal of the anticoagulant effects of dabigatran by intravenous administration of 5.0 g idarucizumab (BI 655075) in patients treated with dabigatran etexilate who have uncontrolled bleeding or require emergency

surgery or procedures. RE-VERSE-AD (A study of the RE-VERSal Effects of Idarucizumab on Active Dabigatran) trial" listed as a category 3 study in the RMP (MEA 001). The RMP version 3.0 has also been submitted. In addition, the Marketing authorisation holder took the opportunity to update the immunogenicity section in 5.1 of SmPC and to bring the PI in line with the latest QRD template version 10."	
Privigen - human normal immunoglobulin - EMEA/H/C/000831/II/0119 MAH: CSL Behring GmbH, Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Brigitte Keller-Stanislawski, "Update of sections 4.8 and 5.1 of the SmPC to include the PATH (IgPro20_3003) study results (safety & efficacy study with chronic inflammatory demyelinating polyneuropathy (CIDP) patients). Minor changes are also introduced to section 4.2 of the SmPC. In addition, the MAH took the opportunity to make some editorial changes to sections 4.3 and 5.2 of the SmPC. The Package leaflet and the RMP (finally agreed version 5.1) were updated accordingly." Opinion adopted on 06.07.2017.	Positive Opinion adopted by consensus on 06.07.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
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 3.0) has also been submitted, proposing to collect safety information regarding tedizolid phosphate by conducting three investigator initiated studies and deleting the original proposed long term safety study. The MAH also took the opportunity	Weekly start timetable.

to make minor editorial corrections throughout the product information." Request for Supplementary Information adopted on 06.07.2017.	
Sylvant - siltuximab - EMEA/H/C/003708/II/0023, Orphan MAH: Janssen-Cilag International NV, Rapporteur: Concepcion Prieto Yerro, PRAC Rapporteur: Brigitte Keller-Stanislawski, "Submission of the final report from study CNTO328SMM2001 listed as a category 3 study in the RMP This is a 'Phase 2, Randomized, Double-blind, Placebo-controlled, Multicenter Study of Siltuximab (Anti IL-6 Monoclonal Antibody) in Subjects with High-risk Smoldering Multiple Myeloma' to evaluate immunogenicity data. No changes to the PI are proposed. The RMP is being updated accordingly." Opinion adopted on 06.07.2017.	Positive Opinion adopted by consensus on 06.07.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Synflorix - pneumococcal polysaccharide conjugate vaccine (adsorbed) - EMEA/H/C/000973/II/0117 MAH: GlaxoSmithkline Biologicals SA, Rapporteur: Kristina Dunder, PRAC Rapporteur: Qun-Ying Yue, "Update of sections 4.2, 4.4 and 5.1 of the SmPC in order to reflect the results from study10PN-PD-DIT-072, a phase III, open, controlled, multi-centric study to evaluate the immunogenicity, safety and reactogenicity of Synflorix in children at an increased risk of pneumococcal infection. The Package Leaflet is updated accordingly. An updated RMP version 16 has also been submitted. This submission fulfils the post-authorisation measure MEA 065." Request for Supplementary Information adopted on 06.07.2017.	Weekly start timetable.
Tagrisso - osimertinib - EMEA/H/C/004124/II/0016 MAH: AstraZeneca AB, Rapporteur: Jorge Camarero Jiménez, PRAC Rapporteur: Sabine Straus, "Provision of the final CSR for Study Aura 17; a phase II, open label, single-arm study to assess the safety and efficacy of AZD9291 in Asia pacific patients with locally advanced/metastatic non-small cell lung cancer whose disease has progressed with previous epidermal growth factor receptor tyrosine kinase inhibitor therapy and whose tumours harbour a T790M mutation within the epidermal growth factor receptor	Positive Opinion adopted by consensus on 06.07.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

gene). An updated RMP version 7.0 was agreed during the procedure."
Opinion adopted on 06.07.2017.

Tamiflu - oseltamivir - EMEA/H/C/000402/II/0128

MAH: Roche Registration Limited, Rapporteur: Outi Mäki-Ikola, PRAC Rapporteur: Kirsti Villikka, "Update of section 4.6 of the SmPC in order to reflect the final study results from non-interventional safety study BV29684, which assessed the safety of oseltamivir exposure in pregnant women, and was listed as a category 3 study in the RMP (MEA099). The RMP version 15.0 has also been updated to reflect the study results."

Tyverb - lapatinib - EMEA/H/C/000795/II/0048/G

MAH: Novartis Europharm Ltd, Rapporteur: Filip Josephson, PRAC Rapporteur: Ulla Wändel Liminga, "1) C.I.4 (type II): Update of sections 4.4, 4.8, and 5.1 of the SmPC in order to add a warning on QTc prolongation and update safety information following the submission of study report EGF114271 (A Phase IV placebo controlled single sequence crossover study to evaluate the effect of repeat oral doses of lapatinib on cardiac repolarization in patients with advanced cancer). The Package Leaflet is updated accordingly.
2) C.I.4 (type II): Update of section 4.8 of the SmPC in order to further elaborate on the undesirable effect 'serious cutaneous reactions' based on the review of the Novartis safety database. The Package Leaflet is 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.
Moreover, the MAH took the opportunity to update Annex II to delete an Annex II condition which has been fulfilled with procedure ANX. 28.2.
The RMP (version 32) is updated accordingly to the scopes presented above and also to introduce template-related changes, study milestones updates, and to upgrade 'food effect' to an important identified risk (from procedure EMEA/H/C/000795/II/0024)."
Request for Supplementary Information adopted on 23.03.2017, 10.11.2016.

Uptravi - selexipag - EMEA/H/C/003774/II/0009 MAH: Actelion Registration Ltd., Rapporteur: Martina Weise, PRAC Rapporteur: Julie Williams, "Update of section 4.5 of the SmPC to add information on the effect of selexipag administration on the pharmacokinetics of midazolam, its metabolite 1-hydroxymidazolam and the CYP3A4 metabolism, based on data from the completed clinical pharmacology study AC-065-114, a single-centre, open-label, randomized, two-treatment crossover Phase 1 study investigating the effect of selexipag on the pharmacokinetics of midazolam and its metabolite 1-hydroxymidazolam in healthy male subjects. An updated RMP (version 5.1) has also been submitted, to add the results of study AC-065-114, reclassify 'hyperthyroidism' as an important identified risk and update the PASS timelines and protocol versions in accordance with the current EXPOSURE protocol version 3 and the EDUCATE protocol version 2." Opinion adopted on 06.07.2017.	Positive Opinion adopted by consensus on 06.07.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Xalkori - crizotinib - EMEA/H/C/002489/II/0049/G MAH: Pfizer Limited, Rapporteur: Alexandre Moreau, PRAC Rapporteur: Ghania Chamouni, "Submission of the final results related of the Non_Interventional Post-Authorisation Safety Study (PASS) A8081049 "A cross-sectional study to evaluate the effectiveness of XALKORI (PF_02341066, also referred to as crizotinib) Therapeutic Management Guide among physicians prescribing XALKORI in Europe" and PASS A8081050 "A cross-sectional study to evaluate the effectiveness of XALKORI Patient Information Brochure among non-small cell lung cancer (NSCLC) patients receiving XALKORI treatment in Europe". In the light of the results from PASS study A8081049, the MAH is proposing to update Annex II to remove the XALKORI TMG from the Educational Materials. The MAH is also taking the opportunity to state "monotherapy" in section 4.1 of the SmPC as requested by CHMP and to bring the PI in line with the latest QRD template." Opinion adopted on 06.07.2017.	Positive Opinion adopted by consensus on 06.07.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Xarelto - rivaroxaban - EMEA/H/C/000944/II/0052/G	

MAH: Bayer AG, Rapporteur: Kristina Dunder, PRAC Rapporteur: Qun-Ying Yue, “Group of variations consisting of:
1) C.1.4. To add the authorised indications “Treatment of deep vein thrombosis (DVT) and pulmonary embolism (PE), and prevention of recurrent DVT and PE in adults” to Xarelto 10 mg based on Einstein Choice trial (A randomised phase III clinical study to evaluate efficacy and safety of Reduced-dosed rivaroxaban and standard-dosed rivaroxaban versus ASA in the long-term prevention of recurrent symptomatic venous thromboembolism in patients with symptomatic deep-vein thrombosis and/or pulmonary embolism) in section 4.1 of the SmPC 10 mg.
Consequently:
- Changes in sections 4.2, 4.8 and 5.1 for Xarelto 10mg, 15mg and 20 mg are made in order to update the posology, efficacy and safety information.
- Annex III is updated to include Xarelto 10 mg into Patient alert card to support management of bleeding when the 10 mg is treated for long-term prevention of recurrent VTE
- RMP (version 10) is updated
2) B.II.e.5.a.1- to add a new pack size of 14 film coated tablets in blister (PP/alu) for Xarelto 10 mg
3) B.II.e.5.a.1- to add a new pack size of 28 film coated tablets in blister (PP/alu) for Xarelto 10 mg
4) B.II.e.5.a.1- to add a new pack size of 98 film coated tablets in blister (PP/alu) for Xarelto 10 mg
5) B.II.e.1.b.1 to change immediate packaging of the finished product for 10 mg film coated tablets to introduce HDPE bottle with screw cap including new presentation (pack containing 100 film coated tablets for 10 mg strength)
6) C.1.4 To add information on interactions with SSRIs and SNRIs in section 4.5 and a related warning in section 4.4 of the SmPC based on post-hoc analyses to investigate bleeding risk for rivaroxaban in patients with and without use of SSRI or SNRIs from the pivotal studies.
In addition, MedDRA terminology is updated in the adverse drug reactions table in section 4.8 of the SmPC
7) C.1.11.z To delete from the summary of safety concerns: “Patients undergoing major

orthopaedic surgery other than elective hip or knee replacement surgery" and "Remedial pro-coagulant therapy for excessive haemorrhage". Part II - Modules SVIII: Summary of the safety concerns, Part III, Section 1 Safety Concerns and overview of planned pharmacovigilance action were amended accordingly. In addition, Part II, Safety Specification, module SIV, Populations not studied in clinical trials: "Patients undergoing major orthopaedic surgery other than elective hip or knee replacement surgery" and "Remedial pro-coagulant therapy for excessive haemorrhage" was updated.

Request for Supplementary Information adopted on 18.05.2017.

Xgeva - denosumab - EMEA/H/C/002173/II/0051

MAH: Amgen Europe B.V., Rapporteur: Kristina Dunder, PRAC Rapporteur: Ulla Wändel Liminga, "To update to the Risk Management Plan (RMP) with an newly categorised important potential risk of hypercalcemia following treatment discontinuation in patients other than those with growing skeletons following an updated safety assessment of the risk of hypercalcaemia following denosumab discontinuation conducted earlier this year. For XGEVA, hypercalcemia following discontinuation of denosumab is already an identified risk in patients with a growing skeleton.
The applicant is taking the opportunity of implementing a minor correction to the RMP for correction or to add clarification."
Request for Supplementary Information adopted on 22.06.2017, 26.01.2017.

Zavesca - miglustat - EMEA/H/C/000435/II/0056, Orphan

MAH: Actelion Registration Ltd., Rapporteur: Kristina Dunder, PRAC Rapporteur: Qun-Ying Yue, "Submission of 8th NPC (Niemann-Pick type C) Registry report and update of Annex II-D to delete the NPC Registry listed as an obligation to the marketing authorisation.

The RMP version 12.1 has also been submitted to reflect the above changes.
In addition, the Marketing authorisation holder took the opportunity to introduce minor changes and bring the Product Information and Annex A in

line with the latest QRD template version 10." Request for Supplementary Information adopted on 21.04.2017.	
WS1117/G **Stocrin-EMEA/H/C/000250/WS1117/0110/G** **Sustiva-EMEA/H/C/000249/WS1117/0139/G** MAH: Bristol-Myers Squibb Pharma EEIG, Lead Rapporteur: Bruno Sepodes, Lead PRAC Rapporteur: Ana Sofia Diniz Martins, "C.I.4 (Type II) - Update of sections 4.4, 4.5 and 5.1 of the SmPC in order to add a warning and update the safety information on QTc prolongation based on the final results from study AI266959; this is an interventional study to determine the concentration-electrocardiographic effects of efavirenz in healthy subjects enriched for cyp2b6 polymorphisms; the Package Leaflet is updated accordingly. The RMP version 8 has also been submitted. C.I.4 (Type II) – Update of sections 4.4 and 4.8 to add catatonia as a Psychiatric symptom following an assessment of catatonia cases reported in the literature and via the United States (US) Food and Drug Administration Adverse Event Reporting System (FAERS)." Request for Supplementary Information adopted on 06.07.2017, 06.04.2017.	Weekly start timetable.
WS1182 **AMGEVITA-EMEA/H/C/004212/WS1182/0001** **SOLYMBIC-EMEA/H/C/004373/WS1182/0001** MAH: Amgen Europe B.V., Lead Rapporteur: Kristina Dunder, Lead PRAC Rapporteur: Ulla Wändel Liminga, "Submission of the final report from study/studies 20130258, An open-label, single-arm extension study to evaluate the long-term safety and efficacy of ABP 501 in subjects with moderate to severe rheumatoid arthritis, listed as a category 3 study in the RMP (MEA002). No changes of the PI are proposed, the RMP is updated accordingly." Request for Supplementary Information adopted on 06.07.2017.	Weekly start timetable.

B.5.4. PRAC assessed procedures

PRAC Led **Aclasta - zoledronic acid -** **EMEA/H/C/000595/II/0069** MAH: Novartis Europharm Ltd, Rapporteur: Kristina Dunder, PRAC Rapporteur: Ulla Wändel Liminga, PRAC-CHMP liaison: Kristina Dunder, "Submission of the final 5-year report from study (ZOL446H2422) listed as a category 3 study in the RMP. This is a non-interventional post-authorisation safety study using health registries to compare safety of Aclasta against oral bisphosphonates and untreated population controls." Request for Supplementary Information adopted on 06.07.2017.	Weekly start timetable.
PRAC Led **Inovelon - rufinamide -** **EMEA/H/C/000660/II/0041, Orphan** MAH: Eisai Ltd, Rapporteur: Alexandre Moreau, PRAC Rapporteur: Ghania Chamouni, PRAC-CHMP liaison: Alexandre Moreau, "Submission of the final clinical study report for study E2080-E044-401, the European registry of anti-epileptic drug use in patients with Lennox-Gastaut Syndrome (LAG), listed as a category 3 study in the RMP, in order to fulfil MEA 002.1. This is a non-interventional EU registry study entering patients (aged ≥4 years) with LGS who required a modification in anti-epileptic therapy (either the addition of another AED or the change of one drug to another) to evaluate the long-term safety of rufinamide." Opinion adopted on 06.07.2017. Request for Supplementary Information adopted on 05.05.2017.	Positive Opinion adopted by consensus on 06.07.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
PRAC Led **NovoEight - turoctocog alfa -** **EMEA/H/C/002719/II/0020** MAH: Novo Nordisk A/S, Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Brigitte Keller-Stanislawski, PRAC-CHMP liaison: Jan Mueller-Berghaus, "Submission of an amended protocol for PASS study NN7008-3553, category 3 study in the RMP. Submission of an updated RMP version 3 to update the timelines of the milestones in order to integrate the required additional pharmacovigilance activities, which include a	Weekly start timetable.

change in the Last Patient Last Visit (LPLV) date and a change in the Clinical Trial Report (CTR) finalisation date. In addition, the duration of the trial has been amended from 4 to 7 years." Request for Supplementary Information adopted on 06.07.2017.	
PRAC Led **Rebif - interferon beta-1a - EMEA/H/C/000136/II/0129** MAH: Merck Serono Europe Limited, PRAC Rapporteur: Qun-Ying Yue, PRAC-CHMP liaison: Filip Josephson, "Submission of an updated RMP version 9.0 in order to upgrade the important potential risk "Immunogenicity/safety risk associated with the formation of neutralizing antibodies" to important identified risk" Opinion adopted on 06.07.2017.	Positive Opinion adopted by consensus on 06.07.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
PRAC Led **Retacrit - epoetin zeta - EMEA/H/C/000872/II/0077** MAH: Hospira UK Limited, Rapporteur: Martina Weise, PRAC Rapporteur: Valerie Strassmann, PRAC-CHMP liaison: Martina Weise, "Submission of the final report from the registry based healthcare database study linked to PASCO (PMS-830-07-0043)) listed as a category 3 study in the RMP. This is an observational study on the incidence of thromboembolic events in patients with renal anaemia treated with erythropoietin-zeta as compared with erythropoietin-alpha and other erythropoiesis-stimulating agents." Opinion adopted on 06.07.2017.	Positive Opinion adopted by consensus on 06.07.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
PRAC Led **Revlimid - lenalidomide - EMEA/H/C/000717/II/0095, Orphan** MAH: Celgene Europe Limited, Rapporteur: Alexandre Moreau, PRAC Rapporteur: Ghania Chamouni, PRAC-CHMP liaison: Alexandre Moreau, "Submission of the final results of the non-interventional, observational category 3 post-authorisation safety study (Study CC-5013-PASS-001) in subjects treated with lenalidomide to further characterise the safety profile of lenalidomide plus dexamethasone in the treatment of relapsed and/or refractory (R/R) MM in a real-world setting."	
PRAC Led **Silapo - epoetin zeta -**	Positive Opinion adopted by consensus on 06.07.2017. The Icelandic and Norwegian CHMP

EMEA/H/C/000760/II/0045 MAH: STADA Arzneimittel AG, Rapporteur: Martina Weise, PRAC Rapporteur: Valerie Strassmann, PRAC-CHMP liaison: Martina Weise, "Submission of the final report from the registry based healthcare database study linked to PASCO (PMS-830-07-0043)) listed as a category 3 study in the RMP. This is an observational study on the incidence of thromboembolic events in patients with renal anaemia treated with erythropoietin-zeta as compared with erythropoietin-alpha and other erythropoiesis-stimulating agents. In addition, an updated RMP (version 11) is submitted to reflect the outcome of the study." Opinion adopted on 06.07.2017.	Members were in agreement with the CHMP recommendation.
PRAC Led **Tysabri - natalizumab - EMEA/H/C/000603/II/0101** MAH: Biogen Idec Ltd, Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Brigitte Keller-Stanislawski, PRAC-CHMP liaison: Jan Mueller-Berghaus, "Submission of the final clinical study report for TYGRIS, a post-marketing safety observational cohort program designed to obtain long-term safety data (approximately 5 years) in subjects with MS treated with natalizumab, and comprising parallel studies 101MS402 (United States and Canada) and 101MS403 (Rest of World). The application included an updated RMP version 23." Opinion adopted on 06.07.2017. Request for Supplementary Information adopted on 06.04.2017.	Positive Opinion adopted by consensus on 06.07.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
PRAC Led **Yervoy - ipilimumab - EMEA/H/C/002213/II/0049** MAH: Bristol-Myers Squibb Pharma EEIG, Rapporteur: Paula Boudewina van Hennik, PRAC Rapporteur: Sabine Straus, PRAC-CHMP liaison: Johann Lodewijk Hillege, "Submission of an updated RMP (version 17) in order to amend the study objectives and milestones for two studies: - study CA184332, a multi-site retrospective observational study of US patients with unresectable or metastatic melanoma receiving ipilimumab (Yervoy) as first line therapy in a community setting, a category 3 study in the RMP (MEA 029): to submit the final study report with 2-years of follow-up	Weekly start timetable.

- study CA184338, a multi-site retrospective observational study of US patients with unresectable or metastatic melanoma receiving ipilimumab (Yervoy) as first line therapy, a category 3 study in the RMP (MEA 030): to submit the final study report with 4-years of follow-up." Request for Supplementary Information adopted on 06.07.2017.	
PRAC Led **WS1163** **Harvoni-EMEA/H/C/003850/WS1163/0051** **Sovaldi-EMEA/H/C/002798/WS1163/0041** MAH: Gilead Sciences International Ltd, Lead PRAC Rapporteur: Julie Williams, PRAC-CHMP liaison: Greg Markey, "To provide updated RMPs for Sovaldi and Harvoni following the CHMP opinion, endorsing a PRAC recommendation, issued on 15 December 2016 (EMA/CHMP/847450/2016) on the Article 20 procedure for Direct-acting antivirals (DAAs) indicated for the treatment of hepatitis C (interferon free). The PRAC requested `hepatitis B reactivation' to be considered as important identified risk for all direct-acting antivirals. In addition, `emergence of hepatocellular carcinoma' and `recurrence of hepatocellular carcinoma' have been included as important potential risks. `Patients with previous HCC' have been reflected as missing information in the RMP of the DAAs, since this population was excluded from existing clinical trials. The requested studies have also been reflected in the RMPs." Opinion adopted on 06.07.2017. Request for Supplementary Information adopted on 05.05.2017.	Positive Opinion adopted by consensus on 06.07.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
PRAC Led **WS1169** **Exviera-EMEA/H/C/003837/WS1169/0028** **Viekirax-EMEA/H/C/003839/WS1169/0032** MAH: AbbVie Limited, Lead Rapporteur: Filip Josephson, Lead PRAC Rapporteur: Dolores Montero Corominas, PRAC-CHMP liaison: Concepcion Prieto Yerro, "To provide updated RMPs for Exviera and Viekirax following the CHMP opinion, endorsing a PRAC recommendation, issued on 15 December 2016 (EMA/CHMP/847450/2016) on the Article 20 procedure for Direct-acting antivirals (DAAs)	Positive Opinion adopted by consensus on 06.07.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

indicated for the treatment of hepatitis C (interferon free). The PRAC requested `hepatitis B reactivation' to be considered as important identified risk for all direct-acting antivirals. In addition, `emergence of hepatocellular carcinoma' and `recurrence of hepatocellular carcinoma' have been included as important potential risks. `Patients with previous HCC' have been reflected as missing information in the RMP of the DAAs, since this population was excluded from existing clinical trials. The requested studies have also been reflected in the RMPs." Opinion adopted on 06.07.2017. Request for Supplementary Information adopted on 05.05.2017.	
PRAC Led **WS1188** **Humalog-EMEA/H/C/000088/WS1188/0157** **Liprolog-EMEA/H/C/000393/WS1188/0120** MAH: Eli Lilly Nederland B.V., Lead Rapporteur: Robert James Hemmings, Lead PRAC Rapporteur: Julie Williams, PRAC-CHMP liaison: Robert James Hemmings, "Submission of the final report of a non-interventional post-authorisation safety study EUPAS 13422. This study is aimed to evaluate the impact of additional risk minimisation measures on healthcare professionals and on patients' understanding and their behaviour regarding the risk of hypoglycaemia and/or hyperglycaemia due to medication errors associated with administration of Humalog 200 U/ml KwikPen." Request for Supplementary Information adopted on 06.07.2017.	Weekly start timetable.
PRAC Led **WS1198** **Ebymect-EMEA/H/C/004162/WS1198/0022** **Edistride-EMEA/H/C/004161/WS1198/0017** **Forxiga-EMEA/H/C/002322/WS1198/0037** **Xigduo-EMEA/H/C/002672/WS1198/0033** MAH: AstraZeneca AB, Lead Rapporteur: Kristina Dunder, Lead PRAC Rapporteur: Qun-Ying Yue, PRAC-CHMP liaison: Kristina Dunder, "Implement the outcome of the article 20 referral regarding lower limb amputations in the RMP. The variation	

is submitted in order to give the rapporteur the possibility to review and assess the way the requested information has been included in the RMPs."	

B.5.5. CHMP-CAT assessed procedures

B.5.6. CHMP-PRAC-CAT assessed procedures

Strimvelis - autologous CD34+ enriched cell fraction that contains CD34+ cells transduced with retroviral vector that encodes for the human ADA cDNA sequence - EMEA/H/C/003854/II/0006, Orphan, ATMP MAH: GlaxoSmithKline Trading Services, Rapporteur: Christiane Niederlaender, PRAC Rapporteur: Sabine Straus	

B.5.7. PRAC assessed ATMP procedures

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

WS1081 **Hexacima-EMEA/H/C/002702/WS1081/0055** **Hexaxim-EMEA/H/W/002495/WS1081/0062** **Hexyon-EMEA/H/C/002796/WS1081/0059** MAH: Sanofi Pasteur SA, Lead Rapporteur: Jan Mueller-Berghaus Request for Supplementary Information adopted on 21.04.2017, 16.02.2017.	Weekly start timetable.
WS1165 **Aflunov-EMEA/H/C/002094/WS1165/0035** **Foclivia-EMEA/H/C/001208/WS1165/0030** MAH: Seqirus S.r.l, Lead Rapporteur: Daniela Melchiorri Request for Supplementary Information adopted on 22.06.2017, 18.05.2017.	Weekly start timetable.
WS1170 **Aflunov-EMEA/H/C/002094/WS1170/0036** **Foclivia-EMEA/H/C/001208/WS1170/0031**	Positive Opinion adopted by consensus on 06.07.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

MAH: Seqirus S.r.l, Lead Rapporteur: Daniela Melchiorri Opinion adopted on 06.07.2017. Request for Supplementary Information adopted on 01.06.2017.	
WS1192 **Hexacima-EMEA/H/C/002702/WS1192/0066** **Hexaxim-EMEA/H/W/002495/WS1192/0072** **Hexyon-EMEA/H/C/002796/WS1192/0070** MAH: Sanofi Pasteur SA, Lead Rapporteur: Kristina Dunder	
WS1200 **Lyrica-EMEA/H/C/000546/WS1200/0089** **Pregabalin** **Pfizer-EMEA/H/C/003880/WS1200/0019** MAH: Pfizer Limited, Lead Rapporteur: Johann Lodewijk Hillege Opinion adopted on 06.07.2017.	Positive Opinion adopted by consensus on 06.07.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

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

Strensiq - asfotase alfa - EMEA/H/C/003794/II/0020, Orphan MAH: Alexion Europe SAS, Rapporteur: Greg Markey Withdrawal request submitted on 10.07.2017.	The MAH withdrew the procedure on 10.07.2017.
Xeljanz - tofacitinib - EMEA/H/C/004214/II/0002 MAH: Pfizer Limited, Rapporteur: Robert James Hemmings Withdrawal request submitted on 06.07.2017.	The MAH withdrew the procedure on 06.07.2017.

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

- glycopyrronium / formoterol fumarate dihydrate - EMEA/H/C/004245 , indicated as a maintenance bronchodilator treatment to relieve symptoms in adult patients with chronic obstructive pulmonary disease

(COPD)	
- bictegravir / emtricitabine / tenofovir alafenamide - EMEA/H/C/004449 , treatment of adults infected with human immunodeficiency virus-1 (HIV-1)	
- deferiprone - EMEA/H/C/004710 , treatment of iron overload in thalassemia major, Generic, Generic of Ferriprox	
- lesinurad / allopurinol - EMEA/H/C/004412, gout	
- pacritinib - EMEA/H/C/004793 , treatment of disease-related splenomegaly and control of symptoms in patients with primary myelofibrosis (PMF), post-polycythemia vera myelofibrosis (PPV-MF), or post-essential thrombocythemia myelofibrosis (PET-MF) who have thrombocytopenia (platelet counts ≤100,000 /µL).	
- 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.	**Accelerated review**
- botulinum toxin type a - EMEA/H/C/004587 , temporary improvement in the appearance of moderate to severe vertical lines between the eyebrows	
- trastuzumab - EMEA/H/C/004463 , treatment of metastatic and early breast cancer and metastatic gastric cancer (MGC)	
- meropenem / vaborbactam - EMEA/H/C/004669 , treatment of infections	

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

- everolimus - EMEA/H/C/002311/X/0045, Orphan MAH: Novartis Europharm Ltd	

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

- anagrelide - EMEA/H/C/004585 , reduction of elevated platelet counts in at risk	

essential thrombocythaemia patients, Generic, Generic of Xagrid
List of Questions adopted on 21.04.2017.

- plitidepsin - EMEA/H/C/004354, Orphan
Applicant: Pharma Mar, S.A., treatment of multiple myeloma
List of Questions adopted on 23.02.2017.

- ulipristal acetate - EMEA/H/C/001027/X/0045
List of Questions adopted on 22.06.2017.

- fulvestrant - EMEA/H/C/004649
, treatment of breast cancer, List of Questions adopted on 21.04.2017.

- velmanase alfa - EMEA/H/C/003922, Orphan
Applicant: Chiesi Farmaceutici S.p.A., indicated for long-term enzyme replacement therapy in patients with alpha-mannosidosis
List of Questions adopted on 26.01.2017.

- semaglutide - EMEA/H/C/004174
, to improve glycaemic control in adults with type 2 diabetes and to prevent cardiovascular events
List of Questions adopted on 21.04.2017.

- rucaparib - EMEA/H/C/004272, Orphan
Applicant: Clovis Oncology UK Ltd, treatment of ovarian cancer
List of Questions adopted on 23.03.2017.

- human herpesvirus 3 - EMEA/H/C/004336
, prevention of herpes zoster (HZ) and HZ-related complications
List of Questions adopted on 21.04.2017.

- human fibrinogen / human thrombin - EMEA/H/C/004446
, treatment of haemostasis
List of Questions adopted on 21.04.2017.

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

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

Actelsar HCT - telmisartan / hydrochlorothiazide - EMEA/H/C/002676/R/0015
MAH: Actavis Group PTC ehf, Generic, Generic of

MicardisPlus, Rapporteur: Alar Irs, PRAC Rapporteur: Carmela Macchiarulo,

Hexacima - diphtheria, tetanus, pertussis (acellular, component), hepatitis B (rDNA), poliomyelitis (inact.) and Haemophilus type B conjugate vaccine (adsorbed) - EMEA/H/C/002702/R/0068
MAH: Sanofi Pasteur SA, Rapporteur: Jan Mueller-Berghaus, Co-Rapporteur: Bart Van der Schueren,

Hexyon - 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/002796/R/0072
MAH: Sanofi Pasteur Europe, Duplicate, Duplicate of Hexacima, Rapporteur: Jan Mueller-Berghaus, Co-Rapporteur: Bart Van der Schueren, PRAC Rapporteur: Brigitte Keller-Stanislawski,

Marixino - memantine - EMEA/H/C/002658/R/0012
MAH: Consilient Health Ltd., Generic, Generic of Ebixa, Rapporteur: Milena Stain, PRAC Rapporteur: Dolores Montero Corominas

Perjeta - pertuzumab - EMEA/H/C/002547/R/0031
MAH: Roche Registration Limited, Rapporteur: Sinan B. Sarac, Co-Rapporteur: Daniela Melchiorri, PRAC Rapporteur: Doris Stenver,

Prepandrix - A/Indonesia/05/2005 (H5N1) like strain used (PR8-IBCDC-RG2) - EMEA/H/C/000822/R/0071
MAH: GlaxoSmithkline Biologicals SA, Rapporteur: Greg Markey, Co-Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Julie Williams,

Privigen - human normal immunoglobulin - EMEA/H/C/000831/R/0122
MAH: CSL Behring GmbH, Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Brigitte Keller-Stanislawski

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,

Tolucombi - telmisartan / hydrochlorothiazide - EMEA/H/C/002549/R/0020
MAH: KRKA, d.d., Novo mesto, Generic, Generic of MicardisPlus, Rapporteur: Alar Irs, PRAC Rapporteur: Carmela Macchiarulo

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

riviact - brivaracetam - EMEA/H/C/003898/II/0010/G
MAH: UCB Pharma S.A., Rapporteur: Filip Josephson, PRAC Rapporteur: Adam Przybylkowski, Extension of Indication to include adjunctive therapy in the treatment of partial-onset seizures with or without secondary generalisation in patients with epilepsy 4 years of age and older for Briviact. As a consequence, sections 4.1, 4.2, 4.7, 5.1 and 5.2 of the SmPC are updated.
In addition, the Marketing authorisation holder (MAH) submitted a 5ml oral syringe and adaptor for the paediatric population.

The Package Leaflet and Labelling are updated in accordance.

Submission of the final Environmental Risk Assessment for the inclusion of the paediatric population in accordance with the new indication sought."

B.6.8. CHMP assessed procedures scope: Pharmaceutical aspects

BeneFIX - nonacog alfa - EMEA/H/C/000139/II/0146
MAH: Pfizer Limited, Rapporteur: Jan Mueller-Berghaus

Benepali - etanercept - EMEA/H/C/004007/II/0026
MAH: Samsung Bioepis UK Limited (SBUK), Rapporteur: Andrea Laslop

Cosentyx - secukinumab - EMEA/H/C/003729/II/0026
MAH: Novartis Europharm Ltd, Rapporteur:

Tuomo Lapveteläinen

Cystadane - betaine anhydrous - EMEA/H/C/000678/II/0029, Orphan
MAH: Orphan Europe SARL, Rapporteur: Harald Enzmann

Elocta - efmoroctocog alfa - EMEA/H/C/003964/II/0016/G
MAH: Swedish Orphan Biovitrum AB (publ), Rapporteur: Jan Mueller-Berghaus

Foscan - temoporfin - EMEA/H/C/000318/II/0042
MAH: biolitec Pharma Ltd, Rapporteur: Paula Boudewina van Hennik

Hizentra - human normal immunoglobulin - EMEA/H/C/002127/II/0086
MAH: CSL Behring GmbH, Rapporteur: Jan Mueller-Berghaus

Intanza - influenza vaccine (split virion, inactivated) - EMEA/H/C/000957/II/0054
MAH: Sanofi Pasteur Europe, Rapporteur: Jorge Camarero Jiménez

Opdivo - nivolumab - EMEA/H/C/003985/II/0037/G
MAH: Bristol-Myers Squibb Pharma EEIG, Rapporteur: Jorge Camarero Jiménez,

Soliris - eculizumab - EMEA/H/C/000791/II/0100, Orphan
MAH: Alexion Europe SAS, Rapporteur: Jorge Camarero Jiménez

Uptravi - selexipag - EMEA/H/C/003774/II/0010
MAH: Actelion Registration Ltd., Rapporteur: Martina Weise

Voncento - human coagulation factor VIII / human Von Willebrand factor - EMEA/H/C/002493/II/0030/G
MAH: CSL Behring GmbH, Rapporteur: Paula Boudewina van Hennik

WS1084/G
Ganfort-EMEA/H/C/000668/WS1084/0028/G
Lumigan-EMEA/H/C/000391/WS1084/0053/G
MAH: Allergan Pharmaceuticals Ireland, Lead Rapporteur: Hanne Lomholt Larsen

WS1226
Humalog-EMEA/H/C/000088/WS1226/0158
Liprolog-EMEA/H/C/000393/WS1226/0121
MAH: Eli Lilly Nederland B.V, Lead Rapporteur: Robert James Hemmings

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

Ameluz - 5-aminolevulinic acid - EMEA/H/C/002204/II/0027/G
MAH: Biofrontera Bioscience GmbH, Rapporteur: Harald Enzmann, "C.I.4
Update of sections 4.2, 4.4, 4.8 and 5.1 of the SmPC in order to update the posology and method of administration of Ameluz for the treatment of actinic keratosis (AK) and field cancerization in combination with daylight and to update the safety information, based on the clinical study results from ALA-AK-CT009; this is a phase III, randomised, interventional, observer-blinded study aimed to compare the efficacy and safety of Ameluz in the treatment of mild to moderate AK with Metvix in combination with daylight photodynamic therapy. Section 5.2 of the SmPC has included a minor editorial change. 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.5.b
Change in the legal status of Ameluz from "medicinal product subject to restricted medical prescription" to "medicinal product subject to medical prescription"."

Aranesp - darbepoetin alfa -
EMEA/H/C/000332/II/0143

MAH: Amgen Europe B.V., Rapporteur: Martina Weise,"Update of section of section 4.8 the SmPC in order to add a warning on injection site bruise and haemorrhage with frequency unknown and to provide additional instructions on the use of the device in the PL following signal procedure EMEA/H/C000332/SDA/090 on cases of incorrect device use / device malfunction"

Epclusa - sofosbuvir / velpatasvir -
EMEA/H/C/004210/II/0012

MAH: Gilead Sciences International Ltd, Rapporteur: Filip Josephson"Update of section 5.3 of the SmPC in order to add non-clinical safety findings based on a 6-month carcinogenicity study conducted with velpatasvir in transgenic mice"

ReFacto AF - moroctocog alfa -
EMEA/H/C/000232/II/0140

MAH: Pfizer Limited, Rapporteur: Hanne Lomholt Larsen,"Submission of the report 'The Immunogenicity of ReFacto AF in UK PUPs Who Started Treatment from 2010' prepared by the United Kingdom Haemophilia Centre Doctors' Organisation (UKHCDO).
This report is being submitted in the context of a post-approval commitment, MEA 115.1 ('The MAH commits to submit the CSR for "A Postauthorization Safety Surveillance Registry or ReFacto AF in Previously Untreated Patients (PUPs) in Usual Care Settings – study number 4435" and to initiate the registry'), as supporting evidence of the ongoing safety evaluation of ReFacto AF in PUPs with haemophilia A and with a specific focus on the development of inhibitors."

Revatio - sildenafil -
EMEA/H/C/000638/II/0077

MAH: Pfizer Limited, Rapporteur: Johann Lodewijk Hillege"Update of section 4.6 of the SmPC in order to revise the statement concerning the detection of sildenafil and its active metabolite in human mik and the potential for impact on the breastfed infact.

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."

Stelara - ustekinumab - EMEA/H/C/000958/II/0058

MAH: Janssen-Cilag International NV, Rapporteur: Greg MarkeyUpdate of section 4.8 of the SmPC in order to include Lower Respiratory Tract Infection as an Adverse Drug Reaction based on a comprehensive evaluation of safety information from the STELARA clinical studies database and post-marketing database, as well as available literature.
The Package Leaflet is updated accordingly."

Strensiq - asfotase alfa - EMEA/H/C/003794/II/0019/G, Orphan

MAH: Alexion Europe SAS, Rapporteur: Greg Markey,Update of section 5.1 of the SmPC in order to update information following final results from studies ENB-006-09 [A Randomized, Open-Label, Multicenter, Multinational, Dose-Ranging, Historical Control Study of the Safety,Efficacy, Pharmacokinetics, and Pharmacodynamics of Asfotase Alfa (Human Recombinant Tissue-Nonspecific Alkaline Phosphatase Fusion Protein) in Children with Hypophosphatasia (HPP)] (and its extension ENB-008-10 [Extension Study of Protocol ENB-006-09 Evaluating the Long-Term Safety and Efficacy of Asfotase Alfa (Human Recombinant Tissue-Nonspecific Alkaline Phosphatase Fusion Protein) in Children with Hypophosphatasia (HPP)]) and ENB-009-10 [A Randomized, Open-Label, Multicenter, Multinational, Dose-Ranging, Concurrent Control Study of the Safety, Efficacy, and Pharmacokinetics of ENB-0040 (Human Recombinant Tissue-Nonspecific Alkaline Phosphatase Fusion Protein) in Adolescents and Adults with Hypophosphatasia (HPP)] listed as an obligation in the Annex II (ANX002). In addition, the Marketing authorisation holder (MAH) took the opportunity to propose editorial changes for section 4.5 to better clarify the information provided."

Stribild - elvitegravir / cobicistat / emtricitabine / tenofovir disoproxil - EMEA/H/C/002574/II/0083

MAH: Gilead Sciences International Ltd, Rapporteur: Robert James HemmingsUpdate of

sections 4.5 of the SmPC in order to add drug-drug interaction data from Study GS-US-292-1316; this is a Phase 1, Open-Label, Fixed Sequence Study Evaluating the Pharmacokinetics and Drug Interaction Potential Between Elvitegravir/Cobicistat/Emtricitabine/Tenofovir Alafenamide Single-Table Regimen and Sertraline in Healthy Subjects.

In addition, the Marketing authorisation holder (MAH) took the opportunity make administrative amendments to section 4.8 of the SmPC."

Tecfidera - dimethyl fumarate - EMEA/H/C/002601/II/0041

MAH: Biogen Idec Ltd, Rapporteur: Martina Weise, "Update of sections 4.4 and 4.8 of the SmPC in order to add anaphylactic reaction as a warning and as an adverse reaction with unknown frequency, based on post-marketing experience. The Package Leaflet is updated accordingly.

In addition, the Biogen Idec Ltd took the opportunity to bring the PI in line with the latest QRD template version 10."

Tecfidera - dimethyl fumarate - EMEA/H/C/002601/II/0042

MAH: Biogen Idec Ltd, Rapporteur: Martina Weise, "Update of sections 4.2, 4.8, 5.1 and 5.2 of the SmPC in order to update the safety and efficacy information in the paediatric population based on the clinical study results from study 109MS202, listed as a category 3 study in the RMP; this is an open-label, multicentre, multidose study designed to assess the effect of Tecfidera on magnetic resonance imaging lesions and pharmacokinetics, safety and tolerability in paediatric population with relapsing-remitting multiple sclerosis.

There are no updates proposed in the package leaflet or RMP."

Tecfidera - dimethyl fumarate - EMEA/H/C/002601/II/0043/G

MAH: Biogen Idec Ltd, Rapporteur: Martina Weise"C.I.13
Submission of non-clinical study report for study PD-15-73: a haemotoxicity study of two test compounds (BIO 0022819 and BIO 0022817) on T-CFC progenitor stem cells and T-cells derived

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from both human and non-human primate bone marrow and peripheral blood mononuclear cells (MNCs). This submission is linked to a category 3 study in the RMP.

C.I.13
Submission of non-clinical study report for study P00012-15-05: a preclinical study to evaluate the toxicity potential and toxicokinetic profile of dimethyl fumarate and hydroxyurea, when co-administered once daily via nasogastric intubation to cynomolgus monkeys for a minimum of 91 days. This submission is linked to a category 3 study in the RMP."

Tecfidera - dimethyl fumarate - EMEA/H/C/002601/II/0044
MAH: Biogen Idec Ltd, Rapporteur: Martina Weise,"Submission of the final study report of an interventional PASS listed as a category 4 study in the RMP: A Multicentre, Open-Label, Single-Arm Study to Evaluate Gastrointestinal Tolerability in Subjects with Relapsing-Remitting Multiple Sclerosis Receiving Dimethyl Fumarate (TOLERATE), study number 109MS407."

Toviaz - fesoterodine - EMEA/H/C/000723/II/0049
MAH: Pfizer Limited, Rapporteur: Concepcion Prieto Yerro, "Update of the SmPC sections 4.6 and 5.3 with revised information from reproductive toxicity studies in mice. In addition, the MAH took the opportunity to bring the PI in line with the latest QRD template version 10.0."

Trevicta - paliperidone - EMEA/H/C/004066/II/0011
MAH: Janssen-Cilag International NV, , Rapporteur: Kristina Dunder"Update of section 4.8 of the SmPC in order to update the safety information after assessment of study R092670-SCA-3004 (A Randomized, Double-Blind, Placebo-Controlled, Parallel-Group Study of Paliperidone Palmitate Evaluating Time to Relapse in Subjects With Schizoaffective Disorder). The Package Leaflet has been updated accordingly"

Vargatef - nintedanib - EMEA/H/C/002569/II/0017
MAH: Boehringer Ingelheim International GmbH, Rapporteur: Sinan B. Sarac"Update of section 4.8 of the SmPC in order to add 'weight decreased' as

a new adverse drug reaction based on a safety review of clinical trials and post-marketing data. The Package Leaflet is updated accordingly. In addition, the Marketing authorisation holder (MAH) took the opportunity to implement a minor correction in the English product information, minor corrections to the Croatian, Danish, Dutch and Finnish translations and to bring section 4 of the Package Leaflet in line with QRD template version 10."

Vargatef - nintedanib - EMEA/H/C/002569/II/0018

MAH: Boehringer Ingelheim International GmbH, Rapporteur: Sinan B. Sarac"Update of section 4.4 of the SmPC to amend the current warning on hepatic function to include that drug liver induced injury was associated with nintendanib administration, to include female sex as a factor of increased risk of liver enzyme elevations, update of section 4.8 of the SmPC to add 'drug-induced liver injury' (DILI) as new ADR and update of section 5.2 of the SmPC to amend the current information related to the mean exposure to nintedanib by race, based on a review of clinical trials and post-marketing data on DILI and on the exposure safety relationship between nintedanib plasma exposure and liver enzyme elevations, as requested by the PRAC as part of PSUSA/00010318/201611. The Package Leaflet is updated accordingly. In addition, the MAH took the opportunity to make some minor changes to section 4.4 and 4.8 of the SmPC."

Xeljanz - tofacitinib - EMEA/H/C/004214/II/0003

MAH: Pfizer Limited, Rapporteur: Robert James Hemmings, "Submission of 2 transported inhibition studies evaluating tofacitinib for its potential to inhibit organic anion transporter (OAT) 1, OAT3 and to interact with Human MRP2 Efflux (ABC) Transporter in fulfilment of the Recommendation dated 26 January 2017."

Xultophy - insulin degludec / liraglutide - EMEA/H/C/002647/II/0021

MAH: Novo Nordisk A/S, Rapporteur: Kristina DunderUpdate of section 5.1 of the SmPC in order to reflect data for transfer from insulin glargine U100 to Xultophy as compared to a basal-bolus regimen. The update is based on data from the clinical trial NN9068-4185: "A clinical trial

comparing efficacy and safety of insulin degludec/liraglutide (IDegLira) versus basal-bolus therapy in subjects with type 2 diabetes mellitus". The MAH has taken the opportunity to make minor editorial and formatting changes throughout the Annexes."	
- sodium oxybate - **EMEA/H/C/000593/II/0067/G** "Update of section 4.8 of the SmPC in order to add the adverse reactions "increased libido" and "seborrhea" with an unknown frequency. Update of section 4.6 of the SmPC in order to amend the information about breast-feeding. The Package Leaflet is updated accordingly."	
Zykadia - ceritinib - **EMEA/H/C/003819/II/0016** MAH: Novartis Europharm Ltd, 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."	
WS1205 **Descovy-EMEA/H/C/004094/WS1205/0020** **Genvoya-EMEA/H/C/004042/WS1205/0034** **Odefsey-EMEA/H/C/004156/WS1205/0018** MAH: Gilead Sciences International Ltd, Lead Rapporteur: Robert James Hemmings, "Update of section 4.5 of the SmPC in order to provide the final study report for the in vitro study AD-120-2045; this is a non-clinical study on the Effect of Xanthine Oxidase Inhibitors on Metabolism of Tenofovir alafenamide fumarate in Primary Human Hepatocytes. This study is listed in their respective Risk	

Management Plans (RMPs) as an additional pharmacovigilance activity (Category 3) (Genvoya: MEA 006; Descovy: MEA 004; Odefsey: MEA 007).

The requested worksharing procedure proposed amendments to the Summary of Product Characteristics."

WS1218
Brimica
Genuair-EMEA/H/C/003969/WS1218/0015
Duaklir
Genuair-EMEA/H/C/003745/WS1218/0015
MAH: AstraZeneca AB, Lead Rapporteur: Nithyanandan Nagercoil"Update of section 5.1 of the SmPC in order to update information following results from study M-40464-33 (A Multiple Dose, Randomised, Double-Blind, Placebo Controlled, Parallel Clinical Trial to Assess the Effect of Aclidinium Bromide/Formoterol Fumarate Fixed-Dose Combination on Lung Hyperinflation, Exercise Capacity and Physical Activity in Patients with Moderate to Severe Chronic Obstructive Pulmonary Disease (COPD))"

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."

WS1225/G
Exviera-EMEA/H/C/003837/WS1225/0031

/G

Viekirax-EMEA/H/C/003839/WS1225/0035/G

MAH: AbbVie Limited, Lead Rapporteur: Filip Josephson"Submission of the final reports for two phase IIIb studies (studies M14-226 and M15-461) listed as category 3 studies in the RMP. These are open-label studies evaluating the safety and efficacy of ombitasvir/paritaprevir/ritonavir and dasabuvir with or without ribavirin in hepatitis C virus infected patients with several renal impairment or end-stage renal disease with or without compensated cirrhosis."

B.6.10. CHMP-PRAC assessed procedures

Adcetris - brentuximab vedotin - EMEA/H/C/002455/II/0049, Orphan

MAH: Takeda Pharma A/S, Rapporteur: Paula Boudewina van Hennik, PRAC Rapporteur: Sabine Straus, "Update of sections 4.2, 4.8, 5.1 and 5.2 of the SmPC with data from study C25002; a phase 1/2 study of brentuximab vedotln (SGN-35) in paediatric patients with relapsed or refractory systemic anaplastic large cell lymphoma or hodgkin lymphoma (listed in the agreed PIP covering the conditions of Hodgkin lymphoma and anaplastic large cell lymphoma for ADCETRIS (EMEA-000980-PIP01-10-M04)). An updated RMP version 11.0 was provided as part of the application."

Adenuric - febuxostat - EMEA/H/C/000777/II/0047

MAH: Menarini International Operations Luxembourg S.A., Rapporteur: Andrea Laslop, PRAC Rapporteur: Jan NeuhauserUpdate of sections 4.4 and 4.5 of the SmPC in order to reflect the results of preclinical study MRPO-2015-PKM-005 "Pharmacokinetic of azathioprine in the rat after one-week daily oral treatment at three different dosages and with the concomitant oral administration of febuxostat or allopurinol" and clinical study REP-POPPK-MRP-2015-PKM-005 "Population Pharmacokinetic analysis from study titled Pharmacokinetic of azathioprine in the rat after one-week daily oral treatment at three different dosages and with the concomitant oral administration of febuxostat or allopurinol",

investigating the drug-drug interaction with azathioprine when co-administered with febuxostat.

The RMP version 6.0 has also been submitted.

In addition, the MAH took the opportunity to correct the typing errors and to bring the PI in line with the latest QRD template version 10."

Cerdelga - eliglustat - EMEA/H/C/003724/II/0013, Orphan
MAH: Genzyme Europe BV, Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Dolores Montero Corominas"Update of section 4.8. of the SmPC in order to amend the safety information based on the analysis of Adverse Events from the following clinical trials: GZGD00304 (Phase 2), GZGD02507 (ENGAGE), GZGD02607 (ENCORE) and GZGD03109 (EDGE) to address post-authorisation MEA011.1 which is included in the current approved Risk Management Plan.

Update of the labelling in order to reflect the instructions on use for the sleeve of the intermediate packaging of the single blister.

The RMP version 4.0 has also been submitted."

Eperzan - albiglutide - EMEA/H/C/002735/II/0033
MAH: GlaxoSmithKline Trading Services Limited, Rapporteur: Kristina Dunder, PRAC Rapporteur: Julie WilliamsUpdate of the Package Leaflet in order to amend the layout and content of the Instructions for Use (IFU). In addition, the RMP version 8 has also been submitted to implement additional pharmacovigilance and risk minimisation activities addressing the safety concern of "medication errors/device issue potentially leading to lack of efficacy or inadequate diabetes control" and to add a PASS study to investigate the effectiveness of the new IFU."

Nulojix - belatacept - EMEA/H/C/002098/II/0045
MAH: Bristol-Myers Squibb Pharma EEIG, Rapporteur: Filip Josephson, PRAC Rapporteur: Ulla Wändel Liminga"Update of sections 4.4 and 4.8 of the SmPC in order to add a warning and update the safety information on the risk of venous thrombosis of the renal allograft when anti-thymocyte globulin (ATG) and belatacept are

coadministered (at the same or nearly the same time) in patients with other predisposing risk factors for thrombosis.

The update is based on a review of the potential increased risk for allograft thrombosis with belatacept given in close temporal relation to Thymoglobulin, as requested during assessment of PSUR 8 (EMEA/H/C/PSUSA/00000311/201606).

In addition, the MAH took the opportunity update section 6.6 “Special precautions for disposal and other handling” of the SmPC and the “Information for healthcare professionals (HCPs)”in the Package Leaflet (PL) with additional safety instructions for the co-administration of Belatacept.

Submission of this variation application fulfils LEG 021 for Nulojix.

Consistently with the above, RMP version 14 has also been submitted, including addition of the potential risk of venous thrombosis of the allograft when ATG and belatacept are coadministered in patients with other predisposing risk factors for thrombosis and a number of administrative changes.”

Olumiant - baricitinib - EMEA/H/C/004085/II/0002

MAH: Eli Lilly Nederland B.V., Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Patrick Batty“Update of sections 4.5 and 5.2 of the SmPC, based on the final study report of in vitro study to investigate the inhibitory effect of baricitinib on the organic anion transporter 2 (OAT2) in fulfilment of PAM (MEA 001). The updated RMP version 3.0 has been submitted as part of this application.”

Opdivo - nivolumab - EMEA/H/C/003985/II/0038

MAH: Bristol-Myers Squibb Pharma EEIG, Rapporteur: Jorge Camarero Jiménez, PRAC Rapporteur: Brigitte Keller-Stanislawski, “Update of section 4.8 of the SmPC with longer follow-up for subjects proceeding to allogeneic transplant following nivolumab treatment, of section 5.1 of the SmPC with efficacy data from longer follow-up based on final results from study CA209205 listed as a PAES in the Annex II; this is a Phase 2, non-comparative, multi-cohort,

single-arm, open-label study of nivolumab (BMS-936558) in cHL subjects after failure of ASCT
Annex II is updated to remove the commitment.
Version 7.5 of the RMP has been submitted."

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."

Spedra - avanafil - EMEA/H/C/002581/II/0027/G
MAH: Menarini International Operations Luxembourg S.A., Rapporteur: Concepcion Prieto Yerro, PRAC Rapporteur: Dolores Montero Corominas"Update of section 4.4. to reflect the results of clinical study TA-402 "A Double-Blind, Randomized, Placebo-Controlled, Single-Dose, Parallel Study to Assess the Effects of Avanafil on Multiple Parameters of Vision, including, but Not Limited to Visual Acuity, Intraocular Pressure, Pupillometry, and Color Vision Discrimination, in Healthy Male Subjects).

Update of section 4.6. of the SmPC in order to reflect the results of clinical study TA-401 "A Randomized, Double-Blind, Placebo-Controlled, Parallel Group, Multicenter Clinical Trial of the Effect of Avanafil on Spermatogenesis in Healthy Adult Males and Adult Males with Mild Erectile Dysfunction". The Package Leaflet is updated accordingly.

The RMP version 5.1has also been submitted.

In addition, the MAH took the opportunity to make an editorial correction on the approved SmPc by adding the missing adverse reaction epistaxis from the tabulated list of adverse reactions reported in section 4.8. Additionally, the MAH took the opportunity of this variation to align the information included in Section 3 "How to take Spedra" in the Package Leaflet to section

4.2 “Posology” in the SmPC.

Some additional minor amendments, due to translation mistakes are proposed for the French Product Information.”

Tresiba - insulin degludec - EMEA/H/C/002498/II/0028

MAH: Novo Nordisk A/S, Rapporteur: Kristina Dunder, PRAC Rapporteur: Qun-Ying Yue, Update of section 5.1 of the SmPC based on new clinical data from a cardiovascular outcome trial EX1250-4080 (DEVOTE) conducted for Tresiba. DEVOTE was a randomised, double-blind and event-driven clinical trial with a median duration of 2 years comparing the cardiovascular safety of Tresiba versus insulin glargine (100 units/mL) in patients with type 2 diabetes mellitus at high risk of cardiovascular events.

The RMP version 8 has also been submitted, with updates consequent to the data in support of the application.”

WS1168

AZILECT-EMEA/H/C/000574/WS1168/0077

Rasagiline ratiopharm-EMEA/H/C/003957/WS1168/0010

MAH: Teva B.V., Lead Rapporteur: Bruno Sepodes, Lead PRAC Rapporteur: Ana Sofia Diniz Martins,“Update of sections 4.4, 4.7 and 4.8 to include a new warning on excessive daytime sleepiness and sudden sleep onset episodes and sudden sleep onset episodes, update of section 4.9 to remove ‘dysphoria’ as a symptom reported following overdose of rasagiline based on a CCDS update. The Package Leaflet is updated accordingly. The RMP version 2.0 has also been submitted. In addition, the Worksharing applicant (WSA) took the opportunity to make editorial changes throughout the PI, to correct the invented name for Rasagiline Ratiopharm in the Czech annexes and to bring the PI in line with the latest QRD template version 10.”

WS1180

Corlentor-EMEA/H/C/000598/WS1180/0047

Ivabradine Anpharm-EMEA/H/C/004187/WS1180/0006

Procoralan-EMEA/H/C/000597/WS1180/0046 MAH: Les Laboratoires Servier, Lead Rapporteur: Johann Lodewijk Hillege, Lead PRAC Rapporteur: Menno van der Elst, “Update to the section 4.8 of the SmPC with new ADRs: Ventricular tachycardia, Ventricular fibrillation and Torsade de pointes. The PL is updated accordingly. The RMP version 6 has also been submitted. In addition the MAH took the opportunity to align the PI with the latest QRD template 10.0.”	
WS1211 **Januvia-EMEA/H/C/000722/WS1211/0059** **Ristaben-EMEA/H/C/001234/WS1211/0051** **TESAVEL-EMEA/H/C/000910/WS1211/0059** **Xelevia-EMEA/H/C/000762/WS1211/0063** MAH: Merck Sharp & Dohme Limited, Lead Rapporteur: Johann Lodewijk Hillege, Lead PRAC Rapporteur: Menno van der Elst“Update of sections 4.2, 4.4 and 5.2 of the SmPC in order to modify the information on dosing, an existing warning and administration instructions, respectively for use of sitagliptin in patients with type 2 diabetes mellitus and renal impairment. Consequently, the RMP version 8 has also been updated accordingly. In addition, the Worksharing applicant (WSA) took the opportunity to update the list of local representatives in the Package Leaflet for Tesavel and to bring the PI in line with the latest QRD template version 10. Minor editorial changes are also introduced in the Product Information.”	
WS1212/G **Efficib-EMEA/H/C/000896/WS1212/0085/G** **Janumet-EMEA/H/C/000861/WS1212/0085/G** **Ristfor-EMEA/H/C/001235/WS1212/0072/G** **Velmetia-EMEA/H/C/000862/WS1212/0088/G** MAH: Merck Sharp & Dohme Limited, Lead Rapporteur: Johann Lodewijk Hillege, Lead PRAC Rapporteur: Menno van der Elst“Update of sections 4.2, and 5.2 of the SmPC in order to modify the information on dosing, and	

administration instructions respectively for use of sitagliptin/metformin in patients with type 2 diabetes mellitus and moderate renal impairment. Consequently, the RMP version 8 has also been updated accordingly.

Section 4.5 of the SmPC is also updated to include information on the concominant use of ranolazine, vandetanib, dolutegravir and cimetidine.

In addition, the Worksharing applicant (WSA) took the opportunity to update the list of local representatives in the Package Leaflet for Efficib and to bring the PI in line with the latest QRD template version 10. Minor editorial changes are also introduced in the Product Information."

B.6.11. PRAC assessed procedures

PRAC Led

Eliquis - apixaban -
EMEA/H/C/002148/II/0043

MAH: Bristol-Myers Squibb / Pfizer EEIG, Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Menno van der Elst, PRAC-CHMP liaison: Johann Lodewijk HillegeSubmission of the final report from study (CV185-365) listed as a category 3 study in the RMP. This is a post authorisation safety study which evaluates the effectiveness of Eliquis (apixaban) risk minimisation tools in the European Economic Area countries. A RMP (version 17.0) has also been submitted to reflect the completion of the study CV185-365."

PRAC Led

Invokana - canagliflozin -
EMEA/H/C/002649/II/0030

MAH: Janssen-Cilag International NV, Rapporteur: Martina Weise, PRAC Rapporteur: Valerie Strassmann, PRAC-CHMP liaison: Martina Weise"Submission of an updated RMP version 7.0 in order to include prior commitments made to PRAC during the PSUR/LEG procedural review of pancreatitis cases and the Article 20 referral procedure reviewing lower limb amputation in relation to the use of SGLT-2 inhibitors. In addition, the updated RMP reflects labelling changes that resulted from a variation to add information regarding fatal DKA cases to the existing DKA warning and the Article 31

procedure reviewing metformin-containing medicines."

PRAC Led

Viread - tenofovir disoproxil - EMEA/H/C/000419/II/0182

MAH: Gilead Sciences International Ltd, Rapporteur: Joseph Emmerich, PRAC Rapporteur: Caroline Laborde, PRAC-CHMP liaison: Joseph Emmerich"Submission of the final report from Study GX-US-174-0172, listed as a category 3 study in the RMP. This is a 5-year observational (non-interventional) renal safety registry conducted to provide further safety data in HBV-infected patients with decompensated liver disease."

PRAC Led

Vokanamet - canagliflozin / metformin - EMEA/H/C/002656/II/0031

MAH: Janssen-Cilag International NV, Rapporteur: Martina Weise, PRAC Rapporteur: Menno van der Elst, PRAC-CHMP liaison: Johann Lodewijk Hillege"Submission of an updated RMP version 7.0 in order to include prior commitments made to PRAC during the PSUR/LEG procedural review of pancreatitis cases and the Article 20 referral procedure reviewing lower limb amputation in relation to the use of SGLT-2 inhibitors. In addition, the updated RMP reflects labelling changes that resulted from a variation to add information regarding fatal DKA cases to the existing DKA warning and the Article 31 procedure reviewing metformin-containing medicines."

PRAC Led

WS1197

Actraphane-EMEA/H/C/000427/WS1197/0072

Actrapid-EMEA/H/C/000424/WS1197/0066

Insulatard-EMEA/H/C/000441/WS1197/0069

Mixtard-EMEA/H/C/000428/WS1197/0073

Protaphane-EMEA/H/C/000442/WS1197/0068

MAH: Novo Nordisk A/S, Lead Rapporteur: Hanne Lomholt Larsen, Lead PRAC Rapporteur: Doris Stenver, PRAC-CHMP liaison: Sinan B. SaracSubmission of an updated RMP version 3.0

according to GVP Module V, in order to remove three important potential risks (immunogenicity, allergic reactions and lack of efficacy) related to the new NN729 manufacturing process from the RMP, remove hypoglycaemia and anaphylactic reactions, remove peripheral neuropathy, refraction disorders, lipodystrophy, urticaria, rash, oedema and diabetic retinopathy and remove missing information concerning special populations. No changes are proposed to the product information."

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

WS0935/G
Filgrastim
Hexal-EMEA/H/C/000918/WS0935/0035/G
Zarzio-EMEA/H/C/000917/WS0935/0036/G
MAH: Sandoz GmbH, Lead Rapporteur: Greg Markey

WS1172
Infanrix hexa-EMEA/H/C/000296/WS1172/0221
MAH: GlaxoSmithkline Biologicals SA, Lead Rapporteur: Bart Van der Schueren

WS1184
Eucreas-EMEA/H/C/000807/WS1184/0063
Icandra-EMEA/H/C/001050/WS1184/0064
Zomarist-EMEA/H/C/001049/WS1184/0064
MAH: Novartis Europharm Ltd, Lead Rapporteur: Kristina Dunder

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 Lead Rapporteur: Jan Mueller-Berghaus

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WS1194

Infanrix hexa-EMEA/H/C/000296/WS1194/0222

MAH: GlaxoSmithkline Biologicals SA, Lead Rapporteur: Bart Van der Schueren

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WS1196/G

Ebymect-EMEA/H/C/004162/WS1196/0023/G

Xigduo-EMEA/H/C/002672/WS1196/0034/G

MAH: AstraZeneca AB, Lead Rapporteur: Kristina Dunder

WS1201/G

Glyxambi-EMEA/H/C/003833/WS1201/0009/G

Jentadueto-EMEA/H/C/002279/WS1201/0041/G

Trajenta-EMEA/H/C/002110/WS1201/0031/G

MAH: Boehringer Ingelheim International GmbH, Lead Rapporteur: Johann Lodewijk Hillege

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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

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WS1204/G

Herceptin-EMEA/H/C/000278/WS1204/0134/G

Kadcyla-EMEA/H/C/002389/WS1204/003

7/G

MAH: Roche Registration Limited, Lead Rapporteur: Jan Mueller-Berghaus

WS1214

Aflunov-EMEA/H/C/002094/WS1214/0039

Foclivia-EMEA/H/C/001208/WS1214/0033

MAH: Seqirus S.r.l, Lead Rapporteur: Daniela Melchiorri

WS1224

Relvar Ellipta-EMEA/H/C/002673/WS1224/0031

Revinty Ellipta-EMEA/H/C/002745/WS1224/0027

MAH: Glaxo Group Ltd, Lead Rapporteur: Concepcion Prieto Yerro

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

Disclosure of information related to plasma master files cannot be released at 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):

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

Qualification of Biomarkers:

HTA:

G.2. Ongoing procedures

G.3. PRIME

Disclosure of some information related to PRIME cannot be released at present time as these contain

commercially confidential information.

G.3.1. List of procedures concluding at 17-20 July 2017 CHMP plenary:

G.3.2. List of procedures starting in July 2017 for August 2017 CHMP adoption of outcomes

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