19 June 2017
EMA/CHMP/309580/2017
Inspections, Human Medicines Pharmacovigilance and Committees Division

Committee for medicinal products for human use (CHMP)

Draft agenda for the meeting on 19-22 June 2017

Chair: Tomas Salmonson – Vice-Chair: Harald Enzmann

19 June 2017, 13:00 – 19:30, room 2A

20 June 2017, 08:30 – 19:30, room 2A

21 June 2017, 08:30 – 19:30, room 2A

22 June 2017, 08:30 – 15:00, 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 19-22 June 2017. See June 2017 CHMP minutes (to be published post July 2017 CHMP meeting).

1.2. Adoption of agenda

CHMP agenda for 19-22 June 2017

1.3. Adoption of the minutes

CHMP minutes for 15-18 May 2017.

2. Oral Explanations

2.1. Pre-authorisation procedure oral explanations

2.1.1. alpha-1-antitrypsin - Orphan - EMEA/H/C/003934

Kamada BioPharma Limited at Fieldfisher LLP; treatment and maintenance therapy of adult patients with congenital deficiency of alpha-1 antitrypsin and lung disease with clinical evidence of emphysema and airway obstruction (FEV1/SVC<70%)

Scope: Oral explanation

Action: Oral explanation to be held on 20 June 2017 at time 09:00

List of Outstanding Issues adopted on 23.03.2017. List of Questions adopted on 21.07.2016.

2.1.2. prasterone - EMEA/H/C/004138

treatment of vulvovaginal atrophy

Scope: Oral explanation

Action Oral explanation to be held on 20 June 2017 at time 11:00

List of Outstanding Issues adopted on 26.01.2017, 13.10.2016. List of Questions adopted on 26.05.2016.

2.1.3. cladribine - EMEA/H/C/004230

treatment of highly active relapsing-remitting multiple sclerosis (MS)

Scope: Oral explanation, report from SAG held 8 June 2017

Action: Oral explanation to be held on 20 June 2017 at time 14:00

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

2.1.4. 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 22 June 2017 at time 09:00

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

2.1.5. atezolizumab - EMEA/H/C/004143

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

Scope: Oral explanation

Action: Oral explanation to be held on 20 June 2017 at time 16:00

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

2.2. Re-examination procedure oral explanations

2.3. Post-authorisation procedure oral explanations

2.4. Referral procedure oral explanations

2.4.1. 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: Oral explanation

Action: Oral explanation to be held on 19 June 2017 at time 16:00

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.

List of Outstanding issues adopted on 21.04.2017. List of Questions adopted on 26.01.2017.

See 10.4

3. Initial applications

3.1. Initial applications; Opinions

3.1.1. efavirenz / emtricitabine / tenofovir disoproxil - EMEA/H/C/004240

treatment of HIV-1 infection

Scope: Opinion

Action: For adoption

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

3.1.2. tivozanib hydrochloride monohydrate - EMEA/H/C/004131

treatment of adult patients with advanced renal cell carcinoma (RCC)

Scope: Opinion

Action: For adoption

List of Outstanding Issues adopted on 26.01.2017. List of Questions adopted on 21.07.2016.

3.1.3. adalimumab - EMEA/H/C/004279

treatment of rheumatoid arthritis, psoriatic arthritis and ankylosing spondylitis

Scope: Opinion

Action: For adoption

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

3.1.4. alpha-1-antitrypsin - Orphan - EMEA/H/C/003934

Kamada BioPharma Limited at Fieldfisher LLP; treatment and maintenance therapy of adult patients with congenital deficiency of alpha-1 antitrypsin and lung disease with clinical evidence of emphysema and airway obstruction (FEV1/SVC<70%)

Scope: Opinion

Action: For adoption

List of Outstanding Issues adopted on 23.03.2017. List of Questions adopted on 21.07.2016.

3.1.5. ribociclib - EMEA/H/C/004213

treatment of breast cancer

Scope: Opinion

Action: For adoption

List of Outstanding Issues adopted on 21.04.2017. List of Questions adopted on 26.01.2017.

3.1.6. cladribine - EMEA/H/C/004230

treatment of highly active relapsing-remitting multiple sclerosis (MS)

Scope: Opinion

Action: For adoption

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

3.1.7. glecaprevir / pibrentasvir - EMEA/H/C/004430

Accelerated assessment

Indicated for the treatment of chronic hepatitis C virus (HCV) infection in adults

Scope: Opinion

Action: For adoption

List of Questions adopted on 19.04.2017.

3.1.8. nitisinone - EMEA/H/C/004281

treatment of hepatorenal tyrosinemia type 1

Scope: Opinion

Action: For adoption

List of Outstanding Issues adopted on 23.03.2017. List of Questions adopted on 21.07.2016.

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

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

Scope: Opinion/Second list of Outstanding Issues

Action: For adoption

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

3.1.10. sofosbuvir / velpatasvir / voxilaprevir - EMEA/H/C/004350

Accelerated assessment

Treatment of chronic hepatitis C virus in adults
(HCV) infection in adults

Scope: Opinion

Action: For adoption

List of Questions adopted on 19.04.2017.

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

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

Ammtek; treatment of neonatal diabetes

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 24.01.2017.

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

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

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 23.02.2017.

3.2.3. entecavir - EMEA/H/C/004458

treatment of chronic hepatitis B virus infection

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 15.12.2016.

3.2.4. lacosamide - EMEA/H/C/004443

treatment of epilepsy

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 26.01.2017.

3.2.5. miglustat - EMEA/H/C/004366

treatment of Gaucher disease

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 15.12.2016.

3.2.6. sirukumab - EMEA/H/C/004165

treatment of rheumatoid arthritis

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 26.01.2017.

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

Accelerated assessment

Santen Oy; treatment of severe vernal keratoconjunctivitis (VKC)

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 19.04.2017.

3.2.8. niraparib - Orphan - EMEA/H/C/004249

Tesaro UK Limited; Treatment of epithelial ovarian, fallopian tube, or primary peritoneal cancer

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. brigatinib - EMEA/H/C/004248

treatment of anaplastic lymphoma kinase (ALK)-positive non-small cell lung cancer (NSCLC)

Scope: Day 120 list of questions

Action: For adoption

3.3.2. ropeginterferon alfa-2b - Orphan - EMEA/H/C/004128

AOP Orphan Pharmaceuticals AG; treatment of polycythemia vera

Scope: Day 120 list of questions

Action: For adoption

3.3.3. caplacizumab - Orphan - EMEA/H/C/004426

Ablynx NV; indicated for the treatment of acquired thrombotic thrombocytopenic purpura (aTTP).

Scope: Day 120 list of questions

Action: For adoption

3.3.4. imatinib - EMEA/H/C/004748

treatment of newly diagnosed and chronic Philadelphia chromosome (bcr-abl) positive (Ph+) chronic myeloid leukaemia (CML), gastrointestinal stromal tumours (GIST), unresectable dermatofibrosarcoma protuberans (DFSP) and recurrent and/or metastatic DFSP

Scope: List of questions

Action: For adoption

3.3.5. nitisinone - EMEA/H/C/004582

treatment of hereditary tyrosinemia type 1

Scope: Day 120 list of questions

Action: For adoption

3.3.6. sodium benzoate - Orphan - EMEA/H/C/004150

Lucane Pharma; treatment of non ketotic hyperglycinemia, urea cycle disorders including carbamoyl-phosphate synthase-1 deficiency, ornithine transcarbamylase deficiency, citrullinaemia type 1, argininosuccinic aciduria, hyperargininaemia, n-acetylglutamate synthase deficiency, ornithine translocase deficiency and lysinuric protein intolerance

Scope: Day 120 list of questions

Action: For adoption

3.3.7. ertugliflozin / metformin hydrochloride - EMEA/H/C/004314

treatment of type 2 diabetes mellitus

Scope: Day 120 list of questions

Action: For adoption

3.3.8. ertugliflozin - EMEA/H/C/004315

type 2 diabetes mellitus

Scope: Day 120 list of questions

Action: For adoption

3.3.9. ertugliflozin / sitagliptin - EMEA/H/C/004313

type 2 diabetes mellitus

Scope: Day 120 list of questions

Action: For adoption

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

3.4.1. rurioctocog alfa pegol - EMEA/H/C/004195

treatment of haemophilia A

Scope: Letter from the applicant dated 12 June 2017 requesting an extension of clock stop to respond to List of Outstanding Issues adopted on 21.04.2017.

Action: For adoption

List of Outstanding Issues adopted on 21.04.2017, 15.12.2016. List of Questions adopted on 21.07.2016.

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

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

Scope: Request for an extension to the clock stop to respond to the list of questions adopted on 21.04.2017

Action: For adoption

List of Questions adopted on 21.04.2017.

3.4.3. entecavir - EMEA/H/C/004377

treatment of chronic hepatitis B virus infection

Scope: Request for an extension to the 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.

3.4.4. pegfilgrastim - EMEA/H/C/004262

treatment of neutropenia

Scope: Update on the timetable for the assessment of the Applicant's responses to the list of questions adopted on 13.10.2016.

Action: For adoption

List of questions adopted on 13.10.2016

3.4.5. trastuzumab - EMEA/H/C/004346

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

Scope: Update on the timetable for the assessment of the Applicant's responses to 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.

3.4.6. pegfilgrastim - EMEA/H/C/004413

treatment of neutropenia

Scope: Request for an extension to the clock stop to respond to the list of questions adopted on 23.03.2017

Action: For adoption

List of Questions adopted on 23.03.2017.

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

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 adoption

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.

3.5.2. Adlumiz - anamorelin - EMEA/H/C/003847

Helsinn Birex Pharmaceuticals Ltd; treatment of anorexia, cachexia or unintended weight loss in adult patients with non-small cell lung cancer (NSCLC)

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

Letter from the applicant dated 01 June 2017 requesting a re-examination of the Opinion adopted on 18 May 2017 and consultation of a Scientific Advisory Group

Action: For adoption

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

Opinion adopted on 18.05.2017. Oral explanation was held on 19.04.2017. List of Outstanding Issues adopted on 23.02.2017, 10.11.2016. List of Questions adopted on 25.02.2016.

3.5.3. Human IGG1 monoclonal antibody specific for human interleukin-1 alpha XBiotech - human IgG1 monoclonal antibody specific for human interleukin-1 alpha - EMEA/H/C/004388

XBiotech Germany GmbH; treatment of metastatic colorectal cancer

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

Letter from the applicant dated 02 June 2017 requesting a re-examination of the Opinion adopted on 18 May 2017 and consultation of a Scientific Advisory Group

Action: For adoption

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 15.12.2016. List of Questions adopted on 21.07.2016.

3.6. Initial applications in the decision-making phase

3.7. Withdrawals of initial marketing authorisation application

3.7.1. levamisole - Orphan - EMEA/H/C/004330

ACE Pharmaceuticals BV; treatment of Steroid Sensitive Nephrotic syndrome

Scope: Withdrawal of initial marketing authorisation application

Action: For information

List of Questions adopted on 15.12.2016.

3.7.2. ngr-htnf - Orphan - EMEA/H/C/004455

MolMed SpA; treatment of advanced malignant pleural mesothelioma

Scope: Withdrawal of initial marketing authorisation application

Action: For information

List of Questions adopted on 21.04.2017.

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

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

4.1.1. Mimpara - cinacalcet - EMEA/H/C/000570/X/0055/G

Amgen Europe B.V.

Rapporteur: Kristina Dunder, Co-Rapporteur: Andrea Laslop, PRAC Rapporteur: Ulla Wändel Liminga

Scope: "Extension application to introduce a new pharmaceutical form associated with new strengths (1 mg, 2.5 mg and 5 mg hard capsules) grouped with a type II variation (C.1.6.a) to include paediatric use in the approved indication.

As a consequence, sections 4.2 and 4.4 of the SmPC are updated to detail posology in paediatric patients and to update the safety information, respectively.

The Package Leaflet and Labelling are updated in accordance.

In addition, the Marketing authorisation holder (MAH) took the opportunity to update the list of local representatives in the Package Leaflet.

Furthermore, the PI is brought in line with the latest QRD template version 10."

Action: For adoption

List of Outstanding Issues adopted on 23.03.2017. List of Questions adopted on 13.10.2016.

4.1.2. SonoVue - sulphur hexafluoride - EMEA/H/C/000303/X/0034/G

Bracco International B.V.

Rapporteur: Alexandre Moreau, PRAC Rapporteur: Claire Ferard

Scope: "Extension application to introduce a new route of administration (intravesical use) grouped with a type II variation (C.I.6.a) to add a new indication (to include use in ultrasonography of the excretory urinary tract in paediatric patients to detect or exclude vesicoureteral reflux).
As a consequence, sections 4.1, 4.2, 4.8, 5.1 and 6 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.
Moreover, the updated RMP version 9.1 has been submitted as part of this application."

Action: For adoption

List of Questions adopted on 26.01.2017.

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. Benlysta - belimumab - EMEA/H/C/002015/X/0046/G

Glaxo Group Ltd

Rapporteur: Kristina Dunder, PRAC Rapporteur: Ulla Wändel Liminga

Scope: "Extension application to introduce a new pharmaceutical form (solution for injection), a new strength (200 mg) and a new route of administration (subcutaneous use) grouped with a type II variation (C.I.4) to include changes in the Product Information."

Action: For adoption

List of Questions adopted on 23.02.2017.

4.2.2. Exjade - deferasirox - EMEA/H/C/000670/X/0054

Novartis Europharm Ltd

Rapporteur: Alexandre Moreau, PRAC Rapporteur: Claire Ferard

Scope: "Extension application for a new pharmaceutical form (Exjade 90, 180 and 360 mg granules)."

Action: For adoption

List of Questions adopted on 23.02.2017.

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. ellaOne - ulipristal acetate - EMEA/H/C/001027/X/0045

Laboratoire HRA Pharma, SA

Rapporteur: Paula Boudewina van Hennik

Scope: "Addition of a new pharmaceutical form (film-coated tablets) to the existing strength 30 mg."

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. Alecensa - alectinib - EMEA/H/C/004164/II/0001

Roche Registration Limited

Rapporteur: Filip Josephson, Co-Rapporteur: Sinan B. Sarac, PRAC Rapporteur: Patrick Batty

Scope: "Extension of Indication to extend the indication of Alecensa (alectinib) to first line treatment of adult patients with anaplastic lymphoma kinase (ALK)-positive advanced non-small cell lung cancer (NSCLC); as a consequence, sections 4.1, 4.8, 5.1 and 5.2 of the SmPC are updated. The Package Leaflet and the RMP are updated in accordance."

Action: For adoption

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

Amgen Europe B.V.

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

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

Action: For adoption

5.1.3. Faslodex - fulvestrant - EMEA/H/C/000540/II/0057

AstraZeneca UK Ltd

Rapporteur: Filip Josephson, Co-Rapporteur: Tuomo Lapveteläinen, PRAC Rapporteur: Ulla Wändel Liminga

Scope: "Extension of Indication to include the treatment of postmenopausal women with locally advanced or metastatic breast cancer who have not received prior endocrine therapy for Faslodex. As a consequence, sections 4.1, 4.8 and 5.1 of the SmPC are updated in order to update the safety and pharmacodynamics information. The Package Leaflet is updated in accordance.

In addition, the Marketing authorisation holder (MAH) took the opportunity to introduce clarifications in the SmPC."

Action: For adoption

Request for Supplementary Information adopted on 23.02.2017.

5.1.4. Faslodex - fulvestrant - EMEA/H/C/000540/II/0059

AstraZeneca UK Ltd

Rapporteur: Filip Josephson, PRAC Rapporteur: Ulla Wändel Liminga

Scope: "Extension of Indication to include the use of Faslodex in combination with palbociclib for the treatment of hormone receptor (HR)-positive, human epidermal growth factor receptor 2 (HER2)-negative locally advanced or metastatic breast cancer in women who have received prior endocrine therapy (see section 5.1). In pre- or perimenopausal women, the combination treatment with palbociclib should be combined with a luteinizing hormone releasing hormone (LHRH) agonist for Faslodex

As a consequence, sections 4.1, 4.2, 4.4, 5.1, 5.3 and 6.6 of the SmPC are updated to update the safety and efficacy information.The Package Leaflet is updated in accordance. RMP version 11 was included in the application."

Action: For adoption

5.1.5. Harvoni - ledipasvir / sofosbuvir - EMEA/H/C/003850/II/0039

Gilead Sciences International Ltd

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

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 2) are updated in accordance."

Action: For adoption

Request for Supplementary Information adopted on 21.04.2017, 26.01.2017.

5.1.6. Kaletra - lopinavir / ritonavir - EMEA/H/C/000368/II/0161/G

AbbVie Ltd.

Rapporteur: Joseph Emmerich, PRAC Rapporteur: Caroline Laborde

Scope: "Extension of Indication to include children aged 14 days and older in the treatment of HIV-1.

As a consequence, sections 4.1, 4.2, 4.3, 4.8, 5.1 and 5.2 of the SmPC are updated. The Package Leaflet and Labelling are updated in accordance. The studies provided in support of the paediatric indication are part of the agreed PIP decision P/0144/2012.

In addition, the Marketing authorisation holder (MAH) further updated section 4.4 to add information regarding the use of Kaletra oral solution with feeding tubes.

The updated RMP v.8 is provided accordingly.

IB-B.II.e.5.a.2-To add a new pack size of 120 ml in (2X 60ml bottles) for Kaletra

80mg/ml/20 mg/ml oral solution (EU/1/01/172/003).

IA-B.IV.1.a.1-To add a new 2 ml oral dose syringe for the 120ml presentation."

Action: For adoption

Request for Supplementary Information adopted on 18.05.2017, 26.01.2017.

5.1.7. Keytruda - pembrolizumab - EMEA/H/C/003820/II/0027

Merck Sharp & Dohme Limited

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

Scope: "Extension of Indication to include 1st line treatment of metastatic non-squamous non-small cell lung cancer (NSCLC) in combination with platinum-pemetrexed chemotherapy based on the results from study KEYNOTE-021 (cohort G); a Phase 1/2, open-label trial of pembrolizumab in combination with chemotherapy or immunotherapy in patients with locally advanced or metastatic NSCLC.
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.
An updated RMP version 9.0 was provided as part of the application."

Action: For adoption

5.1.8. 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 23.03.2017.

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

Bristol-Myers Squibb Pharma EEIG

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

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

Action: For adoption

Request for Supplementary Information adopted on 23.03.2017.

5.1.10. Orencia - abatacept - EMEA/H/C/000701/II/0105

Bristol-Myers Squibb Pharma EEIG

Rapporteur: Outi Mäki-Ikola, Co-Rapporteur: Agnes Gyurasics, PRAC Rapporteur: Kirsti Villikka

Scope: "Extension of Indication to include a new indication for Orencia: treatment of psoriatic arthritis in adults.

As a consequence, sections 4.1, 4.2, 4.8, 5.1 and 5.2 of the SmPC are proposed to be updated. The Package Leaflet is updated in accordance.

In addition, the Marketing authorisation holder (MAH) took the opportunity to update the list of local representatives in the Package Leaflet.

A revised RMP was included in this submission (version 21)."

Action: For adoption

Request for Supplementary Information adopted on 26.01.2017.

5.1.11. Prolia - denosumab - EMEA/H/C/001120/II/0068

Amgen Europe B.V.

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

Scope: "Extension of Indication to include "Treatment of osteoporosis associated with sustained systemic glucocorticoid therapy in women and men at increased risk of fracture. Prevention of osteoporosis in women and men at increased risk of fracture who are starting or have recently started long-term glucocorticoid therapy." for Prolia; as a consequence, sections 4.1 and 5.1 of the SmPC are updated to reflect the new indications or are

consequential to the analysis of the data from the pivotal study. The Package Leaflet is updated in accordance.
The Risk Management Plan version 19.0 has also been updated to capture the new indications.
The variation proposed amendments to the Summary of Product Characteristics and Package Leaflet."

Action: For adoption

5.1.12. 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 23.03.2017.

5.1.13. Soliris - eculizumab - Orphan - EMEA/H/C/000791/II/0090

Alexion Europe SAS

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

Scope: "Extension of Indication of Soliris to include the 'treatment of Refractory generalized Myasthenia Gravis (gMG) patients who are antiacetylcholine receptor (AChR) antibody-positive'.
As a consequence, sections 4.1, 4.2, 4.4, 4.8, 5.1 and 5.2 of the SmPC are updated to include information on the new indication and to include the new methodology to calculate the Adverse Drug Reaction frequencies (section 4.8).
The RMP is updated accordingly (version 14.0)."

Action: For adoption

Request for Supplementary Information adopted on 23.03.2017.

5.1.14. Stivarga - regorafenib - EMEA/H/C/002573/II/0020

Bayer Pharma AG

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

Scope: "Extension of indication of Stivarga to include treatment of adult patients with hepatocellular carcinoma (HCC) who have been previously treated with one systemic therapy.

As a consequence, sections 4.1, 4.2, 4.4, 4.8 and 5.1 of the EU SmPC are updated. The package leaflet and RMP (version 5.0) have been updated accordingly.

Furthermore, the PI is brought in line with the latest QRD template version 10.0."

Action: For adoption

Request for Supplementary Information adopted on 18.05.2017, 23.02.2017.

5.1.15. Victoza - liraglutide - EMEA/H/C/001026/II/0042

Novo Nordisk A/S

Rapporteur: Johann Lodewijk Hillege, Co-Rapporteur: Hanne Lomholt Larsen, PRAC Rapporteur: Menno van der Elst

Scope: "Extension of Indication to include a new indication/population in Section 4.1 of the SmPC for Victoza.

As a consequence, sections 4.2, 4.4, 4.7, 4.8, 5.1 and 6.5 of the SmPC are updated to add warnings and update the safety information based on the findings of the LEADER (EX2211-3748) clinical study results, which constitutes the data set for the application. The Package Leaflet and Labelling (sections 17 and 18) are updated in accordance.

Updates to the liraglutide RMP based on the LEADER study results are also proposed: RMP Version 27 was submitted with the application, showing the proposed RMP changes."

Action: For adoption

Request for Supplementary Information adopted on 18.05.2017, 23.02.2017.

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

Bristol-Myers Squibb Pharma EEIG

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

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

Action: For adoption

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

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. Autologous T cells transduced with retroviral vector encoding an anti-CD19 CD28/CD3-zeta chimeric antigen receptor – Orphan - EMA/H/C/0004480

Kite Pharma UK Ltd; Intended for the treatment of adult patients with diffuse large B-cell lymphoma (DLBCL) who have not responded to their prior therapy, or have had disease progression after autologous stem cell transplant (ASCT)

Scope: Briefing note and CAT 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. 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."

Action: For adoption

9.1.2. Humalog-EMEA/H/C/000088 & Liprolog-EMEA/H/C/000393 - WS1158/0117/G

MAH: Eli Lilly Nederland B.V.

Lead Rapporteur: Robert James Hemmings, Lead PRAC Rapporteur: Julie Williams

Scope: "Type II (B.IV.1.c): to add a pre-filled pen: the Humalog and Liprolog 100 U/ml Junior KwikPen. The Junior KwikPen can administer insulin in half unit increments and contains the insulin lispro 3ml cartridge that is already approved for use. The pack contains 5 pre-filled pens.

Type IAin (B.II.e.5.a.1): to add a new pack size of 10 (2x5) pre-filled pens (multipack) for the Humalog and Liprolog 100 U/ml Junior KwikPen. This presentation contains the insulin lispro 3ml cartridge that is already approved for use.

Type II (C.I.z): to change the SmPC of the already authorised 100 U/ml Humalog and Liprolog presentations: to change section 4.2 to include the paediatric population; to change section 4.4 to remove text that states that the product should only be used in children in preference to soluble insulin when a fast action of insulin might be beneficial. The PL is updated accordingly."

Action: For adoption

Request for Supplementary Information adopted on 05.05.2017.

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

10.2.1. Desloratadine-containing products - desloratadine - EMEA/H/A-5(3)/1431

Rapporteur: Koenraad Norga, Co-Rapporteur: Andrea Laslop

Scope: Discussion/Opinion

Prescription status of desloratadine-containing products

Action: For adoption

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

10.4.1. 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: Oral explanation

Action: Oral explanation to be held on 19 June 2017 at time 16:00

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.

List of Outstanding issues adopted on 21.04.2017. List of Questions adopted on 26.01.2017.

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. Symbioflor 2, Escherichia Coli bacteria (cells and autolysate) - EMEA/H/A-31/1441

Symbiopharm GmbH,

Rapporteur: Harald Enzmann, Co-rapporteur: Milena Stain;

Scope: Opinion

Article 31 triggered by the BfArM in Germany in March 2016 requesting the review of the benefit-risk balance for Symbioflor 2 and associated names following concerns that the effectiveness of the medicine(s) has not been adequately demonstrated.

Action: For adoption

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

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

June 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. Seating plan for CHMP under Estonian EU Presidency, 1 July – 31 December 2017

CHMP Seating Plan 1 July – 31 December 2017, under Estonian EU presidency

Action: For information

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 6-9 June 2017

Action: For information

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

Action: For adoption

14.2.2. Committee for Advanced Therapies (CAT)

CAT draft minutes of meeting held on 15-16 June 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. Committee for Herbal Medicinal Products (HMPC)

Report from the HMPC meeting held on 29-30 May 2017

Action: For information

14.2.4. Paediatric Committee (PDCO)

PIPs reaching D30 at June 2017 PDCO

Action: For information

Report from the PDCO meeting held on 20-23 June 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 discussion

14.2.5. Committee for Orphan Medicinal Products (COMP)

Report from the COMP meeting held on 13-15 June 2017
Action: For information

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

Report from the Coordination Group for Mutual Recognition and Decentralised Procedures – Human (CMDh) on the meeting held on 19-21 June 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 6-9 June 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)

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

Action: For adoption

Reflection paper on the dissolution specification for generic solid oral immediate release products with systemic action (EMA/257304/2017)

Action: For adoption

Overview of comments (EMA/257305/2017)

Action: For information

14.3.3. Name Review Group (NRG)

Table of Decisions of the NRG meeting held on 31 May 2017.

Action: For adoption

14.3.4. Pharmacogenomics Working Party (PGWP)

Chair: Krishna Prasad/Markus Paulmichl

PGWP report on any differences of the metabolism and the clinical implications of the CYP2D6 genetic polymorphisms between codeine, dihydrocodeine, ethylmorphine, oxycodone and tramadol (EMA/CHMP/211025/2017)

Rapporteur: Marc Maliepaard

Action: For adoption

Concept paper on an Addendum on terms and concepts of pharmacogenomic features related to metabolism to the Guideline on the use of pharmacogenetic methodologies in the pharmacokinetic evaluation of medicinal products (EMA/CHMP/37646/2009)

Rapporteurs: Marc Maliepaard, Adrian Llerena

Action: For adoption for 3 months public consultation

14.3.5. Pharmacokinetics Working Party (PKWP)

Chair: Jan Welink/Alfredo Garcia-Arieta

Q&A on Biowaivers for BCS Class III substances

Rapporteur: Carolien Versantvoort

Action: For discussion

14.4. Cooperation within the EU regulatory network

14.4.1. Results from centralised initial MAA EMA-Industry-Rapporteurs survey – 2016-2017

Results from the MAA survey which took place from September 2016 to February 2017

Action: For information

14.5. Cooperation with International Regulators

14.5.1. EMA/FDA strategic document on Gaucher disease

Action: For adoption

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

14.7. CHMP work plan

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

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)*

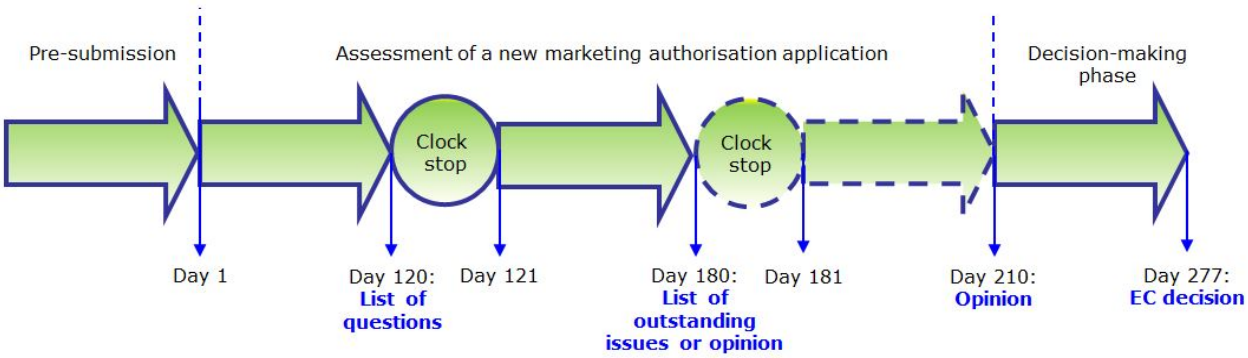
The items listed in this section are those for which marketing authorisation holders (MAHs) or applicants have been invited to the CHMP plenary meeting to address questions raised by the Committee. Oral explanations normally relate to on-going applications (section 3, 4 and 5) or referral procedures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

19 June 2017
EMA/CHMP/309580/2017

Annex to June 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 June 2017: **For adoption**

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

Final Outcome of Rapporteurship allocation for June 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

Firdapse - amifampridine - EMEA/H/C/001032/S/0049, Orphan
MAH: BioMarin Europe Ltd, Rapporteur: Greg Markey, PRAC Rapporteur: Julie Williams

B.2. RENEWALS OF MARKETING AUTHORISATIONS OUTCOMES

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

Constella - linaclotide - EMEA/H/C/002490/R/0032
MAH: Allergan Pharmaceuticals International Limited, Rapporteur: Martina Weise, Co-Rapporteur: Concepcion Prieto Yerro, PRAC Rapporteur: Valerie Strassmann
Request for Supplementary Information adopted on 18.05.2017.

B.2.2. Renewals of Marketing Authorisations for unlimited validity

Ecalta - anidulafungin - EMEA/H/C/000788/R/0033

MAH: Pfizer Limited, Rapporteur: Johann Lodewijk Hillege, Co-Rapporteur: Hanne Lomholt Larsen, PRAC Rapporteur: Sabine Straus
Request for Supplementary Information adopted on 21.04.2017.

Forxiga - dapagliflozin - EMEA/H/C/002322/R/0035
MAH: AstraZeneca AB, Rapporteur: Kristina Dunder, Co-Rapporteur: Martina Weise, PRAC Rapporteur: Qun-Ying Yue

Glubrava - pioglitazone / metformin hydrochloride - EMEA/H/C/000893/R/0054
MAH: Takeda Pharma A/S, Informed Consent of Competact, Rapporteur: Patrick Salmon, Co-Rapporteur: Concepcion Prieto Yerro, PRAC Rapporteur: Almath Spooner

NexoBrid - concentrate of proteolytic enzymes enriched in bromelain - EMEA/H/C/002246/R/0031, Orphan
MAH: MediWound Germany GmbH, Rapporteur: Harald Enzmann, Co-Rapporteur: Robert James Hemmings, PRAC Rapporteur: Valerie Strassmann

Zoledronic acid Hospira - zoledronic acid - EMEA/H/C/002365/R/0026
MAH: Hospira UK Limited, Rapporteur: Kristina Dunder, PRAC Rapporteur: Doris Stenver

B.2.3. Renewals of Conditional Marketing Authorisations

B.3. POST-AUTHORISATION PHARMACOVIGILANCE OUTCOMES

Truvada EMEA/H/C/000594 MEA 045
(Emtricitabine / tenofovir disoproxil) PRAC Rapporteur: Julie Williams; **Action:** For adoption

Signal detection

PRAC recommendations on signals adopted at the PRAC meeting held on 6 - 9 June 2017
PRAC:

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

EMEA/H/C/PSUSA/00000939/201610
(deferasirox)
CAPS:
Exjade (EMEA/H/C/000670) (deferasirox), MAH: Novartis Europharm Ltd, Rapporteur: Alexandre Moreau, PRAC Rapporteur: Claire Ferard, “01 Nov 2015 – 31 Oct 2016”

EMEA/H/C/PSUSA/00001653/201609
(hydrochlorothiazide / irbesartan)
CAPS:
CoAprovel (EMEA/H/C/000222) (irbesartan / hydrochlorothiazide), MAH: Sanofi Clir SNC, Rapporteur: Concepcion Prieto Yerro
Irbesartan Hydrochlorothiazide Zentiva (EMEA/H/C/000783) (irbesartan / hydrochlorothiazide), MAH: sanofi-aventis groupe, Rapporteur: Concepcion Prieto Yerro
Karvezide (EMEA/H/C/000221) (irbesartan / hydrochlorothiazide), MAH: sanofi-aventis groupe, Rapporteur: Concepcion Prieto Yerro
NAPS:
NAPs - EU
, PRAC Rapporteur: Dolores Montero Corominas, “30-Sep-2013 to 30-Sep-2016”

EMEA/H/C/PSUSA/00001758/201609
(interferon alpha-2b)
CAPS:
IntronA (EMEA/H/C/000281) (interferon alfa-2b), MAH: Merck Sharp & Dohme Limited, Rapporteur: Koenraad Norga
NAPS:
NAPs – EU - PRAC Rapporteur: Jean-Michel Dogné, “21-9-2011 – 20-9-2016”

EMEA/H/C/PSUSA/00002666/201611
(rotavirus vaccine pentavalent (live, oral))
CAPS:
RotaTeq (EMEA/H/C/000669) (rotavirus vaccine (live, oral)), MAH: MSD Vaccins, Rapporteur: Greg Markey, PRAC Rapporteur: Julie Williams, “28 November 2015 to 27 November 2016”

EMEA/H/C/PSUSA/00010132/201611
(radium-223 dichloride)
CAPS:
Xofigo (EMEA/H/C/002653) (radium-223), MAH: Bayer AG, Rapporteur: Harald Enzmann, PRAC Rapporteur: Patrick Batty, “15-MAY-2016 - 14-NOV-2016”

EMEA/H/C/PSUSA/00010262/201611 (trametinib) CAPS: **Mekinist** (EMEA/H/C/002643) (trametinib), MAH: Novartis Europharm Ltd, Rapporteur: Paula Boudewina van Hennik, PRAC Rapporteur: Patrick Batty, "30th May 2016 to 29th November 2016"	
EMEA/H/C/PSUSA/00010316/201611 (ketoconazole (centrally authorised product only)) CAPS: **Ketoconazole HRA** (EMEA/H/C/003906) (ketoconazole), MAH: Laboratoire HRA Pharma, Rapporteur: Concepcion Prieto Yerro, PRAC Rapporteur: Željana Margan Koletić, "20 May 2016 to 19 November 2016"	
EMEA/H/C/PSUSA/00010455/201611 (lumacaftor / ivacaftor) CAPS: **Orkambi** (EMEA/H/C/003954) (lumacaftor / ivacaftor), MAH: Vertex Pharmaceuticals (Europe) Ltd., Rapporteur: Nithyanandan Nagercoil, PRAC Rapporteur: Almath Spooner, "20 May 2016 to 19 November 2016"	
EMEA/H/C/PSUSA/00010114/201610 (lurasidone) CAPS: **Latuda** (EMEA/H/C/002713) (lurasidone), MAH: Sunovion Pharmaceuticals Europe Ltd, Rapporteur: Filip Josephson, PRAC Rapporteur: Qun-Ying Yue, "28/10/2015 to 28/10/2016 Update of section 4.8 of the SmPC to add the adverse reaction hyponatremia with a frequency uncommon, to change the frequency of the adverse reaction hypersensitivity, rash and pruritus from not known to uncommon, and to change frequency of angioedema from not known to rare. Under the table, the footnote regarding hypersensitivity is deleted. The Package Leaflet is updated accordingly." **Action:** For adoption	Correction of the opinion and assessment report adopted in May 2017.

B.4. EPARs / WPARs

Adlumiz - anamorelin - EMEA/H/C/003847
Applicant: Helsinn Birex Pharmaceuticals Ltd, treatment of anorexia, cachexia or unintended weight loss in adult patients with non-small cell lung cancer (NSCLC), New active substance (Article 8(3) of Directive No 2001/83/EC)

Blitzima - rituximab - EMEA/H/C/004723
Applicant: Celltrion Healthcare Hungary Kft., treatment of Non-Hodgkin's lymphoma (NHL) and Chronic lymphocytic leukaemia (CLL) and Granulomatosis with polyangiitis and microscopic polyangiitis, Duplicate, Duplicate of Truxima, Similar biological application (Article 10(4) of Directive No 2001/83/EC)

Efavirenz/Emtricitabine/Tenofovir disoproxil Zentiva - efavirenz / emtricitabine / tenofovir disoproxil - EMEA/H/C/004250
Applicant: Zentiva k.s., treatment of HIV-1 infection, Generic, Generic of Atripla, Generic application (Article 10(1) of Directive No 2001/83/EC)

Elmisol (WD) - levamisole - EMEA/H/C/004330, Orphan
Applicant: ACE Pharmaceuticals BV, treatment of Steroid Sensitive Nephrotic syndrome, Known active substance (Article 8(3) of Directive No 2001/83/EC)
WPAR

Human IGG1 monoclonal antibody specific for human interleukin-1 alpha XBiotech - human IgG1 monoclonal antibody specific for human interleukin-1 alpha - EMEA/H/C/004388
Applicant: XBiotech Germany GmbH, treatment of metastatic colorectal cancer, New active substance (Article 8(3) of Directive No 2001/83/EC)

Insulin lispro Sanofi - insulin lispro - EMEA/H/C/004303
Applicant: sanofi-aventis groupe, treatment of diabetes mellitus, Similar biological application (Article 10(4) of Directive No 2001/83/EC)

Kyntheum - brodalumab - EMEA/H/C/003959
Applicant: LEO Pharma A/S, moderate to severe

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

Masipro - masitinib - EMEA/H/C/004159, Orphan
Applicant: AB Science, treatment of mastocytosis, New active substance (Article 8(3) of Directive No 2001/83/EC)

Oxervate - cenegermin - EMEA/H/C/004209, Orphan
Applicant: Dompe farmaceutici S.p.A., treatment of neurotrophic keratitis, New active substance (Article 8(3) of Directive No 2001/83/EC)

Reagila - cariprazine - EMEA/H/C/002770
Applicant: Gedeon Richter Plc., treatment of schizophrenia, New active substance (Article 8(3) of Directive No 2001/83/EC)

Ritemvia - rituximab - EMEA/H/C/004725
Applicant: Celltrion Healthcare Hungary Kft., treatment of Non-Hodgkin's lymphoma (NHL), Granulomatosis with polyangiitis and microscopic polyangiitis, Similar biological application (Article 10(4) of Directive No 2001/83/EC)

Spherox - spheroids of human autologous matrix-associated chondrocytes - EMEA/H/C/002736, ATMP
Applicant: CO.DON AG, treatment of cartilage defects, New active substance (Article 8(3) of Directive No 2001/83/EC)

Trimbow - beclometasone dipropionate anhydrous / formoterol fumarate dihydrate / glycopyrronium bromide - EMEA/H/C/004257
Applicant: Chiesi Farmaceutici S.p.A., symptomatic treatment and reduction of exacerbations in adult patients with chronic obstructive pulmonary disease (COPD) with airflow limitation and who are at risk of exacerbations, Fixed combination application (Article 10b of Directive No 2001/83/EC)

Tuxella - rituximab - EMEA/H/C/004724
Applicant: Celltrion Healthcare Hungary Kft., treatment of Non-Hodgkin's lymphoma (NHL), Chronic lymphocytic leukaemia (CLL) , Granulomatosis with polyangiitis and microscopic polyangiitis, Duplicate, Duplicate of

Truxima, Similar biological application (Article 10(4) of Directive No 2001/83/EC)	
Veltassa - patiromer sorbitex calcium - EMEA/H/C/004180 Applicant: Vifor Fresenius Medical Care Renal Pharma France, treatment of hyperkalaemia, 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

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. The Committee adopted a Request for Supplementary information together with a specific timetable.
Apidra - insulin glulisine - EMEA/H/C/000557/II/0074/G MAH: Sanofi-Aventis Deutschland GmbH, Rapporteur: Greg Markey Opinion adopted on 09.06.2017.	Positive Opinion adopted by consensus on 09.06.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Betaferon - interferon beta-1b - EMEA/H/C/000081/II/0114 MAH: Bayer AG, Rapporteur: Greg Markey	Weekly start timetable.
Colobreathe - colistimethate sodium - EMEA/H/C/001225/II/0031 MAH: Teva B.V., Rapporteur: Nithyanandan Nagercoil	Weekly start timetable.
Elocta - efmoroctocog alfa - EMEA/H/C/003964/II/0012/G MAH: Swedish Orphan Biovitrum AB (publ), Rapporteur: Jan Mueller-Berghaus Request for Supplementary Information adopted on 21.04.2017.	Weekly start timetable.
Elonva - corifollitropin alfa - EMEA/H/C/001106/II/0036/G MAH: Merck Sharp & Dohme Limited, Rapporteur: Paula Boudewina van Hennik Request for Supplementary Information adopted on 09.06.2017.	Weekly start timetable. The Committee adopted a Request for Supplementary information together with a specific timetable.

Extavia - interferon beta-1b - EMEA/H/C/000933/II/0084 MAH: Novartis Europharm Ltd, Informed Consent of Betaferon, Rapporteur: Greg Markey	Weekly start timetable.
Flixabi - infliximab - EMEA/H/C/004020/II/0013/G MAH: Samsung Bioepis UK Limited (SBUK), Rapporteur: Jan Mueller-Berghaus Request for Supplementary Information adopted on 01.06.2017.	Weekly start timetable. The Committee adopted a Request for Supplementary information together with a specific timetable.
Foclivia - influenza virus surface antigens (inactivated) of strain A/Vietnam/1194/2004 (H5N1) - EMEA/H/C/001208/II/0027 MAH: Seqirus S.r.l, Rapporteur: Daniela Melchiorri Request for Supplementary Information adopted on 18.05.2017, 06.04.2017.	Weekly start timetable.
Ganfort - bimatoprost / timolol - EMEA/H/C/000668/II/0027/G MAH: Allergan Pharmaceuticals Ireland, Rapporteur: Hanne Lomholt Larsen Request for Supplementary Information adopted on 09.06.2017.	Weekly start timetable. The Committee adopted a Request for Supplementary information together with a specific timetable.
GONAL-f - follitropin alfa - EMEA/H/C/000071/II/0138/G MAH: Merck Serono Europe Limited, Rapporteur: Nithyanandan Nagercoil	Weekly start timetable.
Hizentra - human normal immunoglobulin - EMEA/H/C/002127/II/0074/G MAH: CSL Behring GmbH, Rapporteur: Jan Mueller-Berghaus Request for Supplementary Information adopted on 23.02.2017.	
Humira - adalimumab - EMEA/H/C/000481/II/0167 MAH: AbbVie Ltd., Rapporteur: Kristina Dunder	Weekly start timetable.
Imatinib Teva - imatinib - EMEA/H/C/002585/II/0026 MAH: Teva B.V., Generic, Generic of Glivec, Rapporteur: Jorge Camarero Jiménez Opinion adopted on 09.06.2017. Request for Supplementary Information adopted on 27.04.2017.	Positive Opinion adopted by consensus on 09.06.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Imvanex - modified vaccinia Ankara virus - EMEA/H/C/002596/II/0027	Weekly start timetable.

MAH: Bavarian Nordic A/S, Rapporteur: Greg Markey	
Insuman - insulin human - EMEA/H/C/000201/II/0117/G MAH: Sanofi-aventis Deutschland GmbH, Rapporteur: Bart Van der Schueren	Weekly start timetable.
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. The Committee adopted a Request for Supplementary information together with a specific 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. The Committee adopted a Request for Supplementary information together with a specific timetable.
Menveo - meningococcal group A, C, W135 and Y conjugate vaccine - EMEA/H/C/001095/II/0065 MAH: GSK Vaccines S.r.l, Rapporteur: Johann Lodewijk Hillege Request for Supplementary Information adopted on 27.04.2017.	Weekly start timetable.
Mysimba - naltrexone hydrochloride / bupropion hydrochloride - EMEA/H/C/003687/II/0013/G MAH: Orexigen Therapeutics Ireland Limited, Rapporteur: Hanne Lomholt Larsen Opinion adopted on 01.06.2017. Request for Supplementary Information adopted on 06.04.2017.	Positive Opinion adopted by consensus on 01.06.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
NovoRapid - insulin aspart - EMEA/H/C/000258/II/0118 MAH: Novo Nordisk A/S, Rapporteur: Kristina Dunder	Weekly start timetable.
Nucala - mepolizumab - EMEA/H/C/003860/II/0007 MAH: GlaxoSmithKline Trading Services, Rapporteur: Nithyanandan Nagercoil Request for Supplementary Information adopted on 18.05.2017, 23.03.2017.	Weekly start timetable.
Praluent - alirocumab - EMEA/H/C/003882/II/0022/G MAH: sanofi-aventis groupe, Rapporteur:	Positive Opinion adopted by consensus on 01.06.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP

Johann Lodewijk Hillege Opinion adopted on 01.06.2017.	recommendation.
Prevenar 13 - pneumococcal polysaccharide conjugate vaccine (13-valent, adsorbed) - EMEA/H/C/001104/II/0156 MAH: Pfizer Limited, Rapporteur: Kristina Dunder Opinion adopted on 01.06.2017.	Positive Opinion adopted by consensus on 01.06.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Prezista - darunavir - EMEA/H/C/000707/II/0083/G MAH: Janssen-Cilag International NV, Rapporteur: Johann Lodewijk Hillege Request for Supplementary Information adopted on 18.05.2017, 26.01.2017.	
Simponi - golimumab - EMEA/H/C/000992/II/0075/G MAH: Janssen Biologics B.V., Rapporteur: Kristina Dunder Opinion adopted on 09.06.2017.	Positive Opinion adopted by consensus on 09.06.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Sustiva - efavirenz - EMEA/H/C/000249/II/0142/G MAH: Bristol-Myers Squibb Pharma EEIG, Rapporteur: Bruno Sepodes Opinion adopted on 01.06.2017.	Positive Opinion adopted by consensus on 01.06.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Synflorix - pneumococcal polysaccharide conjugate vaccine (adsorbed) - EMEA/H/C/000973/II/0116/G MAH: GSK Biologicals SA, Rapporteur: Kristina Dunder Request for Supplementary Information adopted on 18.05.2017.	Weekly start timetable.
TachoSil - human thrombin / human fibrinogen - EMEA/H/C/000505/II/0077/G MAH: Takeda Austria GmbH, Rapporteur: Jan Mueller-Berghaus Opinion adopted on 01.06.2017. Request for Supplementary Information adopted on 06.04.2017.	Positive Opinion adopted by consensus on 01.06.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Thyrogen - thyrotropin alfa - EMEA/H/C/000220/II/0090 MAH: Genzyme Europe BV, Rapporteur: Patrick Salmon Opinion adopted on 01.06.2017. Request for Supplementary Information adopted	Positive Opinion adopted by consensus on 01.06.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

on 06.04.2017.	
Vihuma - simoctocog alfa - EMEA/H/C/004459/II/0001/G MAH: Octapharma AB, Rapporteur: Jan Mueller-Berghaus	Weekly start timetable.
Vimizim - elosulfase alfa - EMEA/H/C/002779/II/0017/G, Orphan MAH: BioMarin Europe Ltd, Rapporteur: Johann Lodewijk Hillege Request for Supplementary Information adopted on 30.03.2017.	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. The Committee adopted a Request for Supplementary information together with a specific timetable.
Zepatier - elbasvir / grazoprevir - EMEA/H/C/004126/II/0007 MAH: Merck Sharp & Dohme Limited, Rapporteur: Greg Markey Opinion adopted on 01.06.2017.	Positive Opinion adopted by consensus on 01.06.2017. 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 05.05.2017.	Weekly start timetable.
WS1145/G **Aflunov- EMEA/H/C/002094/WS1145/0034/G** **Foclivia- EMEA/H/C/001208/WS1145/0029/G** MAH: Seqirus S.r.l, Lead Rapporteur: Daniela Melchiorri Request for Supplementary Information adopted on 11.05.2017.	Weekly start timetable.
WS1150/G **Infanrix hexa- EMEA/H/C/000296/WS1150/0218/G** MAH: GSK Biologicals SA, Lead Rapporteur: Bart Van der Schueren	Weekly start timetable.
WS1166	Weekly start timetable.

Infanrix hexa- **EMEA/H/C/000296/WS1166/0219** MAH: GSK Biologicals SA, Lead Rapporteur: Bart Van der Schueren	

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

Alecensa - alectinib - **EMEA/H/C/004164/II/0003** MAH: Roche Registration Limited, Rapporteur: Filip Josephson, “Update of section 4.8 of the SmPC in order to add "Increased blood alkaline phosphatase" as new Adverse Drug Reaction with a common frequency identified during routine signal detection. The package leaflet is updated accordingly. In addition, the Marketing authorisation holder (MAH) took the opportunity to introduce some formatting changes in the Product Information.” Opinion adopted on 09.06.2017.	Positive Opinion adopted by consensus on 09.06.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
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 Data Sheet. The Package Leaflet is updated accordingly.”	Weekly start timetable.
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.	Weekly start timetable.

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."	
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. The Committee adopted a Request for Supplementary information together with a specific timetable.
Cosentyx - secukinumab - EMEA/H/C/003729/II/0021/G MAH: Novartis Europharm Ltd, Rapporteur: Tuomo Lapveteläinen, "Update of section 5.1 of the SmPC in order to add long term (52 week) data from the CLEAR study (CAIN457A2317) and to add new data from a scalp psoriasis study (CAIN457US01). In addition the MAH has taken the occasion to include a correction in section 4.2 of the SmPC to avoid medication errors -the Package Leaflet has been updated accordingly- and in section 5.1 of the SmPC to align the Psoriatic Arthritis Response Criteria (PsARC) definition to the relevant EMA guideline. The MAH has also implemented the latest QRD template version 10.0." Opinion adopted on 01.06.2017.	Positive Opinion adopted by consensus on 01.06.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

Eperzan - albiglutide - EMEA/H/C/002735/II/0031 MAH: GlaxoSmithKline Trading Services, Rapporteur: Kristina Dunder, “Update of sections 4.4 and 4.8 of the SmPC in order to add angioedema with frequency ‘rare’ and to include a warning concerning hypersensitivity reactions in general. The Package Leaflet is updated accordingly. In addition, the Marketing authorisation holder took the opportunity to implement minor editorial amendments throughout the Product Information.” Request for Supplementary Information adopted on 21.04.2017.	Weekly start timetable.
Eviplera - emtricitabine / rilpivirine / tenofovir disoproxil - EMEA/H/C/002312/II/0082 MAH: Gilead Sciences International Ltd, Rapporteur: Johann Lodewijk Hillege, “Update of section 4.5 of the SmPC with Drug-Drug Interaction information for Eviplera based on the results from Study TMC435-TiDP16-C114; this is a Phase I, 2-panel, open-label, randomized, cross-over trial in healthy subjects to investigate the pharmacokinetic interaction between TMC435 and antiretroviral agents, TMC278 and tenofovir disoproxil fumarate (TDF), at steady-state. The Package Leaflet is updated accordingly. In addition, the Marketing authorisation holder (MAH) took the opportunity to introduce minor administrative changes in the SmPC and to update the list of local representatives in the Package Leaflet for Estonia, Latvia and Lithuania. Minor linguistic amendments (MLAs) have been implemented to the translations of the product information annexes: CS, DE, ES, FR, IS, IT, NL, NO, PT, SE and SK.”	Weekly start timetable.
Glivec - imatinib - EMEA/H/C/000406/II/0108 MAH: Novartis Europharm Ltd, Rapporteur: Jorge Camarero Jiménez, “Submission of the final CSR for study STI571A2405 - the International Study for Chronic Myeloid Leukemia (CML) in childhood and adolescents (I-CML-Ped Study).	Weekly start timetable.

The provision of the study report addresses the post-authorisation measure MEA 162.8." Request for Supplementary Information adopted on 21.04.2017.	
Halaven - eribulin - EMEA/H/C/002084/II/0038 MAH: Eisai Europe Ltd., Rapporteur: Filip Josephson, "Update of the SmPC section 5.1 with additional information on the mechanism of action of eribulin. Furthermore, the MAH has taken the opportunity to include in the package leaflet the name of the manufacturer responsible for batch release to align with the Annex II of the Product Information, and to update information related to the local representatives." Request for Supplementary Information adopted on 01.06.2017.	Weekly start timetable. The Committee adopted a Request for Supplementary information together with a specific timetable.
Harvoni - ledipasvir / sofosbuvir - EMEA/H/C/003850/II/0052 MAH: Gilead Sciences International Ltd, Rapporteur: Filip Josephson, "Submission of the final report from study GS-US-337-0115 listed as a category 3 study in the RMP. This is a phase 3, multicentre, randomized, open-label study to investigate the efficacy and safety of sofosbuvir/ledipasvir fixed-dose combination ± ribavirin for 12 or 24 weeks in Subjects with chronic genotype hepatitis C virus (HCV) and human immunodeficiency virus (HIV)-1 co-infection."	Weekly start timetable.
Humira - adalimumab - EMEA/H/C/000481/II/0168 MAH: AbbVie Ltd., Rapporteur: Kristina Dunder, "Update of section 5.1 of the SmPC in order to update information on the long-term safety, tolerability, and efficacy of adalimumab in subjects with moderate to severe hidradenitis suppurativa after finalization of phase III open-label extension studyM12-555."	Weekly start timetable.
Ibrance - palbociclib - EMEA/H/C/003853/II/0006 MAH: Pfizer Limited, Rapporteur: Filip Josephson, "Update of sections 4.2, 4.8 and 5.1 in order to reflect the results of the study A5481008 (PALOMA-2) and of the Phase 2 portion of A5481010 single-arm study. The MAH took the opportunity to implement minor	

editorial changes to the PIL."	
Imbruvica - ibrutinib - EMEA/H/C/003791/II/0034, Orphan MAH: Janssen-Cilag International NV, Rapporteur: Filip Josephson, "Submission of the final report from non-clinical study 17-008-Sal-X-MU (AMES assays for major human metabolites M21 + M34) listed as a category 3 studies in the RMP. The in vitro metabolism report (FK10269) in Mod. 4.2.2.4 is amended to document the production of the metabolites M21 and M34." Opinion adopted on 09.06.2017.	Positive Opinion adopted by consensus on 09.06.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
IntronA - interferon alfa-2b - EMEA/H/C/000281/II/0110 MAH: Merck Sharp & Dohme Limited, Rapporteur: Koenraad Norga, "Update of section 4.8 of the SmPC in order to add pericarditis with the frequency uncommon based on continuous monitoring of the safety profile; the Package Leaflet is updated accordingly." Opinion adopted on 01.06.2017.	Positive Opinion adopted by consensus on 01.06.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Jardiance - empagliflozin - EMEA/H/C/002677/II/0025 MAH: Boehringer Ingelheim International GmbH, Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Dolores Montero Corominas, "Submission of the final results for a non-interventional study 1245.122 exploring the characteristics of patients initiating empagliflozin or other noninsulin glucose lowering drugs in the United Kingdom in order to fulfil MEA 009. The RMP (version 11.0) is updated accordingly." Request for Supplementary Information adopted on 23.02.2017.	
Kuvan - sapropterin - EMEA/H/C/000943/II/0048/G, Orphan MAH: BioMarin International Limited, Rapporteur: Patrick Salmon, "Update of section 4.9 to add information regarding shortening of QT interval at high doses following review of data of study QTC-001. Submission of the clinical study report EMR700773-004 (pilot study assessing the effect of sapropterin on cognitive abilities, study prematurely terminated due to enrolment issues)	Weekly start timetable.

In addition, the MAH took the opportunity of this procedure to clarify the wording of section 4.2 and section 3 of the PL." Request for Supplementary Information adopted on 11.05.2017, 09.03.2017.	
Lixiana - edoxaban - EMEA/H/C/002629/II/0012 MAH: Daiichi Sankyo Europe GmbH, Rapporteur: Concepcion Prieto Yerro, "Update of sections 4.2 and 5.1 of the SmPC in order to add information deriving from new clinical data for the use of edoxaban as anticoagulant therapy for patients with non-valvular atrial fibrillation undergoing cardioversion. The Package Leaflet is updated accordingly. In addition, the Marketing authorisation holder (MAH) took the opportunity to update the list of local representatives for Portugal in the Package Leaflet and to bring the PI in line with the latest QRD template version 10.0." Request for Supplementary Information adopted on 11.05.2017.	Weekly start timetable.
Multaq - dronedarone - EMEA/H/C/001043/II/0038 MAH: sanofi-aventis groupe, Rapporteur: Johann Lodewijk Hillege, "Submission of the final results of a single historic prospective observational study to evaluate the efficacy of dronedarone in clinical practice (EFFECT-AF study). The product information and RMP remain unchanged." Request for Supplementary Information adopted on 01.06.2017.	Weekly start timetable. The Committee adopted a Request for Supplementary information together with a specific timetable.
Mysimba - naltrexone hydrochloride / bupropion hydrochloride - EMEA/H/C/003687/II/0010 MAH: Orexigen Therapeutics Ireland Limited, Rapporteur: Hanne Lomholt Larsen, "Submission of study report NB-CVOT - Multicenter, Randomized, Double-Blind, Placebo-Controlled Study Assessing the Occurrence of Major Adverse Cardiovascular Events (MACE) in Overweight and Obese Subjects with Cardiovascular Risk Factors Receiving Naltrexone SR/Bupropion SR. The product information remains unchanged." Request for Supplementary Information adopted on 21.04.2017, 23.02.2017.	Weekly start timetable.

Mysimba - naltrexone hydrochloride / bupropion hydrochloride - EMEA/H/C/003687/II/0014 MAH: Orexigen Therapeutics Ireland Limited, Rapporteur: Hanne Lomholt Larsen, "C.I.13: Submission of the final report from study NaltrexBuprop-1004; a Phase 1, Open-Label, Sequential Design Study to Evaluate the Potential Effect of Multiple Oral Doses of Extended-Release Combination of Naltrexone and Bupropion on the Pharmacokinetics of a Single Oral Dose of Metformin in Healthy Subjects. This study does not lead to changes in the product information." Request for Supplementary Information adopted on 21.04.2017.	Weekly start timetable.
Mysimba - naltrexone hydrochloride / bupropion hydrochloride - EMEA/H/C/003687/II/0015 MAH: Orexigen Therapeutics Ireland Limited, Rapporteur: Hanne Lomholt Larsen, "Submission of the final report from study NB-404 A Multicenter, Randomized, Open-Label, Controlled, Method-of-Use Study Assessing the Effect of Naltrexone SR/Bupropion SR on Body Weight and Cardiovascular Risk Factors in Overweight and Obese Subjects. This study does not lead to changes in the product information." Request for Supplementary Information adopted on 21.04.2017.	Weekly start timetable.
Odomzo - sonidegib - EMEA/H/C/002839/II/0010 MAH: Sun Pharmaceutical Industries Europe B.V., Rapporteur: Paula Boudewina van Hennik, "Submission of the final results from the clinical pharmacology study CLDE225A2112, which was a Phase Ib, multi-center, two parallel groups, open-label, drug-drug interaction study to assess the effect of sonidegib on the pharmacokinetics of bupropion and warfarin in patients with advanced solid tumors. This study is listed as a measure in the RMP. The SmPC section 4.5 has been updated to reflect that the results of a drug-drug interaction study in cancer patients demonstrate that the systemic exposure of bupropion (a CYP2B6 substrate) and warfarin (a CYP2C9 substrate) is not altered when co-administered with sonidegib. The PIL has been amended accordingly."	Positive Opinion adopted by consensus on 01.06.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

Opinion adopted on 01.06.2017.	
Revolade - eltrombopag / eltrombopag olamine - EMEA/H/C/001110/II/0042 MAH: Novartis Europharm Ltd, Rapporteur: Concepcion Prieto Yerro, "Submission of the ASPIRE (TRC114968) final study report, a Three-Part Study of Eltrombopag in Thrombocytopenic Subjects with Myelodysplastic Syndromes or Acute Myeloid Leukemia (Part 1: Open-Label, Part 2: Randomized, Double-Blind, Part 3: Extension) assessing the potential risk of haematological changes, optimal dose escalation scheme and eltrombopag pharmacokinetics." Request for Supplementary Information adopted on 16.03.2017.	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 in order to state that Selincro may influence the ability to drive and use machines. Update of section 4.8 of the SmPC in order to add the adverse drug reaction "diarrhoea" with frequency "common". The Package Leaflet is updated accordingly. In addition, the Marketing authorisation holder (MAH) took the opportunity to update the list of local representatives in the Package Leaflet." Request for Supplementary Information adopted on 01.06.2017.	Weekly start timetable. The Committee adopted a Request for Supplementary information together with a specific timetable.
Sprycel - dasatinib - EMEA/H/C/000709/II/0055 MAH: Bristol-Myers Squibb Pharma EEIG, Rapporteur: Sinan B. Sarac, "Update of section 4.8 of the SmPC in order to add nephrotic syndrome as an adverse reaction based on the results of routine pharmacovigilance activities. The Package Leaflet is updated accordingly. In addition, the Marketing authorisation holder (MAH) took the opportunity to update the date of latest renewal (Section 9, SmPC) along with the phone number of the local representative in Croatia and the name of local representative in Ireland listed in the PIL." Opinion adopted on 09.06.2017.	Positive Opinion adopted by consensus on 09.06.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Strensiq - asfotase alfa - EMEA/H/C/003794/II/0018, Orphan MAH: Alexion Europe SAS, Rapporteur: Greg	Positive Opinion adopted by consensus on 09.06.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP

Markey, “Update of section 4.5 of the SmPC in order to update the information on Asfotase alfa interaction with the Alkaline Phosphatase (ALP), used as the detection reagent in many routine laboratory assays, which may leading to abnormal values reports. The Package Leaflet is updated accordingly.” Opinion adopted on 09.06.2017.	recommendation.
Tarceva - erlotinib - EMEA/H/C/000618/II/0051 MAH: Roche Registration Limited, Rapporteur: Sinan B. Sarac, “Submission of the Real World Data Reports (BIOMARQUEURS FRANCE CSR and ESCAP-2011-CPHG CSR) and a new CSR Addendum of the previously submitted pivotal study BR.21, in order to discuss the currently available evidence supporting the use of erlotinib for treatment of patients with locally advanced or metastatic NSCLC without EGFR-activating mutations after failure of at least one prior chemotherapy regimen, as requested in recommendation originating from variation EMEA/H/C/000618/II/0043.”	Weekly start timetable.
Truvada - emtricitabine / tenofovir disoproxil - EMEA/H/C/000594/II/0138/G MAH: Gilead Sciences International Ltd, Rapporteur: Greg Markey, “Submission of the final report from studies GS-US-276-0101 and GS-US-276-0105, listed as a category 3 studies in the RMP. GS-US-276-0101 - This is a A Prospective, Observational Study of Pregnancy Outcomes among Women Exposed to Truvada for PrEP Indication Nested in the Antiretroviral Pregnancy Registry GS-US-276-0105 – This is a A Prospective, Observational, Drug Utilization Study of Subjects Taking Truvada for Pre-exposure Prophylaxis in the USA.” Request for Supplementary Information adopted on 09.06.2017.	Weekly start timetable. The Committee adopted a Request for Supplementary information together with a specific timetable.
Venclyxto - venetoclax - EMEA/H/C/004106/II/0003, Orphan MAH: AbbVie Ltd., Rapporteur: Filip Josephson, “Submission of the final report from study R&D 16/1398: Assessment of Cytochrome P450 mRNA Induction by A-1195425 in Cultured Human Hepatocytes to evaluate CYP1A2 and	Weekly start timetable.

CYP2B6 induction response which is included as a Post Authorisation Measure (Category 3) in RMP 2.0."	
Ventavis - iloprost - EMEA/H/C/000474/II/0055 MAH: Bayer Pharma AG, Rapporteur: Alexandre Moreau, "Update of sections 4.9 of the SmPC in order to update the safety information related to overdose following a cumulative review of overdose cases. The Package Leaflet (PIL) is updated accordingly. In addition, the Marketing authorisation holder (MAH) took the opportunity to update the list of local representatives in the PIL, to align the PIL with the SmPC for children and adolescents and to adjust the labelling of the inner carton without blue box." Opinion adopted on 01.06.2017.	Positive Opinion adopted by consensus on 01.06.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Votubia - everolimus - EMEA/H/C/002311/II/0044, Orphan MAH: Novartis Europharm Ltd, Rapporteur: Harald Enzmann, "Update of sections 4.2, 4.4 and 4.8 of the SmPC for Votubia 2.5 mg, 5 mg and 10 mg tablets and 2 mg, 3 mg and 5 mg dispersible tablets in order to reflect on data from study CRAD001M230, in particular to revise dosing recommendations for patients with hepatic impairment, to update the warning related to infections, to include "sepsis" as an adverse drug reaction with the frequency "uncommon" and to revise frequencies of the following adverse drug reactions: "pharyngitis" ["common" to "very common"], "pneumonitis" ["uncommon" to "common"] and "rash" ["common" to "very common]". In addition, the subsection on Paediatric population in section 4.8 of the SmPC was updated based on results from study CRAD001M230. Furthermore, sections 5.1 and 5.2 of the SmPC for Votubia 2 mg, 3 mg and 5 mg dispersible tablets was updated to add new information on pharmacodynamic and pharmacokinetic properties based on results from study CRAD001M230. The Package Leaflet is updated accordingly."	
Wakix - pitolisant - EMEA/H/C/002616/II/0007, Orphan MAH: BIOPROJET PHARMA, Rapporteur: Joseph Emmerich, "Submission of the final CSR for	Weekly start timetable.

Study P11-11; a multi-centre, single dose trial to evaluate the pharmacokinetics of pitolisant in children from 6 to less than 18 years with narcolepsy (Measure 3 of the agreed PIP)." Request for Supplementary Information adopted on 27.04.2017.	
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 21.04.2017.	
XGEVA - denosumab - EMEA/H/C/002173/II/0054 MAH: Amgen Europe B.V., Rapporteur: Kristina Dunder, PRAC Rapporteur: Ulla Wändel Liminga, "Submission of an updated Risk management plan (RMP) version 25 in order to include that cataracts are no longer considered to be a potential risk associated with denosumab therapy supported by the recently completed Study 20080560 (Multicentre, randomized, double blind, placebo-controlled study in men with nonmetastatic prostate cancer receiving androgen deprivation therapy cataract development and progression study using a slit-lamp-based evaluation system (Lens Opacities Classification System III (LOCS III).), where results showed no difference between the risk of developing cataracts in the denosumab and placebo groups."	Weekly start timetable.

Request for Supplementary Information adopted on 09.06.2017.	
Xofigo - radium-223 - EMEA/H/C/002653/II/0025 MAH: Bayer AG, Rapporteur: Harald Enzmann, "Update of sections 4.2, 5.1, 5.2, and 11 of the SmPC based on the update of the Xofigo Company Core Data Sheet (CCDS) to version 5.0. 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 and to introduce non safety related editorial changes to increase comprehensibility."	Weekly start timetable.
Yondelis - trabectedin - EMEA/H/C/000773/II/0051, Orphan MAH: Pharma Mar, S.A., Rapporteur: Sinan B. Sarac, "Update of sections 4.2, 4.4 and 5.2 of the SmPC in order to update the safety information following submission of study report ET743-OVC-1004 "An Open-Label, Multicenter, Pharmacokinetic Study of Trabected in in Subjects with Advanced Malignancies and Hepatic Dysfunction" listed in the RMP. In addition, the Marketing authorisation holder (MAH) took the opportunity to introduce editorial changes in the SmPC." Request for Supplementary Information adopted on 21.04.2017.	Weekly start timetable.
Zostavax - shingles (herpes zoster) vaccine (live) - EMEA/H/C/000674/II/0112 MAH: MSD Vaccins, Rapporteur: Jan Mueller-Berghaus, "Update of section 5.1 of the SmPC in order to add information on long-term effectiveness of Zostavax on herpes zoster and postherpetic neuralgia in individuals 50 years of age or older following the first interim results from the post-licensure observational study (Protocol 024) listed as category 3 study in the RMP. In addition, the marketing authorisation holder took the opportunity to bring the product information in line with the latest QRD template version 10."	Weekly start timetable.
Zyclara - imiquimod - EMEA/H/C/002387/II/0013 MAH: Meda AB, Rapporteur: Nithyanandan Nagercoil, "Update of sections 4.2, 4.4 and 5.1	

of the SmPC in order to add data on the clinical experience gained with study X-03016-3284 (LEIDA 2) and a meta-analysis of X-03016-3271 and X-03016-3284. The MAH took the opportunity to update the details of local representatives in the PIL."	
WS1105 **IntronA-** **EMEA/H/C/000281/WS1105/0107** **PegIntron-** **EMEA/H/C/000280/WS1105/0128** **ViraferonPeg-** **EMEA/H/C/000329/WS1105/0121** MAH: Merck Sharp & Dohme Limited, Lead Rapporteur: Filip Josephson, "Update of sections 4.2 and 4.8 of the SmPC in order to update the safety information with information on HCV/HBV co-infection, and to add an ADR on hepatitis B reactivation in HCV/HBV co-infected patients as post marketing adverse experience respectively. The Package Leaflet and Labelling are updated accordingly. In addition, the Worksharing applicant (WSA) took the opportunity to bring the PI in line with the latest QRD template version 10 including the implementation of the use of combined SmPCs and PLs for PegIntron and ViraferonPeg and the use of combined SmPCs for Intron A in multidose pen." Request for Supplementary Information adopted on 21.04.2017, 23.02.2017.	Weekly start timetable.
WS1135 **Glyxambi-** **EMEA/H/C/003833/WS1135/0003** **Jardiance-** **EMEA/H/C/002677/WS1135/0030** **Synjardy-** **EMEA/H/C/003770/WS1135/0026** MAH: Boehringer Ingelheim International GmbH, Lead Rapporteur: Johann Lodewijk Hillege, "Update of section 5.2 of the SmPC in order to reflect the results from PK/PD study 1245.87 `An open-label, randomised, multicentre, single-dose, parallel group trial to evaluate pharmacokinetics and pharmacodynamics of empagliflozin in children and adolescents from 10 to less than 18 years of age with type 2 diabetes mellitus', previously assessed as Article 46 submission for Jardiance [EMEA/H/C/002677/P46-007]."	Positive Opinion adopted by consensus on 01.06.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

Opinion adopted on 01.06.2017.	
WS1162 **Glyxambi-** **EMEA/H/C/003833/WS1162/0004** **Jentadueto-** **EMEA/H/C/002279/WS1162/0038** **Trajenta-** **EMEA/H/C/002110/WS1162/0028** MAH: Boehringer Ingelheim International GmbH, Lead Rapporteur: Johann Lodewijk Hillege, "Update of section 5.2 of the SmPC in order to reflect the results from PK/PD study 1218.56 `A randomised, double-blind, placebo-controlled parallel group dose-finding study of linagliptin (1 mg or 5 mg administered orally once daily) over 12 weeks in children and adolescents, from 10 to 17 years of age, with type 2 diabetes mellitus', previously assessed as Article 46 submission for Trajenta [EMEA/H/C/002110/P46/016]." Opinion adopted on 01.06.2017.	Positive Opinion adopted by consensus on 01.06.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1181/G **Exviera-** **EMEA/H/C/003837/WS1181/0030/G** **Viekirax-** **EMEA/H/C/003839/WS1181/0034/G** MAH: AbbVie Ltd., Lead Rapporteur: Filip Josephson, "Submission of the final report for two phase IIIb studies (studies M13-774 and M13-862) to support the 3 direct-acting antiviral regimen administered with and without ribavirin for 12 weeks for hepatitis C virus genotype 1 infected, treatment-experienced and treatment-naïve subjects without cirrhosis, listed as category 3 studies in the RMP."	Weekly start timetable.

B.5.3. CHMP-PRAC assessed procedures

Abasaglar - insulin glargine - **EMEA/H/C/002835/II/0014** MAH: Eli Lilly Regional Operations GmbH, Rapporteur: Robert James Hemmings, PRAC Rapporteur: Carmela Macchiarulo, "Submission of the final report from study I4L-MC-ABER(ABER). This is a Prospective, Randomized, Open-Label Comparison of a Long-Acting Basal Insulin Analog LY2963016 to LANTUS® in Adult Patients with Type 2 Diabetes Mellitus: the ELEMENT 5 Study. This study was conducted in	Weekly start timetable.

non European countries. This study replaces the cancelled studies that were planned to be conducted in China and other countries and that were described in the RMP.
An updated RMP version 1.6 is submitted accordingly."
Request for Supplementary Information adopted on 09.06.2017.

Adcetris - brentuximab vedotin - EMEA/H/C/002455/II/0045, Orphan
MAH: Takeda Pharma A/S, Rapporteur: Paula Boudewina van Hennik, PRAC Rapporteur: Sabine Straus, "Update of section 5.1 of the SmPC in order to add 5-year follow-up overall survival (OS) data from patients included in study SG035-0004, a phase 2 open-label study of brentuximab vedotin in the treatment of patients with relapsed or refractory systemic anaplastic large cell lymphoma (ALCL), in accordance with the specific obligation SOB 028. Annex II of the product information and the RMP (version 9.0) are updated accordingly."

Champix - varenicline - EMEA/H/C/000699/II/0064
MAH: Pfizer Limited, Rapporteur: Hanne Lomholt Larsen, PRAC Rapporteur: Doris Stenver, "Update of sections 4.6 and 5.1 of the SmPC in order to update the safety information based on the final results from study Study A3051078, a varenicline Pregnancy Cohort Study.
This is a prospective cohort study to compare women who use varenicline while pregnant with women who smoke while pregnant with respect to birth outcomes

The Package Leaflet is updated accordingly.

The RMP version 10.1 has also been submitted.

In addition, the Marketing authorisation holder (MAH) took the opportunity to bring the PI in line with the latest QRD template version 10."
Opinion adopted on 09.06.2017.
Request for Supplementary Information adopted on 06.04.2017.

Champix - varenicline - EMEA/H/C/000699/II/0066
MAH: Pfizer Limited, Rapporteur: Hanne Lomholt Larsen, PRAC Rapporteur: Doris Stenver, "Update of section 5.1 of the SmPC in

Weekly start timetable.

order to update the safety information based on final results from Clinical Study A3051148 (A Phase 4, Non-Treatment Follow-Up for Cardiac Assessments Following Use of Smoking Cessation Treatments in Subjects With and Without a History of Psychiatric Disorders), a non-interventional category 3 Post-authorisation safety study (PASS) in the RMP. The RMP version 10.1 has also been submitted."	
Cimzia - certolizumab pegol - EMEA/H/C/001037/II/0060 MAH: UCB Pharma S.A., Rapporteur: Kristina Dunder, PRAC Rapporteur: Ulla Wändel Liminga, "Update of section 4.6 of the SmPC in order to update the information on pregnancy and lactation based on two pharmacokinetics studies evaluation the transfer of Cimzia into breastmilk and via the placenta (UP0016 and UP0017). The Package Leaflet is updated accordingly. The RMP version 12 has also been submitted."	
Edurant - rilpivirine - 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." Request for Supplementary Information adopted on 09.06.2017, 06.04.2017.	Weekly start timetable.
Glyxambi - empagliflozin / linagliptin - EMEA/H/C/003833/II/0005/G MAH: Boehringer Ingelheim International GmbH, Rapporteur: Johann Lodewijk Hillege,	

PRAC Rapporteur: Julie Williams, “C.I.1.a. Update of section 4.4. of the SmPC to add a warning on the risk of lower limb amputations to align the PI wording with the parallel finalised Article 20 referral on the risk of lower limb amputation of SGLT2 inhibitors. The Package Leaflet is updated accordingly. C.I.4.z.Update of sections 4.2., 4.4., 4.5., 4.8., 5.1., of the SmPC to update the PI with data from study 1245.25 Emp-Reg. This is a Phase III, multicentre, international, randomised, parallel group, double blind cardiovascular safety study of BI 10773 (10 mg and 25 mg administered orally once daily) compared to usual care in type 2 diabetes mellitus patients with increased cardiovascular risk. An RMP (v.2.0) has also been submitted.”	
Imbruvica - ibrutinib - EMEA/H/C/003791/II/0033/G, Orphan MAH: Janssen-Cilag International NV, Rapporteur: Filip Josephson, PRAC Rapporteur: Patrick Batty, “C.I.4 (Type II) - Update of sections 4.4 and 5.1 of the SmPC in order to update the safety information related to bleeding related events based on final results from study PCYC-1132-NT listed as a category 3 (MEA 004.1) study in the RMP; this is an in-vitro study to evaluate the effect of ibrutinib on platelet aggregation ;The Package Leaflet is updated accordingly. C.I.4 (Type II) - Update of section 4.4 and 4.5 of the SmPC in order to update the safety information based on final) results from study LYM1003 listed as a category 3 study in the RMP (MEA 009.1); this is a drug-drug interaction study to assess steady state PK of repeated oral doses of ibrutinib alone in patients with B-cell malignancies and when combined with a moderate and strong CYP3A inhibitor; The Package Leaflet is updated accordingly. C.I.4 (Type II) - Update of section 4.5 of the SmPC in order to update the safety information based on final results from study FK12024; this is a DDI study with CYP3A inhibitor posaconazole, in simulated subjects; The Package Leaflet is are updated accordingly. C.I.4 (Type II) - Update of section 4.4 of the SmPC in order to update the safety information	Weekly start timetable.

on antimicrobial prophylaxis following routine pharmacovigilance activity.

C.I.11.z (Type IB) - Submission of an updated RMP in order to extend the closure date of study PCYC-1112-CA (ANX 003.2) to Q2 2019. Yearly updates will be submitted in Q2 2017 and Q2 2018. Annex II has been updated accordingly.

C.I.11.a (Type Iain) - To update the RMP to include an additional action for Study PCI-32765 CAN3001 (MEA017) to provide a "further interim report in 5 years' from time from the cut-off date of the current report (12 November 2015)". This change has been agreed by the CHMP in the outcome of EMA/H/C/ 003791/MEA/017.

The RMP version 6.8 has 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 09.06.2017.

Increlex - mecasermin - EMEA/H/C/000704/II/0044/G, Orphan

MAH: Ipsen Pharma, Rapporteur: Outi Mäki-Ikola, PRAC Rapporteur: Kirsti Villikka, "Update of section 4.4 of the SmPC in order to update the warning regarding antibody response to injected IGF-1.

Submission of an updated RMP version 9 , including the educational materials, to update the instructions for antibody testing and improve wording and advices."
Request for Supplementary Information adopted on 23.02.2017.

Jardiance - empagliflozin - EMEA/H/C/002677/II/0026

MAH: Boehringer Ingelheim International GmbH, Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Dolores Montero Corominas, "Submission of the final results of a non-clinical study on the effect of empagliflozin on blood ketone level at refeeding after a fasting period, comparison between refeeding with glucose or fat in order to fulfil MEA 010. The RMP (version 11.0) is updated accordingly."
Request for Supplementary Information adopted

on 23.02.2017.

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."

Levemir - insulin detemir - EMEA/H/C/000528/II/0084

MAH: Novo Nordisk A/S, Rapporteur: Hanne Lomholt Larsen, PRAC Rapporteur: Doris Stenver, "Submission of the summary analysis report on the incidence of neoplasms with the combination of liraglutide and insulin detemir from the cardiovascular outcome trial for Victoza®, trial EX2211-3748 (LEADER®). As a consequence the following important potential risk of "malignant neoplasms following combination treatment with insulin detemir + liraglutide + metformin" is deleted from the updated RMP version 18."
Request for Supplementary Information adopted on 21.04.2017, 23.02.2017.

Mozobil - plerixafor - EMEA/H/C/001030/II/0030/G, Orphan

MAH: Genzyme Europe BV, Rapporteur: Paula Boudewina van Hennik, PRAC Rapporteur: Sabine Straus, "Submission of the final report from study ARD12858 (MOZ23510) "A pilot, exploratory, randomized, phase 2 safety study evaluating tumor cell (plasma cell) mobilization and apheresis product contamination in plerixafor plus non-pegylated G-CSF mobilized patients and in non pegylated G-CSF alone mobilized patients" listed as a category 3 study in the RMP .

Submission of the final report from study OBS13611 (MOZ18009), a multicenter, noninterventional registry designed to evaluate the long-term outcomes for patients who received plerixafor for stem cell mobilization and completed hematopoietic stem cell transplantation (HSCT) compared with patients who received other mobilization methods and

completed HSCT, listed as a category 3 study in the RMP.

Submission of the final report from study OBS13612 (MOZ19310), monitoring the plerixafor off-label transplant use, in patients and donors in EBMT centers performing autologous transplants and/or allogeneic transplants, listed as a category 3 study in the RMP."
Request for Supplementary Information adopted on 23.03.2017.

Mysimba - naltrexone hydrochloride / bupropion hydrochloride - EMEA/H/C/003687/II/0017
MAH: Orexigen Therapeutics Ireland Limited, Rapporteur: Hanne Lomholt Larsen, PRAC Rapporteur: Martin Huber, "Submission of the final report from phase I study NaltrexBuprop-1001 (TQT) to evaluate the potential effect of Naltrexone and Bupropion extended-release combination on cardiac repolarization in healthy subjects and updated RMP to include study NaltrexBuprop-1001 but also studies recently completed (NB-CVOT, NaltrexBuprop-4001, NaltrexBuprop-1004 and NB-404).
The MAH also took the opportunity to include throughout the RMP references to the PASS protocols currently under discussion at the PRAC."

Neulasta - pegfilgrastim - EMEA/H/C/000420/II/0093/G
MAH: Amgen Europe B.V., Rapporteur: Robert James Hemmings, PRAC Rapporteur: Patrick Batty.

Nuwiq - simoctocog alfa - EMEA/H/C/002813/II/0017/G
MAH: Octapharma AB, Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Ulla Wändel Liminga, "C.I.4: Update of sections 4.2, 4.8 and 5.1 of the SmPC to reflect avaialble data from Previously Untreated Patients (PUP) from GENA-05 (interim report) study. The Package Leaflet is updated accordingly. The RMP version 8.0 has also been submitted.
In addition, the Marketing authorisation holder (MAH) took the opportunity to update the Product Information throughout to bring it in line with the Core summary of product characteristics for human plasma-derived and

recombinant coagulation factor VIII products (EMA/CHMP/BPWP/1619/1999 rev. 2) and with the QRD version 10 (affects labelling only). Moreover the MAH is combining the SmPC for all strengths and updating Annex A with detailed information on the packaging. C.I.11: Submission of an updated RMP version 8.0 in order to align the content in a single harmonised worldwide version for simoctocog alfa (rFVIII)." Request for Supplementary Information adopted on 21.04.2017.	
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." Request for Supplementary Information adopted on 09.06.2017.	Weekly start timetable.
Orkambi - lumacaftor / ivacaftor - EMEA/H/C/003954/II/0021 MAH: Vertex Pharmaceuticals (Europe) Ltd., Rapporteur: Nithyanandan Nagercoil, PRAC Rapporteur: Almath Spooner, "Update of section 4.8 of the SmPC in order to add information on respiratory events based on final results from study Study VX14-809-106 (Study 106), a Phase 3b, open-label study to evaluate safety and tolerability of lumacaftor and ivacaftor combination therapy in subjects 12 years and older with Cystic Fibrosis and advanced lung disease, homozygous for the F508del-CFTR Mutation. Efficacy was evaluated as a secondary	

objective. This study report is being submitted to fulfil MEA 002.
An updated RMP (version 3.2) has also been submitted."

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."

Prolia - denosumab - EMEA/H/C/001120/II/0062

MAH: Amgen Europe B.V., Rapporteur: Kristina Dunder, PRAC Rapporteur: Ulla Wändel Liminga, "Update of the product information (SmPC sections 4.4, 4.8 and PL sections 3 and 4) as well as the Risk Management Plan (RMP) to update the safety information and reflect the multiple vertebral fractures (MVF) following discontinuation of Prolia treatement as a new important risk. This variation follows a concluded analysis of osteoporosis-related fracture data in subjects who discontinued investigational product and remained on study in either the Prolia phase 3 pivotal fracture study (Study 20030216) or its study extension (Study 20060289) to better understand the incidence of fracture following treatment discontinuation. The results of this analysis conclude that multiple vertebral fractures may occur following discontinuation of Prolia treatment, particularly in patients with a history of vertebral fracture.
In addition, the applicant took the opportunity to update the PI in line with the QRD template latest version, amend the PI previous version typographical errors from previous version, and implement minor changes in the Package leaflet local representatives."

Request for Supplementary Information adopted on 21.04.2017, 15.12.2016.	
Prolia - denosumab - EMEA/H/C/001120/II/0063 MAH: Amgen Europe B.V., Rapporteur: Kristina Dunder, PRAC Rapporteur: Ulla Wändel Liminga, "Update of the SmPC section 5.1 to provide information on the clinical study data experience in patients in treatment transition from an oral Bisphosphonate to desunomab, information resulting from the assessment on data of study report 20110153 and a discussion on the issue of long term antiresorptive treatment, in particular when long-term bisphosphonate treatment is followed by denosumab." Request for Supplementary Information adopted on 21.04.2017, 15.12.2016.	
Prolia - denosumab - EMEA/H/C/001120/II/0065 MAH: Amgen Europe B.V., Rapporteur: Kristina Dunder, PRAC Rapporteur: Ulla Wändel Liminga, "To amend the Risk Management Plan (RMP). 1. Modification of the important potential risk of hypercalcemia following treatment discontinuation in patients with growing skeletons to include the adult population 2. Removal of the important potential risk of fracture healing complications. 3. Addition of study 20090601 as a category 4 study pharmacovigilance activity. Request for Supplementary Information adopted on 26.01.2017.	
Prolia - denosumab - EMEA/H/C/001120/II/0069 MAH: Amgen Europe B.V., Rapporteur: Kristina Dunder, PRAC Rapporteur: Ulla Wändel Liminga, "Change(s) in the Summary of Product Characteristics and Package Leaflet due to new clinical/pharmacovigilance data from study 20080560 (Variation category C.I.4) Update of section 4.8 of the SmPC in order to update the safety information as cataracts are no longer considered to be a potential risk and/or adverse reaction associated with denosumab therapy, relevant changes to the SmPC, package leaflet and RMP are proposed supported by the final data report from study/studies (20080560) category 3 study in	Weekly start timetable.

the RMP (Multicentre, randomized, double blind, placebo-controlled study in men with nonmetastatic prostate cancer receiving androgen deprivation therapy cataract development and progression study using a slit-lamp-based evaluation system (Lens Opacities Classification System III (LOCS III).) In addition the RMP has been updated to remove the important potential risk “cataract in men with prostate cancer receiving androgen deprivation therapy”. Request for Supplementary Information adopted on 09.06.2017.	
Remicade - infliximab - EMEA/H/C/000240/II/0204 MAH: Janssen Biologics B.V., Rapporteur: Kristina Dunder, PRAC Rapporteur: Ulla Wändel Liminga, “Submission of the final registry report from the C0168T71 study (a review and analysis of birth outcomes from Swedish, Danish and Finnish medical birth registers) and an evaluation of pregnancy data from multiple sources. Section 4.6 of the SmPC, relevant section of the PL and the RMP version 13.2 has been updated to reflect the study results. The MAH has also taken the opportunity to bring the product in line with the QRD template and update the local representative section of the PL.” Request for Supplementary Information adopted on 23.03.2017.	
Reyataz - atazanavir / atazanavir sulfate - EMEA/H/C/000494/II/0111 MAH: Bristol-Myers Squibb Pharma EEIG, Rapporteur: Joseph Emmerich, PRAC Rapporteur: Caroline Laborde, “Update of sections 4.4 and 4.8 of the SmPC to add a warning on chronic kidney disease observed in HIV infected patients during treatment with atazanavir (with or without ritonavir). This update is based on a review of the MAH safety database, a cohort study of patients with laboratory values from a large US administrative claims database and a review of published scientific literature. The Package Leaflet is updated accordingly. The RMP version 12.0 has also been submitted.”	Weekly start timetable.

Request for Supplementary Information adopted on 09.06.2017.

Saxenda - liraglutide - EMEA/H/C/003780/II/0011

MAH: Novo Nordisk A/S, Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Menno van der Elst, "Based on submission of the LEADER clinical study results (EX2211-3748: liraglutide effect on and action in diabetes, evaluation of cardiovascular outcome results), changes to sections 4.4, and 5.1 of the SmPC are being proposed in order to update the safety information and include a description of the clinical study outcomes. The Package Leaflet and Labelling are updated accordingly. In addition, the Marketing authorisation holder (MAH) took the opportunity to implement minor editorial changes throughout the product information.

The LEADER study was included in the liraglutide RMP as a required pharmacovigilance activity (category 3) to specifically address the important potential risk of cardiovascular disorders in patients with Type 2 Diabetes Mellitus. Updates to the liraglutide RMP based on the study results are also proposed: this variation application fulfils two post-approval commitments in relation to the cardiovascular outcomes trial (MEA 002), as well as to provide additional information on the breast cancer cases found in LEADER (MEA 005). RMP Version 27 was submitted with the application. These liraglutide RMP modifications are in line with the proposed updates to the Saxenda Product Information described above."
Request for Supplementary Information adopted on 23.02.2017.

Voncento - human coagulation factor VIII / human von willebrand factor - EMEA/H/C/002493/II/0017/G

MAH: CSL Behring GmbH, Rapporteur: Paula Boudewina van Hennik, PRAC Rapporteur: Sabine Straus, "C.I.4 (type II): Update of section 4.8 of the SmPC in order to update the frequencies of undesirable effects to reflect the final clinical study data from study CSLCT-BIO-08-53 in haemophilia A paediatric patients. The Package Leaflet is updated accordingly. The submission of the final CSR CSLCT-BIO-08-53

also leads to changes to the RMP (ver. 6.1) in order update the Company Core Safety Information (CCSI).
C.I.11.z (type IB): Submission of a revised RMP in order to remove the commitment to conduct a post-marketing study for haemophilia A patients (CSLCT-BIO-12-78) for Voncento as consequence of new data from study CSLCT-BIO-08-53.
In addition, the Marketing authorisation holder (MAH) took the opportunity to combine different strengths in the SmPC and Package Leaflet."
Request for Supplementary Information adopted on 23.03.2017, 10.11.2016, 01.04.2016, 19.11.2015.

Wakix - pitolisant - EMEA/H/C/002616/II/0004/G, Orphan

MAH: BIOPROJET PHARMA, Rapporteur: Joseph Emmerich, PRAC Rapporteur: Kirsti Villikka,
"Update of sections 4.4, 4.5, 4.6 and 5.2 of the SmPC based on the final CSR of study P15-02 (to assess the mass balance recovery, metabolite profile and metabolite identification of 14C-pitolisant at steady state conditions, in healthy CYP2D6 phenotyped subjects), P14-07 (to evaluate pharmacokinetic interaction of pitolisant with sodium oxybate and modafinil in healthy male volunteers) and P15-15 (to evaluate pharmacokinetic interaction of pitolisant with CYP3A4 substrates (midazolam), CYP2B6 substrates (bupropion), UGT2B7 inhibotors (probenecide)) in fulfilment of PAM (MEA 02, 03 and 04). The Package Leaflet is updated accordingly. In addition, the Marketing authorisation holder (MAH) took the opportunity to make minor editorial change in section 4.8 of the SmPC. Moreover, updated RMP version 5.0 has been submitted as part of this application."
Request for Supplementary Information adopted on 23.03.2017.

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

MAH: Amgen Europe B.V., Rapporteur: Kristina Dunder, PRAC Rapporteur: Ulla Wändel Liminga,
" ☐ To
(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 26.01.2017.	
Xtandi - enzalutamide - EMEA/H/C/002639/II/0035 MAH: Astellas Pharma Europe B.V., Rapporteur: Jorge Camarero Jiménez, PRAC Rapporteur: Eva A. Segovia, "Update of sections 4.4 and 4.8 of the SmPC to reflect the final results of the post authorisation safety study (PASS) CL-9785-0403 which evaluated the risk of seizure among subjects with mCRPC treated with enzalutamide who were at potential increased risk of seizure (UPWARD) and was listed as a category 3 in the RMP. The RMP version 11.0 has also been submitted. In addition, the Marketing authorisation holder (MAH) took the opportunity to make a correction in section 5.1 of the SmPC." Opinion adopted on 09.06.2017. Request for Supplementary Information adopted on 09.03.2017.	
Yervoy - ipilimumab - EMEA/H/C/002213/II/0047/G MAH: Bristol-Myers Squibb Pharma EEIG, Rapporteur: Paula Boudewina van Hennik, PRAC Rapporteur: Sabine Straus, "Update of section 4.4 to revised the current warning on concurrent administration with vemurafenib to enhance awareness on the potential of hypersensitivity reactions when ipilimumab is used sequentially with vemurafenib as requested by the PRAC following the assessment of PSUSA/00009200/201603. Update of sections 4.8 of the SmPC to amend the frequency of the adverse drug reaction 'Vogt-Konyanagi-Haranda syndrome' from 'not know' to 'very rare'. The RMP (version 16) has been updated accordingly.]In addition, the Marketing authorisation holder (MAH) took the opportunity to implement some editorial changes to sections 4.2 and 4.4 of the SmPC to update the dose modification information for	Weekly start timetable.

hepatotoxicity management guidelines in line with National Cancer Institute (NCI) Common Terminology Criteria for Adverse Events (CTCAE) recommendations (version 4)."
Request for Supplementary Information adopted on 09.06.2017.

Zinbryta - daclizumab - EMEA/H/C/003862/II/0007

MAH: Biogen Idec Ltd, Rapporteur: Bruno Sepodes, PRAC Rapporteur: Eva A. Segovia, "Update of sections 4.4 and 4.8 of the SmPC in order to add autoimmune haemolytic anaemia with frequency 'uncommon' and to include a warning concerning symptoms of this adverse drug reaction based on reported post-marketing cases.
The Package Leaflet is updated accordingly. In addition, the Marketing authorisation holder took the opportunity to implement minor editorial amendments throughout the Product Information.
The RMP version 5.1 has been approved."
Opinion adopted on 09.06.2017.
Request for Supplementary Information adopted on 05.05.2017.

Zykadia - ceritinib - EMEA/H/C/003819/II/0015

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

The RMP version 9.0 has also been submitted."

WS1026
Rasilez-EMEA/H/C/000780/WS1026/0110
Rasilez HCT-EMEA/H/C/000964/WS1026/0080

MAH: Novartis Europharm Ltd, Lead Rapporteur: Daniela Melchiorri, Lead PRAC Rapporteur: Carmela Macchiarulo, "Update of section 5.1 of the SmPC in order to reflect the results of study SPP100F2301 (ATMOSPHERE) a multicenter, randomized, doubleblind, parallel group, active-controlled study to evaluate the efficacy and safety of both aliskiren monotherapy and aliskiren/enalapril

combination therapy compared to enalapril monotherapy, on morbidity and mortality in patients with chronic heart failure (NYHA Class II - IV). The RMP (v 13) has also been updated to reflect the study results."
Request for Supplementary Information adopted on 21.04.2017, 15.12.2016.

WS1130/G
Efficib-
EMEA/H/C/000896/WS1130/0081/G
Janumet-
EMEA/H/C/000861/WS1130/0081/G
Ristfor-
EMEA/H/C/001235/WS1130/0068/G
Velmetia-
EMEA/H/C/000862/WS1130/0084/G
MAH: Merck Sharp & Dohme Limited, Lead Rapporteur: Johann Lodewijk Hillege, Lead PRAC Rapporteur: Menno van der Elst,
"C.I.11.b: Submission of an updated RMP in order to add a targeted questionnaire related to lactic acidosis as part of the outcome of the referral procedure EMEA/H/A-31/1432 (finally agreed version 7.1).
C.I.3.b: Update of sections 4.4 of the SmPC in order to add a warning on bullous pemphigoid following the PRAC assessment outcome of EMEA/H/C/PSUSA/2711/201408; the Package Leaflet is being updated accordingly."
Opinion adopted on 09.06.2017.
Request for Supplementary Information adopted on 06.04.2017.

WS1141
Januvia-
EMEA/H/C/000722/WS1141/0056
Ristaben-
EMEA/H/C/001234/WS1141/0048
TESAVEL-
EMEA/H/C/000910/WS1141/0056
Xelevia-EMEA/H/C/000762/WS1141/0060
MAH: Merck Sharp & Dohme Limited, Lead Rapporteur: Johann Lodewijk Hillege, Lead PRAC Rapporteur: Menno van der Elst, "Update of sections 4.4 of the SmPC in order to add a warning on bullous pemphigoid following the PRAC assessment outcome of EMEA/H/C/PSUSA/2711/201408; the Package Leaflet is being updated accordingly. Consequently, the RMP is also updated accordingly (finally agreed version 7.1)."

Opinion adopted on 09.06.2017.
Request for Supplementary Information adopted on 06.04.2017.

B.5.4. PRAC assessed procedures

PRAC Led
Avastin - bevacizumab - EMEA/H/C/000582/II/0095
MAH: Roche Registration Limited, Rapporteur: Sinan B. Sarac, PRAC Rapporteur: Doris Stenver, PRAC-CHMP liaison: Sinan B. Sarac, "Submission of an updated RMP version 28.1 in order to reduce the 20 year long-term follow-up (LTFU) information from the paediatric population after patients complete the minimum 5.5 year follow-up period as defined in the BO20924 (BERNIE) paediatric study protocol and to amend the date of submission of the final report (addendum CSR) for the BO20924 (BERNIE) study from Q1 2017 to Q3 2019."
Opinion adopted on 09.06.2017.
Request for Supplementary Information adopted on 05.05.2017.

PRAC Led
Cervarix - human papillomavirus vaccine [types 16, 18] (recombinant, adjuvanted, adsorbed) - EMEA/H/C/000721/II/0088
MAH: GSK Biologicals SA, Rapporteur: Bart Van der Schueren, PRAC Rapporteur: Jean-Michel Dogné, PRAC-CHMP liaison: Bart Van der Schueren, "Submission of the final report from the pregnancy registry data (study EPI-HPV-067); this study is a Post Authorisation Safety Study (PASS), and information related to the use of Cervarix during pregnancy was identified as important missing information in the Risk Management Plan (RMP)."
Opinion adopted on 09.06.2017.

PRAC Led
Eperzan - albiglutide - EMEA/H/C/002735/II/0029/G
MAH: GlaxoSmithKline Trading Services, PRAC Rapporteur: Julie Williams, PRAC-CHMP liaison: Greg Markey, "II: C.I.11.b - Update of the RMP to amend Study 201805 (category 3 study): "Observational Study of the Risk of Common Malignant Neoplasms and Malignant Neoplasms

of Special Interest (Thyroid and Pancreatic Cancer) in Subjects Prescribed Albiglutide Compared to Those Prescribed Other Antidiabetic Agents", in order to use a different database to study the risk of neoplasms in association with albiglutide exposure II: C.I.11.b – Update of the RMP to add a new category 3 study as an additional pharmacovigilance activity – Study 207351: "Observational Study to Assess Maternal and Fetal Outcomes following exposure to Albiglutide during Pregnancy"" Request for Supplementary Information adopted on 26.01.2017.	
PRAC Led **Iclusig - ponatinib - EMEA/H/C/002695/II/0038, Orphan** MAH: Incyte Biosciences UK Ltd, Rapporteur: Greg Markey, PRAC Rapporteur: Patrick Batty, PRAC-CHMP liaison: Greg Markey, "Submission of an updated RMP (version 17) in order to provide the statistical analysis plan for the study AP24534-14-401 (included in the pharmacovigilance plan of the RMP), as per the PRAC request made in the framework of MEA 015." Opinion adopted on 09.06.2017.	
PRAC Led **Mosquirix - plasmodium falciparum and hepatitis B vaccine (recombinant, adjuvanted) - EMEA/H/W/002300/II/0020** MAH: GSK Biologicals SA, PRAC Rapporteur: Jean-Michel Dogné, PRAC-CHMP liaison: Bart Van der Schueren, "Update of the RMP (version 3.0) in order to 1) add cerebral malaria as an important potential risk, 2) add mortality by gender as missing information, 3) add the WHO Pilot Implementation Programme as a category 3 study, 4) change the study dates for studies Malaria-073 (200596, Phase IIIb randomized, open, controlled study to evaluate the immunogenicity and safety of Mosquirix, when administered as primary vaccination at 6, 7.5 and 9 months of age with or without coadministration of measles and rubella and yellow fever vaccines to children living in sub-Saharan, Africa), EPI-MAL-002(115055, an observational cohort study to estimate the incidence of AESI, of meningitis and of other AE	Weekly start timetable.

leading to hospitalisation or death, in children, prior to implementation of Mosquirix), EPI-MAL-003 (115056, a prospective surveillance study to evaluate the safety, the effectiveness and the impact of Mosquirix in infants and young children in sub-Saharan Africa), EPI-MAL-005 (116682, an epidemiology study to assess Plasmodium falciparum parasite prevalence and malaria control measures in catchment areas of two interventional studies pre- and post-Mosquirix introduction (EPI-MAL-002 and EPI-MAL-003) to assess, in field conditions, vaccine benefit-risk in children in sub-Saharan Africa), EPI-MAL-010 (205071, a longitudinal, cross-sectional ancillary study of the EPI-MAL-005 study to evaluate the genetic diversity in circumsporozoite sequences before and after the implementation of Mosquirix in malaria-positive subjects ranging from 6 months to less than 5 years of age), 5) amend the protocol of study EPI-MAL-002, 6) update the draft protocol of study EPI-MAL-003, 7) provide a new draft of the protocol of study EPI-MAL-010, 8) provide a new protocol for the Pilot Implementation Programme." Request for Supplementary Information adopted on 09.06.2017.	
PRAC Led **Myozyme - alglucosidase alfa - EMEA/H/C/000636/II/0062** MAH: Genzyme Europe BV, Rapporteur: Alexandre Moreau, PRAC Rapporteur: Caroline Laborde, PRAC-CHMP liaison: Alexandre Moreau, "Submission of the final study report of non-interventional, non-imposed PASS study "Myozyme (alglucosidase alfa) Safety Information Packet effectiveness evaluation: a healthcare professional survey" (Myozyme SIP EU HCP Survey, ALGMYC08432). In addition, updated RMP version 8.0 has been submitted as part of this application." Request for Supplementary Information adopted on 09.06.2017.	Weekly start timetable.
PRAC Led **Remicade - infliximab - EMEA/H/C/000240/II/0201/G** MAH: Janssen Biologics B.V., Rapporteur: Kristina Dunder, PRAC Rapporteur: Ulla Wändel Liminga, PRAC-CHMP liaison: Kristina Dunder, "Submission of the clinical study reports for	

C0168T45 and C0168T62 together with an overall summary and evaluation of the complete long term safety follow-up programs for Remicade (as per MEA 79).

Study C0168T45 (RESULTS: REMICADE Safety Under Long term Study) is a Multicenter International Observational Study of the Long-term Safety of Infliximab

Study C0168T62 (RESULTS UC: REMICADE Safety Under Long-term Study in Ulcerative Colitis) is a Multicenter International Study of the Long-term Safety of Infliximab in Ulcerative Colitis.

The RMP (RMP 14.0) has been updated to reflect the completion of these studies."
Request for Supplementary Information adopted on 23.02.2017.

PRAC Led

Suliqua - insulin glargine / lixisenatide - EMEA/H/C/004243/II/0002

MAH: sanofi-aventis groupe, Rapporteur: Kristina Dunder, PRAC Rapporteur: Julie Williams, PRAC-CHMP liaison: Greg Markey, "Submission of the final report from a pharmacoepidemiology study listed as a category 3 study in the RMP. This is retrospective database study of GLP-1 receptor agonists and risk of Acute Pancreatitis, Pancreatic Cancer and Thyroid Cancer, in Particular Medullary Thyroid Cancer, which primary objective was to estimate the incidence rates of acute pancreatitis, pancreatic and thyroid cancer among adult T2DM patients treated with GLP-1 receptor agonists (i.e. Exenatide & liraglutide) versus the ones treated with other antidiabetics."
Opinion adopted on 09.06.2017.

PRAC Led

Xeplion - paliperidone - EMEA/H/C/002105/II/0031

MAH: Janssen-Cilag International NV, Rapporteur: Kristina Dunder, PRAC Rapporteur: Qun-Ying Yue, PRAC-CHMP liaison: Filip Josephson, "Submission of final study report "Post-Authorization Safety Study Using European Union Databases to Assess the Risk of Cardiovascular and Cerebrovascular Adverse Events in Elderly Patients Treated with

Paliperidone Palmitate, Paliperidone Prolonged-Release, and Other Antipsychotics". No changes in the PI are proposed." Request for Supplementary Information adopted on 23.02.2017.	
PRAC Led **Xiapex - collagenase clostridium histolyticum - EMEA/H/C/002048/II/0089** MAH: Swedish Orphan Biovitrum AB (publ), Rapporteur: Martina Weise, PRAC Rapporteur: Martin Huber, PRAC-CHMP liaison: Martina Weise, "Submission of the final clinical study report for study B1531005, a non-interventional study to evaluate the outcomes (clinical treatment success measured by goniometry assessment, recurrence rate measured by goniometry assessment, subject and physician global assessment of treatment satisfaction, complications resulting from the procedure based on the Adverse Event/Serious Adverse Event (AE/SAE)) of 3 various treatment options for Dupuytren's contracture, listed as a category 3 study in the RMP. The RMP (version 13.0) is updated accordingly." Opinion adopted on 09.06.2017. Request for Supplementary Information adopted on 06.04.2017.	
PRAC Led **Xyrem - sodium oxybate - EMEA/H/C/000593/II/0066** MAH: UCB Pharma Ltd., Rapporteur: Bruno Sepodes, PRAC Rapporteur: Ana Sofia Diniz Martins, PRAC-CHMP liaison: Bruno Sepodes, "Submission of the final report from study (C00302) listed as a category 3 study in the RMP. This is a post marketing non-interventional surveillance pharmacoepidemiology study (PMSS) to evaluate long-term safety, tolerability and compliance in administration of Xyrem (sodium oxybate) oral solution in patients who receive treatment with this medication in regular clinical practice." Request for Supplementary Information adopted on 09.06.2017.	Weekly start timetable.

B.5.5. CHMP-CAT assessed procedures

Holoclar - ex vivo expanded autologous human corneal epithelial cells containing	Weekly start timetable.

stem cells - EMEA/H/C/002450/II/0012/G, Orphan, ATMP MAH: Chiesi Farmaceutici S.p.A., Rapporteur: Egbert Flory, Request for Supplementary Information adopted on 11.05.2017.	

B.5.6. CHMP-PRAC-CAT assessed procedures

B.5.7. PRAC assessed ATMP procedures

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

WS1112 **Hexacima-EMEA/H/C/002702/WS1112/0057** **Hexaxim-EMEA/H/W/002495/WS1112/0063** **Hexyon-EMEA/H/C/002796/WS1112/0061** MAH: Sanofi Pasteur Europe, Duplicate, Duplicate of Hexacima, Lead Rapporteur: Jan Mueller-Berghaus Opinion adopted on 01.06.2017. Request for Supplementary Information adopted on 30.03.2017.	Positive Opinion adopted by consensus on 01.06.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1154 **Rotarix-EMEA/H/C/000639/WS1154/0097** MAH: GlaxoSmithKline Biologicals S.A., Lead Rapporteur: Bart Van der Schueren Opinion adopted on 09.06.2017.	Positive Opinion adopted by consensus on 09.06.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1161 **Kisplyx-EMEA/H/C/004224/WS1161/0005** **Lenvima-EMEA/H/C/003727/WS1161/0009** MAH: Eisai Europe Ltd., Lead Rapporteur: Bart Van der Schueren Opinion adopted on 01.06.2017.	Positive Opinion adopted by consensus on 01.06.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
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	Weekly start timetable.

on 18.05.2017.	
WS1170 **Aflunov-** **EMEA/H/C/002094/WS1170/0036** **Foclivia-** **EMEA/H/C/001208/WS1170/0031** MAH: Seqirus S.r.l, Lead Rapporteur: Daniela Melchiorri Request for Supplementary Information adopted on 01.06.2017.	Weekly start timetable. The Committee adopted a Request for Supplementary information together with a specific timetable.
WS1171 **Jentadueto-** **EMEA/H/C/002279/WS1171/0039** **Synjardy-** **EMEA/H/C/003770/WS1171/0027** MAH: Boehringer Ingelheim International GmbH, Lead Rapporteur: Johann Lodewijk Hillege	Weekly start timetable.
WS1174 **Hexacima-** **EMEA/H/C/002702/WS1174/0062** **Hexaxim-** **EMEA/H/W/002495/WS1174/0068** **Hexyon-** **EMEA/H/C/002796/WS1174/0066** MAH: Sanofi Pasteur SA, Lead Rapporteur: Jan Mueller-Berghaus Opinion adopted on 09.06.2017.	Positive Opinion adopted by consensus on 09.06.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1175 **Abseamed-** **EMEA/H/C/000727/WS1175/0064** **Binocrit-** **EMEA/H/C/000725/WS1175/0064** **Epoetin alfa Hexal-** **EMEA/H/C/000726/WS1175/0063** MAH: SANDOZ GmbH, Lead Rapporteur: Alexandre Moreau	Weekly start timetable.

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

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

B.6. START OF THE PROCEDURES TIMETABLES FOR INFORMATION

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

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

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

- adalimumab - EMEA/H/C/004319
, treatment of rheumatoid arthritis, axial spondyloarthritis, psoriasis, hidradenitis suppurativa (HS), Crohn's disease, ulcerative colitis and uveitis.
List of Questions adopted on 23.03.2017.

- carmustine - EMEA/H/C/004326
, treatment of brain tumors, multiple myeloma, Hodgkin's disease and non-Hodgkin's lymphomas, Generic
List of Questions adopted on 13.10.2016.

- dupilumab - EMEA/H/C/004390
, treatment of moderate-to-severe atopic dermatitis
List of Questions adopted on 23.03.2017.

- fluticasone furoate / umeclidinium / vilanterol - EMEA/H/C/004781
, treatment of adult patients with chronic obstructive pulmonary disease (COPD), Duplicate, Duplicate of Trelegy Ellipta
List of Questions adopted on 21.04.2017.

- guselkumab - EMEA/H/C/004271
, treatment of plaque psoriasis
List of Questions adopted on 21.04.2017.

- glecaprevir / pibrentasvir - EMEA/H/C/004430
, indicated for the treatment of chronic hepatitis C virus (HCV) infection in adults
List of Questions adopted on 19.04.2017.

- 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
List of Questions adopted on 15.12.2016.

- naloxone - EMEA/H/C/004325
, Nyxoid is intended for emergency use for known or suspected opioid overdose as manifested by respiratory and/or central nervous system depression
List of Questions adopted on 23.03.2017.

- trastuzumab - EMEA/H/C/004323
, treatment of breast cancer and metastatic gastric cancer
List of Questions adopted on 26.01.2017.

- ritonavir - EMEA/H/C/004549
, treatment of HIV-1,
List of Questions adopted on 21.04.2017.

- tolvaptan - EMEA/H/C/000980/X/0024
List of Questions adopted on 21.04.2017.

- pasireotide - EMEA/H/C/002052/X/0030/G, Orphan
MAH: Novartis Europharm Ltd
List of Questions adopted on 21.04.2017.

- sofosbuvir / velpatasvir / voxilaprevir - EMEA/H/C/004350
, Treatment of chronic hepatitis C virus in adults (HCV) infection in adults
List of Questions adopted on 19.04.2017.

- tacrolimus - EMEA/H/C/004435
, prophylaxis of transplant rejection and treatment of allograft rejection,
List of Questions adopted on 21.04.2017.

- nilotinib - EMEA/H/C/000798/X/0088/G, Orphan
MAH: Novartis Europharm Ltd
List of Questions adopted on 23.03.2017.

- fluticasone furoate / umeclidinium / vilanterol - EMEA/H/C/004363
, treatment of adult patients with chronic obstructive pulmonary disease (COPD)
List of Questions adopted on 21.04.2017.

- ciclosporin - EMEA/H/C/004411, Orphan
Applicant: Santen Oy, treatment of severe vernal keratoconjunctivitis (VKC),
List of Questions adopted on 19.04.2017.

- enzalutamide -
EMEA/H/C/002639/X/0029
List of Questions adopted on 21.07.2016.

- buprenorphine / naloxone -
EMEA/H/C/004407
, treatment for opioid drug dependence,,
treatment for opioid drug dependence
List of Questions adopted on 23.02.2017.

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

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,

Pradaxa - dabigatran etexilate -
EMEA/H/C/000829/R/0105
MAH: Boehringer Ingelheim International GmbH, Rapporteur: Hanne Lomholt Larsen, Co-Rapporteur: Joseph Emmerich, PRAC Rapporteur: Torbjorn Callreus

Selincro - nalmefene -
EMEA/H/C/002583/R/0022
MAH: H. Lundbeck A/S, Rapporteur: Harald Enzmann, Co-Rapporteur: Patrick Salmon, PRAC Rapporteur: Martin Huber

Venclyxto - venetoclax -

EMEA/H/C/004106/R/0005, Orphan
MAH: AbbVie Ltd., Rapporteur: Filip Josephson,
PRAC Rapporteur: Patrick Batty

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

Taltz - ixekizumab -
EMEA/H/C/003943/II/0009
MAH: Eli Lilly Nederland B.V., Rapporteur: Kristina Dunder, Co-Rapporteur: Greg Markey, PRAC Rapporteur: Brigitte Keller-Stanislawski, "Extension of Indication to include alone or in combination with conventional disease-modifying anti-rheumatic drug (cDMARD), the treatment of active psoriatic arthritis in adult patients who have responded inadequately to, or who are intolerant to one or more DMARD therapies. As a consequence, sections 4.1, 4.2, 4.4, 4.5, 4.8, 5.1, and 5.2 of the SmPC are updated to reflect the new safety and efficacy information. The Package Leaflet and RMP have been updated accordingly."
Request for 1 year of market protection for a new indication (Article 14(11) of Regulation (EC) 726/2004)

B.6.8. CHMP assessed procedures scope: Pharmaceutical aspects

Advate - octocog alfa -
EMEA/H/C/000520/II/0085
MAH: Baxter AG, Rapporteur: Jan Mueller-Berghaus,

Fluenz Tetra - influenza vaccine (live attenuated, nasal) -
EMEA/H/C/002617/II/0072
MAH: AstraZeneca AB, Rapporteur: Bart Van der Schueren

Myozyme - alglucosidase alfa -
EMEA/H/C/000636/II/0063/G
MAH: Genzyme Europe BV, Rapporteur: Alexandre Moreau

Plavix - clopidogrel -
EMEA/H/C/000174/II/0127/G
MAH: Sanofi Clir SNC, Rapporteur: Bruno Sepodes

Remicade - infliximab -
EMEA/H/C/000240/II/0205
MAH: Janssen Biologics B.V., Rapporteur: Kristina Dunder

Surgiflo Haemostatic Matrix Kit -Ferrosan - human thrombin -
EMEA/H/D/002301/II/0016
MAH: Presafe Denmark A/S, Rapporteur: Jan Mueller-Berghaus

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
MAH: MCM Vaccine B.V., Rapporteur: Bart Van der Schueren

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

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.”

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.”

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."

Humira - adalimumab - EMEA/H/C/000481/II/0169

MAH: AbbVie Ltd., 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)"

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."

Norvir - ritonavir - EMEA/H/C/000127/II/0147

MAH: AbbVie Ltd., 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."

- 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."

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""

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."

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."

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."

WS1178
Aluvia-EMEA/H/W/000764/WS1178/0102
Kaletra-EMEA/H/C/000368/WS1178/0164
MAH: AbbVie Ltd., 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 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."

B.6.10. CHMP-PRAC assessed procedures

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."

OPDIVO - nivolumab - EMEA/H/C/003985/II/0036/G
MAH: Bristol-Myers Squibb Pharma EEIG, Co-Rapporteur: Paula Boudewina van Hennik, PRAC Rapporteur: Brigitte Keller-Stanislawski, "C.I.4 (Type II) - Update of sections 4.2, 5.1, 5.2 and 6.6 of the SmPC in order to introduce new dosing regimens and schedule.

C.I.4 (Type II) - Update of sections 4.2, 5.1, 5.2 and 6.6 of the SmPC in order to introduce change the infusion time from 60 minutes to 30 minutes

These changes are based on interim results from study CA209153; this is a phase IIIb/IV safety trial of nivolumab in subjects with advanced or metastatic non-small cell Lung cancer who have progressed during or after receiving at least one prior systemic regimen; The Package Leaflet is updated accordingly. The RMP version 10.0 has also been submitted."

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.”

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.”

Vedrop - tocofersolan -
EMEA/H/C/000920/II/0022

MAH: Orphan Europe S.A.R.L., Rapporteur: Greg Markey, PRAC Rapporteur: Patrick Batty“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.”

B.6.11. PRAC assessed procedures

PRAC Led
Revlimid - lenalidomide - EMEA/H/C/000717/II/0095, Orphan
MAH: Celgene Europe Limited, Rapporteur: Alexandre Moreau, PRAC Rapporteur: Claire Ferard, 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.”

B.6.12. CHMP-CAT assessed procedures

B.6.13. 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 ServicesRapporteur: Christiane Niederlaender, PRAC Rapporteur: Sabine Straus

B.6.14. PRAC assessed ATMP procedures

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

WS1200
Lyrica-EMEA/H/C/000546/WS1200/0089
Pregabalin Pfizer-EMEA/H/C/003880/WS1200/0019
MAH: Pfizer Limited, Lead Rapporteur: Johann Lodewijk Hillege“To update section 4.8 of the Summary of Product Characteristing and section 4 of the Package Leaflet following the outcome of the PRAC Post-Authorisation Measure (LEG) resulting from PSUSA procedure EMEA/H/C/PSUSA/00002511/201601, procedure number EMEA/H/C/000546/LEG/052 (Lyrica) and EMEA/H/C/003880/LEG/005 (Pregabalin Pfizer) adopted on 23 March 2017.”

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 19-22 June 2017 CHMP plenary:

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

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