18 April 2017
EMA/244895/2017
Inspections, Human Medicines Pharmacovigilance and Committees Division

Committee for medicinal products for human use (CHMP)

Draft agenda for the meeting on 18-21 April 2017

Chair: Tomas Salmonson – Vice-Chair: Harald Enzmann

18 April 2017, 12:30 – 21:00, room 3A

19 April 2017, 08:30 – 21:00, room 3A

20 April 2017, 08:30 – 21:00, room 3A

21 April 2017, 08:30 – 13:00, room 3A

Health and safety information

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

Disclaimers

Some of the information contained in this agenda is considered commercially confidential or sensitive and therefore not disclosed. With regard to intended therapeutic indications or procedure scopes listed against products, it must be noted that these may not reflect the full wording proposed by applicants and may also vary during the course of the review. Additional details on some of these procedures will be published in the CHMP meeting highlights once the procedures are finalised and start of referrals will also be available.

Of note, this agenda is a working document primarily designed for CHMP members and the work the Committee undertakes.

Note on access to documents

Some documents mentioned in the agenda cannot be released at present following a request for access to documents within the framework of Regulation (EC) No 1049/2001 as they are subject to on-going procedures for which a final decision has not yet been adopted. They will become public when adopted or considered public according to the principles stated in the Agency policy on access to documents (EMA/127362/2006).

30 Churchill Place ● Canary Wharf ● London E14 5EU ● United Kingdom

Telephone +44 (0)20 3660 6000 **Facsimile** +44 (0)20 3660 5520
Send a question via our website www.ema.europa.eu/contact

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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 18-21 April 2017. See April 2017 CHMP minutes (to be published post May 2017 CHMP meeting).

1.2. Adoption of agenda

CHMP agenda for 18-21 April 2017

1.3. Adoption of the minutes

CHMP minutes for 20-23 March 2017.

2. Oral Explanations

2.1. Pre-authorisation procedure oral explanations

2.1.1. - anamorelin - EMEA/H/C/003847

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

Scope: Oral explanation

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

List of Outstanding Issues adopted on 23.02.2017, 10.11.2016. List of Questions adopted on 25.02.2016.

Participation of patients' representatives

2.1.2. - trientine tetrahydrochloride - Orphan - EMEA/H/C/004005

GMP-Orphan SA; Wilson's disease

Scope: Oral explanation

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

List of Outstanding Issues adopted on 13.10.2016. List of Questions adopted on 28.04.2016.

2.1.3. - masitinib - Orphan - EMEA/H/C/004159

AB Science; treatment of mastocytosis

Scope: Oral explanation

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

List of Outstanding Issues adopted on 23.02.2017. List of Questions adopted on 15.09.2016.

2.1.4. - vosaroxin - Orphan - EMEA/H/C/004118

Sunesis Europe Ltd; treatment acute myeloid leukaemia

Scope: Oral explanation, report from SAG

Action: Oral explanation to be held on 19 April 2017 at time 09.00

List of Outstanding Issues adopted on 15.12.2016. List of Questions adopted on 28.04.2016.

2.1.5. - padeliporfin - EMEA/H/C/004182

treatment of prostate cancer

Scope: Oral explanation, report from SAG

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

List of Outstanding Issues adopted on 15.12.2016. List of Questions adopted on 26.05.2016.

2.1.6. - patiromer sorbitex calcium - EMEA/H/C/004180

treatment of hyperkalaemia

Scope: Oral explanation

Action: Oral explanation to be held on 20 April 2017 at time 18:00

List of Outstanding Issues adopted on 23.02.2017. List of Questions adopted on 15.09.2016.

2.1.7. - human IgG1 monoclonal antibody specific for human interleukin-1 alpha - EMEA/H/C/004388

treatment of metastatic colorectal cancer

Scope: Oral explanation

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

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

21.07.2016.

BWP report

2.2. Re-examination procedure oral explanations

2.3. Post-authorisation procedure oral explanations

2.4. Referral procedure oral explanations

3. Initial applications

3.1. Initial applications; Opinions

3.1.1. - inotuzumab ozogamicin - Orphan - EMEA/H/C/004119

Pfizer Limited; treatment B-cell precursor acute lymphoblastic leukaemia (ALL)

Scope: Opinion

Action: For adoption

List of Outstanding Issues adopted on 23.02.2017. List of Questions adopted on 15.09.2016.

BWP report

3.1.2. - cerliponase alfa - Orphan - EMEA/H/C/004065

Accelerated assessment

BioMarin International Limited; treatment of neuronal ceroid lipofuscinosis type 2

Scope: Opinion

Action: For adoption

List of Outstanding Issues adopted on 21.02.2017. List of Questions adopted on 13.12.2016.

BWP report

3.1.3. - etanercept - EMEA/H/C/004192

treatment of rheumatoid arthritis, juvenile idiopathic arthritis, psoriatic arthritis, axial spondyloarthritis, ankylosing spondylitis (AS), non-radiographic axial spondyloarthritis, plaque psoriasis and paediatric plaque psoriasis

Scope: Opinion

Action: For adoption

List of Outstanding Issues adopted on 23.02.2017. List of Questions adopted on 01.04.2016.

BWP report

3.1.4. - febuxostat - EMEA/H/C/004374

treatment of hyperuricaemia

Scope: Opinion

Action: For adoption

List of Outstanding Issues adopted on 23.02.2017. List of Questions adopted on 15.09.2016.

3.1.5. - sarilumab - EMEA/H/C/004254

treatment of active rheumatoid arthritis

Scope: Opinion

Action: For adoption

List of Outstanding Issues adopted on 23.02.2017. List of Questions adopted on 10.11.2016.

BWP report

3.1.6. - rituximab - EMEA/H/C/003903

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

Scope: Opinion

Action: For adoption

List of Outstanding Issues adopted on 23.02.2017. List of Questions adopted on 15.09.2016.

3.1.7. - rituximab - EMEA/H/C/004729

treatment of Non-Hodgkin's lymphoma (NHL), Rheumatoid arthritis and Granulomatosis with polyangiitis and microscopic polyangiitis

Scope: Opinion

Action: For adoption

List of Outstanding Issues adopted on 23.02.2017.

3.1.8. - dimethyl fumarate - EMEA/H/C/002157

treatment of moderate to severe plaque psoriasis in adults in need of systemic drug therapy

Scope: Opinion/List of Outstanding Issues

Action: For adoption

List of Outstanding Issues adopted on 23.02.2017. List of Questions adopted on 28.04.2016.

3.1.9. - nusinersen - Orphan - EMEA/H/C/004312

Accelerated assessment

Biogen Idec Ltd; for the treatment of Spinal Muscular Atrophy (SMA).

Scope: Opinion

Action: For adoption

Oral explanation held on 22.03.2017. List of Outstanding Issues adopted on 23.03.2017. List of Questions adopted on 24.01.2017.

3.1.10. - carglumic acid - EMEA/H/C/004019

treatment of hyperammoniemia

Scope: Opinion

Action: For adoption

List of Outstanding Issues adopted on 23.02.2017. List of Questions adopted on 15.09.2016.

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

3.2.1. - insulin lispro - EMEA/H/C/004303

treatment of diabetes mellitus

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 26.01.2017.

BWP report

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

treatment of haemophilia A

Scope: Day 180 list of outstanding issue

Action: For adoption

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

BWP report

3.2.3. - ribociclib - EMEA/H/C/004213

treatment of breast cancer

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 26.01.2017.

3.2.4. - brodalumab - EMEA/H/C/003959

moderate to severe plaque psoriasis

Scope: Day 180 list of outstanding issue

Action: For adoption

Oral explanation held on 24.01.2017. List of Outstanding Issues adopted on 26.01.2017, 10.11.2016, 15.09.2016. List of Questions adopted on 01.04.2016.

3.2.5. - cladribine - EMEA/H/C/004230

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

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 10.11.2016.

3.2.6. - cenegermin - Orphan - EMEA/H/C/004209

Accelerated assessment

Dompe farmaceutici s.p.a.; treatment of neurotrophic keratitis

Scope: Day 120 list of outstanding issue

Action: For adoption

List of Questions adopted on 21.02.2017.

BWP report

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

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

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 15.12.2016.

3.2.8. - adalimumab - EMEA/H/C/004279

treatment of rheumatoid arthritis, psoriatic arthritis and ankylosing spondylitis

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 10.11.2016.

BWP report

3.2.9. - atezolizumab - EMEA/H/C/004143

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

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 15.09.2016.

BWP report

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

Ipsen Pharma; treatment of carcinoid syndrome

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 10.11.2016.

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

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

Accelerated assessment

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

Scope: Day 90 list of questions

Action: For adoption

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

treatment of adrenal insufficiency

Scope: Day 120 list of questions

Action: For adoption

3.3.3. - anagrelide - EMEA/H/C/004585

reduction of elevated platelet counts in at risk essential thrombocythaemia patients

Scope: Day 120 list of questions

Action: For adoption

3.3.4. - benralizumab - EMEA/H/C/004433

treatment of severe asthma with an eosinophilic phenotype

Scope: Day 120 list of questions

Action: For adoption

BWP report

3.3.5. - betrixaban - EMEA/H/C/004309

treatment of prophylaxis of venous thromboembolism (VTE)

Scope: Day 120 list of questions

Action: For adoption

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

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

Scope: Day 120 list of questions

Action: For adoption

BWP report

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

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

Scope: Day 120 list of questions

Action: For adoption

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

AVI Biopharma International Ltd; treatment of Duchenne muscular dystrophy

Scope: Day 120 list of questions

Action: For adoption

3.3.9. - fulvestrant - EMEA/H/C/004649

Treatment of breast cancer

Scope: Day 120 list of questions

Action: For adoption

3.3.10. - guselkumab - EMEA/H/C/004271

treatment of plaque psoriasis

Scope: Day 120 list of questions

Action: For adoption

BWP report

3.3.11. - bevacizumab - EMEA/H/C/004360

treatment of metastatic breast cancer, unresectable advanced, metastatic or recurrent non-small cell lung cancer, unresectable advanced metastatic or recurrent non-squamous non-small cell lung cancer, advanced and/or metastatic renal cell cancer, advanced epithelial ovarian, fallopian tube, or primary peritoneal cancer, platinum-sensitive epithelial ovarian, fallopian tube or primary peritoneal cancer

Scope: Day 120 list of questions

Action: For adoption

BWP report

3.3.12. - bevacizumab - EMEA/H/C/004728

treatment of metastatic breast cancer, unresectable advanced, metastatic or recurrent non-small cell lung cancer, unresectable advanced metastatic or recurrent non-squamous non-small cell lung cancer, advanced and/or metastatic renal cell cancer, advanced epithelial ovarian, fallopian tube, or primary peritoneal cancer, platinum-sensitive epithelial ovarian, fallopian tube or primary peritoneal cancer

Scope: Day 120 list of questions

Action: For adoption

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

Pfizer Limited; combination therapy with daunorubicin (DNR) and cytarabine (AraC) for the

treatment of adult patients with previously untreated, de novo acute myeloid leukaemia (AML)

Scope: Day 120 list of questions

Action: For adoption

BWP report

3.3.14. - semaglutide - EMEA/H/C/004174

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

Scope: Day 120 list of questions

Action: For adoption

BWP report

3.3.15. - ritonavir - EMEA/H/C/004549

treatment of HIV-1

Scope: Day 120 list of questions

Action: For adoption

3.3.16. - rotigotine - EMEA/H/C/004286

treatment of idiopathic Restless Legs Syndrome and Parkinson's disease

Scope: Day 120 list of questions

Action: For adoption

3.3.17. - Herpes zoster vaccine (recombinant surface antigen, adjuvanted) - EMEA/H/C/004336

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

Scope: Day 120 list of questions

Action: For adoption

BWP report

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

Accelerated assessment

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

Scope: Day 90 list of questions

Action: For adoption

3.3.19. - tacrolimus - EMEA/H/C/004435

prophylaxis of transplant rejection and treatment of allograft rejection

Scope: Day 120 list of questions

Action: For adoption

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

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

Scope: Day 120 list of questions

Action: For adoption

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

Accelerated assessment

Santen Oy; treatment of severe vernal keratoconjunctivitis (VKC)

Scope: Day 90 list of questions

Action: For adoption

3.3.22. - human fibrinogen / human thrombin - EMEA/H/C/004446

treatment of haemostasis

Scope: Day 120 list of questions

Action: For adoption

BWP report

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

MolMed SpA; treatment of advanced malignant pleural mesothelioma

Scope: Day 120 list of questions

Action: For adoption

BWP report

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

3.4.1. - spheroids of human autologous matrix-associated chondrocytes - ATMP - EMEA/H/C/002736

Claimed indication: repair of symptomatic articular cartilage defects of the femoral condyle and the patella of the knee (International Cartilage Repair Society [ICRS] grade III or IV) with defect sizes up to 10 cm^2 in adults

Scope: update on the CAT discussion

Action: For discussion

List of Outstanding Issues adopted on 17.02.2017. List of Questions adopted on 19.04.2013.

BWP report

3.4.2. - 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: Request by the applicant for an extension of clock stop to respond to the List of Outstanding Issues adopted on 23 March 2017. The new timetable has been adopted by CHMP on 12 April via written procedure.

Action: For information

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

3.4.3. - pegfilgrastim - EMEA/H/C/004413

treatment of neutropenia

Scope: Letter from the applicant dated 7 April 2017 requesting an extension of clock stop to respond to Day 120 list of questions adopted on 23.03.2017

Action: For adoption

List of Questions adopted on 23.03.2017.

3.4.4. - cariprazine - EMEA/H/C/002770

treatment of schizophrenia

Scope: cancellation of the consultation of expert Healthcare Professionals (ophthalmology)

Action: For information

List of Outstanding issues adopted on 23.02.2017. List of Questions adopted on 21.07.2016.

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

3.6. Initial applications in the decision-making phase

3.6.1. Lokelma - sodium zirconium cyclosilicate - EMEA/H/C/004029

AstraZeneca AB; for the treatment of hyperkalaemia

Scope: update

Action: For discussion

3.7. Withdrawals of initial marketing authorisation application

3.7.1. - solithromycin - EMEA/H/C/004179

treatment of bacterial infections

Scope: letter from the applicant dated 27.03.2017 informing about the withdrawal of initial marketing authorisation application

Action: For information

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

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

4.1.1. Celsentri - maraviroc - EMEA/H/C/000811/X/0046/G

ViiV Healthcare UK Limited

Rapporteur: Filip Josephson, Co-Rapporteur: Bruno Sepodes, PRAC Rapporteur: Qun-Ying Yue

Scope: "Extension application to introduce new pharmaceutical form (20mg/ml oral solution) and 2 new strengths of film-coated tablets (25mg and 75mg) to the currently approved presentations for Celsentri, grouped with extension of indication to include paediatric use (2 to 18 years).

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.

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

Action: For adoption

List of Outstanding Issues adopted on 23.02.2017. List of Questions adopted on 13.10.2016.

4.1.2. Kuvan - sapropterin - Orphan - EMEA/H/C/000943/X/0047

BioMarin International Limited

Rapporteur: Patrick Salmon, PRAC Rapporteur: Almath Spooner

Scope: "Extension application to introduce a new pharmaceutical form associated with new strength (100 mg and 500 mg powder for oral solution)."

Action: For adoption

List of Questions adopted on 23.02.2017.

4.1.3. Nexium Control - esomeprazole - EMEA/H/C/002618/X/0016

Pfizer Consumer Healthcare Ltd

Rapporteur: Romaldas Mačiulaitis, Co-Rapporteur: Robert James Hemmings, PRAC Rapporteur: Simona Kudeliene

Scope: "Extension application to introduce a new pharmaceutical form (Gastro-resistant capsule, hard)"

Action: For adoption

List of Outstanding Issues adopted on 23.02.2017. List of Questions adopted on 10.11.2016.

4.1.4. Revestive - teduglutide - Orphan - EMEA/H/C/002345/X/0029

Shire Pharmaceuticals Ireland Ltd

Rapporteur: Sinan B. Sarac, PRAC Rapporteur: Torbjorn Callreus

Scope: "Extension application to add a new strength of 1.25mg (paediatric formulation)."

Action: For adoption

List of Questions adopted on 10.11.2016.

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

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

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

Otsuka Pharmaceutical Europe Ltd

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

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

Action: For adoption

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

Novartis Europharm Ltd

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

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

Action: For adoption

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

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

Novartis Europharm Ltd

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

Scope: Letter from the applicant dated 30 March 2017 requesting an extension of 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.

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. Avastin - bevacizumab - EMEA/H/C/000582/II/0092

Roche Registration Limited

Rapporteur: Sinan B. Sarac, Co-Rapporteur: Bjorg Bolstad, PRAC Rapporteur: Doris Stenver

Scope: "Extension of Indication to include the use of Avastin in combination with paclitaxel and carboplatin for the treatment of adult patients with first recurrence of platinum-sensitive epithelial ovarian, fallopian tube or primary peritoneal cancer.
As a consequence, sections 4.1, 4.2, 4.8 and 5.1 of the SmPC are updated with efficacy and safety information from study GOG-0213. The Package Leaflet is updated in accordance. An update RMP is also included (version 27)."

Action: For adoption

Request for Supplementary Information adopted on 15.12.2016.

5.1.2. Cubicin - daptomycin - EMEA/H/C/000637/II/0061

Merck Sharp & Dohme Limited

Rapporteur: Greg Markey, Co-Rapporteur: Svein Rune Andersen, PRAC Rapporteur: Julie Williams

Scope: "Extension of indication to extend the S. aureus bacteraemia indication to include paediatric patients 1 to 17 years of age; as a consequence, sections 4.1, 4.2, 4.4, 4.5, 4.8, 5.1, 5.2 and 6.6 of the SmPC are updated. The Package Leaflet and Labelling are updated accordingly.

In addition, the marketing authorisation holder (MAH) took the opportunity to bring the product information in line with the latest QRD template version 10 and to combine the SmPCs for both strengths (350 and 500 mg). The MAH also updated the RMP, from last approved version 9.1 to the current proposed version 10.0."

Action: For adoption

5.1.3. 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 26.01.2017.

5.1.4. Opdivo - nivolumab - EMEA/H/C/003985/II/0019

Bristol-Myers Squibb Pharma EEIG

Rapporteur: Aranzazu Sancho-Lopez, Co-Rapporteur: Paula Boudewina van Hennik, PRAC Rapporteur: Brigitte Keller-Stanislawski

Scope: "Extension of Indication to include the treatment of locally advanced unresectable or metastatic urothelial carcinoma in adults after failure of prior platinum-containing therapy for Opdivo.
As a consequence, sections 4.1, 4.4, 4.8, 5.1 and 5.2 of the SmPC are updated in order to add the proposed indication, add a warning about the patient populations excluded from the clinical trial, and update the safety information. The Package Leaflet is updated in accordance.
Moreover, the updated RMP version 7.0 has been submitted"

Action: For adoption

Request for Supplementary Information adopted on 23.03.2017, 15.12.2016.

5.1.5. Tasigna - nilotinib - Orphan - EMEA/H/C/000798/II/0084/G

Novartis Europharm Ltd

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

Scope: "This grouped variation application consists of three Type II variation applications as follows:
- Update of the 150 mg SmPC sections 4.1, 4.2, 4.4, 4.8 and 5.1, and Package Leaflet based on the results from study CAMN107I2201 (ENESTfreedom): A Phase II, single-arm study evaluating nilotinib treatment discontinuation (treatment-free remission (TFR)) in newly-diagnosed patients with Philadelphia chromosome-positive chronic myelogenous leukemia in chronic phase (Ph+ CML-CP) who achieved a sustained deep molecular response.
- Update of the 150 mg and 200 mg Tasigna SmPC sections 4.1, 4.2, 4.4, 4.8 and 5.1, and Package Leaflet based on the results from study CAMN107A2408 (ENESTop): A Phase II, single-arm study evaluating nilotinib treatment discontinuation (treatment-free remission (TFR)) in patients with Ph+ CML-CP who achieved a sustained deep molecular response on nilotinib treatment after switching from imatinib treatment.
- Update of the 200 mg Tasigna SmPC sections 4.8 and 5.1, based on the results from study CAMN107A2405 (ENESTcmr): A Phase III open-label, randomised study to evaluate nilotinib or imatinib treatment in patients with Ph+ CML-CP who have not achieved a deep molecular response after previous imatinib therapy.
Additional changes to the labelling are proposed to comply with the latest QRD template

version 10.
An updated RMP, version 16, is also provided in this application."

Report from SAG Oncology

Action: For adoption

Request for Supplementary Information adopted on 23.02.2017, 13.10.2016.

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

8.2. 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. Hemoprostol - misoprostol - Article 58 - EMEA/H/W/002652

Linepharma International Limited; treatment and prevention of Post Partum Haemorrhage

Rapporteur: Paula Boudewina van Hennik, Co-Rapporteur: Nithyanandan Nagercoil, PRAC Rapporteur: Alexandre Moreau

Scope: Withdrawal of Article 58 scientific opinion

Action: For information

Article 58 of Regulation (EC) No 726/2004

10. Referral procedures

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

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

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

10.4. Disagreement between Member States on application for medicinal product (potential serious risk to public health) –under Article 29(4) of Directive 2001/83/EC

10.4.1. 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: List of outstanding issues/Opinion

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.

Action: For adoption

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

10.5.1. Etopophos and associated names– etoposide - EMEA/H/A-30/1417

MAH: Bristol-Myers Squibb group of companies and associated companies

Rapporteur: Greg Markey, Co-Rapporteur: Paula Boudewina van Hennik,

Scope: Opinion

Harmonisation exercise for Etopophos and associated names

Action: For adoption

List of outstanding issues adopted 15.12.2016, 21.07.2016, 25.02.2016 CHMP, List of Questions adopted on 22.10.2015.

10.5.2. Vepesid and associated names - etoposide - EMEA/H/A-30/1425

MAH: Bristol-Myers Squibb group of companies and associated companies

Rapporteur: Greg Markey, Co-Rapporteur: Paula van Hennik

Scope: Opinion

Harmonisation exercise for Vepesid and associated names

Action: For adoption

List of outstanding issues adopted 15.12.2016, 21.07.2016, 25.02.2016 CHMP. List of Questions adopted on 22.10.2015.

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

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

April 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

14.2. Coordination with EMA Scientific Committees

14.2.1. Pharmacovigilance Risk Assessment Committee (PRAC)

Summary of recommendations and advice of PRAC meeting held on 3-6 April 2017

Action: For information

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

Reports (EURD list) for April 2017

Action: For adoption

14.2.2. Committee for Advanced Therapies (CAT)

CAT draft minutes of meeting held on 10-12 April 2017
Action: For information

14.2.3. Committee for Herbal Medicinal Products (HMPC)

Report from the HMPC meeting held on 27-28 March 2017

Action: For information

14.2.4. Paediatric Committee (PDCO)

PIPs reaching D30 at April 2017 PDCO

Action: For information

Report from the PDCO meeting held on 21-24 March 2017

Action: For information

14.2.5. Committee for Orphan Medicinal Products (COMP)

Report from the COMP meeting held on 10-12 April 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 18-20 April 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 28 March April 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. Name Review Group (NRG)

Table of Decisions of the NRG meeting held on 29 March 2017.

Action: For adoption

14.3.3. Biologics Working Party (BWP)

Chair: Sol Ruiz,

Election of BWP Vice-Chair

Action: For adoption

14.3.4. Vaccines Working Party (VWP)

Chair: Mair Powell

Election of VWP Vice-Chair

Action: For adoption

VWP responses to a PDCO list of questions related to the PIP EMEA-001888-PIP01-15, adopted by written procedure on 11 April 2017

Action: For information

14.3.5. Biosimilar Medicinal Product Working Party (BMWP)

Chair: Elena Wolff-Holz/Martina Weise

Nomination of additional PK expert to BMWP

• Current list of members

Action: For adoption

14.4. Cooperation within the EU regulatory network

14.4.1. Revision of the Commission Regulation (EC) No 847/2000 of April 2000 laying down the provisions for implementation of the Criteria for designation of a medicinal product as an orphan medicinal product and definitions of the Concept 'similar medicinal product and 'clinical superiority'

Action: For discussion

14.5. Cooperation with International Regulators

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

14.7. CHMP work plan

14.8. Planning and reporting

14.9. Others

15. Any other business

15.1. AOB topic

15.1.1. Preparedness of the system and capacity increase

Action: For discussion

16. Explanatory notes

The notes below give a brief explanation of the main sections and headings in the CHMP agenda and should be read in conjunction with the agenda or the minutes.

Oral explanations *(section 2)*

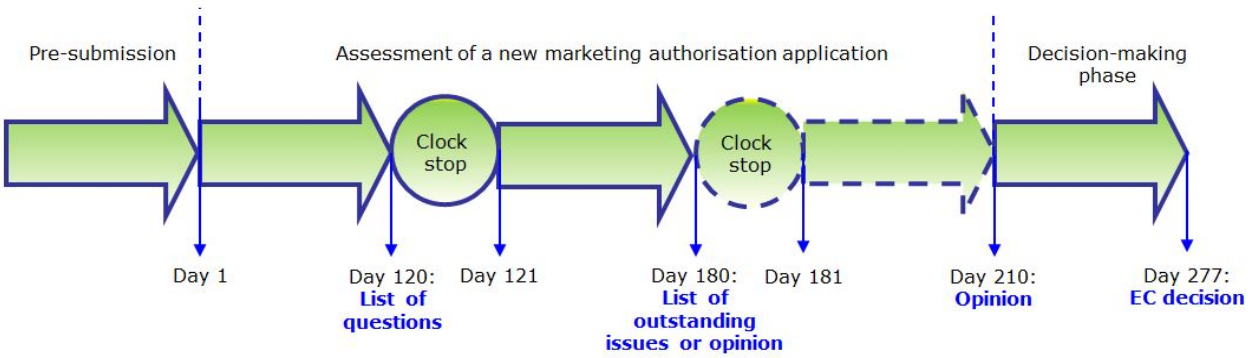
The items listed in this section are those for which marketing authorisation holders (MAHs) or applicants have been invited to the CHMP plenary meeting to address questions raised by the Committee. Oral explanations normally relate to on-going applications (section 3, 4 and 5) or referral procedures (section 10) but can relate to any other issue for which the CHMP would like to discuss with company representatives in person.

Initial applications *(section 3)*

This section lists applications for marketing authorisations of new medicines that are to be discussed by the Committee.

Section 3.1 is for medicinal products nearing the end of the evaluation and for which the CHMP is expected to adopt an **opinion** at this meeting on whether marketing authorisation should be granted. Once adopted, the CHMP opinion will be forwarded to the European Commission for a final legally binding decision valid throughout the EU.

The other items in the section are listed depending on the stage of the evaluation, which is shown graphically below:



The assessment of an application for a new medicine takes up to 210 'active' days. This active evaluation time is interrupted by at least one 'clock-stop' during which time the applicant prepares the answers to questions from the CHMP. The clock stop happens after day 120 and may also happen after day 180, when the CHMP has adopted a list of questions or outstanding issues to be addressed by the company. Related discussions are listed in the agenda under sections 3.2 (**Day 180 List of outstanding issues**) and 3.3 (**Day 120 list of questions**).

CHMP discussions may also occur at any other stage of the evaluation, and these are listed under section 3.4, **update on ongoing new applications for centralised procedures**.

The assessment leads to an opinion from the CHMP by day 210. Following a CHMP opinion the European Commission takes usually 67 days to issue a legally binding decision (i.e. by day 277 of the procedure). CHMP discussions on products that have received a CHMP opinion and are awaiting a decision are listed under section 3.6, **products in the decision making phase**.

Extension of marketing authorisations according to Annex I of Reg. 1234/2008 *(section 4)*

Extensions of marketing authorisations are applications for the change or addition of new strengths,

formulations or routes of administration to existing marketing authorisations. Extension applications follow a 210-day evaluation process, similarly to applications for new medicines (see figure above).

Type II variations - Extension of indication procedures *(section 5)*

Type II variations are applications for a change to the marketing authorisation which requires an update of the product information and which is not covered in section 4. Type II variations include applications for a new use of the medicine (extension of indication), for which the assessment takes up to 90 days. For the applications listed in this section, the CHMP may adopt an opinion or request supplementary information from the applicant.

Ancillary medicinal substances in medical devices *(section 6)*

Although the EMA does not regulate medical devices it can be asked by the relevant authorities (the so-called Notified Bodies) that are responsible for regulating these devices to give a scientific opinion on a medicinal substance contained in a medical device.

Re-examination procedures (new applications) under article 9(2) of regulation no 726/2004 *(section 3.5)*

This section lists applications for new marketing authorisation for which the applicant has requested a re-examination of the opinion previously issued by the CHMP.

Re-examination procedures *(section5.3)*

This section lists applications for type II variations (including extension of indication applications) for which the applicant has requested re-examination of the opinion previously issued by the CHMP.

Withdrawal of application *(section 3.7)*

Applicants may decide to withdraw applications at any stage during the assessment and a CHMP opinion will therefore not be issued. Withdrawals are included in the agenda for information or discussion, as necessary.

Procedure under article 83(1) of regulation (EC) 726/2004 (compassionate use) *(section 7)*

Compassionate use is a way of making available to patients with an unmet medical need a promising medicine which has not yet been authorised (licensed) for their condition. Upon request, the CHMP provides recommendations to all EU Member States on how to administer, distribute and use certain medicines for compassionate use.

Pre-submission issues *(section 8)*

In some cases the CHMP may discuss a medicine before a formal application for marketing authorisation is submitted. These cases generally refer to requests for an accelerated assessment for medicines that are of major interest for public health or can be considered a therapeutic innovation. In case of an accelerated assessment the assessment timetable is reduced from 210 to 150 days.

Post-authorisation issues *(section 9)*

This section lists other issues concerning authorised medicines that are not covered elsewhere in the agenda. Issues include supply shortages, quality defects, some annual reassessments or renewals or type II variations to marketing authorisations that would require specific discussion at the plenary.

Referral procedures *(section 10)*

This section lists referrals that are ongoing or due to be started at the plenary meeting. A referral is a procedure used to resolve issues such as concerns over the safety or benefit-risk balance of a medicine or a class of medicines. In a referral, the EMA is requested to conduct a scientific assessment of a

particular medicine or class of medicines on behalf of the EU. Further information on such procedures can be found here.

Pharmacovigilance issues *(section 11)*

This section lists issues that have been discussed at the previous meeting of the PRAC, the EMA's committee responsible for evaluating and monitoring safety issues for medicines. Feedback is provided by the PRAC. This section also refers to the early notification system, a system used to notify the European regulatory network on proposed EMA communication on safety of medicines.

Inspections Issues *(section 12)*

This section lists inspections that are undertaken for some medicinal products. Inspections are carried out by regulatory agencies to ensure that marketing authorisation holders comply with their obligations. Inspection can relate to good manufacturing practice (GMP), good clinical practice (GCP), good laboratory practice (GLP) or good pharmacovigilance practice (GVP).

Innovation task force *(section 13)*

The Innovation Task Force (ITF) is a body set up to encourage early dialogue with applicants developing innovative medicines. Minutes from the last ITF meeting as well as any related issue that requires discussion with the CHMP are listed in this section of the agenda. Further information on the ITF can be found here.

Scientific advice working party (SAWP) *(section 14.3.1)*

This section refers to the monthly report from the CHMP's Scientific Advice Working Party (SAWP) on scientific advice given to companies during the development of medicines. Further general information on SAWP can be found here.

Satellite groups / other committees *(section 14.2)*

This section refers to the reports from groups and committees making decisions relating to human medicines: the Coordination Group for Mutual Recognition and Decentralised Procedures – Human (CMDh), the Committee for Orphan Medicinal Products (COMP), the Committee for Herbal Medicinal Products (HMPC), Paediatric Committee (PDCO), the Committee for Advanced Therapies (CAT) and the Pharmamacovigilance Risk Assessment Committee (PRAC).

Invented name issues *(section 14.3)*

This section list issues related to invented names proposed by applicants for new medicines. The CHMP has established the Name Review Group (NRG) to perform reviews of the invented names. The group's main role is to consider whether the proposed names could create a public-health concern or potential safety risk. Further information can be found here.

More detailed information on the above terms can be found on the EMA website: www.ema.europa.eu/

EUROPEAN MEDICINES AGENCY
SCIENCE MEDICINES HEALTH

18 April 2017
EMA/244896/2017

Annex to April 2017 CHMP Agenda

Pre submission and post authorisations issues

30 Churchill Place ● Canary Wharf ● London E14 5EU ● United Kingdom

Telephone +44 (0)20 3660 6000 **Facsimile** +44 (0)20 3660 5520
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
April 2017: **For adoption**

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

Final Outcome of Rapporteurship allocation for
April 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

Defitelio - defibrotide -
EMEA/H/C/002393/S/0020, Orphan
MAH: Gentium S.r.l., Rapporteur: Nithyanandan Nagercoil, PRAC Rapporteur: Julie Williams

Kolbam - cholic acid -
EMEA/H/C/002081/S/0020, Orphan
MAH: Retrophin Europe Ltd, Rapporteur: Robert James Hemmings, PRAC Rapporteur: Patrick Batty
Request for Supplementary Information adopted on 23.03.2017.

Raxone - idebenone -
EMEA/H/C/003834/S/0005, Orphan
MAH: Santhera Pharmaceuticals (Deutschland) GmbH, Rapporteur: John Joseph Borg, PRAC Rapporteur: Carmela Macchiarulo
Request for Supplementary Information adopted on 23.03.2017, 26.01.2017.

SCENESSE - afamelanotide -
EMEA/H/C/002548/S/0011, Orphan
MAH: Clinuvel (UK) Limited, Rapporteur: Harald Enzmann, Co-Rapporteur: Robert James Hemmings, PRAC Rapporteur: Valerie Strassmann

B.2. RENEWALS OF MARKETING AUTHORISATIONS OUTCOMES

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

alli - orlistat - EMEA/H/C/000854/R/0054
MAH: Glaxo Group Ltd, Informed Consent of Xenical, Rapporteur: Greg Markey, Co-Rapporteur: Eleftheria Nikolaidi, PRAC Rapporteur: Patrick Batty
Request for Supplementary Information adopted on 23.02.2017.

Capecitabine medac - capecitabine -
EMEA/H/C/002568/R/0017
MAH: Medac Gesellschaft fuer klinische Spezialpraeparate m.b.H, Generic, Generic of Xeloda, Rapporteur: Filip Josephson, PRAC Rapporteur: Martin Huber

Increlex - mecasermin -
EMEA/H/C/000704/R/0042, Orphan
MAH: Ipsen Pharma, Rapporteur: Outi Mäki-Ikola, Co-Rapporteur: Agnes Gyurasics, PRAC Rapporteur: Kirsti Villikka
Request for Supplementary Information adopted on 23.02.2017.

B.2.2. Renewals of Marketing Authorisations for unlimited validity

Atriance - nelarabine -
EMEA/H/C/000752/R/0037, Orphan
MAH: Novartis Europharm Ltd, Rapporteur: Sinan B. Sarac, Co-Rapporteur: Paula Boudewina van Hennik, PRAC Rapporteur: Torbjorn Callreus
Request for Supplementary Information adopted on 23.02.2017.

ECALTA - anidulafungin -
EMEA/H/C/000788/R/0033
MAH: Pfizer Limited, Rapporteur: Johann Lodewijk Hillege, Co-Rapporteur: Hanne Lomholt Larsen, PRAC Rapporteur: Sabine Straus

Revestive - teduglutide - EMEA/H/C/002345/R/0038, Orphan
MAH: Shire Pharmaceuticals Ireland Ltd, Rapporteur: Sinan B. Sarac, Co-Rapporteur: Harald Enzmann, PRAC Rapporteur: Torbjorn Callreus
Request for Supplementary Information adopted on 23.03.2017.

B.2.3. Renewals of Conditional Marketing Authorisations

Translarna - ataluren - EMEA/H/C/002720/R/0032, Orphan
MAH: PTC Therapeutics International Limited, Rapporteur: Johann Lodewijk Hillege, Co-Rapporteur: Concepcion Prieto Yerro, PRAC Rapporteur: Sabine Straus

Zalmoxis - allogeneic t cells genetically modified with a retroviral vector encoding for a truncated form of the human low affinity nerve growth factor receptor (ðlngfr) and the herpes simplex i virus thymidine kinase (hsv-tk mut2) - EMEA/H/C/002801/R/0003, Orphan, ATMP
MAH: MolMed SpA, Rapporteur: Johannes Hendrikus Ovelgonne, PRAC Rapporteur: Brigitte Keller-Stanislawski

B.3. POST-AUTHORISATION PHARMACOVIGILANCE OUTCOMES

Signal detection

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

Albiglutide (EMEA/H/C/002735) (Eperzan);
MAH: GlaxoSmithKline Trading Services Limited; Rapporteur: Kristina Dunder, Co-Rapporteur: Svein Rune Andersen, PRAC Rapporteur: Julie Williams,
Signal of acute kidney injury: **For adoption**

Lenflunomide; teriflunomide, (Arava, Lenflunomide Winthrop, Lenflunomide Medac, Aubagio); Rapporteur: , Co-Rapporteur:, PRAC Rapporteur: Sabine Straus,
Signal of falsely decreased ionized calcium levels: **For adoption**

Temozolomide, (Temodal) MAH: Merck Sharp & Dohme Limited
Rapporteur: Harald Enzmann, Co-Rapporteur: Filip Josephson, PRAC Rapporteur: Martin Huber,
Signal of meningoencephalitis Herpetic: **For adoption**

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

EMEA/H/C/PSUSA/00000208/201609
(anagrelide)
CAPS:
Xagrid (EMEA/H/C/000480) (anagrelide), MAH: Shire Pharmaceutical Contracts Ltd., Rapporteur: Alexandre Moreau
NAPS:
Thromboreductin 0,5 mg 16/0027/04-S SK - AOP ORPHAN PHARMACEUTICALS AG
Thromboreductin 0,5 mg capsules 20060389 BG - AOP ORPHAN PHARMACEUTICALS AG
Thromboreductin 0,5 mg cietās kapsulas 04-0076 LV - AOP ORPHAN PHARMACEUTICALS AG
Thromboreductin 0,5 mg kapsule UP/I-530-09/08-01/408 HR - AOP ORPHAN PHARMACEUTICALS AG
Thromboreductin 0,5 mg kemény kapszula OGYI-T-9545/01 HU - AOP ORPHAN PHARMACEUTICALS AG
Thromboreductin 0,5 mg kietosios kapsules LT/1/03/2755/001 LT - AOP ORPHAN PHARMACEUTICALS AG
THROMBOREDUCTIN 0,5 mg trde kapsule 5363-I-2767/10 SI - AOP ORPHAN PHARMACEUTICALS AG
THROMBOREDUCTIN 0,5 mg tvrdé tobolky 16/123/04-C CZ - AOP ORPHAN PHARMACEUTICALS AG
Thromboreductin, 0,5 mg kapsułki twarde 10331 PL - AOP ORPHAN PHARMACEUTICALS AG
THROMBOREDUCTIN, capsule, 0,5 mg 7585/2015/01 RO - AOP ORPHAN PHARMACEUTICALS AG
Thromboreductin® 0,5 mg Kapseln 1-24286 AT - AOP ORPHAN PHARMACEUTICALS AG
, PRAC Rapporteur: Caroline Laborde, “Update of section 4.4 of the SmPC to add a warning on pulmonary hypertension and section 4.8 to change the frequency of pulmonary hypertension to ‘uncommon’. The Package leaflet is updated accordingly.”

EMEA/H/C/PSUSA/00000931/201609
(daptomycin)
CAPS:

Cubicin (EMEA/H/C/000637) (daptomycin), MAH: Merck Sharp & Dohme Limited, Rapporteur: Greg Markey, PRAC Rapporteur: Julie Williams, “12 Sep 2016 to 11 Sep 2016”

EMEA/H/C/PSUSA/00000954/201609
(denosumab (indicated for osteoporosis and for bone loss associated with hormone ablation in prostate cancer))
CAPS:
Prolia (EMEA/H/C/001120) (denosumab), MAH: Amgen Europe B.V., Rapporteur: Kristina Dunder, PRAC Rapporteur: Ulla Wändel Liminga, “27/09/2015-26/09/2016”

EMEA/H/C/PSUSA/00002653/201609
(rivaroxaban)
CAPS:
Xarelto (EMEA/H/C/000944) (rivaroxaban), MAH: Bayer Pharma AG, Rapporteur: Kristina Dunder, PRAC Rapporteur: Qun-Ying Yue, “16/09/2015 - 15/09/2016”

EMEA/H/C/PSUSA/00002880/201608
(telbivudine)
CAPS:
Sebivo (EMEA/H/C/000713) (telbivudine), MAH: Novartis Europharm Ltd, Rapporteur: Joseph Emmerich, PRAC Rapporteur: Caroline Laborde, “01 Sep 2013 – 31 Aug 2016”

EMEA/H/C/PSUSA/00003001/201609
(trabectedin)
CAPS:
Yondelis (EMEA/H/C/000773) (trabectedin), MAH: Pharma Mar, S.A., Rapporteur: Sinan B. Sarac, PRAC Rapporteur: Torbjorn Callreus, “From 18 September 2015 to 17 September 2016”

EMEA/H/C/PSUSA/00003149/201608
(zoledronic acid (indicated for cancer and fractures))
CAPS:
Zoledronic acid Hospira (EMEA/H/C/002365) (zoledronic acid), MAH: Hospira UK Limited, Rapporteur: Filip Josephson
Zoledronic acid medac (EMEA/H/C/002359) (zoledronic acid), MAH: medac Gesellschaft fur klinische Spezialpraparate mbH, Rapporteur: Alar Irs
Zometa (EMEA/H/C/000336) (zoledronic acid), MAH: Novartis Europharm Ltd, Rapporteur:

Sinan B. Sarac
NAPS:
ACID ZOLEDRONIC SANDOZ 4 mg/100 ml soluţie perfuzabilă 5011/2012/01 RO - S.C. SANDOZ S.R.L.
ACID ZOLEDRONIC SANDOZ 4 mg/100 ml soluţie perfuzabilă 5011/2012/02 RO - S.C. SANDOZ S.R.L.
ACID ZOLEDRONIC SANDOZ 4 mg/100 ml soluţie perfuzabilă 5011/2012/03 RO - S.C. SANDOZ S.R.L.
ACIDE ZOLEDRONIQUE SANDOZ 4 mg/100 ml, solution pour perfusion 34009 224 088 9 7 FR - SANDOZ
ACIDE ZOLEDRONIQUE SANDOZ 4 mg/100 ml, solution pour perfusion 34009 582 943 0 3 FR - SANDOZ
ACIDE ZOLEDRONIQUE SANDOZ 4 mg/100 ml, solution pour perfusion 34009 582 944 7 1 FR - SANDOZ
ACIDE ZOLEDRONIQUE SANDOZ 4 mg/100 ml, solution pour perfusion 34009 586 953 0 8 FR - SANDOZ
Ácido zoledrónico Sandoz 4 mg/100 ml solución para perfusión 76505 ES - SANDOZ FARMACÉUTICA, S.A.
Ácido Zoledrónico Sandoz 5466867 PT - SANDOZ FARMACÊUTICA LDA.
Kyselina zoledrónová Sandoz 4 mg/100 ml infúzny roztok 87/0281/12-S SK - SANDOZ PHARMACEUTICALS D.D.
STEOZOL 4 mg/5 ml concentrado para solución para perfusión en jeringa precargada 76675 ES - CHEMI S.P.A.
STEOZOL 4 mg/5 ml pykno dialyma gia paraskeyn dialumatos pros egchysh 84684 GR - ITF HELLAS AE
STEOZOL 4 mg/5ml Konzentrat zur Herstellung einer Infusionslösung in einer Fertigspritze 84927.00.00 DE - CHEMI S.P.A.
STEOZOL 4mg/5ml concentrato per soluzione per infusione 040982011 IT - ITALFARMACO S.P.A
STEOZOL 4mg/5ml concentrato per soluzione per infusione 040982023 IT - ITALFARMACO S.P.A
STEOZOL 4mg/5ml concentrato per soluzione per infusione 040982035 IT - ITALFARMACO S.P.A
STEOZOL 4mg/5ml concentrato per soluzione per infusione 040982047 IT -

ITALFARMACO S.P.A
Zoledronic Acid Sandoz 4 mg / 100 ml infuusioneste, liuos 29715 FI - SANDOZ A/S
Zoledronic acid Sandoz 4 mg/100 ml 021979 CY - SANDOZ GMBH
Zoledronic acid Sandoz 4 mg/100 ml infusionsvätska, lösning 46101 SE - SANDOZ A/S
Zoledronic acid Sandoz 4 mg/100 ml oplossing voor infusie BE424715 BE - SANDOZ N.V.
Zoledronic acid Sandoz 4 mg/100 ml solution for infusion PL 04416/1307 UK - SANDOZ LTD
ZOLEDRONOVA KISELINA SANDOZ MG/100 ML INFUZIONEN RAZTVOR 20120593 BG - SANDOZ PHARMACEUTICALS D.D.
Zoledronsav Sandoz 4 mg/100 ml oldatos infúzió OGYI-T-22283/04 HU - SANDOZ HUNGÁRIA KFT
Zoledronsav Sandoz 4 mg/100 ml oldatos infúzió OGYI-T-22283/05 HU - SANDOZ HUNGÁRIA KFT
Zoledronsav Sandoz 4 mg/100 ml oldatos infúzió OGYI-T-22283/06 HU - SANDOZ HUNGÁRIA KFT
Zoledronsav Sandoz 4 mg/100 ml oldatos infúzió OGYI-T-22283/10 HU - SANDOZ HUNGÁRIA KFT
Zoledronsäure - 1 A Pharma 4 mg/100 ml Infusionslösung 84970.00.00 DE - 1 A PHARMA GMBH
Zoledronsäure Hexal 4 mg/100 ml – Infusionslösung 1-31351 AT - HEXAL PHARMA GMBH
Zoledronsäure Sandoz 4 mg/100 ml – Infusionslösung 1-31349 AT - SANDOZ GMBH
Zoledronska kislina Sandoz 4 mg/100 ml raztopina za infundiranje H/12/01713/001 SI - SANDOZ PHARMACEUTICALS D.D.
Zoledronska kislina Sandoz 4 mg/100 ml raztopina za infundiranje H/12/01713/002 SI - SANDOZ PHARMACEUTICALS D.D.
Zoledronska kislina Sandoz 4 mg/100 ml raztopina za infundiranje H/12/01713/003 SI - SANDOZ PHARMACEUTICALS D.D.
Zoledronska kislina Sandoz 4 mg/100 ml raztopina za infundiranje H/12/01713/004 SI - SANDOZ PHARMACEUTICALS D.D.
Zoledronsyre Sandoz 4 mg/100 ml

infusjonsvæske, oppløsning 11-8739 NO - SANDOZ A/S
Zoledronsyre Sandoz 48818 DK - SANDOZ A/S
, PRAC Rapporteur: Doris Stenver, “01 September 2015 to 31 August 2016”

EMEA/H/C/PSUSA/00009119/201609
(denosumab (indicated for skeletal related events associated with bone metastases and for giant cell tumour of bone))
CAPS:
XGEVA (EMEA/H/C/002173) (denosumab), MAH: Amgen Europe B.V., Rapporteur: Kristina Dunder, PRAC Rapporteur: Ulla Wändel Liminga, “27 September 2015 - 26 September 2016”

EMEA/H/C/PSUSA/00010074/201609
(bedaquiline)
CAPS:
SIRTURO (EMEA/H/C/002614) (bedaquiline), MAH: Janssen-Cilag International NV, Rapporteur: Filip Josephson, PRAC Rapporteur: Qun-Ying Yue, “06 March 2016 to 05 September 2016”

EMEA/H/C/PSUSA/00010133/201609
(regorafenib)
CAPS:
Stivarga (EMEA/H/C/002573) (regorafenib), MAH: Bayer Pharma AG, Rapporteur: Paula Boudewina van Hennik, PRAC Rapporteur: Sabine Straus, “27-Mar-2015 to 26-Sep-2016.”

EMEA/H/C/PSUSA/00010135/201609
(teriflunomide)
CAPS:
AUBAGIO (EMEA/H/C/002514) (teriflunomide), MAH: sanofi-aventis groupe, Rapporteur: Martina Weise, PRAC Rapporteur: Martin Huber, “13-Mar-2015 - 12-Sep-2016”

EMEA/H/C/PSUSA/00010311/201609
(dulaglutide)
CAPS:
Trulicity (EMEA/H/C/002825) (dulaglutide), MAH: Eli Lilly Nederland B.V., Rapporteur: Greg Markey, PRAC Rapporteur: Carmela Macchiarulo, “Based on the PRAC Rapporteur review of data on safety and efficacy, the PRAC considers that the risk-benefit balance of medicinal products containing dulaglutide remains unchanged but recommends that the terms of the marketing

authorisation should be varied as follows: Update of section 4.8 of the SmPC to add 'hypersensitivity' as adverse reaction with a frequency 'uncommon'. 'Anaphylactic reaction' and 'angioedema' are also being added, with a frequency 'rare'. The Package leaflet is updated accordingly."	
EMEA/H/C/PSUSA/00010366/201609 (naltrexone / bupropion) CAPS: **Mysimba** (EMEA/H/C/003687) (naltrexone hydrochloride / bupropion hydrochloride), MAH: Orexigen Therapeutics Ireland Limited, Rapporteur: Hanne Lomholt Larsen, PRAC Rapporteur: Martin Huber, "27-Mar-2016 to 26-Sep-2016"	
EMEA/H/C/PSUSA/00010403/201609 (pembrolizumab) CAPS: **Keytruda** (EMEA/H/C/003820) (pembrolizumab), MAH: Merck Sharp & Dohme Limited, Rapporteur: Daniela Melchiorri, PRAC Rapporteur: Sabine Straus, "04/03/2016 to 03/09/2016"	
EMEA/H/C/PSUSA/00010413/201609 (guanfacine) CAPS: **Intuniv** (EMEA/H/C/003759) (guanfacine), MAH: Shire Pharmaceuticals Ireland Ltd, Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Eva A. Segovia, "18/03/2016 to 17/09/2016"	

B.4. EPARs / WPARs

Axumin - fluciclovine (18F) - EMEA/H/C/004197 Applicant: Blue Earth Diagnostics Ltd, diagnostic agent for PET of adult men with suspected recurrence of prostate cancer, New active substance (Article 8(3) of Directive No 2001/83/EC)	
Dinutuximab beta Apeiron - dinutuximab beta - EMEA/H/C/003918, Orphan Applicant: APEIRON Biologics AG, treatment of neuroblastoma, New active substance (Article 8(3) of Directive No 2001/83/EC)	

Elmiron - pentosan polysulfate sodium - EMEA/H/C/004246, Orphan Applicant: bene-Arzneimittel GmbH, treatment of Interstitial Cystitis, Well-established use application (Article 10a of Directive No 2001/83/EC)	
Ivabradine Accord - ivabradine - EMEA/H/C/004241 Applicant: Accord Healthcare Ltd, treatment of angina pectoris, Generic, Generic of Procoralan, Generic application (Article 10(1) of Directive No 2001/83/EC)	
Refixia - nonacog beta pegol - EMEA/H/C/004178, Orphan Applicant: Novo Nordisk A/S, treatment of haemophilia B, New active substance (Article 8(3) of Directive No 2001/83/EC)	
Solithromycin Triskel EU Services - solithromycin - EMEA/H/C/004179 Applicant: Triskel EU Services Ltd, treatment of bacterial infections, New active substance (Article 8(3) of Directive No 2001/83/EC) **WPAR**	
Trumenba - meningococcal group b vaccine (recombinant, adsorbed) - EMEA/H/C/004051 Applicant: Pfizer Limited, prevent invasive meningococcal disease caused by Neisseria meningitidis serogroup B, New active substance (Article 8(3) of Directive No 2001/83/EC)	

B.5. TYPE II VARIATION, WORKSHARING PROCEDURE OUTCOMES

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

B.5.1. CHMP assessed procedures scope: Pharmaceutical aspects

Advate - octocog alfa - EMEA/H/C/000520/II/0083/G MAH: Baxter AG, Rapporteur: Jan Mueller-Berghaus	Weekly start timetable.
Aflunov - prepandemic influenza vaccine (H5N1) (surface antigen, inactivated, adjuvanted) - EMEA/H/C/002094/II/0032 MAH: Seqirus S.r.l, Rapporteur: Daniela Melchiorri Request for Supplementary Information adopted	Weekly start timetable. The Committee adopted a Request for Supplementary information together with a specific timetable.

on 06.04.2017.	
Afstyla - lonoctocog alfa - EMEA/H/C/004075/II/0001 MAH: CSL Behring GmbH, Rapporteur: Jan Mueller-Berghaus	Weekly start timetable.
Armisarte - pemetrexed - EMEA/H/C/004109/II/0008/G MAH: Actavis Group PTC ehf, Rapporteur: Alar Irs Request for Supplementary Information adopted on 09.03.2017.	Weekly start timetable.
Azopt - brinzolamide - EMEA/H/C/000267/II/0064/G MAH: Alcon Laboratories (UK) Ltd, Rapporteur: Concepcion Prieto Yerro Request for Supplementary Information adopted on 06.04.2017.	Weekly start timetable. The Committee adopted a Request for Supplementary information together with a specific timetable.
Cerezyme - imiglucerase - EMEA/H/C/000157/II/0099/G MAH: Genzyme Europe BV, Rapporteur: Johann Lodewijk Hillege Opinion adopted on 30.03.2017. Request for Supplementary Information adopted on 16.02.2017.	Positive Opinion adopted by consensus on 30.03.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Cetrotide - cetrorelix acetate - EMEA/H/C/000233/II/0056 MAH: Merck Serono Europe Limited, Rapporteur: Martina Weise Opinion adopted on 06.04.2017.	Positive Opinion adopted by consensus on 06.04.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Cetrotide - cetrorelix acetate - EMEA/H/C/000233/II/0058 MAH: Merck Serono Europe Limited, Rapporteur: Martina Weise	
CIAMBRA - pemetrexed - EMEA/H/C/003788/II/0002/G MAH: Menarini International Operations Luxembourg S.A., Generic, Generic of Alimta, Rapporteur: Juris Pokrotnieks Request for Supplementary Information adopted on 06.04.2017.	Weekly start timetable. The Committee adopted a Request for Supplementary information together with a specific timetable.
Cimzia - certolizumab pegol - EMEA/H/C/001037/II/0058/G MAH: UCB Pharma S.A., Rapporteur: Kristina Dunder Opinion adopted on 06.04.2017. Request for Supplementary Information adopted	Positive Opinion adopted by consensus on 06.04.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

on 23.02.2017.	
Darzalex - daratumumab - EMEA/H/C/004077/II/0005/G, Orphan MAH: Janssen-Cilag International NV, Rapporteur: Sinan B. Sarac Opinion adopted on 06.04.2017. Request for Supplementary Information adopted on 23.02.2017.	Positive Opinion adopted by consensus on 06.04.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
ELOCTA - efmoroctocog alfa - EMEA/H/C/003964/II/0012/G MAH: Swedish Orphan Biovitrum AB (publ), Rapporteur: Jan Mueller-Berghaus	Weekly start 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 06.04.2017.	Weekly start timetable. The Committee adopted a Request for Supplementary information together with a specific timetable.
Galafold - migalastat - EMEA/H/C/004059/II/0005, Orphan MAH: Amicus Therapeutics UK Ltd, Rapporteur: Johann Lodewijk Hillege Opinion adopted on 06.04.2017.	Positive Opinion adopted by consensus on 06.04.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Herceptin - trastuzumab - EMEA/H/C/000278/II/0121 MAH: Roche Registration Limited, Rapporteur: Jan Mueller-Berghaus Request for Supplementary Information adopted on 09.02.2017.	Weekly start timetable.
IDELVION - albutrepenonacog alfa - EMEA/H/C/003955/II/0003/G, Orphan MAH: CSL Behring GmbH, Rapporteur: Jan Mueller-Berghaus Opinion adopted on 06.04.2017. Request for Supplementary Information adopted on 23.02.2017.	Positive Opinion adopted by consensus on 06.04.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Imbruvica - ibrutinib - EMEA/H/C/003791/II/0032/G, Orphan MAH: Janssen-Cilag International NV, Rapporteur: Filip Josephson	Weekly start timetable.
Increlex - mecasermin - EMEA/H/C/000704/II/0046/G, Orphan MAH: Ipsen Pharma, Rapporteur: Outi Mäki-Ikola	Positive Opinion adopted by consensus on 30.03.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

Opinion adopted on 30.03.2017. Request for Supplementary Information adopted on 23.02.2017.	
Inhixa - enoxaparin sodium - EMEA/H/C/004264/II/0005/G MAH: Techdow Europe AB, Duplicate, Duplicate of Thorinane, Rapporteur: Andrea Laslop Opinion adopted on 30.03.2017. Request for Supplementary Information adopted on 23.02.2017.	Positive Opinion adopted by consensus on 30.03.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Inhixa - enoxaparin sodium - EMEA/H/C/004264/II/0009/G MAH: Techdow Europe AB, Duplicate, Duplicate of Thorinane, Rapporteur: Andrea Laslop	Weekly start timetable.
Keytruda - pembrolizumab - EMEA/H/C/003820/II/0026/G MAH: Merck Sharp & Dohme Limited, Rapporteur: Daniela Melchiorri	Weekly start timetable.
Mysimba - naltrexone hydrochloride / bupropion hydrochloride - EMEA/H/C/003687/II/0013/G MAH: Orexigen Therapeutics Ireland Limited, Rapporteur: Hanne Lomholt Larsen Request for Supplementary Information adopted on 06.04.2017.	Weekly start timetable. The Committee adopted a Request for Supplementary information together with a specific timetable.
Onivyde - irinotecan hydrochloride trihydrate - EMEA/H/C/004125/II/0002, Orphan MAH: Baxalta Innovations GmbH, Rapporteur: Filip Josephson Request for Supplementary Information adopted on 23.02.2017.	
OPDIVO - nivolumab - EMEA/H/C/003985/II/0031/G MAH: Bristol-Myers Squibb Pharma EEIG, Rapporteur: Aranzazu Sancho-Lopez Opinion adopted on 30.03.2017.	Positive Opinion adopted by consensus on 30.03.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Orkambi - lumacaftor / ivacaftor - EMEA/H/C/003954/II/0018/G MAH: Vertex Pharmaceuticals (Europe) Ltd., Rapporteur: Nithyanandan Nagercoil Opinion adopted on 30.03.2017.	Positive Opinion adopted by consensus on 30.03.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
RotaTeq - rotavirus vaccine (live, oral) - EMEA/H/C/000669/II/0069/G MAH: MSD Vaccins, Rapporteur: Greg Markey	Weekly start timetable.
Simponi - golimumab -	Weekly start timetable. The Committee

EMEA/H/C/000992/II/0074/G MAH: Janssen Biologics B.V., Rapporteur: Kristina Dunder Request for Supplementary Information adopted on 30.03.2017.	adopted a Request for Supplementary information together with a specific timetable.
TachoSil - human thrombin / human fibrinogen - EMEA/H/C/000505/II/0077/G MAH: Takeda Austria GmbH, Rapporteur: Jan Mueller-Berghaus Request for Supplementary Information adopted on 06.04.2017.	Weekly start timetable. The Committee adopted a Request for Supplementary information together with a specific timetable.
Thyrogen - thyrotropin alfa - EMEA/H/C/000220/II/0090 MAH: Genzyme Europe BV, Rapporteur: Patrick Salmon Request for Supplementary Information adopted on 06.04.2017.	Weekly start timetable. The Committee adopted a Request for Supplementary information together with a specific timetable.
Vectibix - panitumumab - EMEA/H/C/000741/II/0084 MAH: Amgen Europe B.V., Rapporteur: Robert James Hemmings	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. The Committee adopted a Request for Supplementary information together with a specific timetable.
Xofigo - radium-223 - EMEA/H/C/002653/II/0022/G MAH: Bayer AG, Rapporteur: Harald Enzmann Request for Supplementary Information adopted on 19.01.2017.	Weekly start timetable.
Zevalin - ibritumomab tiuxetan - EMEA/H/C/000547/II/0046/G MAH: Spectrum Pharmaceuticals B.V., Rapporteur: Sinan B. Sarac Request for Supplementary Information adopted on 30.03.2017, 19.01.2017.	Weekly start timetable. The Committee adopted a Request for Supplementary information together with a specific timetable.
WS1124 **Fertavid- EMEA/H/C/001042/WS1124/0034** **Puregon- EMEA/H/C/000086/WS1124/0092** MAH: Merck Sharp & Dohme Limited, Lead Rapporteur: Nithyanandan Nagercoil	Weekly start timetable.

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

Adempas - riociguat - EMEA/H/C/002737/II/0018/G, Orphan MAH: Bayer Pharma AG, Rapporteur: Johann Lodewijk Hillege, "C.I.13 Submission of the final clinical study report of study 12166: A multicentre, non-randomized, non-blinded, non-controlled study to investigate the impact of multiple doses of riociguat on safety, tolerability, pharmacokinetics and pharmacodynamics in patients with pulmonary hypertension in a 12 week 3 times a day individual dose titration scheme. C.I.13 Submission of the final clinical study report of study 16097: An open-label phase IIIb study of riociguat in patients with in-operable chronic thromboembolic pulmonary hypertension (CTEPH) or recurrent or persisting pulmonary hypertension after surgical treatment who are not satisfactorily treated and cannot participate in any other CTEPH trial." Opinion adopted on 30.03.2017. Request for Supplementary Information adopted on 26.01.2017.	Positive Opinion adopted by consensus on 30.03.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Cimzia - certolizumab pegol - EMEA/H/C/001037/II/0057/G MAH: UCB Pharma S.A., Rapporteur: Kristina Dunder - B.II.e.6.a (Type IB) - to introduce an additional presentation which combines a Pre-filled Syringe (PFS) within a single-use, needle-safe Dose-dispenser Cartridge (DDC) (functional secondary packaging), together with one new pack size, - B.II.e.5.a).1 (Type IAin) - two additional pack sizes, - C.I.4 (Type II) - amend the Product Information (PI) to add the Dose-dispenser Cartridge presentations." Request for Supplementary Information adopted on 23.02.2017.	
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	Weekly start timetable.

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."	
Galafold - migalastat - EMEA/H/C/004059/II/0009, Orphan MAH: Amicus Therapeutics UK Ltd, Rapporteur: Johann Lodewijk Hillege, "Update of section 5.1 of the SmPC to add new mutations in Table 2: Galafold (migalastat) amenability table and to Table 3: Mutations not amenable to Galafold (migalastat). In addition, the Marketing authorisation holder (MAH) took the opportunity to introduce some minor editorial changes to the tables and to update the list of local representatives in the Package Leaflet."	
Glivec - imatinib - EMEA/H/C/000406/II/0108 MAH: Novartis Europharm Ltd, Rapporteur: Aranzazu Sancho-Lopez, "Submission of the final CSR for study STI571A2405 - the International Study for Chronic Myeloid Leukemia (CML) in childhood and adolescents (I-CML-Ped Study). The provision of the study report addresses the post-authorisation measure MEA 162.8."	Weekly start timetable.
GONAL-f - follitropin alfa - EMEA/H/C/000071/II/0136 MAH: Merck Serono Europe Limited, Rapporteur: Nithyanandan Nagercoil, "Update of the SmPC section 4.8 to indicate that thromboembolism can occur both in association with and separate from ovarian hyperstimulation syndrome (OHSS). 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 10.0." Opinion adopted on 30.03.2017. Request for Supplementary Information adopted on 02.02.2017.	Positive Opinion adopted by consensus on 30.03.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Harvoni - ledipasvir / sofosbuvir - EMEA/H/C/003850/II/0035 MAH: Gilead Sciences International Ltd,	

Rapporteur: Filip Josephson, “Update of sections 4.8, 5.1 and 5.2 of the SmPC in order to add emerging clinical data available from studies SOLAR-1 and SOLAR-2.” Request for Supplementary Information adopted on 23.02.2017, 10.11.2016.	
Hetlioz - tasimelteon - EMEA/H/C/003870/II/0008, Orphan MAH: Vanda Pharmaceuticals Ltd., Rapporteur: Greg Markey, “Update of section 4.5 of the SmPC with the deletion of the CYP2C19 statement and the removal of the commitment to conduct a human CYP2C19 Drug-Drug Interaction Study to evaluate the single-dose pharmacokinetics of tasimelteon 20 mg alone and in combination with a CYP2C19 inhibitor, omeprazole, at steady-state from the Risk Management Plan (RMP).” Request for Supplementary Information adopted on 06.04.2017.	Weekly start timetable. The Committee adopted a Request for Supplementary information together with a specific timetable.
IDELVION - albutrepenonacog alfa - EMEA/H/C/003955/II/0005, Orphan MAH: CSL Behring GmbH, Rapporteur: Jan Mueller-Berghaus, “Update of section 4.8 of the SmPC in order to update the safety information by removing a description of a low titer inhibitor based on information from ongoing study CSL654-3003. The Package Leaflet is updated accordingly.”	Weekly start timetable.
Increlex - mecasermin - EMEA/H/C/000704/II/0040, Orphan MAH: Ipsen Pharma, Rapporteur: Outi Mäki-Ikola, “Update of section of 4.1 of the SmPC in order to re-word the recommendation to confirm diagnosis with an IGF-1 generation test used for diagnosis of Severe Primary IGFD” Request for Supplementary Information adopted on 26.01.2017.	
Incruse - umeclidinium bromide - EMEA/H/C/002809/II/0013 MAH: Glaxo Group Ltd, Rapporteur: Concepcion Prieto Yerro, “Update of section 4.8 of the SmPC and relevant section of the PL to add hypersensitivity reactions including rash, urticaria, pruritus as uncommon and anaphylaxis and angioedema as rare adverse reactions. The MAH is taking the opportunity to update the Local representative section in the PL.”	Weekly start timetable.

Request for Supplementary Information adopted on 15.12.2016.	
Lenvima - lenvatinib - EMEA/H/C/003727/II/0008, Orphan MAH: Eisai Europe Ltd., Rapporteur: Bart Van der Schueren, "Submission of Clinical Study Report for Study E78080-J081-208" Request for Supplementary Information adopted on 06.04.2017.	Weekly start timetable. The Committee adopted a Request for Supplementary information together with a specific timetable.
Mimpara - cinacalcet - EMEA/H/C/000570/II/0056 MAH: Amgen Europe B.V., Rapporteur: Kristina Dunder, "Submission of the final report for Study 20090686, a study designed to determine the efficacy of cinacalcet compared with vitamin D therapy for management of secondary HPT (hyperparathyroidism) in haemodialysis subjects. This variation fulfils LEG 031." Request for Supplementary Information adopted on 23.02.2017.	Weekly start 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 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	Weekly start timetable.

Subjects. This study does not lead to changes in the product information."	
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."	Weekly start timetable.
Noxafil - posaconazole - EMEA/H/C/000610/II/0048 MAH: Merck Sharp & Dohme Limited, Rapporteur: Greg Markey, "Update of sections 4.4 and 4.5 of the SmPC in order to strengthen the current warning on interaction of posaconazole with vincristine. 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." Request for Supplementary Information adopted on 23.02.2017.	
SIRTURO - bedaquiline - EMEA/H/C/002614/II/0021, Orphan MAH: Janssen-Cilag International NV, Rapporteur: Filip Josephson, "Update of section 4.4 of the SmPC in order to add delamanid as an example of a drug that prolongs the QT interval following the review of the global safety database for all serious cases received from 28 December 2012 to 30 September 2016. The Package Leaflet is updated accordingly. In addition, the Marketing authorisation holder (MAH) took the opportunity to update the list of local representatives in the Package Leaflet." Request for Supplementary Information adopted on 06.04.2017.	Weekly start timetable. The Committee adopted a Request for Supplementary information together with a specific timetable.
Sutent - sunitinib - EMEA/H/C/000687/II/0064 MAH: Pfizer Limited, Rapporteur: Daniela Melchiorri, "Update of section 4.1 of the SmPC in order to remove statement 'Experience with	

SUTENT as first-line treatment is limited (see section 5.1)' based on the final CSR of study A6181202 in fulfilment of MEA 037.2." Request for Supplementary Information adopted on 26.01.2017.	
Travatan - travoprost - EMEA/H/C/000390/II/0053 MAH: Alcon Laboratories (UK) Ltd, Rapporteur: Concepcion Prieto Yerro, "Following the submission of final CSR for study C-01-79 and a review of supporting clinical studies and post-marketing data, update to SmPC section 4.8 is proposed. The package leaflet is updated accordingly. In addition, MAH took the opportunity to update number of the Spanish representative in the PL." Request for Supplementary Information adopted on 19.01.2017.	Weekly start timetable.
Victrelis - boceprevir - EMEA/H/C/002332/II/0041 MAH: Merck Sharp & Dohme Limited, Rapporteur: Joseph Emmerich, "Update of sections 4.3 and 4.5 of the SmPC in order to add a new contraindication for the interaction of lurasidone following data obtained from the MAH continuous safety monitoring. The Package Leaflet is updated accordingly. In addition, the MAH took the opportunity to implement QRD template version 10, including implementation of the 2D barcode in the PI."	
Viread - tenofovir disoproxil - EMEA/H/C/000419/II/0173 MAH: Gilead Sciences International Ltd, Rapporteur: Joseph Emmerich, PRAC Rapporteur: Caroline Laborde, "Submission of final long-term safety and efficacy data (480 weeks) from two completed Phase 3 studies in HBeAg-negative and HBeAg-positive patients with chronic hepatitis B (CHB), Studies GS-US-174-0102 and GS-US-174-0103." Opinion adopted on 30.03.2017. Request for Supplementary Information adopted on 26.01.2017.	Positive Opinion adopted by consensus on 30.03.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Xagrid - anagrelide - EMEA/H/C/000480/II/0075 MAH: Shire Pharmaceutical Contracts Ltd., Rapporteur: Alexandre Moreau, "Update of sections 4.4, 4.5 and 5.1 of the SmPC in order	Weekly start timetable.

to change the terminology of myeloproliferative disorders to neoplasms, add text regarding platelet count rebound above baseline following dosage interruption, incorporate a section in drug interactions on Cyp 1A2 inducers and update information on the mode of action. The Package Leaflet is updated accordingly. In addition, the MAH took the opportunity to update the list of local representatives in the Package Leaflet, correct typographical errors and bring the PI in line with the latest QRD template. No changes were proposed to the RMP." Request for Supplementary Information adopted on 26.01.2017.	
Xarelto - rivaroxaban - EMEA/H/C/000944/II/0050 MAH: Bayer Pharma 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."	
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.	Weekly start timetable.

In addition, the Marketing authorisation holder (MAH) took the opportunity to introduce editorial changes in the SmPC."	
Zepatier - elbasvir / grazoprevir - EMEA/H/C/004126/II/0005 MAH: Merck Sharp & Dohme Limited, Rapporteur: Greg Markey, "Update of section 4.5 of the SmPC in order to update information regarding drug-drug interaction (DDI) of elbasvir/grazoprevir when co-administrated with sunitinib (tyrosine kinase inhibitor). The Package Leaflet is updated accordingly. In addition, the Marketing authorisation holder (MAH) took the opportunity to include some editorial changes." Opinion adopted on 06.04.2017. Request for Supplementary Information adopted on 23.02.2017.	Positive Opinion adopted by consensus on 06.04.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1055 **Ebymect-EMEA/H/C/004162/WS1055/0016** **Edistride-EMEA/H/C/004161/WS1055/0012** **Forxiga-EMEA/H/C/002322/WS1055/0031** **Qtern-EMEA/H/C/004057/WS1055/0004** **Xigduo-EMEA/H/C/002672/WS1055/0027** MAH: AstraZeneca AB, Lead Rapporteur: Kristina Dunder, "Update of section 4.8 of the SmPC in order to update the safety information related to rash. The Package Leaflet is updated accordingly. Additional editorial changes were made in sections 5.1, 5.2 of the SmPC to Qtern." Request for Supplementary Information adopted on 19.01.2017.	Weekly start timetable.
WS1070 **Bretaris Genuair-EMEA/H/C/002706/WS1070/0032** **Eklira Genuair-EMEA/H/C/002211/WS1070/0032** MAH: AstraZeneca AB, Lead Rapporteur: Nithyanandan Nagercoil, "Update of section 4.3 of the SmPC in order to modify the contraindication section deleting reference to hypersensitivity to atropine or its derivative providing justification for the claim that the chemical structure of aclidinium is unrelated to that of atropine or its derivatives. The Package	

Leaflet is updated accordingly." Request for Supplementary Information adopted on 23.02.2017.	
WS1072 **Eucreas-** **EMEA/H/C/000807/WS1072/0060** **Galvus-EMEA/H/C/000771/WS1072/0051** **Icandra-** **EMEA/H/C/001050/WS1072/0061** **Jalra-EMEA/H/C/001048/WS1072/0051** **Xiliarx-EMEA/H/C/001051/WS1072/0050** **Zomarist-** **EMEA/H/C/001049/WS1072/0061** MAH: Novartis Europharm Ltd, Lead Rapporteur: Kristina Dunder, "Update of section 5.1 of the SmPC, subsection 'cardiovascular risk', with results from a new meta-analysis evaluating the cardiovascular safety of vildagliptin. In addition, the Worksharing applicant (WSA) took the opportunity to bring the annexes in line with the latest QRD template version 10, and to merge the two SmPCs into one single SmPC for Eucreas, Icandra and Zomarist."	Weekly start timetable.
WS1077/G **Aluvia-** **EMEA/H/W/000764/WS1077/0101/G** **Kaletra-** **EMEA/H/C/000368/WS1077/0163/G** **Norvir-** **EMEA/H/C/000127/WS1077/0143/G** MAH: AbbVie Ltd., Lead Rapporteur: Joseph Emmerich, "Update of sections 4.3 and 4.5 of the SmPC in order to add information regarding the interaction of lopinavir/ritonavir and ritonavir with lurasidone and ranolazine. In addition, sections 4.4 and 4.5 of the SmPC are updated to add information regarding the interaction with injectable triamcinolone. The Labelling is updated accordingly." Request for Supplementary Information adopted on 23.02.2017.	
WS1105 **IntronA-** **EMEA/H/C/000281/WS1105/0107** **PegIntron-** **EMEA/H/C/000280/WS1105/0128** **ViraferonPeg-** **EMEA/H/C/000329/WS1105/0121**	Weekly start timetable.

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 23.02.2017.	
WS1110 **Kinzalkomb-** **EMEA/H/C/000415/WS1110/0100** **MicardisPlus-** **EMEA/H/C/000413/WS1110/0102** **PritorPlus-** **EMEA/H/C/000414/WS1110/0110** MAH: Boehringer Ingelheim International GmbH, Lead Rapporteur: Daniela Melchiorri, “Update of sections 4.5 and 4.8 to align the hydrochlorothiazide component information with that of the originator. The Package Leaflet is updated accordingly. In addition, WorkSharing Applicant (WSA) took the opportunity of this procedure to bring the Product Information in line with the latest QRD version 9.1 and 10.0 templates, including combining the SmPC of the different strengths, as well as implement minor editorial changes and reformatting of some sections of the SmPC. The details of local representative (Portugal for MicardisPlus and United Kingdom for PritorPlus and Kinzalkomb) in the PL have been updated.” Opinion adopted on 06.04.2017. Request for Supplementary Information adopted on 16.02.2017.	Positive Opinion adopted by consensus on 06.04.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1148/G **Hexacima-** **EMEA/H/C/002702/WS1148/0059/G** **Hexaxim-** **EMEA/H/W/002495/WS1148/0065/G** **Hexyon-**	Weekly start timetable.

EMEA/H/C/002796/WS1148/0063/G MAH: Sanofi Pasteur Europe, Duplicate, Duplicate of Hexacima, Lead Rapporteur: Jan Mueller-Berghaus, "C.I.4 - Update of section 5.1 of the SmPC in order to include data on persistence of immunity following final results from studies: A3L47: Laboratory analysis on sera stored at Sanofi Pasteur Global Clinical Immunology laboratory and collected within the context of trial PNA19 and A3L49: Phase III, multi-center study in children vaccinated with Hep B vaccine at birth followed by three infant primary series doses of Hexaxim® or Infanrix® hexa in A3LI2 study in Thailand C.I.3.z – To include the information about sequential schedule of hexavalent and pentavalent vaccines in primary series in section 4.2 of SmPC following the assessment of A3L39 study."	

B.5.3. CHMP-PRAC assessed procedures

Champix - varenicline - EMEA/H/C/000699/II/0064 MAH: Pfizer Limited, Rapporteur: Hanne Lomholt Larsen, PRAC Rapporteur: Doris Stenver, "Update of sections 4.5 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." Request for Supplementary Information adopted on 06.04.2017.	Weekly start timetable.
Edurant - rilpivirine - EMEA/H/C/002264/II/0024 MAH: Janssen-Cilag International NV, Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Sabine Straus, "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	Weekly start timetable.

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 06.04.2017.

Emtricitabine/Tenofovir disoproxil Mylan - emtricitabine / tenofovir disoproxil - EMEA/H/C/004050/II/0001

MAH: Mylan S.A.S, Generic, Generic of Truvada, Rapporteur: Romaldas Mačiulaitis, PRAC Rapporteur: Patrick Batty, "Update of the SmPC following the assessment of the extension of indication for the reference product, Truvada, for pre-exposure prophylaxis. The Package Leaflet, Annex II and Labelling are updated in accordance."
Request for Supplementary Information adopted on 23.03.2017.

Erivedge - vismodegib - EMEA/H/C/002602/II/0032

MAH: Roche Registration Limited, Rapporteur: Kristina Dunder, PRAC Rapporteur: Ulla Wändel Liminga, "Update of section 5.3 of the SmPC in order to reflect non-clinical carcinogenicity studies (MEA 003):
- Study 13-0322 is a 26-Week Oral Gavage Carcinogenicity Study with Vismodegib in Hemizygous CByB6F1-Tg(HRAS)2Jic Mice.
- Study 13-0323 is a 104-Week and 52-Week with a 12-Week Recovery Phase Oral Gavage Carcinogenicity Study with Vismodegib in Sprague Dawley Rats.
The RMP (RMP 12.0) has been consequently updated. Furthermore, additional routine changes (including some resulting from the assessment of RMP version 11) have been introduced."
Request for Supplementary Information adopted

on 23.02.2017.

Flixabi - infliximab -
EMEA/H/C/004020/II/0009
MAH: Samsung Bioepis UK Limited (SBUK),
Rapporteur: Jan Mueller-Berghaus, PRAC
Rapporteur: Ulla Wändel Liminga, "Submission of the final study report of study SB2-G31-RA: A Randomised, Double-blind, Parallel Group, Multicentre Clinical Study to Evaluate the Efficacy, Safety, Pharmacokinetics and Immunogenicity of SB2 Compared to Remicade® in Subjects with Moderate to Severe Rheumatoid Arthritis despite Methotrexate Therapy.
The RMP (v. 4) has been updated to reflects the results from the 78 weeks CSR, to exclude 2 of the 5 registries of the pharmacovigilance plan and update in the due date for the prospective observational cohort study of Flixabi in AS (Ankylosing Spondylitis) and CD (Crohn's Disease) patients."
Request for Supplementary Information adopted on 15.12.2016.

Ganfort - bimatoprost / timolol -
EMEA/H/C/000668/II/0026
MAH: Allergan Pharmaceuticals Ireland,
Rapporteur: Hanne Lomholt Larsen, PRAC
Rapporteur: Torbjorn Callreus, "Update of section 4.8 as per the PRAC recommendation following the PSUSA assessment. The Package Leaflet has been updated accordingly.
In addition, the MAH took the opportunity to update the Product Information in line with the QRD template version 10.0, implement the unique identifier – 2D bar code and correct typographical errors.
As per the PRAC recommendation, the updated RMP version 3.2 is also agreed."
Opinion adopted on 06.04.2017.

Keytruda - pembrolizumab -
EMEA/H/C/003820/II/0018/G
MAH: Merck Sharp & Dohme Limited,
Rapporteur: Daniela Melchiorri, PRAC
Rapporteur: Sabine Straus, "Update of section 5.1 of the SmPC to reflect the data from the post-authorisation efficacy studies (PAES) in melanoma; studies P001, P002 and P006.
Annex II has been revised to reflect that these three final CSRs have been submitted.

An updated RMP version 6.0 was provided as part of the application. The following summarizes the changes to the updated RMP:

• □The fin

melanoma studies P001/002/006 and removed as PAES commitments from the RMP;

• □Me

validation report for anti-MK-3475 neutralizing antibody assay were included as Completed Pharmacovigilance Activities;

• □The M

term safety' as missing information in the list of ongoing safety concerns."

Request for Supplementary Information adopted on 26.01.2017.

Keytruda - pembrolizumab - EMEA/H/C/003820/II/0025

MAH: Merck Sharp & Dohme Limited, Rapporteur: Daniela Melchiorri, PRAC Rapporteur: Sabine Straus, "Update of sections 4.2, 4.4 and 4.8 of the SmPC to add a warning for the risk of severe skin reactions and to communicate that Stevens - Johnson syndrome (SJS) and toxic epidermal necrolysis (TEN), including fatal cases, have been reported in patients treated with pembrolizumab. The Package Leaflet has been updated accordingly. The application included an updated RMP version 8.0, and a proposed DHPC and communication plan."

Lemtrada - alemtuzumab - EMEA/H/C/003718/II/0017

MAH: Genzyme Therapeutics Ltd, Duplicate, Duplicate of Lemtrada (WD), Rapporteur: Hanne Lomholt Larsen, PRAC Rapporteur: Torbjorn Callreus, "Update of sections 4.2, 4.4, 4.8 and 5.1 of the SmPC in order to update the safety and long term use information in the posology following final results from study CAMMS03409 - An Extension Protocol For Multiple Sclerosis Patients Who Participated in Genzyme-Sponsored Studies of Alemtuzumab (ongoing at the time of the initial MAA) to evaluate the long term safety and efficacy of alemtuzumab in MS patients who received alemtuzumab during prior company-sponsored studies. The RMP version 3.0 has also been submitted. The PL has been updated accordingly.

In addition, the Marketing authorisation holder (MAH) took the opportunity to bring the PI in line with the latest QRD template version 10.0 and to introduce editorial corrections in the PI."

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 23.02.2017.

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)."

OPDIVO - nivolumab - EMEA/H/C/003985/II/0024

MAH: Bristol-Myers Squibb Pharma EEIG, Rapporteur: Aranzazu Sancho-Lopez, PRAC

Rapporteur: Brigitte Keller-Stanislawski,
“Update of section 5.1 of the SmPC in order to reflect the final overall survival and response data, including duration of response with longer follow-up, following completion of PAES CA209037 (Randomized, Open-Label, Phase 3 Trial of nivolumab vs Investigator's Choice in Advanced (Unresectable or Metastatic) Melanoma Patients Progressing Post Anti-CTLA-4 Therapy) and its addendum on predictability of efficacy with biomarkers.
This application fulfils ANX 001 and 003.1.
Annex II has been updated accordingly.
RMP version 5.5 has been submitted within this application.”
Request for Supplementary Information adopted on 23.03.2017, 26.01.2017.

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) follwoing 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 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 15.12.2016.

Zavesca - miglustat -
EMEA/H/C/000435/II/0056, Orphan
MAH: Actelion Registration Ltd., Rapporteur: Kristina Dunder, PRAC Rapporteur: Qun-Ying Yue, “Submission of 8th NPC (Niemann-Pick type C) Registry report and update of Annex II-D to delete the NPC Registry listed as an obligation to the marketing authorisation.

The RMP version 12.1 has also been submitted to reflect the above changes.
In addition, the Marketing authorisation holder took the opportunity to introduce minor changes and bring the Product Information and Annex A in line with the latest QRD template version 10.”

WS0992/G
Relvar Ellipta-
EMEA/H/C/002673/WS0992/0022/G
Revinty Ellipta-
EMEA/H/C/002745/WS0992/0017/G
MAH: Glaxo Group Ltd, Lead Rapporteur: Concepcion Prieto Yerro, Lead PRAC Rapporteur: Dolores Montero Corominas, “Type II C.I.4:- Update of sections 4.4, 4.8 and 5.1 of the SmPC in order to update the safety information and include data from the HZC113782 (SUMMIT) study (designed to investigate whether FF/VI-Furoate/Vilanterol could improve survival in patients with moderate COPD- chronic obstructive pulmonary disease who had, or were at increased risk for CV-cardiovascular disease). The Package Leaflet and Labelling are updated accordingly. The RMP v.8.1 is updated accordingly.

Type II C.I.4:- Update of section 4.8 of the SmPC in order to add "paradoxical

bronchospasm" to the list of adverse reactions. The Package Leaflet and Labelling are updated accordingly.

Type IB C.I.z:- Update of section 5.1 the SmPC in order to correct an error identified in the pharmacodynamic section."
Request for Supplementary Information adopted on 23.02.2017, 13.10.2016.

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 15.12.2016.

WS1101
Relvar Ellipta-
EMEA/H/C/002673/WS1101/0029
Revinty Ellipta-
EMEA/H/C/002745/WS1101/0025
MAH: Glaxo Group Ltd, Lead Rapporteur: Concepcion Prieto Yerro, Lead PRAC Rapporteur: Dolores Montero Corominas, "Update of section 5.1 of the SmPC in order to update the safety information in relation to results of HZC115151 study (A 12-month, open label, randomised, effectiveness study to evaluate fluticasone furoate (FF, GW685698)/vilanterol (VI, GW642444) Inhalation Powder delivered once daily via a Novel Dry Powder Inhaler (NDPI) compared with the existing COPD maintenance therapy alone in subjects with Chronic Obstructive Pulmonary Disease (COPD)(an Annex II condition) of the Relvar Ellipta and

Revinty Ellipta (92/22mcg strength only). Consequently the RMP version 8.3 is updated." Request for Supplementary Information adopted on 23.02.2017.	
WS1117/G **Stocrin-** **EMEA/H/C/000250/WS1117/0110/G** **Sustiva-** **EMEA/H/C/000249/WS1117/0139/G** MAH: Bristol-Myers Squibb Pharma EEIG, Lead Rapporteur: Bruno Sepodes, Lead PRAC Rapporteur: Ana Sofia Diniz Martins, "C.I.4 (Type II) - Update of sections 4.4, 4.5 and 5.1 of the SmPC in order to add a warning and update the safety information on QTc prolongation based on the final results from study AI266959; this is an interventional study to determine the concentration-electrocardiographic effects of efavirenz in healthy subjects enriched for cyp2b6 polymorphisms; the Package Leaflet is updated accordingly. The RMP version 8 has also been submitted. C.I.4 (Type II) – Update of sections 4.4 and 4.8 to add catatonia as a Psychiatric symptom following an assessment of catatonia cases reported in the literature and via the United States (US) Food and Drug Administration Adverse Event Reporting System (FAERS)." Request for Supplementary Information adopted on 06.04.2017.	Weekly start timetable.
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 version 7 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.	Weekly start timetable.

C.I.3.b: Update of sections 4.4 of the SmPC in order to add Bullous pemphigoid as a warning following the PRAC assessment outcome EMEA/H/C/PSUSA/2711/201408; the Labelling is being updated accordingly." Request for Supplementary Information adopted on 06.04.2017.	
WS1133/G **Atripla-** **EMEA/H/C/000797/WS1133/0121/G** **Descovy-** **EMEA/H/C/004094/WS1133/0015/G** **Eviplera-** **EMEA/H/C/002312/WS1133/0081/G** **Genvoya-** **EMEA/H/C/004042/WS1133/0029/G** **Odefsey-** **EMEA/H/C/004156/WS1133/0011/G** **Stribild-** **EMEA/H/C/002574/WS1133/0080/G** **Truvada-** **EMEA/H/C/000594/WS1133/0136/G** **Viread-** **EMEA/H/C/000419/WS1133/0174/G** MAH: Gilead Sciences International Ltd, Lead Rapporteur: Robert James Hemmings, Lead PRAC Rapporteur: Amelia Cupelli, "The group of Workshare variations includes: Updates of sections 4.4 and 4.5 of the SmPC for the tenofovir disoproxil fumarate (TDF)-containing products (Viread, Truvada, Atripla, Eviplera, Stribild) which includes the results from Study GS-US-342-1167 and Study GS-US-342-1326. Update of section 4.5 for the tenofovir alafenamide (TAF)-containing products (Genvoya, Descovy, Odefsey) which include the results from Study GS-US-342-1167. Study GS-US-342-1167 is a Phase I Study to Evaluate the Pharmacokinetic Drug-Drug Interactions between Sofosbuvir/GS-5815 Fixed Dose Combination (FDC) Tablets and Antiretrovirals Efavirenz/Emtricitabine/Tenofovir Disoproxil Fumarate (EFV/FTC/TDF; Atripla), Emtricitabine/Riplivirine/Tenofovir Disoproxil Fumarate (FTC/RPV/TDF; Complera), Dolutegravir (DTG; Tivicay) o	

Elvitegravir/Cobicistat/Emtricitabine/Tenofovir Alafenamide Fumarate (EVG/COBI/FTC/TAF) in Healthy Subjects. Study GS-US-342-1326, a Phase I Study to Evaluate the Pharmacokinetic Drug-Drug Interaction Potential between Sofosbuvir/GS-5816 (SOF/GS-5816) Fixed-Dose Combination (FDC) Tablet and HIV Antiretroviral Regimens Elvitegravir/Cobicistat/Emtricitabine/Tenofovir Disoproxil Fumarate (EVG/COBI/FTC/TDF), Ritonavir-boosted Darunavir (DRV/r) plus Emtricitabine/Tenofovir Disoproxil Fumarate (FTC/TDF), Ritonavir-boosted Atazanavir (ATV/r) plus FTC/TDF, Ritonavir/boosted Lopinavir (LPV/r) plus FTC/TDF or Raltegravir plus FTC/TDF. The Package Leaflet and Risk Managent Plan (RMP) are updated accordingly.”	
WS1134 **Truvada-EMEA/H/C/000594/WS1134/0137** **Viread-EMEA/H/C/000419/WS1134/0175** MAH: Gilead Sciences International Ltd, Lead Rapporteur: Joseph Emmerich, Lead PRAC Rapporteur: Caroline Laborde, “Update of section 4.5 of the SmPC for Viread and Truvada with interactions between emtricitabine (FTC), tenofovir disoproxil fumarate (TDF), ledipasvir, sofosbuvir and dolutegavir based on new clinical pharmacology data from study GS-US-377-1501. This is a Phase 1, open-label, multiple-dose study that evaluated the pharmacokinetic drug-drug interaction potential between Harvoni (ledipasvir [LDV]/sofosbuvir [SOF]) and FTC/TDF+dolutegravir (DTG). The RMP version 22 for Viread and version 14 for Truvada have also been submitted.”	
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	Weekly start timetable.

PRAC Rapporteur: Menno van der Elst, “Update of sections 4.4 of the SmPC in order to add Bullous pemphigoid as a warning following the PRAC assessment outcome EMEA/H/C/PSUSA/2711/201408; the Labelling is being updated accordingly. Consequently, the RMP version 7 is updated accordingly.”
Request for Supplementary Information adopted on 06.04.2017.

B.5.4. PRAC assessed procedures

PRAC Led
Bydureon - exenatide - EMEA/H/C/002020/II/0042
MAH: AstraZeneca AB, Rapporteur: Kristina Dunder, PRAC Rapporteur: Qun-Ying Yue, , “Submission of the updated RMP version 25 following closure and final summary of Exenatide Pregnancy Registry (a prospective, observational study conducted in the United States that actively collected information on exposure to antidiabetic medication during pregnancy and the associated pregnancy outcomes in patients with Type 2 diabetes mellitus). Moreover, the MAH included additional minor updates to the RMP.”
Request for Supplementary Information adopted on 23.02.2017.

PRAC Led
Halaven - eribulin - EMEA/H/C/002084/II/0033
MAH: Eisai Europe Ltd., Rapporteur: Filip Josephson, PRAC Rapporteur: Ulla Wändel Liminga, , “Update of the RMP version 4.2 following the revision of the protocol for a post-authorisation study to capture data on the frequency of resolution and time to resolution of eribulin-induced or aggravated peripheral neuropathy from a phase 3 study, E7389-A001-303 (ACCRU) to an observational study, E7389-M044-504 (IRENE). The submission of the corresponding study report to the EMA / PRAC remains unchanged and is planned during 2019.”
Request for Supplementary Information adopted on 23.02.2017, 15.12.2016.

PRAC Led
Respreeza - human alpha1-proteinase

inhibitor - EMEA/H/C/002739/II/0013 MAH: CSL Behring GmbH, PRAC Rapporteur: Eva A. Segovia, , “Submission of an updated RMP version 3.1 in order to include the final safety data from study CE1226_3001, which were assessed in the type II variation EMEA/H/C/002739/II/0002 and adjustments in the Non-Clinical Safety specification part (Part II, Module SII) with non-clinical information from local tolerability trials.” Opinion adopted on 06.04.2017.	
PRAC Led **Tysabri - natalizumab - EMEA/H/C/000603/II/0101** MAH: Biogen Idec Ltd, Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Brigitte Keller-Stanislawski, , “Submission of the final clinical study report for TYGRIS, a post-marketing safety observational cohort program designed to obtain long-term safety data (approximately 5 years) in subjects with MS treated with natalizumab, and comprising parallel studies 101MS402 (United States and Canada) and 101MS403 (Rest of World). The application included an updated RMP version 23.” Request for Supplementary Information adopted on 06.04.2017.	Weekly start timetable.
PRAC Led **Tysabri - natalizumab - EMEA/H/C/000603/II/0102** MAH: Biogen Idec Ltd, Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Brigitte Keller-Stanislawski, , “Submission of the final clinical study report for study STRATIFY-2 (101JC402), an observational, longitudinal cohort study designed to gather post-marketing data on the incidence of progressive multifocal leukoencephalopathy (PML) in natalizumab-treated subjects with MS. An updated RMP version 23 was provided accordingly.” Opinion adopted on 06.04.2017.	
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, , “Submission of the final clinical study report for study B1531005, a non-interventional study to evaluate the outcomes	Weekly start timetable.

(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."
Request for Supplementary Information adopted on 06.04.2017.

PRAC Led
WS0943
Saxenda-EMEA/H/C/003780/WS0943/0009
Victoza-EMEA/H/C/001026/WS0943/0041
MAH: Novo Nordisk A/S, Lead Rapporteur: Johann Lodewijk Hillege, Lead PRAC Rapporteur: Menno van der Elst, , "Submission of the final results from the main "Liraglutide safety and surveillance program using the Optum Research Database" study and sub-study on breast cancer - RMP category 3 study."
Request for Supplementary Information adopted on 26.01.2017, 15.09.2016.

PRAC Led
WS1088
Eucreas-EMEA/H/C/000807/WS1088/0057
Galvus-EMEA/H/C/000771/WS1088/0048
Icandra-EMEA/H/C/001050/WS1088/0058
Jalra-EMEA/H/C/001048/WS1088/0048
Xiliarx-EMEA/H/C/001051/WS1088/0047
Zomarist-EMEA/H/C/001049/WS1088/0058
MAH: Novartis Europharm Ltd, Lead PRAC Rapporteur: Qun-Ying Yue, "Following the outcome of an Article 31 referral procedure for metformin and metformin-containing products (Procedure EMEA/H/A-31/1432), the Applicant was requested toupdate the Risk Management Plan (RMP) for galvus, Jalra, Xiliarx, Eucreas, Icandra and Zomarist to implement a targeted questionnaire for cases of lactic acidosis."
Request for Supplementary Information adopted on 23.02.2017.

B.5.5. CHMP-CAT assessed procedures

Imlygic - talimogene laherparepvec - EMEA/H/C/002771/II/0008, ATMP MAH: Amgen Europe B.V., Rapporteur: Olli Tenhunen, CHMP Coordinators: Tuomo Lapveteläinen Request for Supplementary Information adopted on 17.02.2017.	

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

WS0921 **Ebymect-EMEA/H/C/004162/WS0921/0019** **Edistride-EMEA/H/C/004161/WS0921/0015** **Forxiga-EMEA/H/C/002322/WS0921/0034** **Qtern-EMEA/H/C/004057/WS0921/0005** **Xigduo-EMEA/H/C/002672/WS0921/0030** MAH: AstraZeneca AB, Lead Rapporteur: Kristina Dunder Opinion adopted on 06.04.2017.	Positive Opinion adopted by consensus on 06.04.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1030 **ANORO-EMEA/H/C/002751/WS1030/0015** **Incruse-EMEA/H/C/002809/WS1030/0014** **Laventair-EMEA/H/C/003754/WS1030/0017** **Relvar Ellipta-EMEA/H/C/002673/WS1030/0028** **Revinty Ellipta-EMEA/H/C/002745/WS1030/0024** MAH: Glaxo Group Ltd, Lead Rapporteur: Nithyanandan Nagercoil Request for Supplementary Information adopted on 23.02.2017.	Weekly start timetable.
WS1081 **Hexacima-EMEA/H/C/002702/WS1081/0055** **Hexaxim-EMEA/H/W/002495/WS1081/0062**	Weekly start timetable.

Hexyon- **EMEA/H/C/002796/WS1081/0059** MAH: Sanofi Pasteur SA, Lead Rapporteur: Jan Mueller-Berghaus Request for Supplementary Information adopted on 16.02.2017.	
WS1085 **Ceprotin-** **EMEA/H/C/000334/WS1085/0098** **HyQvia-EMEA/H/C/002491/WS1085/0034** **Kiovig-EMEA/H/C/000628/WS1085/0076** MAH: Baxalta Innovations GmbH, Lead Rapporteur: Jan Mueller-Berghaus Request for Supplementary Information adopted on 23.02.2017.	Weekly start timetable.
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 Request for Supplementary Information adopted on 30.03.2017.	Weekly start timetable. The Committee adopted a Request for Supplementary information together with a specific timetable.
WS1126 **Gardasil-** **EMEA/H/C/000703/WS1126/0070** **Silgard-EMEA/H/C/000732/WS1126/0061** MAH: MSD Vaccins, Lead Rapporteur: Kristina Dunder Request for Supplementary Information adopted on 16.03.2017.	Weekly start timetable.
WS1131 **Januvia-** **EMEA/H/C/000722/WS1131/0055** **Ristaben-** **EMEA/H/C/001234/WS1131/0047** **TESAVEL-** **EMEA/H/C/000910/WS1131/0055** **Xelevia-EMEA/H/C/000762/WS1131/0059** MAH: Merck Sharp & Dohme Limited, Lead Rapporteur: Johann Lodewijk Hillege Opinion adopted on 30.03.2017.	Positive Opinion adopted by consensus on 30.03.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1146 **Hexacima-**	Weekly start timetable.

EMEA/H/C/002702/WS1146/0058
Hexaxim-
EMEA/H/W/002495/WS1146/0064
Hexyon-
EMEA/H/C/002796/WS1146/0062
MAH: Sanofi Pasteur SA, Lead Rapporteur: Jan Mueller-Berghaus

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

- fluticasone furoate / umeclidinium / vilanterol - EMEA/H/C/004781
, treatment of adult patients with chronic obstructive pulmonary disease (COPD),

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

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

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

- amifampridine -
EMEA/H/C/001032/S/0049, Orphan
MAH: BioMarin Europe Ltd

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

Forxiga - dapagliflozin -
EMEA/H/C/002322/R/0035
MAH: AstraZeneca AB, Rapporteur: Kristina Dunder, 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: Filip Josephson, PRAC Rapporteur: Doris Stenver

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

Alecensa - alectinib - EMEA/H/C/004164/II/0001
MAH: Roche Registration Limited, Rapporteur: Filip Josephson, Co-Rapporteur: Sinan B. Sarac, PRAC Rapporteur: Julie WilliamsExtension 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."

BLINCYTO - blinatumomab - EMEA/H/C/003731/II/0011, Orphan
MAH: Amgen Europe B.V., Rapporteur: Alexandre Moreau, Co-Rapporteur: Daniela Melchiorri, PRAC Rapporteur: Jana MladáExtension 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.

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

MAH: Merck Sharp & Dohme Limited, Rapporteur: Daniela Melchiorri, Co-Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Sabine Straus"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."

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

MAH: Amgen Europe B.V., Rapporteur: Kristina Dunder, Co-Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Ulla Wändel Liminga, "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."

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

MAH: Bristol-Myers Squibb Pharma EEIG, Rapporteur: Paula Boudewina van Hennik, PRAC Rapporteur: Sabine Straus"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."

B.6.8. 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

Apidra - insulin glulisine - EMEA/H/C/000557/II/0074/G
MAH: Sanofi-aventis Deutschland GmbH, Rapporteur: Greg Markey,

Betaferon - interferon beta-1b - EMEA/H/C/000081/II/0114
MAH: Bayer AG, Rapporteur: Greg Markey,

Elonva - corifollitropin alfa - EMEA/H/C/001106/II/0036/G
MAH: Merck Sharp & Dohme Limited, Rapporteur: Paula Boudewina van Hennik,

Extavia - interferon beta-1b - EMEA/H/C/000933/II/0084
MAH: Novartis Europharm Ltd, Informed Consent of Betaferon, Rapporteur: Greg Markey,

Flixabi - infliximab - EMEA/H/C/004020/II/0013/G
MAH: Samsung Bioepis UK Limited (SBUK), Rapporteur: Jan Mueller-Berghaus,

Ganfort - bimatoprost / timolol - EMEA/H/C/000668/II/0027/G
MAH: Allergan Pharmaceuticals Ireland, Rapporteur: Hanne Lomholt Larsen

Memantine ratiopharm - memantine - EMEA/H/C/002671/II/0008
MAH: ratiopharm GmbH, Generic, Generic of Ebixa, Rapporteur: Bart Van der Schueren,

Praluent - alirocumab -
EMEA/H/C/003882/II/0022/G
MAH: sanofi-aventis groupe, Rapporteur: Johann Lodewijk Hillege,

Prevenar 13 - pneumococcal polysaccharide conjugate vaccine (13-valent, adsorbed) -
EMEA/H/C/001104/II/0156
MAH: Pfizer Limited, Rapporteur: Kristina Dunder

Simponi - golimumab -
EMEA/H/C/000992/II/0075/G
MAH: Janssen Biologics B.V., Rapporteur: Kristina Dunder,

Sustiva - efavirenz -
EMEA/H/C/000249/II/0142/G
MAH: Bristol-Myers Squibb Pharma EEIG, Rapporteur: Bruno Sepodes

Xultophy - insulin degludec / liraglutide -
EMEA/H/C/002647/II/0019
MAH: Novo Nordisk A/S, Rapporteur: Kristina Dunder,

Zepatier - elbasvir / grazoprevir -
EMEA/H/C/004126/II/0007
MAH: Merck Sharp & Dohme Limited, Rapporteur: Greg Markey

B.6.9. 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. 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.”

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.”

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 CLEAR study (CAIN457A2317) and to add new data from a scalp psoriasis study (CAIN457US01) . In addition the MAH has taken the occasion to include correction in section 4.2 to avoid medication errors and the Package leaflet has been updated accordingly, and in section 5.1 to align the information PsARC definition to the EMA guideline. The MAH has also implemented the latest QRD version 10.0.”

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.”

Harvoni - ledipasvir / sofosbuvir - EMEA/H/C/003850/II/0052

MAH: Gilead Sciences International Ltd, Rapporteur: Filip JosephsonSubmission 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.”

Imbruvica - ibrutinib - EMEA/H/C/003791/II/0034, Orphan

MAH: Janssen-Cilag International NV,

Rapporteur: Filip JosephsonSubmission 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."

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 as an adverse reaction based on continuous monitoring of the safety profile; the Package Leaflet is updated accordingly."

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."

Odomzo - sonidegib - EMEA/H/C/002839/II/0010
MAH: Novartis Europharm Ltd, 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."

Prolia - denosumab - EMEA/H/C/001120/II/0069
MAH: Amgen Europe B.V., Rapporteur: Kristina Dunder, PRAC Rapporteur: Ulla Wändel LimingaChange(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 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”.”

Selincro - nalmefene - EMEA/H/C/002583/II/0020/G

MAH: H. Lundbeck A/S, Rapporteur: Martina Weise,“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.”

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.”

Strensiq - asfotase alfa - EMEA/H/C/003794/II/0018, Orphan

MAH: Alexion Europe SAS, Rapporteur: Greg 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."

Tarceva - erlotinib - EMEA/H/C/000618/II/0051

MAH: Roche Registration Limited, Rapporteur: Sinan B. SaracSubmission 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."

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."

Venclyxto - venetoclax - EMEA/H/C/004106/II/0003, Orphan

MAH: AbbVie Ltd., Rapporteur: Filip JosephsonSubmission 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 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."

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 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 Study 20080560 completed recently where results showed no difference between the risk of developing cataracts in the denosumab and placebo groups."

Zostavax - shingles (herpes zoster) vaccine (live) - EMEA/H/C/000674/II/0112

MAH: MSD Vaccins, Rapporteur: Jan Mueller-BerghausUpdate 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."

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].”

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].”

B.6.10. 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

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."

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 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 primary objective of this study relates to understanding the potential safety risk for cardiovascular events. It is a non-treatment extension to study A3051123, to collect data on cardiovascular safety for all participants in the A3051123 trial for an additional 28 weeks, allowing for a total of 52 weeks of cardiovascular safety data collection.

The RMP version 10.1 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 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.”

Neulasta - pegfilgrastim - EMEA/H/C/000420/II/0093/G

MAH: Amgen Europe B.V., Rapporteur: Robert James Hemmings, PRAC Rapporteur: Julie Williams,“- B.IV.1.a.3 (type II) – To add a new device which may have a significant impact on the delivery of the product: the on-body injector (Onpro kit) to be used with Neulasta, 6mg solution for injection, pre-filled syringe - B.II.e.5.c (type II) – To change the fill volume from 0.6 to 0.64 mL for Neulasta, 6 mg, solution for injection pre-filled syringe co-packed with the new on-body injector (Onpro

kit)
In addition the MAH took the opportunity to introduce editorial changes to module 3.2.P.2.4 Container Closure System
Update of sections 3, 4.2, 5.1, 6.4, 6.5, 6.6 and 8 of the SmPC. The Labelling, Package Leaflet and the RMP (version 4.2) are updated accordingly. In addition the Marketing authorisation holder (MAH) took the opportunity to update the list of local representatives in the Package Leaflet, to include some editorial changes and correct some typos throughout the product information, and to bring the product information in line with the latest QRD template version 10."

Odomzo - sonidegib - EMEA/H/C/002839/II/0011
MAH: Novartis Europharm Ltd, 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."

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."

B.6.11. PRAC assessed procedures

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 Approval 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)."

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."

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 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."

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."

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 SepodesSubmission 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."

PRAC Led
WS1163
Harvoni-
EMEA/H/C/003850/WS1163/0051
Sovaldi-EMEA/H/C/002798/WS1163/0041
MAH: Gilead Sciences International Ltd, Lead Rapporteur: Filip Josephson, Lead PRAC Rapporteur: Julie Williams, PRAC-CHMP liaison: Greg Markey"To provide updated RMPs for Sovaldi and Harvoni following the CHMP opinion, endorsing a PRAC recommendation, issued on 15 December 2016 (EMA/CHMP/847450/2016) on the Article 20 procedure for Direct-acting antivirals (DAAs) indicated for the treatment of hepatitis C (interferon free). The PRAC requested `hepatitis B reactivation' to be considered as important identified risk for all direct-acting antivirals. In addition, `emergence of hepatocellular carcinoma' and `recurrence of hepatocellular carcinoma' have been included as important potential risks. `Patients with previous HCC' have been reflected as missing information in the RMP of the DAAs, since this population was excluded from existing clinical trials. The requested studies have also been reflected in the RMPs."

PRAC Led
WS1169
Exviera-EMEA/H/C/003837/WS1169/0028
Viekirax-
EMEA/H/C/003839/WS1169/0032
MAH: AbbVie Ltd., Lead Rapporteur: Filip Josephson, Lead PRAC Rapporteur: Dolores Montero Corominas, PRAC-CHMP liaison: Concepcion Prieto Yerro"To provide updated

RMPs for Exviera and Viekirax following the CHMP opinion, endorsing a PRAC recommendation, issued on 15 December 2016 (EMA/CHMP/847450/2016) on the Article 20 procedure for Direct-acting antivirals (DAAs) indicated for the treatment of hepatitis C (interferon free). The PRAC requested `hepatitis B reactivation' to be considered as important identified risk for all direct-acting antivirals. In addition, `emergence of hepatocellular carcinoma' and `recurrence of hepatocellular carcinoma' have been included as important potential risks. `Patients with previous HCC' have been reflected as missing information in the RMP of the DAAs, since this population was excluded from existing clinical trials. The requested studies have also been reflected in the RMPs."

B.6.12. CHMP-CAT assessed procedures

B.6.13. CHMP-PRAC-CAT assessed procedures

B.6.14. PRAC assessed ATMP procedures

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

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

WS1170
Aflunov-
EMEA/H/C/002094/WS1170/0036
Foclivia-
EMEA/H/C/001208/WS1170/0031
MAH: Seqirus S.r.l, Lead Rapporteur: Daniela Melchiorri

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

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 18-21 April 2017 CHMP plenary:

G.3.2. List of procedures starting in April 2017 for May 2017 CHMP adoption of outcomes

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