24 March 2017
EMA/CHMP/196172/2017
Inspections, Human Medicines Pharmacovigilance and Committees Division

Committee for medicinal products for human use (CHMP)

Minutes of the meeting on 20-23 February 2017

Chair: Tomas Salmonson – Vice-Chair: Harald Enzmann

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

Table of contents

1. Introduction

1.1. Welcome and declarations of interest of members, alternates and experts

In accordance with the Agency's policy on handling of declarations of interests of scientific committees' members and experts, based on the declarations of interest submitted by the Committee members, alternates and experts and based on the topics in the agenda of the current meeting, the Committee Secretariat announced that no restriction in the involvement of meeting participants in upcoming discussions was identified as included in the list of participants and restrictions. See (current) February 2017 CHMP minutes for the list of participants and restrictions in relation to declarations of interests applicable to the items of the agenda for the CHMP plenary session held 20-23 February 2017 (to be published post March 2017 CHMP meeting).

Participants in this meeting were asked to declare any changes, omissions or errors to their declared interests and/or additional restrictions concerning the matters for discussion. No new or additional interests or restrictions were declared.

Discussions, deliberations and voting took place in full respect of the restricted involvement of Committee members and experts in line with the relevant provisions of the Rules of Procedure and as included in the list of participants. All decisions taken at this meeting were made in the presence of a quorum of members (i.e. 22 or more members were present in the room). All decisions, recommendations and advice were agreed by consensus, unless otherwise specified.

1.2. Adoption of agenda

CHMP agenda for 20-23 February 2017.

The CHMP adopted the agenda.

1.3. Adoption of the minutes

CHMP minutes for 23-26 January 2017.

The CHMP adopted the minutes.

2. Oral Explanations

2.1. Pre-authorisation procedure oral explanations

2.1.1. - dinutuximab beta - Orphan - EMEA/H/C/003918

APEIRON Biologics AG; treatment of neuroblastoma

Scope: Oral explanation to be held on 22 February 2017 at time 09:00

Action: For adoption

List of Outstanding Issues adopted on 13.10.2016, 26.05.2016. List of Questions adopted on 24.09.2015.

Oral explanation to be held on 22 February 2017 at time 09:00 was cancelled.

See also 3.2.10

2.1.2. - nonacog beta pegol - Orphan - EMEA/H/C/004178

Novo Nordisk A/S; treatment of haemophilia B

Scope: Oral explanation

Action: Oral explanation to be held on 21 February 2017 at time 09:00

List of Outstanding Issues adopted on 15.09.2016. List of Questions adopted on 26.05.2016.

An oral explanation was held on 21 February 2017 at 09:00. The members further discussed the presented data as well as the wording of the indication.

The CHMP adopted the BWP report.

2.1.3. - pentosan polysulfate sodium - Orphan - EMEA/H/C/004246

bene-Arzneimittel GmbH; treatment of Interstitial Cystitis

Scope: Oral explanation

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

List of Outstanding Issues adopted on 10.11.2016. List of Questions adopted on 23.06.2016.

An oral explanation was held on 20 February 2017 at time 16:00.

See 3.2.13

2.2. Re-examination procedure oral explanations

No items

2.3. Post-authorisation procedure oral explanations

No items

2.4. Referral procedure oral explanations

No items

3. Initial applications

3.1. Initial applications; Opinions

3.1.1. Emtricitabine/tenofovir disoproxil Krka d.d. - emtricitabine / tenofovir disoproxil - EMEA/H/C/004686

KRKA, d.d., Novo mesto; treatment of HIV-1 infection

Scope: Opinion

Action: For adoption

Generic application (Article 10(1) of Directive No 2001/83/EC), Duplicate of Emtricitabine/Tenofovir disoproxil Krka

The Committee confirmed that all issues previously identified in this application had been addressed.

The Committee adopted a positive opinion recommending the granting of a marketing authorisation by consensus together with the CHMP assessment report and translation timetable.

The Icelandic and Norwegian Members were in agreement with the CHMP recommendation.

The legal status was agreed as medicinal product subject to restricted medical prescription.

The summary of opinion was circulated for information.

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

AstraZeneca AB; for the treatment of hyperkalaemia

Scope: Opinion

Action: For adoption

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

List of Outstanding Issues adopted on 26.01.2017, 10.11.2016. List of Questions adopted on 28.04.2016.

The Committee confirmed that all issues previously identified in this application had been addressed.

The Committee adopted a positive opinion recommending the granting of a marketing authorisation by consensus together with the CHMP assessment report and translation timetable.

Furthermore, the CHMP considered that sodium zirconium cyclosilicate is a new active substance, as claimed by the applicant.

The Icelandic and Norwegian Members were in agreement with the CHMP recommendation.

The legal status was agreed as medicinal product subject to medical prescription.

The summary of opinion was circulated for information.

3.1.3. Natpar - parathyroid hormone - Orphan - EMEA/H/C/003861

Shire Pharmaceuticals Ireland Ltd; treatment of hypoparathyroidism

Scope: Opinion

Action: For adoption

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

List of Outstanding Issues adopted on 15.09.2016, 21.07.2016, 28.04.2016, 24.09.2015. List of Questions adopted on 26.03.2015.

The Committee confirmed that all issues previously identified in this application had been addressed.

The Committee adopted a positive opinion recommending the granting of a conditional marketing authorisation by consensus together with the CHMP assessment report and translation timetable.

The Icelandic and Norwegian Members were in agreement with the CHMP recommendation.

The legal status was agreed as medicinal product subject to restricted medical prescription.

The CHMP noted the letter of recommendation dated 23 February 2017.

The summary of opinion was circulated for information.

The CHMP adopted the BWP report.

3.1.4. Pemetrexed Hospira UK Limited - pemetrexed - EMEA/H/C/004488

Hospira UK Limited; treatment of malignant pleural mesothelioma and non-small cell lung cancer

Scope: Opinion

Action: For adoption

Generic application (Article 10(1) of Directive No 2001/83/EC), Generic of Alimta, Duplicate of Pemetrexed ditromethamine Hospira (WD)

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

The Committee confirmed that all issues previously identified in this application had been addressed.

The Committee adopted a positive opinion recommending the granting of a marketing authorisation by consensus together with the CHMP assessment report and translation timetable.

The Icelandic and Norwegian Members were in agreement with the CHMP recommendation.

The legal status was agreed as medicinal product subject to restricted medical prescription.

The summary of opinion was circulated for information.

3.1.5. Roteas - edoxaban - EMEA/H/C/004339

Daiichi Sankyo Europe GmbH; prevention of stroke; embolism and treatment of venous thromboembolism

Scope: Opinion

Action: For adoption

Informed consent application (Article 10c of Directive No 2001/83/EC)

The Committee confirmed that all issues previously identified in this application had been addressed.

The Committee adopted a positive opinion recommending the granting of a marketing authorisation by consensus together with the CHMP assessment report and translation timetable.

The Icelandic and Norwegian Members were in agreement with the CHMP recommendation.

The legal status was agreed as medicinal product subject to medical prescription.

The summary of opinion was circulated for information.

3.1.6. Varuby - rolapitant - EMEA/H/C/004196

Tesaro UK Limited; prevention of nausea and vomiting

Scope: Opinion

Action: For adoption

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

List of Outstanding Issues adopted on 10.11.2016. List of Questions adopted on 21.07.2016.

The Committee confirmed that all issues previously identified in this application had been addressed.

The Committee adopted a positive opinion recommending the granting of a marketing authorisation by consensus together with the CHMP assessment report and translation timetable.

The Icelandic and Norwegian Members were in agreement with the CHMP recommendation.

The legal status was agreed as medicinal product subject to medical prescription.

The summary of opinion was circulated for information.

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

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

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

Scope: Day 180 list of outstanding issue

Action: For adoption

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

The Committee was reminded of the status of this application and its remaining outstanding issues

The Committee adopted a 2nd list of outstanding issues with a specific timetable.

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

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

Scope: Day 180 list of outstanding issue,

Action: For adoption

List of Questions adopted on 15.07.2016.

The Committee was reminded of the status of this application and its remaining outstanding issues. The CHMP was updated on the discussions at the CAT.

The Committee agreed to the list of outstanding issues with a specific timetable as adopted by the CAT at their February meeting.

The CHMP adopted the BWP report.

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

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

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 15.09.2016.

The Committee was reminded of the status of this application and its remaining outstanding issues.

The Committee adopted a list of outstanding issues with a specific timetable.

The CHMP adopted the similarity assessment report.

The CHMP agreed to consult the Oncology WP and adopted a list of questions to this group.

The CHMP adopted the BWP report.

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

Accelerated assessment

BioMarin International Limited; treatment of neuronal ceroid lipofuscinosis type 2

Scope: Day 180 list of outstanding issue, list of experts for the ad-hoc expert group meeting on 7 March 2017 were adopted via written procedure

Action: For adoption

List of Questions adopted on 13.12.2016.

The Committee was reminded of the status of this application and its remaining outstanding issues.

The Committee adopted a list of outstanding issues

The CHMP agreed to the request by the applicant for an extension to the clock stop to respond to the list of outstanding issues with a specific timetable.

The CHMP agreed to keep the procedure on an accelerated assessment timetable at this time.

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

treatment of cartilage defects

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 19.04.2013.

The Committee was reminded of the status of this application and its remaining outstanding issues.

The Committee agreed to the list of outstanding issues adopted by the CAT at their February meeting with amendments and a specific timetable.

The CHMP adopted the BWP report.

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

GMP-Orphan SA; Wilson's disease

Scope: Day 180 list of outstanding issue/Oral Explanation

Action: For adoption

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

The Committee was reminded of the status of this application and its remaining outstanding issues. After discussion, the CHMP agreed that no oral explanation was required at this time. The CHMP noted the answers from PKWP.

The Committee adopted a 2nd list of outstanding issues with a specific timetable.

3.2.7. - 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: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 01.04.2016.

The Committee was reminded of the status of this application and its remaining outstanding issues.

The Committee adopted a list of outstanding issues with a specific timetable.

The CHMP adopted the BWP report.

3.2.8. - iloperidone - EMEA/H/C/004149

treatment of schizophrenia

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 28.04.2016.

The Committee was reminded of the status of this application and its remaining outstanding issues.

The Committee adopted a list of outstanding issues with a specific timetable.

The CHMP agreed to consult a SAG and adopted a list of questions to this group.

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

treatment of hyperuricaemia

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 15.09.2016.

The Committee was reminded of the status of this application and its remaining outstanding issues.

The Committee adopted a list of outstanding issues with a specific timetable.

3.2.10. - dinutuximab beta - Orphan - EMEA/H/C/003918

APEIRON Biologics AG; treatment of neuroblastoma

Scope: Oral explanation, Day 180 list of outstanding issue

Action: For adoption

List of Outstanding Issues adopted on 13.10.2016, 26.05.2016. List of Questions adopted on 24.09.2015.

List of Questions adopted via written procedure on 14.02.2017.

The Committee was reminded of the status of this application and its remaining outstanding issues.

The CHMP agreed that the oral explanation to be held on 22 February 2017 at time 09:00 was not required at this time.

The Committee adopted a 3rd list of outstanding issues with a specific timetable.

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

treatment of active rheumatoid arthritis

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 10.11.2016.

The Committee was reminded of the status of this application and its remaining outstanding issues.

The Committee adopted a list of outstanding issues with a specific timetable.

The CHMP adopted the BWP report.

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

AB Science; treatment of mastocytosis

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 15.09.2016.

The Committee was reminded of the status of this application and its remaining outstanding issues.

The Committee adopted a list of outstanding issues with a specific timetable.

3.2.13. - pentosan polysulfate sodium - Orphan - EMEA/H/C/004246

bene-Arzneimittel GmbH; treatment of Interstitial Cystitis (IC)

Scope: Oral explanation, Day 180 list of outstanding issue, report form the ad-hoc expert group

Action: For adoption

List of Outstanding Issues adopted on 10.11.2016. List of Questions adopted on 23.06.2016.

The Committee noted the report from the ad-hoc expert group. According to the ad-hoc expert group, there is a medical need for effective medical treatments in this condition. It was considered that the IC definition at the time when most of the pivotal studies were performed would, as per todays classification, fall under bladder pain syndrome (BPS) type 2 – 3. Considering today's classification and medical practice an indication today would therefore need to refer to BPS. The experts further emphasized that a categorization reflecting current clinical practice would refer to symptomatic severity. The ad-hoc experts broadly agreed that study results from a population of BPS 2-3, a population difficult to treat, would be generalizable to patients suffering from other BPS categories. It was also emphasized by the experts that more robust data generation and definition of the patient population which benefits most from the drug would be welcome to spare the patient from trying unsuccessful treatments but also the burden of invasive diagnostics.

An oral explanation was held on 20 February 2017 at time 16:00.

The Committee discussed the indication proposal.

The Committee adopted a 2nd list of outstanding issues with a specific timetable.

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

treatment of schizophrenia

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 21.07.2016.

The Committee was reminded of the status of this application and its remaining outstanding issues.

The Committee adopted a list of outstanding issues with a specific timetable.

The CHMP agreed to consult expert Healthcare Professionals (ophthalmology) in writing. The CHMP adopted the list of questions to the experts.

3.2.15. - 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: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 15.09.2016.

The Committee was reminded of the status of this application and its remaining outstanding issues.

The Committee adopted a list of outstanding issues with a specific timetable.

The CHMP adopted the similarity assessment report

The CHMP adopted the BWP report.

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

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

Scope: Day 180 list of outstanding issue

Action: For adoption

The Committee was reminded of the status of this application and its remaining outstanding issues.

The Committee adopted a list of outstanding issues with a specific timetable.

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

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

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 28.04.2016.

The Committee was reminded of the status of this application and its remaining outstanding issues.

The Committee adopted a list of outstanding issues with a specific timetable.

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

treatment of hyperammoniemia

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 15.09.2016.

The Committee was reminded of the status of this application and its remaining outstanding issues.

The Committee adopted a list of outstanding issues with a specific timetable.

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

treatment of hyperkalaemia

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 15.09.2016.

The Committee was reminded of the status of this application and its remaining outstanding issues.

The Committee adopted a list of outstanding issues with a specific timetable.

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

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

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

Scope: Day 120 list of questions

Action: For adoption

The Committee discussed the issues identified in this application.

The Committee adopted the CHMP recommendation and scientific discussion together with the list of questions.

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

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

Scope: Day 120 list of questions

Action: For adoption

The Committee discussed the issues identified in this application.

The Committee adopted the CHMP recommendation and scientific discussion together with the list of questions.

The CHMP adopted the BWP report.

3.3.3. - trastuzumab - EMEA/H/C/002575

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

Scope: Day 120 list of questions

Action: For adoption

The Committee discussed the issues identified in this application.

The Committee adopted the CHMP recommendation and scientific discussion together with the list of questions.

The CHMP adopted the BWP report.

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

Accelerated assessment

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

Scope: Day 90 list of questions

Action: For adoption

The Committee discussed the issues identified in this application.

The Committee adopted the CHMP recommendation and scientific discussion together with the list of questions .

The CHMP agreed to keep the procedure on an accelerated assessment timetable at this time.

The CHMP adopted the BWP report.

3.3.5. - insulin glargine - EMEA/H/C/004280

treatment of diabetes mellitus

Scope: Day 120 list of questions

Action: For adoption

The Committee discussed the issues identified in this application.

The Committee adopted the CHMP recommendation and scientific discussion together with the list of questions.

3.3.6. - beclometasone dipropionate anhydrous / formoterol fumarate dihydrate / glycopyrronium bromide - EMEA/H/C/004257

for the symptomatic treatment and reduction of exacerbations in adult patients with chronic obstructive pulmonary disease (COPD) with airflow limitation and who are at risk of exacerbations

Scope: Day 120 list of questions

Action: For adoption

The Committee discussed the issues identified in this application. .

The Committee adopted the CHMP recommendation and scientific discussion together with the list of questions.

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

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

Scope: Day 120 list of questions

Action: For adoption

The Committee discussed the issues identified in this application.

The Committee adopted the CHMP recommendation and scientific discussion together with the list of questions.

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

treatment for opioid drug dependence

Scope: Day 120 list of questions

Action: For adoption

The Committee discussed the issues identified in this application.

The Committee adopted the CHMP recommendation and scientific discussion together with the list of questions.

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

3.4.1. - prasterone - EMEA/H/C/004138

treatment of vulvovaginal atrophy

Scope: Letter from the applicant dated 14 February 2017 requesting an extension of clock stop to respond to the List of Outstanding Issues adopted on 26 January 2017

Action: For adoption

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

The CHMP agreed to the request by the applicant for an extension to the clock stop to respond to the list of outstanding issues adopted in January 2017.

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

treatment of HIV-1 infection

Scope: Request by the applicant for an extension to the clock stop to respond to the list of questions adopted on 15.12.2016

Action: For adoption

List of Questions adopted on 15.12.2016.

The CHMP agreed to the request by the applicant for an extension to the clock stop to respond to the list of questions adopted on 15.12.2016.

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

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

Scope: Letter from the applicant dated 8 February 2017 requesting an extension of clock stop to respond to the List of Questions adopted on 26 January 2017

Action: For adoption

List of Questions adopted on 26.01.2017

The CHMP agreed to the request by the applicant for an extension to the clock stop to respond to the list of questions adopted in January 2017.

3.4.4. - miglustat - EMEA/H/C/004366

treatment of Gaucher disease

Scope: Letter from the applicant dated 8 February 2017 requesting an extension of clock stop to respond to the List of Questions adopted on 15 December 2016.

Action: For adoption

List of Questions adopted on 15.12.2016.

The CHMP agreed to the request by the applicant for an extension to the clock stop to respond to the list of questions adopted in December 2016.

3.4.5. - tigecycline - EMEA/H/C/004419

Treatment of:

- complicated skin and soft tissue infections, excluding diabetic foot infections

- complicated intra-abdominal infections
should be used only in situations where it is known or suspected that other alternatives are not suitable

Scope: Letter from the applicant dated 7 February 2017 requesting an extension of clock stop to respond to the List of Questions adopted on 13 October 2016

Action: For adoption

List of Questions adopted on 13.10.2016

The CHMP agreed to the request by the applicant for an extension to the clock stop to respond to the list of questions adopted in October 2016.

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

treatment of metastatic colorectal cancer

Scope: A clock stop extension on the List of Outstanding Issues adopted on 15.12.2016 has been granted via written procedure on 06 February 2017.

Action: For information

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

The CHMP noted the new timetable following the adoption of the request by the applicant by written procedure.

3.4.7. - neratinib - EMEA/H/C/004030

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

Scope: Request by the applicant for clock stop extension to respond to the list of questions adopted on 15 December 2016.

Action: For adoption

The CHMP agreed to the request for clock stop extension to respond to the list of questions adopted on 15 December 2016 with a specific timetable.

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

No items

3.6. Initial applications in the decision-making phase

No items

3.7. Withdrawals of initial marketing authorisation application

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. Esbriet - pirfenidone - Orphan - EMEA/H/C/002154/X/0035/G

Roche Registration Limited

Rapporteur: Greg Markey, Co-Rapporteur: David Lyons, PRAC Rapporteur: Julie Williams

Scope: "Extension application to introduce a new pharmaceutical form associated with 3 new strengths (267mg, 534mg and 801mg film-coated tablets).

Action: For adoption

List of Outstanding Issues adopted on 15.12.2016. List of Questions adopted on 15.09.2016.

The Committee confirmed that all issues previously identified in this application had been addressed.

The Committee adopted a positive opinion by consensus together with the CHMP Assessment Report and translation timetable.

The Icelandic and Norwegian CHMP members were in agreement with the CHMP recommendations.

The summary of opinion was circulated for information.

4.1.2. Pergoveris - follitropin alfa / lutropin alfa - EMEA/H/C/000714/X/0047

Merck Serono Europe Limited

Rapporteur: Nithyanandan Nagercoil, PRAC Rapporteur: Julie Williams

Scope: "Extension application to introduce a new pharmaceutical form (solution for injection) associated with 3 strengths of (300 IU + 150 IU)/ 0.48 ml, (450 IU + 225 IU)/ 0.72 ml and (900 IU + 450 IU)/ 1.44 ml."

Action: For adoption

List of Questions adopted on 10.11.2016.

The Committee confirmed that all issues previously identified in this application had been addressed.

The Committee adopted a positive opinion by consensus together with the CHMP Assessment Report and translation timetable.

The Icelandic and Norwegian CHMP members were in agreement with the CHMP recommendations.

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

4.2.1. 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 strenghts 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 Questions adopted on 13.10.2016.

The Committee noted the issues identified in this application.

The Committee adopted the CHMP recommendation and scientific discussion together with the list of outstanding issues and a specific timetable.

4.2.2. 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 Questions adopted on 10.11.2016.

The Committee discussed the issues identified in this application, relating to further clarification on the analytical method used in the bioequivalence study as well as the data and samples flow management.

The Committee adopted the CHMP recommendation and scientific discussion together with the list of outstanding issues and a specific timetable.

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

Glaxo Group Ltd

Rapporteur: Kristina Dunder, PRAC Rapporteur: Ulla Wändel Liminga

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

Action: For adoption

The Committee noted the issues identified in this application.

The Committee adopted the CHMP recommendation and scientific discussion together with the list of questions.

4.3.2. Exjade - deferasirox - Orphan - EMEA/H/C/000670/X/0054

Novartis Europharm Ltd

Rapporteur: Alexandre Moreau, PRAC Rapporteur: Caroline Laborde

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

Action: For adoption

The Committee noted the issues identified in this application, which were related to Quality, RMP and SmPC.

The Committee adopted the CHMP recommendation and scientific discussion together with the list of questions.

4.3.3. 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

The Committee discussed the issues identified in this application.

The Committee adopted the CHMP recommendation and scientific discussion together with the list of questions.

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

No items

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

No items

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

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

5.1.1. Darzalex - daratumumab - Orphan - EMEA/H/C/004077/II/0002

Janssen-Cilag International NV

Rapporteur: Sinan B. Sarac, PRAC Rapporteur: Marcia Sofia Sanches de Castro Lopes Silva

Scope: "Extension of Indication for Darzalex in combination with lenalidomide and dexamethasone, or bortezomib and dexamethasone, for the treatment of adult patients with multiple myeloma who have received at least one prior therapy. As a consequence, sections 4.1, 4.2, 4.4, 4.5, 5.1 and 5.2 of the SmPC are updated in order to update the information on the target patient population, posology, warnings, interactions, efficacy and pharmacokinetics. A new warning is introduced in section 4.4 regarding neutropenia/thrombocytopenia induced by background therapy.

Furthermore, the CHMP is of the opinion that all specific obligations have been fulfilled following submission of the final results of studies MMY3003 and MMY3004 and in light of the data generated and the evidence of compliance with the specific obligations, the CHMP recommends the granting of a marketing authorisation in accordance with Article 14(1) of Regulation No 726/2004.

The Package Leaflet and Risk Management Plan (RMP version 2.1) are updated in accordance.

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

Request for 1 year of market protection for a new indication (Article 14(11) of Regulation (EC) 726/2004)

Action: For adoption

Request for Supplementary Information adopted on 15.12.2016.

The Committee confirmed that all issues previously identified in this application had been

addressed.

The Committee adopted a positive opinion by consensus together with the CHMP Assessment Report and translation timetable.

The Icelandic and Norwegian CHMP members were in agreement with the CHMP recommendations.

The summary of opinion was circulated for information.

The CHMP agreed by consensus on an additional 1 year of market protection for a new indication.

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

AstraZeneca UK Ltd

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

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

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

Action: For adoption

The Committee discussed the issues identified in this application. The main discussion related to the indication wording.

The Committee adopted a request for supplementary information with a specific timetable.

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

Roche Registration Limited

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

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

Action: For adoption

The Committee noted the issues identified in this application.

The Committee adopted a request for supplementary information with a specific timetable.

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

Santhera Pharmaceuticals (Deutschland) GmbH

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

Scope: "Extension of indication to include treatment of patients with Duchenne muscular dystrophy in whom respiratory function has started to decline and who are currently not taking concomitant glucocorticoids. Raxone can be used in patients previously treated with glucocorticoids or in patients in whom glucocorticoid treatment is not desired, not tolerated or is contraindicated."

Action: For adoption

Request for Supplementary Information adopted on 15.09.2016.

The Committee discussed the issues identified in this application, which were related to efficacy findings and methodological issues.

The Committee adopted a 2nd request for supplementary information with a specific timetable.

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

Gilead Sciences International Ltd

Rapporteur: Filip Josephson, Co-Rapporteur: Alar Irs, PRAC Rapporteur: Patrick Batty

Scope: "Extension of indication to add treatment of chronic hepatitis C in adolescents aged 12 to <18 years.
As a consequence, sections 4.1, 4.2, 4.4, 4.8, 5.1 and 5.2 of the SmPC are updated in order to add information on posology, warnings, safety, efficacy and pharmacokinetics.
The Package Leaflet and Risk Management Plan (RMP version 5.0) are updated in accordance.
In addition, the Marketing authorisation holder (MAH) took the opportunity to update the list of local representatives in the Package Leaflet.
Furthermore, the Product Information is brought in line with the latest QRD template version 10."

Action: For adoption

The Committee discussed the issues identified in this application, which were related to the duration of treatment and follow-up study.

The Committee adopted a request for supplementary information with a specific timetable.

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

Bayer Pharma AG

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

Scope: "Extension of indication of Stivarga to include treatment of adult patients with

hepatocellular carcinoma (HCC) who have been previously treated with one systemic therapy.
As a consequence, sections 4.1, 4.2, 4.4, 4.8 and 5.1 of the EU SmPC are updated. The package leaflet and RMP (version 5.0) have been updated accordingly.
Furthermore, the PI is brought in line with the latest QRD template version 10.0."

Action: For adoption

The Committee discussed the issues identified in this application, relating to the proposed target population. The CHMP agreed that further discussion on the wording of the indication was required.

The Committee adopted a request for supplementary information with a specific timetable.

5.1.7. 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."

Action: For adoption

Request for Supplementary Information adopted on 13.10.2016.

The Committee discussed the issues identified in this application, in particular the long-term follow-up data from patients that stopped treatment and whether data indicated any resistance development during off-therapy. The CHMP agreed to request further clarification from the MAH.

The CHMP agreed to consult the SAG Oncology and adopted a list of questions to this group.

The Committee adopted a request for supplementary information with a specific timetable.

5.1.8. Truvada - emtricitabine / tenofovir disoproxil - EMEA/H/C/000594/II/0131

Gilead Sciences International Ltd

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

Scope: "Extension of Indication to include treatment of HIV-1 infected adolescents, with NRTI resistance or toxicities precluding the use of first line agents, aged 12 to < 18 years for Truvada.
As a consequence, sections 4.1, 4.2, 4.4, 4.8, 5.1 and 5.2 of the SmPC are updated.
The Package Leaflet and the Risk Management plan (v.13) are updated in accordance."

Action: For adoption

Request for Supplementary Information adopted on 10.11.2016.

The Committee confirmed that all issues previously identified in this application had been addressed.

The Committee adopted a positive opinion by consensus together with the CHMP Assessment Report and translation timetable.

The Icelandic and Norwegian CHMP members were in agreement with the CHMP recommendations.

The summary of opinion was circulated for information.

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

Novo Nordisk A/S

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

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

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

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

Action: For adoption

The Committee discussed the issues identified in this application, which were related to the safety and efficacy issues.

The Committee adopted a request for supplementary information with a specific timetable.

5.1.10. Mekinist, Tafinlar – trametinib, dabrafenib - EMEA/H/C/WS0996

Novartis Europharm Ltd

Lead Rapporteur: Filip Josephson, Lead Co-Rapporteur: Paula Boudewina van Hennik, PRAC Rapporteur: Ulla Wändel Liminga

Scope: "Extension of indication to include the combination treatment with trametinib and dabrafenib of adult patients with advanced non-small cell lung cancer (NSCLC) with a BRAF V600 mutation. As a consequence, sections 4.1, 4.2, 4.4, 4.8, 5.1, 5.2 and 5.3 of the Mekinist and Tafinlar SmPC are updated. The Package Leaflet and RMP are updated accordingly. In addition, the Worksharing applicant (WSA) took the opportunity to align the SmPCs of Mekinist and Tafinlar. Furthermore, the Product Information is brought in line with the latest QRD template version 10."

Action: For adoption

Request for Supplementary Information adopted on 10.11.2016.

The Committee confirmed that all issues previously identified in this application had been addressed.

The Committee adopted a positive opinion by consensus together with the CHMP Assessment Report and translation timetable.

The Icelandic and Norwegian CHMP members were in agreement with the CHMP recommendations.

The summary of opinion was circulated for information.

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

5.2.1. BeneFIX - nonacog alfa - EMEA/H/C/000139/II/0141

MAH: Pfizer Limited

Rapporteur: Jan Mueller-Berghaus

Scope: "Update of section 5.2 of the SmPC in order to update the information based on PK data from study B1821048. This study reported PK from an extended collection time beyond the 72 hours previously used in all other BeneFIX studies, to 96 hours, after BeneFIX administration."

Letter dated 17.02.2017 informing about the withdrawal of the variation application.

Action: For information

The CHMP noted the letter from the MAH informing about their decision to withdraw the variation application.

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

No items

6. Ancillary medicinal substances in medical devices

6.1. Ancillary medicinal substances in medical devices; Opinions/ Day 180 list of outstanding issues / Day 120 list of questions

No items

6.2. Update of Ancillary medicinal substances in medical devices

No items

7. Procedure under Article 83(1) of Regulation (EC) 726/2004 (Compassionate Use)

7.1. Procedure under Article 83(1) of Regulation (EC) 726/2004 (Compassionate Use)

No items

8. Pre-submission issues

8.1. Pre-submission issue

No items

8.2. Priority Medicines (PRIME)

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

8.2.1. List of applications received

Action: For information

Note: Products requesting eligibility under PRIME scheme are listed in the Annex G.

The CHMP noted the list of applications received.

8.2.2. Recommendation for PRIME eligibility

Action: For adoption

Note: Recommendation for PRIME are listed in the Annex G.

The CHMP adopted the recommendation for PRIME eligibility. The CHMP reviewed 5 recommendations for eligibility to PRIME: 2 were granted and 3 were denied. The individual outcomes are listed in PRIME Monthly Report on EMA website.

9. Post-authorisation issues

9.1. Post-authorisation issues

9.1.1. Vectibix - panitumumab - EMEA/H/C/000741/II/0080

MAH: Amgen Europe B.V.

Rapporteur: Robert James Hemmings, PRAC Rapporteur: Julie Williams

Scope: "Update of Annex II in order to provide the results of biomarker analyses from the Vectibix clinical programme including Study 20080763 (according to Supplementary Statistical Analysis Plan dated 20 September 2013), Study 20070820 and Study 20060447.

The data submitted are in fulfilment of Annex II obligation ANX017.

The Risk Management Plan (version 21.0) has been updated accordingly.

The requested variation proposed amendments to Annex II and the Risk Management Plan."

Request for Supplementary Information adopted on 10.11.2016.

Action: For adoption

The Committee confirmed that all issues previously identified in this application had been addressed.

The Committee adopted a positive opinion by consensus together with the CHMP Assessment Report and translation timetable.

The Icelandic and Norwegian CHMP members were in agreement with the CHMP recommendations.

The summary of opinion was circulated for information.

See also B.5.3 Annex to Minutes

9.1.2. Tagrisso - osimertinib - EMEA/H/C/004124/II/0009/G

MAH: AstraZeneca AB

Rapporteur: Aranzazu Sancho-Lopez, PRAC Rapporteur: Sabine Straus,

Scope: “Update of SmPC sections 4.2, 4.4, 4.8, 5.1 and 5.2 based on the results from study D5160C00003 (AURA3) and the updated CSRs for studies D5160C00001 (AURAex) and D5160C00002 (AURA2). The Package Leaflet has been updated accordingly. In addition, the MAH took the opportunity to make editorial changes in the SmPC and Package Leaflet.
The application included an updated RMP version 6.0.
The provision of the CSR from study AURA3 addresses the Specific Obligation for Tagrisso and hence the MAH requests the conversion from a Conditional Marketing Authorisation to a Marketing Authorisation not subject to Specific Obligations.”
Request for Supplementary Information adopted on 15.12.2016.

Action: For adoption

The Committee confirmed that all issues previously identified in this application had been addressed.

The Committee adopted a positive opinion by consensus together with the CHMP Assessment Report and translation timetable.

The Icelandic and Norwegian CHMP members were in agreement with the CHMP recommendations.

The summary of opinion was circulated for information.

See also B.5.3 Annex to Minutes

10. Referral procedures

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

10.1.1. Sodium-glucose co-transporter 2 (SGLT2) inhibitors: Canaglifozin – INVOKANA (CAP); canagliflozin, metformin – VOKANAMET (CAP); dapaglifozin – EDISTRIDE (CAP), FORXIGA (CAP); dapaglifozin, metformin – XIGDUO (CAP), EBYMECT (CAP); empaglifozin – JARDIANCE (CAP); empaglifozin, metformin – SYNJARDY (CAP) - EMEA/H/A-20/1442

Applicant: Janssen-Cilag International N.V. (Invokana, Vokanamet); AstraZeneca AB (Edistride, Forxiga, Xigduo, Ebymect); Boehringer Ingelheim International GmbH (Jardiance; Synjardy)

Rapporteurs for the Article 20 referral: PRAC Rapporteur: Valerie Strassmann; PRAC Co-rapporteur: Menno van der Elst

Individual products Rapporteur: Martina Weise, Co-Rapporteur: Kristina Dunder (Invokana), Rapporteur: Martina Weise, Co-Rapporteur: Bjorg Bolstad (Vokanamet), Rapporteurs: Kristina Dunder, Co-Rapporteur: Martina Weise (Edistride), Rapporteurs: Kristina Dunder, Co-Rapporteur: Martina Weise (Forxiga), Rapporteur: Kristina Dunder, Co-Rapporteur: Agnes Gyurasics (Xigduo),Rapporteur: Kristina Dunder, Co-Rapporteur: Agnes Gyurasics (Ebymect), Rapporteur: Johann Lodewijk Hillege, Co-Rapporteur: Bart Van der Schueren (Jardiance), Rapporteur: Johann Lodewijk Hillege, Co-Rapporteur: Daniela Melchiorri (Synjardy).

Scope: Opinion

Review of the benefit-risk balance of sodium-glucose co-transporter-2 (SGLT2) inhibitors following notification by the European Commission of a referral under Article 20 of Regulation (EC) No 726/2004, based on pharmacovigilance data

Action: For adoption

The PRAC adopted at its February 2017 meeting an opinion by consensus with the following recommendation:

PRAC is warning that an increase in cases of lower limb amputation has been observed in patients taking the type 2 diabetes medicine canagliflozin compared with those taking placebo in two clinical trials, CANVAS and CANVAS-R. The studies, which are still ongoing, involved patients at high risk of heart problems.

Patients with diabetes (especially those with poorly controlled diabetes and pre-existing problems with the heart and blood vessels) are at increased risk of infection and ulcers which can lead to amputations. The mechanism by which canagliflozin may increase the risk of amputation is still unclear.

An increased risk has not been seen in studies with other medicines in the same class, dapagliflozin and empagliflozin. However, data available to date are limited and the risk may also apply to these other medicines.

The CHMP adopted an opinion by consensus based on the PRAC recommendation.

The Icelandic and Norwegian Members were in agreement with the CHMP recommendation.

The CHMP noted the EMA public health communication.

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

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

Rapporteur: Koenraad Norga, Co-Rapporteur: Andrea Laslop,

Scope: List of Outstanding Issues/Opinion

Prescription status of desloratadine-containing products

Action: For adoption

The CHMP agreed to consult the PRAC and adopted a list of questions to them.

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

No items

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

10.4.1. Paracomb 500mg/150mg film coated tablets - Paracetamol/Ibuprofen 500 mg/150 mg Paracetamol and Ibuprofen - EMEA/H/1447

Vale Pharmaceutical Ltd

Rapporteur: Nithyanandan Nagercoil, Co-Rapporteur: Romaldas Maciulaitis

RMS: UK, CMS: AT, BE, DE, FR, HR, IE, LU, NL, PT, ES

Decentralised Procedure numbers: UK/H/6034-5/001/DC, UK/H/6176/001/DC

Scope: List of Outstanding Issues/Opinion

Disagreement regarding justification for a fixed dose combination, the demonstration of an additional benefit and of an acceptable safety profile

Action: For adoption

The Committee noted a number of issues relating to the application.

The CHMP adopted a list of outstanding issues with a specific timetable.

Submission of responses: 06.04.2017

Re-start of the procedure: 20.04.2017

Rapporteur/co-rapporteur joint assessment report(s) circulated to CHMP: 03.05.2017

Comments: 08.05.2017

Updated Rapporteur/co-rapporteur joint assessment report(s) circulated to CHMP: 11.05.2017

CHMP opinion: May 2017 CHMP

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

10.5.1. Haldol and associated names - haloperidol - EMEA/H/A-30/1393

Janssen-Cilag Group of companies and associated companies

Rapporteur: Martina Weise, Co-Rapporteur: Katarina Vučić,

Scope: Opinion

Harmonisation exercise for Haldol/Haldol Decanoate and associated names (haloperidol). Review triggered by the EC due to the need to harmonise the product information across all Member States, including the therapeutic indication, the posology, the contra-indications, special warnings and precautions for use and pregnancy and lactation and other sections.

Action: For adoption

List of outstanding issues adopted 13.10.2016, 01.04.2016, 26.03.2015. List of Questions

adopted on 26.06.2014

The Committee adopted a positive opinion by consensus recommending that the concerned marketing authorisations should be varied together with the CHMP assessment report and translation timetable.

The Icelandic and Norwegian Members were in agreement with the CHMP recommendation.

The CHMP noted the EMA question and answer document

10.5.2. Haldol decanoate and associated names – haloperidol - EMEA/H/A-30/1405

Janssen-Cilag Group of companies and associated companies

Rapporteur: Martina Weise, Co-Rapporteur: Katarina Vučić,

Scope: Opinion

Harmonisation exercise for Haldol/Haldol Decanoate and associated names (haloperidol). Review triggered by the EC due to the need to harmonise the product information across all Member States, including the therapeutic indication, the posology, the contra-indications, special warnings and precautions for use and pregnancy and lactation and other sections.

Action: For adoption

List of outstanding issues adopted 13.10.2016, 01.04.2016, 26.03.2015. List of Questions adopted on 26.06.2014

The Committee adopted a positive opinion by consensus recommending that the concerned marketing authorisations should be varied together with the CHMP assessment report and translation timetable.

The Icelandic and Norwegian Members were in agreement with the CHMP recommendation.

The CHMP noted the EMA question and answer document

10.5.3. Saroten and associated names - amitriptyline - EMEA/H/A-30/1430

Lundbeck group of companies and associated companies, Bayer Vital and PNG Gerolymatos Medical

Rapporteur: Eleftheria Nikolaidi, Co-Rapporteur: Alar Irs,

Scope: Opinion

Harmonisation exercise for Saroten and associated names (amitriptyline). Review triggered by Greece due to the need to harmonise the product information across all Member States, including the therapeutic indication, the posology, the contraindications, the adverse effects and the recommendations for use.

Action: For adoption

List of outstanding issues adopted on 15.12.2016, 13.10.2016, 23.06.2016, 01.04.2016. List of Questions adopted 17.12.2015

The members discussed the different sections of the SmPC mainly on the indication and posology and agreed that all remaining issues were considered resolved.

The Committee adopted a positive opinion by consensus recommending that the concerned marketing authorisations should be varied together with the CHMP assessment report and translation timetable.

The Icelandic and Norwegian Members were in agreement with the CHMP recommendation.

The CHMP noted the EMA question and answer document.

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

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

Symbiopharm GmbH,

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

Scope: Ad-hoc expert group report from meeting held on 13.01.2017

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

Action: For information

The CHMP noted the report from the expert group meeting held 13.01.2017. The experts explained that the knowledge on probiotics has evolved over time in particular into the direction of more individualized treatment adapted to specific subtypes of the disease and particular patient symptom profiles. The experts acknowledged that there is no treatment for Irritable Bowel Syndrome (IBS) effective in all patients but a treatment algorithm should be followed to putting the treatment with the best evidence of therapeutic benefit first to spare patients unnecessary burden. The experts reiterated that no treatment in IBS is effective in every patient but emphasized that the repertoire of drugs to be chosen from needs to have at least a sensible margin of efficacy to avoid unsuccessful treatment approaches producing unnecessary harm to the patient by symptom continuation.

The CHMP also noted the letter received from the MAH on 22 February 2017.

10.6.2. Vancomycin containing products – (vancomycin) - EMEA/H/A-31/1440

Rapporteur: Concepcion Prieto-Yerro, Co-rapporteur: Alar Irs

Scope: List of Outstanding Issues

Action: For adoption

Review of the benefit-risk balance following notification by the Spanish Agency of Medicines and Medical Devices of a referral under Article 31 of Directive 2001/83/EC.

The CHMP discussed dosing in children and treatment duration. The CHMP noted the input from Modelling and Simulation Working Group and other groups (PKWP, IDWP and PDCO).

The CHMP adopted a 3rd list of outstanding issues with a specific timetable.

Submission of responses: 06.04.2017

Re-start of the procedure: 20.04.2017

Rapporteur/co-rapporteur joint assessment report circulated to CHMP: 26.04.2017

Comments: 03.05.2017

Updated Rapporteur/co-rapporteur joint assessment report circulated to CHMP: 05.05.2017

CHMP LoOI/Opinion: May 2017 CHMP

10.6.3. Paracetamol modified and prolonged release formulations (NAP) - EMEA/H/A-31/1445

GlaxoSmithKline Consumer Healthcare AB (Alvedon, 665 mg modified-release tablet), various

PRAC led referral **-** PRAC Rapporteur: Laurence de Fays, Co-Rapporteur: Ulla Wändel Liminga

Scope: list of experts for the ad-hoc expert group meeting

Action: For adoption

Review of the benefit-risk balance of paracetamol modified release following notification by Sweden of a referral under Article 31 of Directive 2001/83/EC, based on pharmacovigilance data

The CHMP adopted the list of experts for the ad-hoc expert group.

10.6.4. Human coagulation (plasma-derived) factor VIII: human coagulation factor VIII (antihemophilic factor A) (NAP); human coagulation factor VIII (inhibitor bypassing fraction) (NAP); human coagulation factor VIII, human von Willebrand factor - Voncento (CAP) Recombinant factor VIII: antihemophilic factor (recombinant) (NAP); moroctocog alfa – Refacto AF (CAP) octocog alfa – Advate (CAP), Helixate Nexgen (CAP), Iblias (CAP), Kogenate (CAP), Kovaltry (CAP) - EMEA/H/A-31/1448

Baxter AG (Advate), Bayer Pharma AG (Helixate Nexgen, Iblias, Kogenate, Kovaltry), CSL Behring GmbH (Voncento), Pfizer Limited (Refacto AF), various

PRAC led referral - PRAC Rapporteur: Rafe Suvarna; PRAC Co-rapporteur: Brigitte Keller-Stanislawski

Scope: updated List of experts for ad hoc expert group meeting

Action: For adoption

Review of the benefit-risk balance of factor VIII following notification by Germany of a referral under Article 31 of Directive 2001/83/EC, based on pharmacovigilance data

The CHMP adopted the updated list of experts.

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

No items

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

No items

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

No items

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

No items

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

10.11.1. Cardioxane - Dexrazoxane - EMEA/H/A-13/1453

Clinigen Group

Rapporteur: Alexandre Moreau, Co-Rapporteur: Greg Markey

RMS: FR, CMS: CZ, DE, ES, IT, NL, PL & UK

Decentralised Procedure numbers: FR/H/283/01/II/27G

Scope: Start of procedure and appointment of Rapporteurs

Article 13 triggered by the ANSM in France in January 2017 requesting the CHMP's opinion whether the proposed lifting of the contraindication for a subset of anthracycline treated children is justified.

Action: For adoption

The CHMP noted the notification from ANSM triggering a referral under Article 13.

The CHMP appointed Alexandre Moreau (FR) (interest level 1) as Rapporteur and Greg Markey (UK) (interest level 1) as Co-Rapporteur.

The CHMP agreed on a short procedure with a specific timetable.

Start of the procedure (CHMP): 23.02.2017

Rapporteur's preliminary assessment report circulated to CHMP: 08.03.2017

Co-Rapporteur's preliminary assessment report circulated to CHMP: 08.03.2017

Comments: 13.03.2017

Rapporteur's updated assessment report circulated to CHMP: 16.03.2017

Co-Rapporteur's updated assessment report circulated to CHMP: 16.03.2017

CHMP opinion: March 2017 CHMP

11. Pharmacovigilance issue

11.1. Early Notification System

February 2017 Early Notification System on envisaged CHMP/CMDh outcome accompanied by communication to the general public.

Action: For information

The CHMP noted the February 2017 Early Notification System.

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

The CHMP noted the minutes.

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

Scope: ITF briefing meeting

Meeting date: 27 February 2017

Action: For adoption

The CHMP agreed to the ITF briefing meeting.

Scope: ITF briefing meeting

Meeting date: 13 March 2017

Action: For adoption

The CHMP agreed to the ITF briefing meeting.

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

No items

13.4. Nanomedicines activities

No items

14. Organisational, regulatory and methodological matters

14.1. Mandate and organisation of the CHMP

14.1.1. CHMP meetings to be held in Valletta 28 February - 3 March 2017 under the Maltese Presidency of the Council of the European Union

Scope: Information about the draft agenda topics of the upcoming Strategic Review and Learning meeting 28 February - 2 March 2017

The CHMP noted the agenda and update on the Strategic Review and Learning meeting.

Action: For discussion

Scope: Information about the draft agenda topics of the upcoming meeting on - Making Article 58 and other European Medicines Agency outputs more relevant for non-EU regulators to be held in Valetta 2 March - 3 March 2017

Action: For discussion

The CHMP noted the information.

14.1.2. Survey to committee members on the service provided by the Scientific Committees Service

Scope: Findings of the survey to Committee Members

Action: For information

A short survey was sent in July 2016 to all Committee members to gauge the quality of the service offered to the Committees. Overall, the members were satisfied with the service provided by the Secretariats. The CHMP noted the findings of the survey to Committee Members.

14.1.3. Report on Data-sharing initiative in Alzheimer's disease

Scope: A joint SAWP/CNSWP initiative where EMA promoted a series of meetings with developers to critically appraise the methods used in recent Alzheimer's disease programs and to share information in order to inform regulatory guidance.

Action: For information

The CHMP noted the report on the joint SAWP/CNSWP initiative.

14.1.4. CHMP and ORGAM meeting dates 2019-2021

Action: For information

The CHMP adopted the meeting dates for 2019-2021.

14.1.5. Presentation on Experience of PAES

Scope: Review of experience on imposition of PAES.

Action: For information

The CHMP noted the report on experience with PAES.

14.1.6. ATMP guideline on safety and efficacy follow-up and risk management (EMA/CHMP/65416/2016)

Scope: Call for CHMP sponsors for the development of guidance and template

Action: For information

The CHMP noted the call for CHMP sponsors for the ATMPs RMP guidance template and appointed sponsors.

14.1.7. Overview on current activities in Africa

Scope: Summary on regional initiatives, WHO, Bill & Melinda Gates Foundation and other activities

Action: For information

The CHMP noted the overview on current activities in Africa.

14.1.8. CEPI Meeting on Regulatory science gaps

Scope: Participation of Vaccines Working Party member Nele Berthels in CEPI - Finishing the job on Ebola vaccines: CEPI Meeting on Regulatory science gaps in Washington DC, US, 22nd March 2017

Action: For adoption

The CHMP agreed to the participation of Nele Berthels in this event.

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 06-09 February 2017

Action: For information

The CHMP noted the Summary of recommendations and advice of PRAC.

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

Action: For adoption

The CHMP adopted the List of Union Reference Dates and frequency of submission of Periodic Safety Update Reports (EURD list) for February 2017.

14.2.2. Committee for Advanced Therapies (CAT)

CAT draft minutes of meeting held on 15-17 February 2017
Action: For information

The CHMP noted the draft minutes.

14.2.3. Committee for Herbal Medicinal Products (HMPC)

Report from the HMPC meeting held on 30-31 January 2017

Action: For information

The CHMP noted the report.

14.2.4. Paediatric Committee (PDCO)

PIPs reaching D30 at February 2017 PDCO

Action: For information

The CHMP noted the document.

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

Action: For information

The CHMP noted the document.

Joint CHMP/PDCO session

Agenda for joint session

Action: For information

The CHMP/PDCO Joint session took place on Wednesday 22 February from 14:00 – 15:15 in room 3A

14.2.5. Committee for Orphan Medicinal Products (COMP)

Report from the COMP meeting held on 14-16 February 2017
Action: For information

The CHMP noted the report.

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 20-22 February 2017

Action: For information

The CHMP noted the report.

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

14.3.1. Scientific Advice Working Party (SAWP)

Call for interest for nomination of a replacement SAWP member and his alternate following resignation of Mr Thomas Lang. The required area of expertise is statistics.

Action: For information

The letters of candidacy together with the CV of both member and alternate, as per the SAWP Mandate requirements [see Article 2(10)], should be sent, deadline 13 March 2017.

The CHMP noted the call.

Report from the SAWP meeting held on 6-9 February 2017. Table of conclusions

Action: For information

Scientific advice letters: See Annex G

Disclosure of information related to scientific advice letters cannot be released at present time as these contain commercially confidential information.

The CHMP noted the report.

14.3.2. Name Review Group (NRG)

Table of Decisions of the NRG meeting held on 1 February 2017.

Action: For adoption

The CHMP adopted the document.

14.3.3. Blood Products Working Party (BPWP)

Vice - Chair: Karri Penttilä,

Scope: Call for nomination of a new Chairperson of the Blood Products Working Party (BPWP).

Action: For information

Nominations should be sent by 13 March 2017. Elections will take place at March 2017 CHMP.

The CHMP noted the information.

14.3.4. Biologics Working Party (BWP)

Chair: Sol Ruiz

Scope: Election of a new Chairperson of the Biologics Working Party (BWP).

Action: For adoption

The CHMP elected Sol Ruiz (ES) as chair to the BWP.

14.3.5. Gastroenterology Drafting Group (GDG)

Chair: Elmer Schabel

Scope: Election of a new Chairperson to Gastroenterology Drafting Group (GDG)

Action: For adoption

The CHMP elected Mark Ainsworth (DK) as chair to the Gastroenterology Drafting Group.

14.3.6. Pharmacogenomics Working Party (PGWP)

Chair: Krishna Prasad/Markus Paulmichl,

Nomination of two new additional experts: Sir Munir Pirmohamed (UK) and Wilko Weichert (DE)

Action: For adoption

Postponed

14.3.7. Vaccines Working Party (VWP)

Chair: Mair Powell

Nomination of new additional experts: Daniel Brasseur (BE)

Action: For adoption

Postponed

Scope: Letter from the PRAC requesting consultation of the VWP on signal on potential vaccination failure in children with Tick borne encephalitis vaccine (inactivated)

Action: For adoption

The CHMP noted the letter from the PRAC and endorsed the consultation of the VWP.

14.3.8. Rheumatology/Immunology Working Party (RIWP)

Chair: Jan Mueller-Berghaus,

Scope: Review of the composition of drafting groups and temporary working parties in view of goals to be achieved in 2017.

Action: For discussion

Postponed

Scope: Call for nomination of RIWP core member after resignation of Nils Feltelius

Action: For information

Nominations for a RIWP core member should be sent by 31 March 2017.

The CHMP noted the call.

14.3.9. Oncology Working Party (ONCWP)

Chair: Pierre Demolis/Paolo Foggi,

Scope: Concept paper on a proposal to replace the reflection paper on the regulatory guidance for the use of health – related quality of life (HRQL) measures in the evaluation of medicinal products with a new PRO guideline.

Action: For adoption for public consultation

Postponed

14.3.10. Antimicrobial Advice ad hoc Expert Group (AMEG)

Scope: Extension of the AMEG task to update the categorisation of antimicrobials and the proposed early hazard characterisation.

Action: For adoption

The CHMP adopted the extension of the AMEG task to update the categorisation of antimicrobials and the proposed early hazard characterisation.

14.3.11. Quality Working Party (QWP)

Scope: Incompatibility of meropenem and ciprofloxacin leading to possible precipitation when co-administered intravenously – QWP responses to PRAC questions

Action: For adoption

The CHMP adopted the QWP responses to the PRAC.

Scope: ICH Q3D implementation strategy

Action: For information

The CHMP noted the implementation strategy. The document is addressing specific considerations to enable the practical implementation of ICH Q3D Guideline for Elemental Impurities in the European Union. It is intended to provide guidance for Applicants/MAHs, drug product, drug substance and excipient manufacturers, as well as regulators. In addition to new applications, it will also apply to variations to existing marketed authorized medicinal products.

14.3.12. Safety Working Party (SWP)

Chair: Jan Willem Van der Laan / Sonja Beken,

Scope: Nomination of John Jensen (DKMA) as new member of the ERA Drafting Group.

Action: For adoption

The CHMP appointed John Jensen (DKMA) as new member to the ERA Drafting Group.

14.4. Cooperation within the EU regulatory network

No items

14.5. Cooperation with International Regulators

14.5.1. EMA/FDA strategic document on Gaucher disease

Action: For information

The CHMP noted the strategic document. Comments on the agreed principles should be provided by 3 February. The adoption is planned for the March 2017 CHMP meeting.

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

No items

14.7. CHMP work plan

14.7.1. CHMP 2017 Work Plan

Action: For adoption

The CHMP adopted the CHMP work plan for 2017. The work plan will be published on EMA website.

14.8. Planning and reporting

No items

14.9. Others

No items

15. Any other business

15.1. AOB topic

15.1.1. Operation and Relocation Preparedness - Workstream 2 - Operational Preparedness

Action: For discussion

Postponed

16. List of participants

List of participants including any restrictions with respect to involvement of members/alternates/ experts following evaluation of declared interests for the 20 – 23 February 2017 meeting.

Name	Role	Member State or affiliation	Outcome restriction following evaluation of e-DoI	Topics on agenda for which restrictions apply
Tomas Salmonson	Chair	Sweden	No interests declared	
Andrea Laslop	Member	Austria	No interests declared	
Milena Stain	Alternate	Austria	No interests declared	
Bart Van der Schueren	Member	Belgium	No interests declared	
Christophe Focke	Alternate	Belgium	No restrictions applicable to this meeting	
Mila Vlaskovska	Member	Bulgaria	No interests declared	
Katarina Vučić	Member	Croatia	No interests declared	
Selma Arapovic Dzakula	Alternate	Croatia	No interests declared	
Panayiotis Triantafyllis	Member	Cyprus	No interests declared	
Ondřej Slanař	Member	Czech Republic	No interests declared	
Radka Montoniová	Alternate	Czech Republic	No interests declared	
Sinan B. Sarac	Member	Denmark	No interests declared	
Hanne Lomholt Larsen	Alternate	Denmark	No interests declared	
Alar Irs	Member	Estonia	No restrictions applicable to this meeting	
Outi Mäki-Ikola	Member	Finland	No restrictions applicable to this meeting	
Tuomo Lapveteläinen	Alternate	Finland	No interests declared	
Alexandre Moreau	Member	France	No interests declared	
Joseph Emmerich	Alternate	France	No interests declared	
Harald Enzmann	Member (Vice-Chair)	Germany	No interests declared	

Name	Role	Member State or affiliation	Outcome restriction following evaluation of e-DoI	Topics on agenda for which restrictions apply
Martina Weise	Alternate	Germany	No restrictions applicable to this meeting	
Eleftheria Nikolaidi	Member	Greece	No interests declared	
Maria-Dimokleia Ziotopoulou	Alternate	Greece	No interests declared	
Agnes Gyurasics	Member	Hungary	No interests declared	
Hrefna Gudmundsdottir	Alternate	Iceland	No interests declared	
David Lyons	Member	Ireland	No restrictions applicable to this meeting	
Patrick Salmon	Alternate	Ireland	No interests declared	
Daniela Melchiorri	Member	Italy	No restrictions applicable to this meeting	
Juris Pokrotnieks	Member	Latvia	No restrictions applicable to this meeting	
Romaldas Mačiulaitis	Member	Lithuania	No participation in final deliberations and voting on:	5.1.3. Pegasys - peginterferon alfa-2a - EMEA/H/C/000395/II/0091 4.1.1. Esbriet - pirfenidone - Orphan - EMEA/H/C/002154/X/0035/G
Jacqueline Genoux-Hames	Member	Luxembourg	No interests declared	
John Joseph Borg	Member	Malta	No interests declared	
Johann Lodewijk Hillege	Member	Netherlands	No interests declared	

Name	Role	Member State or affiliation	Outcome restriction following evaluation of e-DoI	Topics on agenda for which restrictions apply
Paula Boudewina van Hennik	Alternate	Netherlands	No interests declared	
Bjorg Bolstad	Alternate	Norway	No restrictions applicable to this meeting	
Piotr Fiedor	Member	Poland	No interests declared	
Bruno Sepodes	Member	Portugal	No interests declared	
Fatima Ventura	Alternate	Portugal	No restrictions applicable to this meeting	
Nela Vilceanu	Member	Romania	No interests declared	
Eva Malikova	Alternate	Slovakia	No interests declared	
Stanislav Primožič	Member	Slovenia	No interests declared	
Concepcion Prieto Yerro	Member	Spain	No interests declared	
Aranzazu Sancho-Lopez	Alternate	Spain	No restrictions applicable to this meeting	
Kristina Dunder	Member	Sweden	No interests declared	
Filip Josephson	Alternate	Sweden	No interests declared	
Greg Markey	Member	United Kingdom	No interests declared	
Nithyanandan Nagercoil	Alternate	United Kingdom	No restrictions applicable to this meeting	
Robert James Hemmings	Co-opted member	United Kingdom	No restrictions applicable to this meeting	
Koenraad Norga	Co-opted member	Belgium	No participation in final deliberations and voting on:	
Jan Mueller-Berghaus	Co-opted member	Germany	No interests declared	

Name	Role	Member State or affiliation	Outcome restriction following evaluation of e-DoI	Topics on agenda for which restrictions apply
Jean-Louis Robert	Co-opted member	Luxembourg	No interests declared	
Sol Ruiz	Co-opted member	Spain	No interests declared	
Sabine Mayrhofer	Expert - in person*	Germany	No interests declared	
Sophie Barbou des Courieres	Expert in person	France	No interests declared	
Frederique D Herbe	Expert in person	France	No interests declared	
Camille Thomassin	Expert in person	France	No restrictions applicable to this meeting	
Muriel Uzzan	Expert in person	France	No interests declared	
Sara Franco	Expert in person	France	No interests declared	
Maria Escudero Galindo	Expert in person	Spain	No participation in discussions, final deliberations and voting on:	3.2.9. febuxostat - EMEA/H/C/004374 3.3.5. insulin glargine - EMEA/H/C/004280
Elisabeth Penninga	Expert in person	Denmark	No interests declared	
Mette Tranholm	Expert in person	Denmark	No interests declared	
Joan Deckers	Expert in person	Netherlands	No interests declared	

Name	Role	Member State or affiliation	Outcome restriction following evaluation of e-DoI	Topics on agenda for which restrictions apply
Gertruida (Trudy) Knol	Expert in person	Netherlands	No interests declared	
Peter Salomons	Expert in person	Netherlands	No interests declared	
Stan van Belkum	Expert in person	Netherlands	No interests declared	
Hoebert Sybe Hiemstra	Expert in person	Netherlands	No interests declared	
Karen van der Velden	Expert in person	Netherlands	No restrictions applicable to this meeting	
Fons Wesseling	Expert in person	Netherlands	No interests declared	
Diederick Slijkerman	Expert in person	Netherlands	No interests declared	
Caroline Norén	Expert in person	Sweden	No interests declared	
Jenny F Jansson	Expert in person	Sweden	No interests declared	
Alenoosh Abedi	Expert in person	Sweden	No interests declared	
Anna Mäkinen Salmi	Expert in person	Sweden	No restrictions applicable to this meeting	
Pierre Demolis	Expert via phone	France	No interests declared	
George Aislaitner	Expert - via phone	Greece	No interests declared	
Anja Schiel	Expert - via phone	Norway	No interests declared	
Olivier Le blaye	Expert - via phone	France	No restrictions applicable to this meeting	
Maura O'Donovan	Expert - via phone	Portugal	No interests declared	
Cecilia Chisholm	Expert - via phone	United Kingdom	No interests declared	

Name	Role	Member State or affiliation	Outcome restriction following evaluation of e-DoI	Topics on agenda for which restrictions apply
Antonio Lopez Navas	Expert - via phone	Spain	No interests declared	
Marie-Christine Bielsky	Expert - via phone	United Kingdom	No restrictions applicable to this meeting	
Jan Span	Expert - via phone	Netherlands	No interests declared	
Macarena Rodriguez Mendizabal	Expert - via phone	Spain	No interests declared	
Lisa Rosner	Expert - via phone	Austria	No interests declared	
Bengt Ljungberg	Expert - via phone	Sweden	No interests declared	
Doris Johanna Hovgaard	Expert - via phone	Denmark	No interests declared	
Kristina Bech Jensen	Expert - via phone	Denmark	No interests declared	
Anne Jorunn Stokka	Expert - via phone	Norway	No restrictions applicable to this meeting	
Per Harald Fuglerud	Expert - via phone	Norway	No restrictions applicable to this meeting	
Tania Meier	Expert via Adobe	Germany	No interests declared	
Janet Schriever	Expert via Adobe	Germany	No interests declared	
Susanne Steinecker	Expert via Adobe	Germany	No interests declared	
Bernhardt Sachs	Expert via Adobe	Germany	No interests declared	
Sylvia Kuehn	Expert via Adobe	Germany	No restrictions applicable to this meeting	
Joerg Zinserling	Expert via Adobe	Germany	No interests declared	
Gabriele Schlosser-Weber	Expert via Adobe	Germany	No interests declared	

Name	Role	Member State or affiliation	Outcome restriction following evaluation of e-DoI	Topics on agenda for which restrictions apply
Michael Bühlen	Expert via Adobe	Germany	No interests declared	
Olli Tenhunen	Expert via Adobe	Finland	No interests declared	

Representative from the European Commission attended the meeting

Meeting run with support from relevant EMA staff

* Experts were only evaluated against the product(s) they have been invited to talk about.

17. 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)*

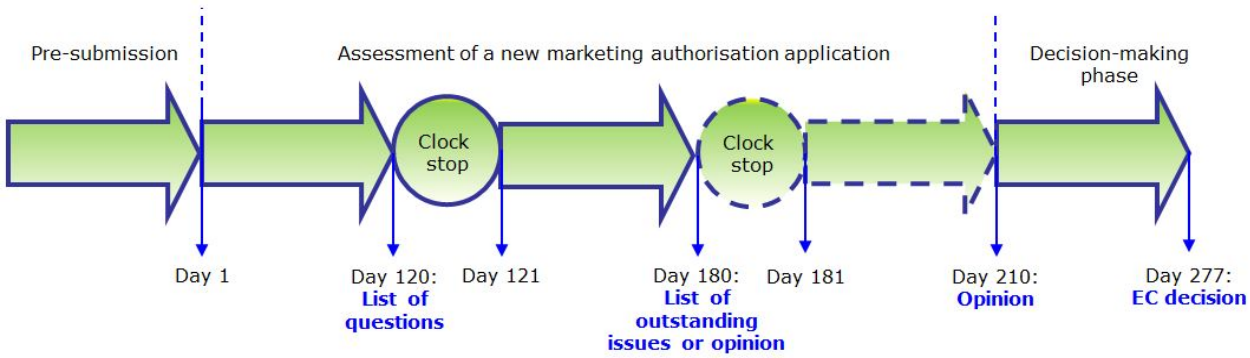
The items listed in this section are those for which marketing authorisation holders (MAHs) or applicants have been invited to the CHMP plenary meeting to address questions raised by the Committee. Oral explanations normally relate to on-going applications (section 3, 4 and 5) or referral procedures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

24 March 2017
EMA/CHMP/211452/2017

ANNEX TO FEBRUARY 2017 CHMP Minutes
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 February 2017: **For adoption**	Adopted.

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

Final Outcome of Rapporteurship allocation for February 2017: **For adoption**	Adopted.

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

Glybera - alipogene tiparvovec - EMEA/H/C/002145/S/0057, Orphan, ATMP MAH: uniQure biopharma B.V., Rapporteur: Christiane Niederlaender, PRAC Rapporteur: Julie Williams Request for Supplementary Information adopted on 20.01.2017.	Positive Opinion adopted by consensus together with the CHMP assessment report. The Marketing Authorisation remains under exceptional circumstances. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP opinion.
Increlex - mecasermin - EMEA/H/C/000704/S/0041, Orphan MAH: Ipsen Pharma, Rapporteur: Outi Mäki-Ikola, PRAC Rapporteur: Kirsti Villikka	Positive Opinion adopted by consensus together with the CHMP assessment report. The Marketing Authorisation remains under exceptional circumstances. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP opinion.
Lojuxta - lomitapide - EMEA/H/C/002578/S/0023 MAH: Aegerion Pharmaceuticals Limited, Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Menno van der Elst Request for Supplementary Information adopted	2nd Request for Supplementary Information adopted

on 23.02.2017, 10.11.2016.	
Obizur - susoctocog alfa - EMEA/H/C/002792/S/0006 MAH: Baxalta Innovations GmbH, Rapporteur: Nithyanandan Nagercoil, PRAC Rapporteur: Brigitte Keller-Stanislawski	Positive Opinion adopted by consensus together with the CHMP assessment report. The Marketing Authorisation remains under exceptional circumstances. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP opinion.

B.2. RENEWALS OF MARKETING AUTHORISATIONS OUTCOMES

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

Kalydeco - ivacaftor - EMEA/H/C/002494/R/0052, Orphan MAH: Vertex Pharmaceuticals (Europe) Ltd., Rapporteur: Concepcion Prieto Yerro, PRAC Rapporteur: Dolores Montero Corominas Request for Supplementary Information adopted on 26.01.2017.	Positive Opinion adopted by consensus together with the CHMP assessment report and translation timetable. Based on the review of the available information the CHMP was of the opinion that an additional five-year renewal was required. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

B.2.2. Renewals of Marketing Authorisations for unlimited validity

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.	Request for Supplementary Information adopted
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.	Request for Supplementary Information adopted
Bretaris Genuair - aclidinium - EMEA/H/C/002706/R/0031 MAH: AstraZeneca AB, Duplicate, Duplicate of Eklira Genuair, Rapporteur: Nithyanandan Nagercoil, Co-Rapporteur: Piotr Fiedor, PRAC Rapporteur: Julie Williams	Positive Opinion adopted by consensus together with the CHMP assessment report and translation timetable. Based on the review of the available information, the CHMP was of the opinion that the renewal of the marketing authorisation can be granted with unlimited validity.

	The Icelandic and Norwegian CHMP Members were in agreement with the CHMP Opinion.
Eklira Genuair - aclidinium - EMEA/H/C/002211/R/0031 MAH: AstraZeneca AB, Rapporteur: Nithyanandan Nagercoil, Co-Rapporteur: Piotr Fiedor, PRAC Rapporteur: Julie Williams	Positive Opinion adopted by consensus together with the CHMP assessment report and translation timetable. Based on the review of the available information, the CHMP was of the opinion that the renewal of the marketing authorisation can be granted with unlimited validity. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP Opinion.
Flebogamma DIF - human normal immunoglobulin - EMEA/H/C/000781/R/0048 MAH: Instituto Grifols, S.A., Rapporteur: Jan Mueller-Berghaus, Co-Rapporteur: Concepcion Prieto Yerro, PRAC Rapporteur: Brigitte Keller-Stanislawski	Positive Opinion adopted by consensus together with the CHMP assessment report and translation timetable. Based on the review of the available information, the CHMP was of the opinion that the renewal of the marketing authorisation can be granted with unlimited validity. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP Opinion.
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.	Request for Supplementary Information adopted
Jakavi - ruxolitinib - EMEA/H/C/002464/R/0032 MAH: Novartis Europharm Ltd, Rapporteur: Filip Josephson, Co-Rapporteur: Robert James Hemmings, PRAC Rapporteur: Ulla Wändel Liminga	Positive Opinion adopted by consensus together with the CHMP assessment report and translation timetable. Based on the review of the available information, the CHMP was of the opinion that the renewal of the marketing authorisation can be granted with unlimited validity. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP Opinion.
Pergoveris - follitropin alfa / lutropin alfa - EMEA/H/C/000714/R/0050 MAH: Merck Serono Europe Limited, Rapporteur: Nithyanandan Nagercoil, Co-Rapporteur: Hanne Lomholt Larsen, PRAC Rapporteur: Julie Williams Request for Supplementary Information adopted on 15.12.2016.	Positive Opinion adopted by consensus together with the CHMP assessment report and translation timetable. Based on the review of the available information, the CHMP was of the opinion that the renewal of the marketing authorisation can be granted with unlimited validity. The Icelandic and Norwegian CHMP Members

	were in agreement with the CHMP Opinion.
Rasilez - aliskiren - EMEA/H/C/000780/R/0112 MAH: Novartis Europharm Ltd, Rapporteur: Daniela Melchiorri, Co-Rapporteur: Melinda Sobor, PRAC Rapporteur: Carmela Macchiarulo Request for Supplementary Information adopted on 23.02.2017.	Request for Supplementary Information adopted
Siklos - hydroxycarbamide - EMEA/H/C/000689/R/0030, Orphan MAH: Addmedica, Rapporteur: Koenraad Norga, Co-Rapporteur: Eleftheria Nikolaidi, PRAC Rapporteur: Jean-Michel Dogné Request for Supplementary Information adopted on 26.01.2017.	Positive Opinion adopted by consensus together with the CHMP assessment report and translation timetable. Based on the review of the available information, the CHMP was of the opinion that the renewal of the marketing authorisation can be granted with unlimited validity. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP Opinion.
Zinforo - ceftaroline fosamil - EMEA/H/C/002252/R/0031 MAH: AstraZeneca AB, Rapporteur: Greg Markey, Co-Rapporteur: Alar Irs, PRAC Rapporteur: Julie Williams	Positive Opinion adopted by consensus together with the CHMP assessment report and translation timetable. Based on the review of the available information, the CHMP was of the opinion that the renewal of the marketing authorisation can be granted with unlimited validity. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP Opinion.
Zoledronic acid medac - zoledronic acid - EMEA/H/C/002359/R/0018 MAH: medac Gesellschaft fur klinische Spezialpraparate mbH, Generic, Generic of Zometa, Rapporteur: Alar Irs, PRAC Rapporteur: Doris Stenver	Positive Opinion adopted by consensus together with the CHMP assessment report and translation timetable. Based on the review of the available information, the CHMP was of the opinion that the renewal of the marketing authorisation can be granted with unlimited validity. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP Opinion

B.2.3. Renewals of Conditional Marketing Authorisations

Darzalex - daratumumab - EMEA/H/C/004077/R/0003, Orphan MAH: Janssen-Cilag International NV, Rapporteur: Sinan B. Sarac, PRAC Rapporteur: Marcia Sofia Sanches de Castro Lopes Silva	Positive Opinion adopted by consensus together with the CHMP assessment report and translation timetable. The CHMP was of the opinion that the renewal for this conditional Marketing Authorisation can be granted.

	The Marketing Authorisation remains conditional. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP Opinion.
Pandemic influenza vaccine H5N1 MedImmune - pandemic influenza vaccine (H5N1) (live attenuated, nasal) - EMEA/H/C/003963/R/0003 MAH: MedImmune LLC, Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Jan Neuhauser	Positive Opinion adopted by consensus together with the CHMP assessment report and translation timetable. The CHMP was of the opinion that the renewal for this conditional Marketing Authorisation can be granted. The Marketing Authorisation remains conditional. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP Opinion.

B.3. POST-AUTHORISATION PHARMACOVIGILANCE OUTCOMES

Signal detection PRAC recommendations on signals adopted at the PRAC meeting held on on 06-09 February 2017 PRAC:	
Opdivo **(EMEA/H/C/003985)** (Nivolumab), MAH: Bristol-Myers Squibb Pharma EEIG, Rapporteur: Aranzazu Sancho-Lopez, Co-Rapporteur: Paula Boudewina van Hennik, Signal of pemphigoid: **For adoption**	Adopted.
PSUR procedures for which PRAC adopted a recommendation for variation of the terms of the MA at its on February 2017 meeting:	
EMEA/H/C/PSUSA/00002665/201607 (rotavirus vaccine monovalent (live, oral)) CAPS: **Rotarix** (EMEA/H/C/000639) (human rotavirus, live attenuated), MAH: GlaxoSmithKline Biologicals S.A., Rapporteur: Bart Van der Schueren, PRAC Rapporteur: Jean-Michel Dogné, "12.07.2015-11.07.2016"	The CHMP, having considered in accordance with Article 28 of Regulation (EC) No 726/2004 the PSUR on the basis of the PRAC recommendation and the PRAC assessment report as appended, recommends by consensus, the variation to the terms of the marketing authorisation(s) for the above mentioned medicinal product(s), concerning the following change(s): Update of section 4.8 of the SmPC to modify the description of the risk of intussusception. The Icelandic and the Norwegian CHMP members agree with the above-mentioned recommendation of the CHMP.
EMEA/H/C/PSUSA/00010035/201607 (ingenol mebutate) CAPS: **Picato** (EMEA/H/C/002275) (ingenol mebutate), MAH: LEO Laboratories Ltd, Rapporteur:	The CHMP, having considered in accordance with Article 28 of Regulation (EC) No 726/2004 the PSUR on the basis of the PRAC recommendation and the PRAC assessment report as appended recommends by consensus, the variation to the

Nithyanandan Nagercoil, PRAC Rapporteur: Julie Williams, “1 February 2016 to 31 July 2016 – DLP 31 July 2016.”	terms of the marketing authorisation for the above mentioned medicinal product, concerning the following change: Update of sections 4.4 of the SmPC to add a warning on keratoacanthoma and of section 5.1 of the SmPC to include information on the high incidence of keratoacanthoma observed in study LP0105-1020 (large treatment area study). The Package leaflet is updated accordingly.
EMEA/H/C/PSUSA/00010404/201607 (atazanavir / cobicistat) CAPS: **EVOTAZ** (EMEA/H/C/003904) (atazanavir / cobicistat), MAH: Bristol-Myers Squibb Pharma EEIG, Rapporteur: Bruno Sepodes, PRAC Rapporteur: Caroline Laborde, “14 January 2016 - 28 July 2016”	The CHMP, having considered in accordance with Article 28 of Regulation (EC) No 726/2004 the PSUR on the basis of the PRAC recommendation and the PRAC assessment report as appended, recommends by consensus, the variation to the terms of the marketing authorisation(s) for the above mentioned medicinal product(s), concerning the following change(s): Update of section 4.6 of the SmPC to add information that atazanavir is detected in human milk. The Package leaflet is updated accordingly. The Icelandic and the Norwegian CHMP members agree with the above-mentioned recommendation of the CHMP.
EMEA/H/C/PSUSA/00010447/201607 (brivaracetam) CAPS: **Briviact** (EMEA/H/C/003898) (brivaracetam), MAH: UCB Pharma S.A., Rapporteur: Filip Josephson, PRAC Rapporteur: Adam Przybylkowski, “15/01/2016-14/07/2016”	The CHMP, having considered in accordance with Article 28 of Regulation (EC) No 726/2004 the PSUR on the basis of the PRAC recommendation and the PRAC assessment report as appended , recommends by consensus the variation to the terms of the marketing authorisation(s) for the above mentioned medicinal product(s), concerning the following change(s): Based on the PRAC Rapporteur review of data on safety and efficacy, the PRAC considers that the risk-benefit balance of medicinal product containing brivaracetam remains unchanged but recommends that the terms of the marketing authorisations should be varied as follows: Update of section 4.8 of the SmPC to add safety information regarding the risks of Type I hypersensitivity and an update of section 4 of the Package Leaflet accordingly. Clarifications on the third BRV metabolite (hydroxy acid) have also been provided in section 5.2 of the SmPC. The Icelandic and the Norwegian CHMP members agree with the above-mentioned recommendation of the CHMP.

B.4. EPARs / WPARs

Afstyla - lonoctocog alfa - EMEA/H/C/004075 Applicant: CSL Behring GmbH, treatment of haemophilia A, Rapporteur: Jan Mueller-Berghaus, Co-Rapporteur: Tuomo Lapveteläinen, New active substance (Article 8(3) of Directive No 2001/83/EC)	adopted.
AMGEVITA - adalimumab - EMEA/H/C/004212 Applicant: Amgen Europe B.V., treatment of rheumatoid arthritis, juvenile idiopathic arthritis, axial spondyloarthritis, psoriatic arthritis, psoriasis, paediatric plaque psoriasis, hidradenitis suppurativa, Crohn's disease, paediatri Crohn's disease and Ulcerative colitis, Similar biological application (Article 10(4) of Directive No 2001/83/EC)	adopted.
Daptomycin Hospira - daptomycin - EMEA/H/C/004310 Applicant: Hospira UK Limited, treatment of complicated skin and soft-tissue infections, Generic, Generic of Cubicin, Generic application (Article 10(1) of Directive No 2001/83/EC)	adopted.
Enpaxiq - pacritinib - EMEA/H/C/004193, Orphan Applicant: CTI Life Sciences Limited, treatment of myelofibrosis, New active substance (Article 8(3) of Directive No 2001/83/EC) **WPAR**	adopted.
Jylamvo - methotrexate - EMEA/H/C/003756 Applicant: Therakind Limited, treatment of rheumatological and dermatological diseases, Hybrid application (Article 10(3) of Directive No 2001/83/EC)	adopted.
Rolufta - umeclidinium - EMEA/H/C/004654 Applicant: GlaxoSmithKline Trading Services Limited, treatment of chronic obstructive pulmonary disease (COPD), Informed consent application (Article 10c of Directive No 2001/83/EC)	adopted.
SOLYMBIC - adalimumab - EMEA/H/C/004373 Applicant: Amgen Europe B.V., treatment of rheumatoid arthritis, juvenile idiopathic arthritis,	adopted.

axial spondyloarthritis, psoriatic arthritis, psoriasis, paediatric plaque psoriasis, hidradenitis suppurativa, Crohn's disease, paediatri Crohn's disease and Ulcerative colitis, Duplicate, Duplicate of AMGEVITA, Similar biological application (Article 10(4) of Directive No 2001/83/EC)	
Tadalafil Lilly - tadalafil - EMEA/H/C/004666 Applicant: Eli Lilly Nederland B.V., treatment of erectile dysfunction and treatment of the signs and symptoms of benign prostate hyperplasia, Informed Consent of Cialis, Informed consent application (Article 10c of Directive No 2001/83/EC)	adopted.
Xeljanz - tofacitinib - EMEA/H/C/004214 Applicant: Pfizer Limited, treatment of active rheumatoid arthritis, New active substance (Article 8(3) of Directive No 2001/83/EC)	adopted.
Yargesa - miglustat - EMEA/H/C/004016 Applicant: JensonR+ Limited, treatment of Gaucher disease, Generic, Generic of Zavesca, Generic application (Article 10(1) of Directive No 2001/83/EC)	adopted.

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

Adcetris - brentuximab vedotin - EMEA/H/C/002455/II/0041/G, Orphan MAH: Takeda Pharma A/S, Rapporteur: Paula Boudewina van Hennik Opinion adopted on 16.02.2017.	Positive Opinion adopted by consensus on 16.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Advate - octocog alfa - EMEA/H/C/000520/II/0082/G MAH: Baxter AG, Rapporteur: Jan Mueller-Berghaus Request for Supplementary Information adopted on 23.02.2017.	Weekly start timetable. The Committee adopted a Request for Supplementary information together with a specific timetable.
Bemfola - follitropin alfa - EMEA/H/C/002615/II/0011 MAH: Gedeon Richter Plc., Rapporteur: Paula Boudewina van Hennik	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

Opinion adopted on 23.02.2017.	
Cerezyme - imiglucerase - EMEA/H/C/000157/II/0099/G MAH: Genzyme Europe BV, Rapporteur: Johann Lodewijk Hillege Request for Supplementary Information adopted on 16.02.2017.	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 Request for Supplementary Information adopted on 23.02.2017.	Weekly start timetable. The Committee adopted a Request for Supplementary information together with a specific timetable.
Colobreathe - colistimethate sodium - EMEA/H/C/001225/II/0023 MAH: Teva B.V., Rapporteur: Nithyanandan Nagercoil Opinion adopted on 23.02.2017. Request for Supplementary Information adopted on 17.11.2016, 12.05.2016, 03.03.2016.	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Cosentyx - secukinumab - EMEA/H/C/003729/II/0017 MAH: Novartis Europharm Ltd, Rapporteur: Tuomo Lapveteläinen Opinion adopted on 16.02.2017.	Positive Opinion adopted by consensus on 16.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Darzalex - daratumumab - EMEA/H/C/004077/II/0004, Orphan MAH: Janssen-Cilag International NV, Rapporteur: Sinan B. Sarac Opinion adopted on 23.02.2017.	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Darzalex - daratumumab - EMEA/H/C/004077/II/0005/G, Orphan MAH: Janssen-Cilag International NV, Rapporteur: Sinan B. Sarac Request for Supplementary Information adopted on 23.02.2017.	Weekly start timetable. The Committee adopted a Request for Supplementary information together with a specific timetable.
Emtricitabine/Tenofovir disoproxil Zentiva - emtricitabine / tenofovir disoproxil - EMEA/H/C/004137/II/0001 MAH: Zentiva k.s., Generic, Generic of Truvada, Rapporteur: Alar Irs Request for Supplementary Information adopted on 23.02.2017.	Weekly start timetable. The Committee adopted a Request for Supplementary information together with a specific timetable.
Fabrazyme - agalsidase beta - EMEA/H/C/000370/II/0093 MAH: Genzyme Europe BV, Rapporteur: Johann	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP

Lodewijk Hillege Opinion adopted on 23.02.2017. Request for Supplementary Information adopted on 19.01.2017.	recommendation.
HBVAXPRO - hepatitis B vaccine (rDNA) - EMEA/H/C/000373/II/0055 MAH: MSD Vaccins, Rapporteur: Jan Mueller-Berghaus Opinion adopted on 02.02.2017.	Positive Opinion adopted by consensus on 02.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Hemoprostol - misoprostol - EMEA/H/W/002652/II/0006/G MAH: Linepharma International Limited, Rapporteur: Paula Boudewina van Hennik Opinion adopted on 23.02.2017.	Positive Opinion adopted by consensus on 23.02.2017. 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. The Committee adopted a Request for Supplementary information together with a specific timetable.
Herceptin - trastuzumab - EMEA/H/C/000278/II/0127/G MAH: Roche Registration Limited, Rapporteur: Jan Mueller-Berghaus Opinion adopted on 23.02.2017.	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Hizentra - human normal immunoglobulin - EMEA/H/C/002127/II/0074/G MAH: CSL Behring GmbH, Rapporteur: Jan Mueller-Berghaus Request for Supplementary Information adopted on 23.02.2017.	The Committee adopted a Request for Supplementary information together with a specific timetable.
IDELVION - albutrepenonacog alfa - EMEA/H/C/003955/II/0003/G, Orphan MAH: CSL Behring GmbH, Rapporteur: Jan Mueller-Berghaus Request for Supplementary Information adopted on 23.02.2017.	Weekly start timetable. The Committee adopted a Request for Supplementary information together with a specific timetable.
Increlex - mecasermin - EMEA/H/C/000704/II/0046/G, Orphan MAH: Ipsen Pharma, Rapporteur: Outi Mäki-Ikola Request for Supplementary Information adopted on 23.02.2017.	The Committee adopted a Request for Supplementary information together with a specific timetable.
Inhixa - enoxaparin sodium - EMEA/H/C/004264/II/0005/G MAH: Techdow Europe AB, Duplicate, Duplicate of Thorinane, Rapporteur: Andrea Laslop	Weekly start timetable. The Committee adopted a Request for Supplementary information together with a specific timetable.

Request for Supplementary Information adopted on 23.02.2017.	
Kanuma - sebelipase alfa - EMEA/H/C/004004/II/0006/G, Orphan MAH: Alexion Europe SAS, Rapporteur: Bart Van der Schueren Opinion adopted on 23.02.2017. Request for Supplementary Information adopted on 10.11.2016.	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Lantus - insulin glargine - EMEA/H/C/000284/II/0107/G MAH: Sanofi-aventis Deutschland GmbH, Rapporteur: Johann Lodewijk Hillege Opinion adopted on 16.02.2017.	Positive Opinion adopted by consensus on 16.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Nimenrix - meningococcal group A, C, W135 and Y conjugate vaccine - EMEA/H/C/002226/II/0062 MAH: Pfizer Limited, Rapporteur: Greg Markey Opinion adopted on 09.02.2017.	Positive Opinion adopted by consensus on 09.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
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.	The Committee adopted a Request for Supplementary information together with a specific timetable.
OPDIVO - nivolumab - EMEA/H/C/003985/II/0022/G MAH: Bristol-Myers Squibb Pharma EEIG, Rapporteur: Aranzazu Sancho-Lopez Opinion adopted on 02.02.2017. Request for Supplementary Information adopted on 01.12.2016.	Positive Opinion adopted by consensus on 02.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Orencia - abatacept - EMEA/H/C/000701/II/0106/G MAH: Bristol-Myers Squibb Pharma EEIG, Rapporteur: Outi Mäki-Ikola Request for Supplementary Information adopted on 02.02.2017.	Weekly start timetable. The Committee adopted a Request for Supplementary information together with a specific timetable.
Pheburane - sodium phenylbutyrate - EMEA/H/C/002500/II/0014 MAH: Lucane Pharma, Rapporteur: David Lyons Opinion adopted on 02.02.2017. Request for Supplementary Information adopted on 10.11.2016.	Positive Opinion adopted by consensus on 02.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

Prevenar 13 - pneumococcal polysaccharide conjugate vaccine (13-valent, adsorbed) - EMEA/H/C/001104/II/0147/G MAH: Pfizer Limited, Rapporteur: Kristina Dunder Opinion adopted on 02.02.2017. Request for Supplementary Information adopted on 15.12.2016.	Positive Opinion adopted by consensus on 02.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Privigen - human normal immunoglobulin - EMEA/H/C/000831/II/0111 MAH: CSL Behring GmbH, Rapporteur: Jan Mueller-Berghaus Opinion adopted on 09.02.2017.	Positive Opinion adopted by consensus on 09.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Retacrit - epoetin zeta - EMEA/H/C/000872/II/0075 MAH: Hospira UK Limited, Rapporteur: Martina Weise Opinion adopted on 23.02.2017.	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Rivastigmine 1A Pharma - rivastigmine - EMEA/H/C/001181/II/0022/G MAH: 1 A Pharma GmbH, Informed Consent of Exelon, Rapporteur: Alexandre Moreau Opinion adopted on 09.02.2017.	Positive Opinion adopted by consensus on 09.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Rivastigmine Hexal - rivastigmine - EMEA/H/C/001182/II/0023/G MAH: Hexal AG, Informed Consent of Exelon, Rapporteur: Alexandre Moreau Opinion adopted on 09.02.2017.	Positive Opinion adopted by consensus on 09.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Rivastigmine Sandoz - rivastigmine - EMEA/H/C/001183/II/0024/G MAH: Sandoz GmbH, Informed Consent of Exelon, Rapporteur: Alexandre Moreu Opinion adopted on 09.02.2017.	Positive Opinion adopted by consensus on 09.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
RoActemra - tocilizumab - EMEA/H/C/000955/II/0067/G MAH: Roche Registration Limited, Rapporteur: Jan Mueller-Berghaus Opinion adopted on 23.02.2017.	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Silapo - epoetin zeta - EMEA/H/C/000760/II/0044 MAH: STADA Arzneimittel AG, Rapporteur: Martina Weise Opinion adopted on 23.02.2017.	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Tenofovir disoproxil Zentiva - tenofovir disoproxil - EMEA/H/C/004120/II/0001 MAH: Zentiva k.s., Generic, Generic of Viread, Rapporteur: John Joseph Borg	Weekly start timetable. The Committee adopted a Request for Supplementary information together with a specific timetable.

Request for Supplementary Information adopted on 23.02.2017.	
Tysabri - natalizumab - EMEA/H/C/000603/II/0098/G MAH: Biogen Idec Ltd, Rapporteur: Jan Mueller-Berghaus Opinion adopted on 02.02.2017. Request for Supplementary Information adopted on 24.11.2016.	Positive Opinion adopted by consensus on 02.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Velphoro - mixture of polynuclear iron(III)-oxyhydroxide, sucrose and starches - EMEA/H/C/002705/II/0009/G MAH: Vifor Fresenius Medical Care Renal Pharma France, Rapporteur: Johann Lodewijk Hillege Opinion adopted on 23.02.2017.	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS0954 **Filgrastim Hexal-EMEA/H/C/000918/WS0954/0033** **Zarzio-EMEA/H/C/000917/WS0954/0034** MAH: SANDOZ GmbH, Lead Rapporteur: Greg Markey Opinion adopted on 23.02.2017. Request for Supplementary Information adopted on 10.11.2016.	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1043/G **Helixate NexGen-EMEA/H/C/000276/WS1043/0182/G** **KOGENATE Bayer-EMEA/H/C/000275/WS1043/0189/G** MAH: Bayer Pharma AG, Lead Rapporteur: Jan Mueller-Berghaus Opinion adopted on 02.02.2017. Request for Supplementary Information adopted on 24.11.2016.	Positive Opinion adopted by consensus on 02.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1068/G **Infanrix hexa-EMEA/H/C/000296/WS1068/0216/G** MAH: GSK Biologicals SA, Lead Rapporteur: Bart Van der Schueren Opinion adopted on 23.02.2017.	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

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

Azarga - brinzolamide / timolol - EMEA/H/C/000960/II/0034	Weekly start timetable. The Committee adopted a Request for Supplementary information

MAH: Alcon Laboratories (UK) Ltd, Rapporteur: Hanne Lomholt Larsen, “Update of sections 4.2, 4.4, 4.6 and 4.8 of the SmPC following a review of the safety profile taking into consideration data from clinical studies and post-marketing experience. The Package Leaflet has been updated accordingly. In addition, the MAH took the opportunity to implement minor editorial changes in the SmPC and Package Leaflet and to update the contact details for the local representative in Spain in the Package Leaflet.” Request for Supplementary Information adopted on 16.02.2017.	together with a specific timetable.
Cerdelga - eliglustat - EMEA/H/C/003724/II/0008, Orphan MAH: Genzyme Europe BV, Rapporteur: Johann Lodewijk Hillege, “Update of section 5.1 of the SmPC to include 2, 3 and 4 years composite stability endpoint data based on the final results of the ENCORE study.” Opinion adopted on 23.02.2017. Request for Supplementary Information adopted on 15.12.2016, 13.10.2016.	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Cervarix - human papillomavirus vaccine [types 16, 18] (recombinant, adjuvanted, adsorbed) - EMEA/H/C/000721/II/0075 MAH: GSK Biologicals SA, Rapporteur: Bart Van der Schueren, “Update of section 5.1 of the SmPC with wording on the clinical efficacy of Cervarix in women aged 26 years and older, based on the submission of the final report for study HPV-015 (MEA 083); this is a phase III, double-blind, randomized, controlled study to evaluate the safety, immunogenicity and efficacy of HPV16/18 L1/AS04 vaccine administered intramuscularly according to a three-dose schedule (0, 1, 6 month) in healthy adult female subjects aged 26 years and above.” Opinion adopted on 23.02.2017. Request for Supplementary Information adopted on 13.10.2016, 23.06.2016, 25.02.2016.	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Cervarix - human papillomavirus vaccine [types 16, 18] (recombinant, adjuvanted, adsorbed) - EMEA/H/C/000721/II/0080 MAH: GSK Biologicals SA, Rapporteur: Bart Van der Schueren, “Submission of final Study report for study HPV-060. Study HPV-060 is an extension of the study HPV-014 (EXT 014 Y5-10). Study HPV-014 with 4 years post-vaccination	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

data was submitted as a commitment in November 2009 (EMEA/H/C/721/FU2 20.5) The purpose of this variation is to fulfil the Post-Authorization Measure (PAM) (MEA-082) with the long term follow up (10 years post-vaccination) data from study HPV-060. GlaxoSmithKline Biologicals (GSK Biologicals) considers that there is no need to change the SmPC at this stage." Opinion adopted on 23.02.2017. Request for Supplementary Information adopted on 10.11.2016, 15.09.2016.	
Cimzia - certolizumab pegol - EMEA/H/C/001037/II/0057/G MAH: UCB Pharma S.A., Rapporteur: Kristina Dunder - 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.	The Committee adopted a Request for Supplementary information together with a specific timetable.
Descovy - emtricitabine / tenofovir alafenamide - EMEA/H/C/004094/II/0013 MAH: Gilead Sciences International Ltd, Rapporteur: Robert James Hemmings, "Submission of 96 week data from Study GS-US-311-1089 in order to support an update of the virological outcomes and measures of bone mineral density in Section 5.1 of the Summary of Product Characteristics (SmPC)." Opinion adopted on 16.02.2017.	Positive Opinion adopted by consensus on 16.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Dynastat - parecoxib - EMEA/H/C/000381/II/0068/G MAH: Pfizer Limited, Duplicate, Duplicate of Xapit, Rapporteur: David Lyons, "C.I.4 – Update of section 4.4 of the SmPC in order to update the safety information related to alcohol use and gastrointestinal (GI) risk. C.I.4 - Update of section 4.6 of the SmPC in order to update the safety information related to oligohydramnios if the product is used during second or third trimester of pregnancy. 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, to bring the PI in line with the latest QRD template version 10.0 and to correct some mistakes." Opinion adopted on 23.02.2017.	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

Fycompa - perampanel - EMEA/H/C/002434/II/0034/G MAH: Eisai Europe Ltd., Rapporteur: Robert James Hemmings, "Update of sections 4.5 and 5.1 of the SmPC in order to add information on the conversion of patients to Fycompa monotherapy (E2007-G000-504, hereby Study 504) and to include the effect of withdrawal of concomitant enzyme-inducing antiepileptic drugs (EIAEDs) on plasma concentrations of perampanel (A supportive analysis, CPMS-E2007-0013R). In addition, the Marketing authorisation holder (MAH) took the opportunity to bring the PI in line with the latest QRD template version 10." Opinion adopted on 23.02.2017. Request for Supplementary Information adopted on 15.12.2016.	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Giotrif - afatinib - EMEA/H/C/002280/II/0022 MAH: Boehringer Ingelheim International GmbH, Rapporteur: Filip Josephson, "Update of section 5.1 of the SmPC in order to update the information about the major mechanism of acquired resistance to afatinib. In addition, the Marketing authorisation holder (MAH) took the opportunity to add the side effects 'itching' and 'dry skin' with frequency very common to the package leaflet to bring it in line with the SmPC." Opinion adopted on 23.02.2017.	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
GONAL-f - follitropin alfa - EMEA/H/C/000071/II/0136 MAH: Merck Serono Europe Limited, Rapporteur: Nithyanandan Nagercoil, "Update of the SmPC sections 4.4 and 4.8 to revise the frequency of thromboembolic events from 'very rare' to 'rare'. 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." Request for Supplementary Information adopted on 02.02.2017.	Weekly start timetable. The Committee adopted a Request for Supplementary information together with a specific timetable.
Harvoni - ledipasvir / sofosbuvir - EMEA/H/C/003850/II/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	The Committee adopted a Request for Supplementary information together with a specific timetable.

SOLAR-1 and SOLAR-2." Request for Supplementary Information adopted on 23.02.2017, 10.11.2016.	
Harvoni - ledipasvir / sofosbuvir - EMEA/H/C/003850/II/0046 MAH: Gilead Sciences International Ltd, Rapporteur: Filip Josephson, "Submission of the final clinical study report of the study GS-US-337-1118: an Open-Label, Multicenter Study To Evaluate The Efficacy And Safety Of Sofosbuvir/Ledipasvir Fixed-Dose Combination ± Ribavirin For 12 or 24 Weeks In Chronic Genotype 1 HCV Infected Subjects Who Participated In A Prior Gilead-Sponsored HCV Treatment Study" Opinion adopted on 23.02.2017.	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Iressa - gefitinib - EMEA/H/C/001016/II/0026 MAH: AstraZeneca AB, Rapporteur: Filip Josephson, "Submission of final study report for IMPRESS study (D791LC00001) and discussion to address one of the 'PRAC Recommendations as per procedure regarding the gefitinib Periodic Safety Update Report (PSUR: EMA/PRAC/4284/2016). No Changes in the PI and in the RMP are proposed" Opinion adopted on 23.02.2017. Request for Supplementary Information adopted on 13.10.2016.	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Jardiance - empagliflozin - EMEA/H/C/002677/II/0025 MAH: Boehringer Ingelheim International GmbH, Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Dolores Montero Corominas, "Submission of the final results for a non-interventional study 1245.122 exploring the characteristics of patients initiating empagliflozin or other noninsulin glucose lowering drugs in the United Kingdom in order to fulfil MEA 009. The RMP (version 11.0) is updated accordingly." Request for Supplementary Information adopted on 23.02.2017.	The Committee adopted a Request for Supplementary information together with a specific timetable.
Keytruda - pembrolizumab - EMEA/H/C/003820/II/0013 MAH: Merck Sharp & Dohme Limited, Rapporteur: Daniela Melchiorri, "Update of section 4.4 of the SmPC to amend existing warnings on immune-related adverse reactions. In addition, the MAH took the opportunity to revise the instructions for handling and storage	Positive Opinion adopted by consensus on 02.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

after reconstitution in SmPC sections 6.3 and 6.6 for increased clarity. The Package Leaflet has been updated accordingly." Opinion adopted on 02.02.2017. Request for Supplementary Information adopted on 17.11.2016.	
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. The Committee adopted a Request for Supplementary information together with a specific timetable.
Mosquirix - plasmodium falciparum and hepatitis B vaccine (recombinant, adjuvanted) - EMEA/H/W/002300/II/0018 MAH: GSK Biologicals SA, Rapporteur: Jan Mueller-Berghaus, "Submission of the final study report of study Malaria-057, a phase 2, open, randomised, controlled, multi-centre study to evaluate the safety and immunogenicity of 7 infant immunisation schedules of the RTS,S/AS01E candidate vaccine against P. falciparum in one study centre with 2 sites in Blantyre, Malawi. No changes to the product information or the RMP were proposed. The data is submitted to fulfil post-approval measure MEA 010." Opinion adopted on 16.02.2017.	Positive Opinion adopted by consensus on 16.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
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. The Committee adopted a Request for Supplementary information together with a specific timetable.

Mysimba - naltrexone hydrochloride / bupropion hydrochloride - EMEA/H/C/003687/II/0011 MAH: Orexigen Therapeutics Ireland Limited, Rapporteur: Hanne Lomholt Larsen, "Submission of study report NaltrexBuprop-4001 - A Multicenter, Randomized, Double-blind, Placebo controlled, Phase 4 Study to Assess the Effect of Naltrexone Hydrochloride and Bupropion Hydrochloride Extended Release Combination on the Occurrence of Major Adverse Cardiovascular Events in Overweight and Obese Subjects with Cardiovascular Disease. The product information remains unchanged." Opinion adopted on 23.02.2017.	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
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.	The Committee adopted a Request for Supplementary information together with a specific timetable.
Nulojix - belatacept - EMEA/H/C/002098/II/0038 MAH: Bristol-Myers Squibb Pharma EEIG, Rapporteur: Filip Josephson, PRAC Rapporteur: Ulla Wändel Liminga, "Update of sections 4.8 and 5.1 of the SmPC in order to update the safety information following the completion of the IM103-008 and IM103_027post-authorization efficacy studies. The Package Leaflet and and Risk Management Plan (Version 12) are updated accordingly." Opinion adopted on 23.02.2017. Request for Supplementary Information adopted on 15.09.2016.	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
OPDIVO - nivolumab - EMEA/H/C/003985/II/0023 MAH: Bristol-Myers Squibb Pharma EEIG, Rapporteur: Aranzazu Sancho-Lopez, "Update of sections 4.8 and 5.1 of the SmPC in order to update the safety and pharmacological information with the 24 months data from the	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

completed NSCLC studies CA209017 and CA209057." Opinion adopted on 23.02.2017. Request for Supplementary Information adopted on 19.01.2017.	
Pradaxa - dabigatran etexilate - EMEA/H/C/000829/II/0097 MAH: Boehringer Ingelheim International GmbH, Rapporteur: Hanne Lomholt Larsen, "Submission of final study report of study 1160.173 "A prospective, open label study to evaluate the pharmacokinetics of dabigatran in non-valvular atrial fibrillation (NVAF) patients with severely impaired renal function on dabigatran etexilate 75 mg BID therapy"." Opinion adopted on 02.02.2017. Request for Supplementary Information adopted on 01.12.2016.	Positive Opinion adopted by consensus on 02.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Praluent - alirocumab - EMEA/H/C/003882/II/0018 MAH: sanofi-aventis groupe, Rapporteur: Johann Lodewijk Hillege, "Submission of study PDY13670 a Phase 1 study of the effects of subcutaneous doses of alirocumab on lipid and lipoprotein metabolism in adults with mildly elevated LDL-cholesterol." Opinion adopted on 23.02.2017.	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Prevenar 13 - pneumococcal polysaccharide conjugate vaccine (13-valent, adsorbed) - EMEA/H/C/001104/II/0145 MAH: Pfizer Limited, Rapporteur: Kristina Dunder, "Update of the SmPC section 5.1 to add information related to antibiotic susceptibility. Editorial changes have also been proposed throughout the SmPC." Opinion adopted on 23.02.2017. Request for Supplementary Information adopted on 15.12.2016, 13.10.2016.	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Prevenar 13 - pneumococcal polysaccharide conjugate vaccine (13-valent, adsorbed) - EMEA/H/C/001104/II/0149 MAH: Pfizer Limited, Rapporteur: Kristina Dunder, "Submission of the final clinical study report (CSR) of study B1851018, a Phase 4 study evaluating the impact of 13vPnC in reducing AOM and NP colonisation caused by S pneumoniae in healthy children, in accordance with the Pharmacovigilance plan outlined in the EU RMP (version 11.0)."	The Committee adopted a Request for Supplementary information together with a specific timetable.

Request for Supplementary Information adopted on 23.02.2017.	
Rapamune - sirolimus - EMEA/H/C/000273/II/0163/G MAH: Pfizer Limited, Rapporteur: Kristina Dunder, "Update of section 4.4 of the SmPC to update the current warning on angioedema to include a possible dose-dependent effect between sirolimus and angioedema based on post-marketing data. Update of section 4.8 of the SmPC to include 'neuroendocrine carcinoma of the skin' as new adverse drug reaction (ADR) with a frequency 'not known' and to replace the ADR 'skin cancer' by 'non-melanoma skin cancer' and 'malignant melanoma' with a 'common' and 'uncommon' frequency respectively, based on post-marketing data. The Package Leaflet is updated accordingly. In addition, the Marketing authorisation holder (MAH) took the opportunity to combine the 0.5 mg, 1 mg and 2 mg tablets SmPC, to fully detail all components of the Rapamune printing ink in section 6.1 of the SmPC and in the Package Leaflet, to align the wording in section 4 of the Package Leaflet with section 4.8 of the SmPC regarding Clostridium difficile, to update the list of local representatives for the Czech Republic, Norway and Sweden in the Package Leaflet and to bring the product information in line with the latest QRD template version 10." Opinion adopted on 16.02.2017. Request for Supplementary Information adopted on 01.12.2016.	Positive Opinion adopted by consensus on 16.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Revestive - teduglutide - EMEA/H/C/002345/II/0037, Orphan MAH: Shire Pharmaceuticals Ireland Ltd, Rapporteur: Sinan B. Sarac, "Update of section 4.2 of the SmPC in order to amend the recommendation that treatment effect should be evaluated after 12 months (instead of the current recommended 6 months) based on literature references." Request for Supplementary Information adopted on 23.02.2017.	The Committee adopted a Request for Supplementary information together with a specific timetable.
Simponi - golimumab - EMEA/H/C/000992/II/0072 MAH: Janssen Biologics B.V., Rapporteur: Kristina Dunder, "Update of section 4.4 of the SmPC in order to include reports of Merkel cell	Positive Opinion adopted by consensus on 02.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

carcinoma in patients treated with TNF blocking agents including Simponi. In addition the frequency of this ADR has been reclassified from “not known” to “rare” in section 4.8 of the SmPC. The Package Leaflet is updated accordingly. Finally the Marketing Authorisation Holder (MAH) took the opportunity to make a small correction in section 5.1 of the SmPC.” Opinion adopted on 02.02.2017.	
Thalidomide Celgene - thalidomide - EMEA/H/C/000823/II/0050, Orphan MAH: Celgene Europe Limited, Rapporteur: Alexandre Moreau, “Submission of a final clinical study report for Study CC-2001-CP-001 together with the population pharmacokinetics (PK) meta-analysis CC-2001-MPK-001 and bioanalytical report CC-2001-CP-001-BA undertaken to evaluate thalidomide PK in multiple myeloma subjects in order to fulfil legally binding measure LEG 027.3.” Opinion adopted on 23.02.2017.	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Vimpat - lacosamide - EMEA/H/C/000863/II/0066/G MAH: UCB Pharma S.A., Rapporteur: Filip Josephson, “Update of section 4.2 of the SmPC in order to update the safety information regarding the use of lacosamide in patients with hepatic impairment, section 4.8 to add a new adverse drug reaction (hepatic enzyme increased (> 2x ULN)) and section 4.9 regarding lacosamide overdose based on postmarketing reports. 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 for Lithuania, Latvia, Estonia, Portugal and Finland.” Opinion adopted on 23.02.2017. Request for Supplementary Information adopted on 15.12.2016.	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Zaltrap - aflibercept - EMEA/H/C/002532/II/0035 MAH: Sanofi-Aventis Groupe, Rapporteur: Filip Josephson, “Update the Product Information (SmPC, section 5.1 Pharmacodynamic properties) to reflect the results of the biomarker programme encompassing the EFC10262, EFC10668 and EFC11338 studies in order to fulfil the Annex II condition of Zaltrap, aflibercept 25 mg/ml, Concentrate for solution for infusion	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

(EMEA/H/C/002532).” Opinion adopted on 23.02.2017.	
Zavicefta - ceftazidime / avibactam - EMEA/H/C/004027/II/0002 MAH: AstraZeneca AB, Rapporteur: Robert James Hemmings, “Update of section 4.4 of the SmPC to revise the paragraph on limitations of clinical data for hospital acquired pneumonia (HAP) indication, section 4.8 of the SmPC to change the frequency from uncommon to common for thrombocytopia and pruritus and section 5.1 of the SmPC to add a new section for HAP/VAP pathogens. The SmPC update is based on the availability of the final CSR for REPROVE (D4281C00001) an updated modelling and simulation report (CAZMS - 09). The Package Leaflet (section 4) is updated accordingly. Study D4281C00001 is a PAES detailed in Annex II.D, therefore an update of Annex II.D is also proposed. In addition, The MAH took the opportunity to add ‘Dilute before use’ to section 5 of the outer Packaging - Carton and to update the list of local representatives in the Package Leaflet.” Opinion adopted on 23.02.2017.	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Zeffix - lamivudine - EMEA/H/C/000242/II/0068 MAH: Glaxo Group Ltd, Duplicate, Duplicate of Epivir, Rapporteur: Joseph Emmerich, “Update of sections 4.4 and 4.6 of the SmPC to reflect pregnancy clinical outcome data from the Antiretroviral Pregnancy Registry (APR); in addition, an introductory paragraph for pregnancy has been added to section 4.6 of the SmPC in line with the Epivir Product Information (lamivudine for Human Immunodeficiency Virus Indication) (variation EMEA/H/C/000107/II/84).” Opinion adopted on 02.02.2017.	Positive Opinion adopted by consensus on 02.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Zelboraf - vemurafenib - EMEA/H/C/002409/II/0039 MAH: Roche Registration Limited, Rapporteur: Filip Josephson, “Update of sections 4.2, 4.8, 5.1 and 5.2 of the SmPC in order to include the paediatric clinical data from the Zelboraf NO25390 (BRIM-P) after request during assessment as per procedure EMEA/H/C/002409/P46/033.” Opinion adopted on 23.02.2017.	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Zepatier - elbasvir / grazoprevir -	Weekly start timetable. The Committee adopted

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." Request for Supplementary Information adopted on 23.02.2017.	a Request for Supplementary information together with a specific timetable.
Zoely - nomegestrol acetate / estradiol - EMEA/H/C/001213/II/0037 MAH: Teva B.V., Rapporteur: Joseph Emmerich, "Update of sections 4.4 and 4.5 of the SmPC with revised information regarding interactions with concomitant medications and risk of reduced efficacy. Further, the current paragraph 'laboratory tests' was moved from section 4.5 to section 4.4 of the SmPC. The Package Leaflet has been updated accordingly." Opinion adopted on 09.02.2017. Request for Supplementary Information adopted on 10.11.2016.	Positive Opinion adopted by consensus on 09.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Zoely - nomegestrol acetate / estradiol - EMEA/H/C/001213/II/0038 MAH: Teva B.V., Rapporteur: Joseph Emmerich, "Update of sections 4.4 and 4.5 of the SmPC concerning Hepatitis C and the risk of elevated ALT due to treatment with the HCV combination regimen ombitasvir/paritaprevir/ritonavir co-administered with ethinylestradiol-containing products. The Package Leaflet has been updated accordingly. In addition, the MAH took the opportunity to make minor editorial changes in the SmPC and Package Leaflet." Opinion adopted on 09.02.2017. Request for Supplementary Information adopted on 10.11.2016.	Positive Opinion adopted by consensus on 09.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1010 **Descovy-EMEA/H/C/004094/WS1010/0006** **Genvoya-EMEA/H/C/004042/WS1010/0017** **Odefsey-EMEA/H/C/004156/WS1010/0004** MAH: Gilead Sciences International Ltd, Lead	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

Rapporteur: Robert James Hemmings, “Update of section 5.2 of the SmPC in order to provide the final results from Study GS-US-320-1615 “A Phase 1, Open-Label, Parallel-Group, Single Dose Study to Evaluate the Pharmacokinetics of Tenofovir Alafenamide (TAF) in Subjects with Normal Hepatic Function and Subjects with Severe Hepatic Impairment”. In addition, the Worksharing applicant (WSA) took the opportunity to update section 4.2 of the SmPC for Descovy to allow dosing in patients with severe hepatic impairment. The information from the CSR for Study GS-US-320-1615 does lead to the addition or deletion of a safety concern in the corresponding RMPs.” Opinion adopted on 23.02.2017. Request for Supplementary Information adopted on 10.11.2016.	
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.	The Committee adopted a Request for Supplementary information together with a specific 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	The Committee adopted a Request for Supplementary information together with a specific timetable.

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.	
WS1091 **Clopidogrel** **Zentiva-EMEA/H/C/000975/WS1091/0056** **Clopidogrel/Acetylsalicylic acid** **Zentiva-EMEA/H/C/001144/WS1091/0048** **DuoPlavin-EMEA/H/C/001143/WS1091/0047** **Iscover-EMEA/H/C/000175/WS1091/0129** **Plavix-EMEA/H/C/000174/WS1091/0125** MAH: Sanofi Clir SNC, Lead Rapporteur: Bruno Sepodes, "Update of section 4.1 to clarify the indication and specify that clopidogrel is indicated for the secondary prevention of atherothrombotic events. In addition, the MAH took the opportunity to update the details of the German local representative in the Clopidogrel/Acetylsalicylic acid Zentiva Package Leaflet." Opinion adopted on 23.02.2017.	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1105 **IntronA-EMEA/H/C/000281/WS1105/0107** **PegIntron-EMEA/H/C/000280/WS1105/0128** **ViraferonPeg-EMEA/H/C/000329/WS1105/0121** MAH: Merck Sharp & Dohme Limited, Lead Rapporteur: Filip Josephson, "Update of sections 4.2 and 4.8 of the SmPC in order to update the safety information with information on HCV/HBV co-infection, and to add an ADR on hepatitis B reactivation in HCV/HBV co-infected patients as post marketing adverse experience respectively. The Package Leaflet and Labelling are updated accordingly. In addition, the Worksharing applicant (WSA) took the opportunity to bring the PI in line with the latest QRD template version 10 including the implementation of the use of combined SmPCs and PLs for PegIntron and ViraferonPeg and the use of combined SmPCs for Intron A in multidose pen."	Weekly start timetable. The Committee adopted a Request for Supplementary information together with a specific timetable.

Request for Supplementary Information adopted on 23.02.2017.	
WS1107/G **Prezista-EMEA/H/C/000707/WS1107/0085/G** **Rezolsta-EMEA/H/C/002819/WS1107/0017/G** MAH: Janssen-Cilag International NV, Lead Rapporteur: Johann Lodewijk Hillege, "Update of sections 4.3 and 4.5 of the SmPC with contra-indication and information of drug-drug interactions of boosted darunavir with elbasvir/grazoprevir (Zepatier) and with lurasidone (Latuda). The PL was updated accordingly. Update of section 4.5 of the Prezista SmPC regarding the drug-drug interaction of boosted darunavir with corticosteroids in line with the PRAC Recommendation for Rezolsta. In addition, the MAH took the opportunity of this variation, for both products, to add information regarding alfuzosin in section 4.5 in line with section 3, to add inhibition of CYP2D6 for the alfa 1 adrenoreceptor antagonist and to correct the frequency of the adverse event osteonecrosis. Section 4.5 of Prezista was also updated to align information between the different formulations and with Rezolsta. An error was correct in section 5.2. The MAH also took the opportunity to update the Product Information with the lasts QRD templates version 9.1 and 10. The contact of the Dutch local representative in the PL was updated." Opinion adopted on 23.02.2017.	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
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 section 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 PI	Weekly start timetable. The Committee adopted a Request for Supplementary information together with a specific timetable.

in line with the latest QRD template, including combining the SmPC of the different strengths, as well as implement minor editorial changes and reformatting of some sections of the SmPC. The Portuguese local representative in the PL has been updated." Request for Supplementary Information adopted on 16.02.2017.	
WS1114 **Exviera-EMEA/H/C/003837/WS1114/0025** **Viekirax-EMEA/H/C/003839/WS1114/0030** MAH: AbbVie Ltd., Lead Rapporteur: Filip Josephson, "Update of section 4.2 of the SmPC to add that treatment duration of 8 weeks may be considered in previously untreated genotype 1b-infected patients with minimal to moderate fibrosis, supported by the results of the GARNET study (M15-684). Consequently the section 5.1 of the SmPC is updated to reflect the results of this study. The Package Leaflet is updated accordingly." Opinion adopted on 23.02.2017.	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

B.5.3. CHMP-PRAC assessed procedures

Adcetris - brentuximab vedotin - EMEA/H/C/002455/II/0043, Orphan MAH: Takeda Pharma A/S, Rapporteur: Paula Boudewina van Hennik, PRAC Rapporteur: Sabine Straus, "Update of sections 4.8 and 5.1 of the SmPC in order to add data from study C25007. The RMP (version 8.0) was updated accordingly. The submission of the clinical study report fulfils SOB 011 of the conditional marketing authorisation for Adcetris." Request for Supplementary Information adopted on 23.02.2017.	The Committee adopted a Request for Supplementary information together with a specific timetable.
ELOCTA - efmoroctocog alfa - EMEA/H/C/003964/II/0010 MAH: Swedish Orphan Biovitrum AB (publ), Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Rafe Suvarna, "Submission of the final Clinical Study Report of study 997HA307 to investigate PK of the 1000 and 3000 IU vial strengths and evaluate safety of rFVIIIFc. Study 997HA307 is listed as an additional PhV activity (category 3 study, MEA 003) in the Risk Management Plan, therefore an updated RMP is	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

included (ver. 1.5)." Opinion adopted on 23.02.2017.	
Epclusa - sofosbuvir / velpatasvir - EMEA/H/C/004210/II/0003 MAH: Gilead Sciences International Ltd, Rapporteur: Filip Josephson, PRAC Rapporteur: Ana Sofia Diniz Martins, "Update of section 5.1 of the SmPC in order to reflect on emerging clinical data from study GS-US-342-1202 investigating efficacy and safety in subjects with chronic hepatitis C virus (HCV) and human immunodeficiency virus (HIV)-1 coinfection. In addition, minor editorial changes are implemented throughout the Product Information." Opinion adopted on 23.02.2017. Request for Supplementary Information adopted on 15.12.2016.	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
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.	The Committee adopted a Request for Supplementary information together with a specific timetable.
EVARREST - human fibrinogen / human thrombin - EMEA/H/C/002515/II/0027/G MAH: Omrix Biopharmaceuticals N. V., Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Brigitte Keller-Stanislawski, "Group of variations consisting of: 1) Submission of the final results for study	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

BIOS-13-005 updating the efficacy and safety information 2) Submission of the final results for study BIOS-13-004 updating the efficacy and safety information 3) Submission of the final results for study 400-12-002 updating the efficacy and safety information 4) Submission of the final results for study 400-12-005 updating the safety information 5) Update of section 5.1 of the SmPC to include further information on main existing efficacy studies Sections 4.8, 5.1 of the SmPC are affected by this group of variations. In addition, the Product Information has been updated in accordance with the QRD template, version 10 and Guideline on core SmPC for plasma-derived fibrin/sealant/haemostatic products (EMA/CHMP/BPWP/598816/2010 rev.1). Section 4.2 has been updated regarding the paediatric information for children under the aged of 1 month, according to the EMA waiver. A revised RMP (version 3) is also introduced, including consequential and routine changes." Opinion adopted on 23.02.2017. Request for Supplementary Information adopted on 10.11.2016.	
Herceptin - trastuzumab - EMEA/H/C/000278/II/0126 MAH: Roche Registration Limited, Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Brigitte Keller-Stanislawski, "C.I.13. Submission of the final study report for the PrefHer study (MO22982); a category 3 study in the RMP to fulfill a required additional pharmacovigilance activity. The PrefHer study is a Phase II, randomized, multicenter, open-label, two-cohort, two-arm, crossover study designed to investigate patient preference for Herceptin intravenous (IV) or Herceptin subcutaneous (SC) administered using the three-weekly (q3w) dosing regimen via the single-use injection device (SID) or from the vial via hand-held syringe, and to compare Health Care professional (HCP) satisfaction and perceived time savings with the two methods of administration in patients with HER2-positive early breast cancer (EBC) in the neoadjuvant/adjuvant setting.	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

The study also evaluated the safety and efficacy (event-free survival) of Herceptin SC and IV. The crossover design of the study also allowed an evaluation of the safety and tolerability of switching between the Herceptin IV and the Herceptin SC formulations, and vice versa." Opinion adopted on 23.02.2017.	
Imbruvica - ibrutinib - EMEA/H/C/003791/II/0025, Orphan MAH: Janssen-Cilag International NV, Rapporteur: Filip Josephson, PRAC Rapporteur: Julie Williams, "Update of the SmPC section 4.4 to remove the warning and precaution regarding the effect of Ibrutinib on the QT interval and section 5.1 to provide additional information regarding the pharmacodynamic effect of Ibrutinib on QT/QTc intervals and cardiac electrophysiology. No changes to the Annex III Package Leaflet are proposed." Opinion adopted on 23.02.2017. Request for Supplementary Information adopted on 10.11.2016, 15.09.2016.	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Increlex - mecasermin - EMEA/H/C/000704/II/0044/G, Orphan MAH: Ipsen Pharma, Rapporteur: Outi Mäki-Ikola, PRAC Rapporteur: Kirsti Villikka, "Update of section 4.4 of the SmPC in order to update the warning regarding antibody response to injected IGF-1. Submission of an updated RMP version 9 , including the educational materials, to update the instructions for antibody testing and improve wording and advices." Request for Supplementary Information adopted on 23.02.2017.	The Committee adopted a Request for Supplementary information together with a specific timetable.
Jardiance - empagliflozin - EMEA/H/C/002677/II/0026 MAH: Boehringer Ingelheim International GmbH, Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Dolores Montero Corominas, "Submission of the final results of a non-clinical study on the effect of empagliflozin on blood ketone level at refeeding after a fasting period, comparison between refeeding with glucose or fat in order to fulfil MEA 010. The RMP (version 11.0) is updated accordingly." Request for Supplementary Information adopted on 23.02.2017.	The Committee adopted a Request for Supplementary information together with a specific timetable.

Jevtana - cabazitaxel - EMEA/H/C/002018/II/0034 MAH: Sanofi-Aventis Groupe, Rapporteur: Alexandre Moreau, PRAC Rapporteur: Caroline Laborde, "Update of sections 4.2, 4.8 and 5.1 of the SmPC in order to add information from completed study EFC11785 (Randomized, open-label multicenter study comparing cabazitaxel at 20 mg/m2 and at 25 mg/m2 every 3 weeks in combination with prednisone for the treatment of metastatic castration-resistant prostate cancer previously treated with a docetaxel-containing regimen). In addition, the MAH is proposing to modify the wording in section 4.1 of the indication from "hormone refractory" to "castration resistant" prostate cancer to reflect current terminology of the disease in the clinical practice. The RMP is updated accordingly and in accordance with the request from the latest PSUR procedure (EMEA/C/H/002018/PSUSA/000476/201506)" Opinion adopted on 23.02.2017. Request for Supplementary Information adopted on 10.11.2016, 15.09.2016.	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
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 "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.	The Committee adopted a Request for Supplementary information together with a specific timetable.
Orkambi - lumacaftor / ivacaftor - EMEA/H/C/003954/II/0017 MAH: Vertex Pharmaceuticals (Europe) Ltd., Rapporteur: Nithyanandan Nagercoil, PRAC Rapporteur: Almath Spooner, "Update of sections 4.8 and 5.1 of the SmPC in order to reflect the long-term safety and efficacy data from Study VX12 809 105. Study VX12 809 105 is a Phase 3, rollover study to evaluate the safety and efficacy of long term treatment with LUM/IVA in subjects	The Committee adopted a Request for Supplementary information together with a specific timetable.

aged 12 years and older with cystic fibrosis, homozygous or heterozygous for the F508del CFTR mutation (MEA 001). A new version of the RMP (ver. 2.7) included in this submission has been updated to include the final data from Study 105. In addition, the MAH took the opportunity to bring the PI in line with the latest QRD template version 10." Request for Supplementary Information adopted on 23.02.2017.	
Perjeta - pertuzumab - EMEA/H/C/002547/II/0028 MAH: Roche Registration Limited, Rapporteur: Sinan B. Sarac, PRAC Rapporteur: Doris Stenver, "Final Clinical Study Report the TRYPHAENA study(BO22280) A randomised, multicentre, multinational Phase II study to evaluate pertuzumab in combination with trastuzumab, given either concomitantly or sequentially with standard anthracyclinebased chemotherapy or concomitantly with a nonanthracycline-based chemotherapy regimen, as neoadjuvant therapy for patients with locally advanced, inflammatory or early stage HER2-positive breast cancer. The RMP (v 8) has been updated to reflect the completion of the study." Opinion adopted on 23.02.2017.	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Saxenda - liraglutide - EMEA/H/C/003780/II/0011 MAH: Novo Nordisk A/S, Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Menno van der Elst, "Based on submission of the LEADER clinical study results (EX2211-3748: liraglutide effect on and action in diabetes, evaluation of cardiovascular outcome results), changes to sections 4.4, and 5.1 of the SmPC are being proposed in order to update the safety information and include a description of the clinical study outcomes. The Package Leaflet and Labelling are updated accordingly. In addition, the Marketing authorisation holder (MAH) took the opportunity to implement minor editorial changes throughout the product information. The LEADER study was included in the liraglutide RMP as a required pharmacovigilance activity (category 3) to specifically address the important potential risk of cardiovascular disorders in patients with Type 2 Diabetes Mellitus. Updates	The Committee adopted a Request for Supplementary information together with a specific timetable.

to the liraglutide RMP based on the study results are also proposed: this variation application fulfils two post-approval commitments in relation to the cardiovascular outcomes trial (MEA 002), as well as to provide additional information on the breast cancer cases found in LEADER (MEA 005). RMP Version 27 was submitted with the application. These liraglutide RMP modifications are in line with the proposed updates to the Saxenda Product Information described above." Request for Supplementary Information adopted on 23.02.2017.	
Senshio - ospemifene - EMEA/H/C/002780/II/0012/G MAH: Shionogi Limited, Rapporteur: Paula Boudewina van Hennik, PRAC Rapporteur: Julie Williams, "- Update of section 4.5 of the SmPC in relation to CYP3A4 based on the results of study E1508I0242and in fulfilment of PAM 008. - Update of section 5.2 of the SmPC with information on ospemifene metabolism and excretion based on the results of study E1508I0242 in fulfilment of PAM 013 and PAM 014. - Update of section 5.2 of the SmPC with information on ospemifene distribution based on the results of studies OSP-PF-046-N and OSP-PF-047-N in fulfilment of PAM 006 and PAM 007. - Update of section 5.2 of the SmPC based on the results of the bile salt export pump (BSEP) transporter study OSP-PF-041-N in fulfilment of PAM 009. As a consequence, an updated RMP version 1.2 is provided accordingly." Opinion adopted on 02.02.2017. Request for Supplementary Information adopted on 10.11.2016.	Positive Opinion adopted by consensus on 02.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
TAGRISSO - osimertinib - EMEA/H/C/004124/II/0009/G MAH: AstraZeneca AB, Rapporteur: Aranzazu Sancho-Lopez, PRAC Rapporteur: Sabine Straus, "Update of SmPC sections 4.2, 4.4, 4.8, 5.1 and 5.2 based on the results from study D5160C00003 (AURA3) and the updated CSRs for studies D5160C00001 (AURAex) and D5160C00002 (AURA2). The Package Leaflet has been updated accordingly. In addition, the MAH took the opportunity to make editorial changes in the SmPC and Package Leaflet. The provision of	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

the CSR from study AURA3 addressed the remaining Specific Obligation for Tagrisso and hence it is recommended to convert the Marketing Authorisation from a Conditional Marketing Authorisation to a Marketing Authorisation not subject to Specific Obligations. Annex II has been updated in accordance. An updated RMP version 6.0 was agreed during the procedure." Opinion adopted on 23.02.2017. Request for Supplementary Information adopted on 15.12.2016.	
TECFIDERA - dimethyl fumarate - EMEA/H/C/002601/II/0035 MAH: Biogen Idec Ltd, Rapporteur: Martina Weise, PRAC Rapporteur: Martin Huber, "To update section 4.8 (Undesirable effects) of the SmPC under the sub-heading 'Tabulated summary of adverse reactions', to include 'liver function abnormalities' as an adverse event, observed in the post-marketing setting, and under the sub-heading 'Hepatic transaminases' to clarify events not observed in placebo-controlled studies. The package leaflet has been updated accordingly (section 4 under heading 'Possible side effects'). The MAH has also taken the opportunity to make minor administrative changes in the package leaflet and to review and update the status timelines of clinical and nonclinical study reports in the Risk Management Plan (v8)." Request for Supplementary Information adopted on 23.02.2017, 15.12.2016.	The Committee adopted a Request for Supplementary information together with a specific timetable.
Torisel - temsirolimus - EMEA/H/C/000799/II/0063, Orphan MAH: Pfizer Limited, Rapporteur: Harald Enzmann, PRAC Rapporteur: Martin Huber, "Submission of final results from Study 3066K1-4438-WW (B1771007) titled "A Randomized Phase 4 Study Comparing 2 Intravenous Temsirolimus (TEMSR) Regimens in Subjects with Relapsed, Refractory Mantle Cell Lymphoma" and fulfilment of obligation to conduct post authorisation measure ANX 027.2. The MAH also evaluated the toxic effects of interest [e.g., bleeding, infection- and mucositis-related events] for study 3066K1-4438-WW (Post-Authorisation Measure, MEA 028) together with a review discussing	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

potential new safety concerns arising from the results. The Risk Management Plan (RMP, v.3.0) is updated accordingly to add myocardial infarction and cardiovascular events in patient with coexisting cardiovascular conditions as important potential risks, and anaemia, thrombocytopenia, hypercholesterolemia, and hypertriglyceridemia as important identified risks. In addition, the Marketing authorisation holder (MAH) took the opportunity to update the list of local representatives in the Package Leaflet." Opinion adopted on 23.02.2017. Request for Supplementary Information adopted on 13.10.2016.	
Tysabri - natalizumab - EMEA/H/C/000603/II/0095 MAH: Biogen Idec Ltd, Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Brigitte Keller-Stanislawski, "Update of section sections 4.2, 4.3, 4.8, 5.1 and 5.2 of the SmPC based on the results of paediatric studies 101MS028 and 101MS328, in accordance with paediatric investigation plan (EMEA-001095-PIP-12). The Package Leaflet has been updated accordingly. The MAH also took the opportunity to make minor amendments in the SmPC and to update the contact details of the local representative in Denmark in the Package Leaflet. An updated RMP version 21 was agreed as part of the procedure." Opinion adopted on 23.02.2017. Request for Supplementary Information adopted on 10.11.2016, 15.09.2016, 23.06.2016.	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Vectibix - panitumumab - EMEA/H/C/000741/II/0080 MAH: Amgen Europe B.V., Rapporteur: Robert James Hemmings, PRAC Rapporteur: Julie Williams, "Update of Annex II in order to provide the results of biomarker analyses from the Vectibix clinical programme including Study 20080763 (according to Supplementary Statistical Analysis Plan dated 20 September 2013), Study 20070820 and Study 20060447. The data submitted are in fulfilment of Annex II obligation ANX017. The Risk Management Plan (version 21.0) has been updated accordingly. The requested variation proposed amendments to Annex II and the Risk Management Plan." Opinion adopted on 23.02.2017.	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

Request for Supplementary Information adopted on 10.11.2016.	
Xadago - safinamide - EMEA/H/C/002396/II/0014 MAH: Zambon SpA, Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Almath Spooner, "Submission of study VDD4193 (Safinamide: In Vitro Metabolic Stability in Human Cryopreserved Hepatocytes, by Fatty Acid Amide Hydrolase enzyme (FAAH), Recombinant Human N-Acylethanolamine Acid Amidase (NAAA) and Recombinant Human Acid Ceramidase (ASAHl)) conducted in order to identify specific substances blocking the amidases (inhibitors of amidases) involved in the metabolism of safinamide. The study fulfils the MEA 001.2." Opinion adopted on 23.02.2017. Request for Supplementary Information adopted on 26.01.2017.	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Zykadia - ceritinib - EMEA/H/C/003819/II/0006/G MAH: Novartis Europharm Ltd, Rapporteur: Aranzazu Sancho-Lopez, PRAC Rapporteur: Ulla Wändel Liminga, "Update of section 4.5 of the SmPC based on the final results of the clinical pharmacology study LDK378A2113 and results of a sub-group evaluating the impact of gastric PH-elevating agents on the steady-state PK, efficacy, and safety of ceritinib in ALK-positive NSCLC patients. The provision of the final CSR for study CLDK378A2113 addresses the post-authorisation measure (PAM) MEA 003. In addition, an updated RMP version 8.0 was agreed during the procedure." Opinion adopted on 23.02.2017. Request for Supplementary Information adopted on 10.11.2016, 21.07.2016, 26.05.2016.	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
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	The Committee adopted a Request for Supplementary information together with a specific timetable.

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.	
WS0993 **Adcirca-EMEA/H/C/001021/WS0993/0025** **Cialis-EMEA/H/C/000436/WS0993/0085** MAH: Eli Lilly Nederland B.V., Lead Rapporteur: Concepcion Prieto Yerro, Lead PRAC Rapporteur: Dolores Montero Corominas, "Update of section 4.4 of the SmPC in order to add a new warning on the risk of non-arteritic anterior ischemic optic neuropathy (NAION) based on the final results of study H6D-MC- LVHQ (category 3 study). In addition the Worksharing applicant (WSA) took the opportunity to update the RMP (version 8.0) accordingly. In addition the Worksharing applicant (WSA) took the opportunity to align the Package Leaflet with the SmPC of Adcirca and Cialis regarding the adverse drug reaction (ADR) 'priapism' and of Cialis only for the ADR 'prolonged erection', to make corrections in the German annexes and to align the product information with the latest QRD template version 10. The Icelandic and the Norwegian CHMP members agree with the above-mentioned recommendation of the CHMP on variation to the terms of the marketing authorisation." Opinion adopted on 23.02.2017. Request for Supplementary Information adopted on 10.11.2016, 15.09.2016.	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1101 **Relvar** **Ellipta-EMEA/H/C/002673/WS1101/0029** **Revinty**	The Committee adopted a Request for Supplementary information together with a specific timetable.

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.	

B.5.4. PRAC assessed procedures

PRAC Led **Adempas - riociguat - EMEA/H/C/002737/II/0014, Orphan** MAH: Bayer Pharma AG, PRAC Rapporteur: Julie Williams, , "Submission of a revised RMP in order to add Off-label use in patients with idiopathic pulmonary pneumonia, with or without pulmonary hypertension as an important identified risk." Opinion adopted on 23.02.2017. Request for Supplementary Information adopted on 10.11.2016.	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
PRAC Led **ATryn - antithrombin alfa - EMEA/H/C/000587/II/0027** MAH: GTC Biotherapeutics UK Limited, Rapporteur: Alexandre Moreau, PRAC Rapporteur: Caroline Laborde, , "Introduction of the first version of the RMP following request in 6th Annual Re-assessment EMEA/H/C/000587/S/0021 and second renewal EMEA/H/C/000587/R/0024. The product information has been updated accordingly to list the risk minimisation measures prior commercialisation of ATryn in each member state. In addition, the due date for the submission of the specific obligation reflected in section E of	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

Annex II of the PI to extend the indication to the peri-partum period has been extended from 31st March 2017 to 31st March 2020." Opinion adopted on 23.02.2017. Request for Supplementary Information adopted on 15.12.2016, 15.09.2016.	
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.	The Committee adopted a Request for Supplementary information together with a specific timetable.
PRAC Led **Eperzan - albiglutide - EMEA/H/C/002735/II/0028/G** MAH: GlaxoSmithKline Trading Services, PRAC Rapporteur: Julie Williams, , "II: C.I.11.b - Submission of a revised RMP in order to introduce the additional risk minimisation measures addressing the important potential risk of medication errors. Annex II of the Product Information is updated accordingly. II: C.I.11.b – Update of the RMP to add a new category 3 study as an additional pharmacovigilance activity - Study 204879: A Randomized, Open-label, Active-Controlled, Parallel-Group, Exploratory Study on the Effects of Repeated Doses of Albiglutide compared to Exenatide on Gastric Myoelectrical Activity and Gastric Emptying in Subjects with Type 2 Diabetes Mellitus II: C.I.11.b - Update of the RMP to add a new category 3 study as an additional pharmacovigilance activity – Study 201840 - An Exploratory Randomized, 2-Part, Single-blind, 2-Period Crossover Study Comparing the Effect of Albiglutide with Exenatide on Regional Brain Activity Related to Nausea in Healthy Volunteers II: C.I.11.b – Update of the RMP to add a new category 3 study as an additional	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

pharmacovigilance activity – Cross-sectional survey to assess the effectiveness of the proposed additional educational materials using Patient Connect” Opinion adopted on 23.02.2017. Request for Supplementary Information adopted on 15.12.2016.	
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.	The Committee adopted a Request for Supplementary information together with a specific timetable.
PRAC Led **Remicade - infliximab - EMEA/H/C/000240/II/0201/G** MAH: Janssen Biologics B.V., Rapporteur: Kristina Dunder, PRAC Rapporteur: Ulla Wändel Liminga, , “Submission of the clinical study reports for C0168T45 and C0168T62 together with an overall summary and evaluation of the complete long term safety follow-up programs for Remicade (as per MEA 79). Study C0168T45 (RESULTS: REMICADE Safety Under Long term Study) is a Multicenter International Observational Study of the Long-term Safety of Infliximab Study C0168T62 (RESULTS UC: REMICADE Safety Under Long-term Study in Ulcerative Colitis) is a Multicenter International Study of the Long-term Safety of Infliximab in Ulcerative Colitis. The RMP (RMP 14.0) has been updated to reflect the completion of these studies.” Request for Supplementary Information adopted on 23.02.2017.	The Committee adopted a Request for Supplementary information together with a specific timetable.
PRAC Led	Positive Opinion adopted by consensus on

Thyrogen - thyrotropin alfa - EMEA/H/C/000220/II/0088 MAH: Genzyme Europe BV, Rapporteur: Patrick Salmon, PRAC Rapporteur: Almath Spooner, , "To transfer the RMP to the latest RMP template. As a consequence, gastrointestinal symptoms, constitutional symptoms, and injection site reactions have been downgraded to identified risks, not categorized as important and therefore have been deleted. In addition, "perceived lower TSH elevation after thyrotropin alfa administration" does not correspond to a safety risk for the patients treated with Thyrogen and was also deleted from the list of important potential risks. Finally, study results and completion date of T4 study have been included and as a consequence, "Use of Thyrogen for remnant ablation in patients originally diagnosed with T4N0-1M0-1 thyroid cancer" was removed from the missing information section. RMP version 9.0 is being submitted." Opinion adopted on 23.02.2017. Request for Supplementary Information adopted on 15.12.2016.	23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
PRAC Led **Trobalt - retigabine - EMEA/H/C/001245/II/0045** MAH: Glaxo Group Ltd, PRAC Rapporteur: Doris Stenver, , "Submission of a revised RMP (version 18) in order to remove a postauthorisation study (PASS) RTG116158, an open label study evaluating the effects of ezogabine/retigabine added to existing anti-epileptic drug(s) on urinary voiding function in subjects with partial onset seizures. In addition, routines change have also been introduced." Opinion adopted on 23.02.2017. Request for Supplementary Information adopted on 10.11.2016.	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
PRAC Led **Xeplion - paliperidone - EMEA/H/C/002105/II/0031** MAH: Janssen-Cilag International NV, Rapporteur: Kristina Dunder, PRAC Rapporteur: Qun-Ying Yue, , "Submission of final study report "Post-Authorization Safety Study Using European Union Databases to Assess the Risk of Cardiovascular and Cerebrovascular Adverse Events in Elderly Patients Treated with	The Committee adopted a Request for Supplementary information together with a specific timetable.

Paliperidone Palmitate, Paliperidone Prolonged-Release, and Other Antipsychotics”. No changes in the PI are proposed.” Request for Supplementary Information adopted on 23.02.2017.	
PRAC Led **Zaltrap - aflibercept - EMEA/H/C/002532/II/0034** MAH: Sanofi-Aventis Groupe, Rapporteur: Filip Josephson, PRAC Rapporteur: Ulla Wändel Liminga, , “Submission of the final results of the Drug Utilisation Study monitoring the use of Zaltrap in cancer patients including potential off-label use and evaluating the potential for intravitreal use. This fulfils the post authorisation commitment MEA 03.” Opinion adopted on 23.02.2017.	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
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.	The Committee adopted a Request for Supplementary information together with a specific timetable.

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, Request for Supplementary Information adopted on 17.02.2017.	The Committee adopted a Request for Supplementary information together with a specific timetable.

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

WS0934/G **Suboxone-EMEA/H/C/000697/WS0934/0034/G** MAH: Indivior UK Limited, Lead Rapporteur: Martina Weise Request for Supplementary Information adopted on 16.02.2017.	Weekly start timetable. The Committee adopted a Request for Supplementary information together with a specific timetable.
WS0984 **AZILECT-EMEA/H/C/000574/WS0984/0073** **Rasagiline ratiopharm-EMEA/H/C/003957/WS0984/0007** MAH: Teva B.V., Lead Rapporteur: Bruno Sepodes Opinion adopted on 09.02.2017. Request for Supplementary Information adopted on 27.10.2016.	Positive Opinion adopted by consensus on 09.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1027 **Genvoya-EMEA/H/C/004042/WS1027/0019** **Stribild-EMEA/H/C/002574/WS1027/0071** **Tybost-EMEA/H/C/002572/WS1027/0030** MAH: Gilead Sciences International Ltd, Lead Rapporteur: Robert James Hemmings Opinion adopted on 02.02.2017. Request for Supplementary Information adopted on 17.11.2016.	Positive Opinion adopted by consensus on 02.02.2017. 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	Weekly start timetable. The Committee adopted a Request for Supplementary information together with a specific timetable.

on 23.02.2017.	
WS1046 **Ambirix-EMEA/H/C/000426/WS1046/0082** **Twinrix Adult-EMEA/H/C/000112/WS1046/0116** **Twinrix Paediatric-EMEA/H/C/000129/WS1046/0117** MAH: GSK Biologicals SA, Lead Rapporteur: Robert James Hemmings Opinion adopted on 16.02.2017.	Positive Opinion adopted by consensus on 16.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1067/G **Infanrix hexa-EMEA/H/C/000296/WS1067/0215/G** MAH: GSK Biologicals SA, Lead Rapporteur: Bart Van der Schueren Opinion adopted on 16.02.2017.	Positive Opinion adopted by consensus on 16.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1069/G **Infanrix hexa-EMEA/H/C/000296/WS1069/0214/G** MAH: GSK Biologicals SA, Lead Rapporteur: Bart Van der Schueren Opinion adopted on 23.02.2017.	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1080 **Copalia-EMEA/H/C/000774/WS1080/0091** **Copalia HCT-EMEA/H/C/001159/WS1080/0057** **Dafiro-EMEA/H/C/000776/WS1080/0093** **Dafiro HCT-EMEA/H/C/001160/WS1080/0058** **Exforge-EMEA/H/C/000716/WS1080/0090** **Exforge HCT-EMEA/H/C/001068/WS1080/0056** MAH: Novartis Europharm Ltd, Lead Rapporteur: Hanne Lomholt Larsen Opinion adopted on 23.02.2017.	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1081 **Hexacima-EMEA/H/C/002702/WS1081/0055** **Hexaxim-EMEA/H/W/002495/WS1081/0062** **Hexyon-EMEA/H/C/002796/WS1081/0059** MAH: Sanofi Pasteur SA, Lead Rapporteur: Jan	Weekly start timetable. The Committee adopted a Request for Supplementary information together with a specific timetable.

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. The Committee adopted a Request for Supplementary information together with a specific timetable.
WS1090/G **OFEV-EMEA/H/C/003821/WS1090/0012/G** **Vargatef-EMEA/H/C/002569/WS1090/0014/G** MAH: Boehringer Ingelheim International GmbH, Lead Rapporteur: Sinan B. Sarac Opinion adopted on 09.02.2017.	Positive Opinion adopted by consensus on 09.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1094/G **Eucreas-EMEA/H/C/000807/WS1094/0058/G** **Galvus-EMEA/H/C/000771/WS1094/0049/G** **Icandra-EMEA/H/C/001050/WS1094/0059/G** **Jalra-EMEA/H/C/001048/WS1094/0049/G** **Xiliarx-EMEA/H/C/001051/WS1094/0048/G** **Zomarist-EMEA/H/C/001049/WS1094/0059/G** MAH: Novartis Europharm Ltd, Lead Rapporteur: Kristina Dunder Opinion adopted on 16.02.2017.	Positive Opinion adopted by consensus on 16.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1098/G **Olazax-EMEA/H/C/001087/WS1098/0019/G** **Olazax Disperzi-EMEA/H/C/001088/WS1098/0020/G** MAH: Glenmark Pharmaceuticals s.r.o., Generic, Duplicate, Generic of Zyprexa, Zyprexa Velotab, Duplicate of Olanzapine Glenmark, Olanzapine Glenmark Europe, Lead Rapporteur: Alexandre Moreau Opinion adopted on 09.02.2017.	Positive Opinion adopted by consensus on 09.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

WS1100 **Adcirca-EMEA/H/C/001021/WS1100/0028** **Cialis-EMEA/H/C/000436/WS1100/0088** MAH: Eli Lilly Nederland B.V., Lead Rapporteur: Concepcion Prieto Yerro"This variation is being submitted to update the tadalafil (Adcirca and Cialis) Summary of Product Characteristics to introduce a warning and precaution regarding cases of sudden hearing loss which have been reported after the use of tadalafil, as requested following the outcome of the assessment of a cumulative review on the topic (Post-Authorisation measures 020 and 046 for Cialis and Adcirca). Section 4.4 of the SmPC and section 2 of the Package Leaflet were therefore updated." Opinion adopted on 23.02.2017.	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1102 **Hirobriz Breezhaler-EMEA/H/C/001211/WS1102/0039** **Onbrez Breezhaler-EMEA/H/C/001114/WS1102/0038** **Oslif Breezhaler-EMEA/H/C/001210/WS1102/0038** MAH: Novartis Europharm Ltd, Lead Rapporteur: Hanne Lomholt Larsen Opinion adopted on 23.02.2017.	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1104 **Epclusa-EMEA/H/C/004210/WS1104/0008** **Harvoni-EMEA/H/C/003850/WS1104/0047** **Sovaldi-EMEA/H/C/002798/WS1104/0039** MAH: Gilead Sciences International Ltd, Lead Rapporteur: Filip Josephson Opinion adopted on 16.02.2017.	Positive Opinion adopted by consensus on 16.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1118/G **Helixate NexGen-EMEA/H/C/000276/WS1118/0185/G** **KOGENATE Bayer-EMEA/H/C/000275/WS1118/0193/G** MAH: Bayer Pharma AG, Duplicate, Duplicate of KOGENATE Bayer, Lead Rapporteur: Jan Mueller-Berghaus	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

Opinion adopted on 23.02.2017.	
WS1119/G **Iblias-EMEA/H/C/004147/WS1119/0004/G** **Kovaltry-EMEA/H/C/003825/WS1119/0007/G** MAH: Bayer Pharma AG, Lead Rapporteur: Kristina Dunder Opinion adopted on 23.02.2017.	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1127 **Zypadhera-EMEA/H/C/000890/WS1127/0033** **Zyprexa-EMEA/H/C/000115/WS1127/0122** **Zyprexa Velotab-EMEA/H/C/000287/WS1127/0092** MAH: Eli Lilly Nederland B.V., Duplicate, Duplicate of Olansek, Lead Rapporteur: Outi Mäki-IkolaTo update section 4.8 of the SmPC and section 4 of the PIL to implement the signal recommendations on 'Olanzapine – Restless legs syndrome (EPITT no 18659)' adopted at the 24-27 October 2016 PRAC. The package leaflet is updated accordingly. In addition the EL annexes are brought in line with the EN text." Opinion adopted on 23.02.2017.	Positive Opinion adopted by consensus on 23.02.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

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

BeneFIX - nonacog alfa - EMEA/H/C/000139/II/0141 MAH: Pfizer Limited, Rapporteur: Jan Mueller-Berghaus, "Update of section 5.2 of the SmPC in order to update the information based on PK data from study B1821048. This study reported PK from an extended collection time beyond the 72 hours previously used in all other BeneFIX studies, to 96 hours, after BeneFIX administration." Withdrawal request submitted on 21.02.2017.	The MAH withdrew the procedure on 21.02.2017.
Imlygic - talimogene laherparepvec - EMEA/H/C/002771/II/0010, ATMP MAH: Amgen Europe B.V., Rapporteur: Olli Tenhunen, , "Submission of the primary analysis (PA) report for Study 20120324 (A Phase 2, Multicenter, Single-arm Trial to Evaluate the Biodistribution and Shedding of Talimogene	The MAH withdrew the procedure on 09.02.2017.

Laherparepvec in Subjects with Unresected, Stage IIIB to IVM1c Melanoma) which is listed as a category 3 pharmacovigilance activity in the Risk Management Plan (RMP)."
Withdrawal request submitted on 09.02.2017.

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

B.6. START OF THE PROCEDURES TIMETABLES FOR INFORMATION

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

- brigatinib - EMEA/H/C/004248
, treatment of anaplastic lymphoma kinase (ALK)-positive non-small cell lung cancer (NSCLC)

- ropeginterferon alfa-2b - EMEA/H/C/004128, Orphan
Applicant: AOP Orphan Pharmaceuticals AG, treatment of polycythemia vera

- caplacizumab - EMEA/H/C/004426, Orphan
Applicant: Ablynx NV, indicated for the treatment of acquired thrombotic thrombocytopenic purpura (aTTP).

- nitisinone - EMEA/H/C/004582
, treatment of hereditary tyrosinemia type 1,

- sodium benzoate - EMEA/H/C/004150, Orphan
Applicant: Lucane Pharma, treatment of non ketotic hyperglycinemia, urea cycle disorders including carbamoyl-phosphate synthase-1 deficiency, ornithine transcarbamylase deficiency, citrullinaemia type 1, argininosuccinic aciduria, hyperargininaemia, n-acetylglutamate synthase deficiency, ornithine translocase deficiency and lysinuric protein intolerance

- ertugliflozin / metformin hydrochloride - EMEA/H/C/004314
, treatment of type 2 diabetes mellitus

- ertugliflozin - EMEA/H/C/004315
, type 2 diabetes mellitus

- ertugliflozin / sitagliptin - EMEA/H/C/004313
, type 2 diabetes mellitus

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

ellaOne - ulipristal acetate - EMEA/H/C/001027/X/0045
MAH: Laboratoire HRA Pharma, SA, Rapporteur: Paula Boudewina van Hennik"Addition of a new pharmaceutical form (film-coated tablets) to the existing strength 30 mg."

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

- insulin lispro - EMEA/H/C/004303
, treatment of diabetes mellitus
List of Questions adopted on 26.01.2017.

- ribociclib - EMEA/H/C/004213
, treatment of breast cancer
List of Questions adopted on 26.01.2017.

- cladribine - EMEA/H/C/004230
, treatment of highly active relapsing-remitting multiple sclerosis (MS)
List of Questions adopted on 10.11.2016.

Revestive - teduglutide - EMEA/H/C/002345/X/0029, Orphan
MAH: Shire Pharmaceuticals Ireland Ltd
List of Questions adopted on 10.11.2016.

- midostaurin - EMEA/H/C/004095, Orphan
treatment of mastocytosis and treatment of acute myeloid leukaemia
List of Questions adopted on 15.12.2016.

- adalimumab - EMEA/H/C/004279
, treatment of rheumatoid arthritis, psoriatic arthritis and ankylosing spondylitis
List of Questions adopted on 10.11.2016.

- nusinersen - EMEA/H/C/004312, Orphan
for the treatment of Spinal Muscular Atrophy (SMA).
List of Questions adopted on 24.01.2017.

- atezolizumab - EMEA/H/C/004143
, treatment of locally advanced or metastatic urothelial carcinoma,
treatment of non-small cell lung carcinoma (NSCLC)
List of Questions adopted on 15.09.2016.

- telotristat ethyl - EMEA/H/C/003937,

Orphan

, treatment of carcinoid syndrome

List of Questions adopted on 10.11.2016.

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

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

Betmiga - mirabegron -
EMEA/H/C/002388/R/0026

MAH: Astellas Pharma Europe B.V., Rapporteur: Concepcion Prieto Yerro, Co-Rapporteur: Nithyanandan Nagercoil, PRAC Rapporteur: Dolores Montero Corominas

Constella - linaclotide -
EMEA/H/C/002490/R/0032

MAH: Allergan Pharmaceuticals International Ltd, Rapporteur: Harald Enzmann, Co-Rapporteur: Concepcion Prieto Yerro, PRAC Rapporteur: Valerie Strassmann

Eylea - aflibercept -
EMEA/H/C/002392/R/0033

MAH: Bayer Pharma AG, Rapporteur: Alexandre Moreau, Co-Rapporteur: Nithyanandan Nagercoil, PRAC Rapporteur: Caroline Laborde

Glybera - alipogene tiparvovec -
EMEA/H/C/002145/R/0062, Orphan, ATMP

MAH: uniQure biopharma B.V., Rapporteur: Christiane Niederlaender, Co-Rapporteur: Egbert Flory, PRAC Rapporteur: Julie Williams

Ibandronic acid Accord - ibandronic acid -
EMEA/H/C/002638/R/0013

MAH: Accord Healthcare Ltd, Generic, Generic of Bondronat, Rapporteur: Alar Irs, PRAC Rapporteur: Doris Stenver

Memantine Merz - memantine hydrochloride -
EMEA/H/C/002711/R/0012

MAH: Merz Pharmaceuticals GmbH, Rapporteur: Concepcion Prieto Yerro, Co-Rapporteur: Bruno Sepodes, PRAC Rapporteur: Dolores Montero Corominas

Picato - ingenol mebutate -
EMEA/H/C/002275/R/0023

MAH: LEO Laboratories Ltd, Rapporteur:

Nithyanandan Nagercoil, Co-Rapporteur: Alexandre Moreau, PRAC Rapporteur: Julie Williams

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.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

Keytruda - pembrolizumab - EMEA/H/C/003820/II/0023/G
MAH: Merck Sharp & Dohme Limited, Rapporteur: Daniela Melchiorri, Co-Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Sabine Straus“Extension of Indication to add treatment of urothelial carcinoma in patients previously treated with chemotherapy based on the results from study KEYNOTE-045; a phase 3, randomized, active-controlled, multi-site, open-label trial evaluating pembrolizumab administered at 200 mg Q3W versus investigators' choice of paclitaxel, docetaxel, or vinflunine in patients previously treated with chemotherapy.
Extension of Indication to add treatment of urothelial carcinoma in patients ineligible for cisplatin (not previously treated) based on the results from study KEYNOTE-52; a phase 2, single-arm, multisite, open-label trial of pembrolizumab at 200 mg Q3W in the treatment of patients with locally advanced or metastatic urothelial carcinoma who are not eligible for cisplatin-containing chemotherapy.

As a consequence, sections 4.1, 4.2, 4.4, 4.8 and 5.1 of the SmPC have been updated and the Package Leaflet has been updated accordingly.
Further, the MAH is proposing a change to section 4.3 of the SmPC to add that only patients with severe hypersensitivity should be excluded from therapy, and a change to section 4.4 of the SmPC adding possible hypersensitivity and anaphylaxis as part of infusion reactions.
The application included an updated RMP version 7.0."

Stribild - elvitegravir / cobicistat / emtricitabine / tenofovir disoproxil - EMEA/H/C/002574/II/0079

MAH: Gilead Sciences International Ltd, Rapporteur: Robert James Hemmings, Co-Rapporteur: Joseph Emmerich, PRAC Rapporteur: Patrick Batty,"Extension of Indication to include the treatment of HIV-1 infected adolescents, with NRTI resistance or toxicities precluding the use of first line agents, aged 12 to < 18 years and weighing ≥ 35 kg; as a consequence, sections 4.1, 4.2, 4.4, 4.8, 5.1 and 5.2 of the SmPC are updated based on pharmacokinetics, safety and efficacy data through 48 weeks of treatment with Stribild in Study GS-US-236-0112.
The Package Leaflet and Risk Management Plan (v.12) are updated in accordance.
The requested variation proposed amendments to the Summary of Product Characteristics, Package Leaflet and Risk Management Plan (RMP).
In addition, the Marketing authorisation holder (MAH) took the opportunity to make minor linguistic amendments"

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

MAH: Gilead Sciences International Ltd, Rapporteur: Greg Markey, Co-Rapporteur: Alexandre Moreau, PRAC Rapporteur: Julie Williams"Extension of Indication to include pre-exposure prophylaxis of HIV in adolescents aged 12 to < 18 years at high risk; as a consequence, sections 4.1, 4.2, 4.4, 4.8, 5.1 and 5.2 of the SmPC are updated based on extrapolation of data for emtricitabine, tenofovir disoproxil fumarate, and Truvada in HIV-infected and uninfected subjects.

The Package Leaflet and Risk Management Plan (v.15) are updated in accordance.
The requested variation proposed amendments to the Summary of Product Characteristics, Package Leaflet and Risk Management Plan (RMP).
In addition, the Marketing authorisation holder (MAH) took the opportunity to make minor linguistic amendments"

Zydelig - idelalisib - EMEA/H/C/003843/II/0032/G

MAH: Gilead Sciences International Ltd, Rapporteur: Filip Josephson, Co-Rapporteur: Paula Boudewina van Hennik, PRAC Rapporteur: Patrick BattyC.I.6. Extension of Indication: Extension of the approved chronic lymphocytic leukemia (CLL) indication for Zydelig to include its use in combination with bendamustine and rituximab based on the results of the primary analysis of pivotal Study GS-US-312-0115 "a Phase 3, randomized, double-blind, controlled study evaluating the efficacy and safety of idelalisib (GS-1101) in combination with bendamustine and rituximab for previously treated chronic lymphocytic leukemia" as a consequence, sections 4.1, 4.8, and 5.1 of the SmPC are updated. The Package Leaflet is updated in accordance.
In addition, the Marketing authorisation holder (MAH) took the opportunity to update the list of local representatives in the Package Leaflet.
The RMP version 2.2 has also been submitted.
C.I.13: Submission of the final report from study 101-08, a phase 2, single-arm study evaluated idelalisib monotherapy and in combination with rituximab in elderly subjects with previously untreated CLL or small lymphocytic lymphoma. Inclusion of this report provides additional safety data to support the evaluation of the use of idelalisib in patients with CLL. Submission of this report is also made in fulfilment of PAM008.
C.I.13: Submission of the final report from study GS-US-312-0123, a phase 3 randomized study evaluated idelalisib in combination with bendamustine and rituximab in subjects with previously untreated CLL. Inclusion of this report is supportive of a complete safety evaluation concerning the use of this combination in patients with CLL."

WS1078
Komboglyze-EMEA/H/C/002059/WS1078/0035
Onglyza-EMEA/H/C/001039/WS1078/0041
MAH: AstraZeneca AB, Lead Rapporteur: Johann Lodewijk Hillege, “Extension of Indication to include the use of a triple combination therapy (saxagliptin, metformin and dapagliflozin) as adjunct to diet and exercise to improve glycaemic control in adult patients aged 18 years and older with type 2 diabetes mellitus, when metformin together with dapagliflozin, do not provide adequate glycaemic control. Editorial changes are made throughout the Summary Products Characteristics and Package Leaflets. Furthermore, the Product Information is brought in line with the latest QRD template version 10 for Onglyza.”

B.6.8. CHMP assessed procedures scope: Pharmaceutical aspects

Advate - octocog alfa - EMEA/H/C/000520/II/0083/G
MAH: Baxter AG, Rapporteur: Jan Mueller-Berghaus

Aflunov - prepandemic influenza vaccine (H5N1) (surface antigen, inactivated, adjuvanted) - EMEA/H/C/002094/II/0032
MAH: Seqirus S.r.l, Rapporteur: Daniela Melchiorri,

Afstyla - lonoctocog alfa - EMEA/H/C/004075/II/0001
MAH: CSL Behring GmbH, Rapporteur: Jan Mueller-Berghaus

Azopt - brinzolamide - EMEA/H/C/000267/II/0064/G
MAH: Alcon Laboratories (UK) Ltd, Rapporteur: Concepcion Prieto Yerro

Cetrotide - cetrorelix acetate - EMEA/H/C/000233/II/0056
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

ELOCTA - efmoroctocog alfa - EMEA/H/C/003964/II/0012/G
MAH: Swedish Orphan Biovitrum AB (publ), Rapporteur: Jan Mueller-Berghaus

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

Galafold - migalastat - EMEA/H/C/004059/II/0005, Orphan
MAH: Amicus Therapeutics UK Ltd, Rapporteur: Johann Lodewijk Hillege

Imbruvica - ibrutinib - EMEA/H/C/003791/II/0032/G, Orphan
MAH: Janssen-Cilag International NV, Rapporteur: Filip Josephson

Inhixa - enoxaparin sodium - EMEA/H/C/004264/II/0009/G
MAH: Techdow Europe AB, Duplicate, Duplicate of Thorinane, Rapporteur: Andrea Laslop

Keytruda - pembrolizumab - EMEA/H/C/003820/II/0026/G
MAH: Merck Sharp & Dohme Limited, Rapporteur: Daniela Melchiorri

Mysimba - naltrexone hydrochloride / bupropion hydrochloride - EMEA/H/C/003687/II/0013/G
MAH: Orexigen Therapeutics Ireland Limited, Rapporteur: Hanne Lomholt Larsen

OPDIVO - nivolumab - EMEA/H/C/003985/II/0031/G
MAH: Bristol-Myers Squibb Pharma EEIG, Rapporteur: Aranzazu Sancho-Lopez, Procedure Manager: Caroline Blanc

Orkambi - lumacaftor / ivacaftor - EMEA/H/C/003954/II/0018/G
MAH: Vertex Pharmaceuticals (Europe) Ltd., Rapporteur: Nithyanandan Nagercoil

RotaTeq - rotavirus vaccine (live, oral) - EMEA/H/C/000669/II/0069/G
MAH: MSD Vaccins, Rapporteur: Greg Markey

Simponi - golimumab -

EMEA/H/C/000992/II/0074/G
MAH: Janssen Biologics B.V., Rapporteur: Kristina Dunder

TachoSil - human thrombin / human fibrinogen - EMEA/H/C/000505/II/0077/G
MAH: Takeda Austria GmbH, Rapporteur: Jan Mueller-Berghaus

Thyrogen - thyrotropin alfa - EMEA/H/C/000220/II/0090
MAH: Genzyme Europe BV, Rapporteur: Patrick Salmon

Vectibix - panitumumab - EMEA/H/C/000741/II/0084
MAH: Amgen Europe B.V., Rapporteur: Robert James Hemmings

Vimizim - elosulfase alfa - EMEA/H/C/002779/II/0017/G, Orphan
MAH: BioMarin Europe Ltd, Rapporteur: Johann Lodewijk Hillege

WS1124
Fertavid-EMEA/H/C/001042/WS1124/0034
Puregon-EMEA/H/C/000086/WS1124/0092
MAH: Merck Sharp & Dohme Limited, Lead Rapporteur: Nithyanandan Nagercoil

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

Eperzan - albiglutide - EMEA/H/C/002735/II/0031
MAH: GlaxoSmithKline Trading Services, Rapporteur: Kristina Dunder“Update of sections 4.4 and 4.8 of the SmPC in order to add angioedema with frequency ‘rare’ and to include a warning concerning hypersensitivity reactions in general.
The Package Leaflet is updated accordingly.
In addition, the Marketing authorisation holder took the opportunity to implement minor editorial amendments throughout the Product Information.”

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."

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)."

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."

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"

Mysimba - naltrexone hydrochloride / bupropion hydrochloride - EMEA/H/C/003687/II/0014

MAH: Orexigen Therapeutics Ireland Limited, Rapporteur: Hanne Lomholt Larsen"C.I.13: Submission of the final report from study NaltrexBuprop-1004; a Phase 1, Open-Label, Sequential Design Study to Evaluate the Potential Effect of Multiple Oral Doses of Extended-Release Combination of Naltrexone and Bupropion on the Pharmacokinetics of a Single Oral Dose of Metformin in Healthy Subjects. This study does not lead to changes in the product information."

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."

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."

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."

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

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."

WS1148/G
Hexacima-EMEA/H/C/002702/WS1148/0059/G
Hexaxim-EMEA/H/W/002495/WS1148/0065/G
Hexyon-EMEA/H/C/002796/WS1148/0063/G
MAH: Sanofi Pasteur Europe, Duplicate, Duplicate of Hexacima, Lead Rapporteur: Jan Mueller-Berghaus

B.6.10. 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."

Edurant - rilpivirine -
EMEA/H/C/002264/II/0024
MAH: Janssen-Cilag International NV, Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Sabine StrausUpdate of sections 4.2, 4.4, 4.6, 5.1 and 5.2 of the SmPC in order to include information: use of rilpivirine in combination with a background regimen for the treatment of HIV-1 infection during pregnancy and postpartum, without dose adjustment following final results from study TMC114HIV3015 listed as a category 3 study in the RMP. This is a single arm, open-label trial to assess the pharmacokinetics of darunavir/ritonavir, etravirine, and rilpivirine in HIV-1-infected pregnant women.
The Package Leaflet is updated accordingly. The RMP version 7.0 has also been submitted.
In addition, the Marketing authorisation holder (MAH) took the opportunity to introduce the latest renewal date in section 9 of the SmPC and the physical address of the Netherlands Local Operating Company in the PIL section 6."

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 typo As per the PRAC recommendation, the updated RMP version 3.2 is also proposed."	
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."	
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)."	
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."	
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)."	
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. 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."	
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/WS1141/005**	

6
Ristaben-EMEA/H/C/001234/WS1141/0048
TESAVEL-EMEA/H/C/000910/WS1141/0056
Xelevia-EMEA/H/C/000762/WS1141/0060
MAH: Merck Sharp & Dohme Limited, Lead Rapporteur: Johann Lodewijk Hillege, Lead PRAC Rapporteur: Menno van der Elst"Update of sections 4.4 of the SmPC in order to add 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."

B.6.11. PRAC assessed procedures

PRAC Led
Respreeza - human alpha1-proteinase inhibitor - EMEA/H/C/002739/II/0013
MAH: CSL Behring Gmb,H PRAC Rapporteur: Eva A. Segovia, "Submission of an updated RMP version 3.1 in order to include the final safety data from CE1226_3001, which were assessed in a type II variation (Procedure No. EMEA/H/C/002739/II/0002) and adjustments in the Non-Clinical Safety specification part (Part II, Module SII)."

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."

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."

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 (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."

PRAC Led

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.”	

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

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

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

---

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

---

WS1140
Glyxambi-EMEA/H/C/003833/WS1140/0002
Jentadueto-EMEA/H/C/002279/WS1140/0037
Trajenta-EMEA/H/C/002110/WS1140/0027
MAH: Boehringer Ingelheim International GmbH, Lead Rapporteur: Johann Lodewijk Hillege“To add the ADR bullous pemphigoid to the Warnings and Precautions sections in the SmPC (section 4.4) and in the package leaflet (section 2).”

WS1146
Hexacima-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.7. DOCUMENTS TABLED IN MMD AFTER THE CHMP PLENARY

B.7.1. Line listing for Variation Type I and Variation Type II (MMD only) post authorisation procedures from the beginning of the year.

B.7.2. Line listing overview of all applications under the centralised procedure (MMD only). line listing - products - authorised, under evaluation, suspended.xls

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 20-23 February 2017 CHMP plenary:

Oncology		
	Conditioning treatment for haematopoietic stem cell transplantationin patients with Acute myeloid leukaemia	The CHMP denied eligibility to PRIME and adopted the critical summary report.
Haematology-haemostaseology		
1.	**Adeno-associated viral vector containing factor IX gene (PF-06838435/SPK-9001)**; ATMP;; Treatment of haemophilia B	The CHMP granted eligibility to PRIME and adopted the critical summary report.
Immunology-Rheumatology-Transplantation		
2.	Treatment of antiphospholipid syndrome	The CHMP denied eligibility to PRIME and adopted the critical summary report.
Infectious disease		
3.	Prevention of postoperative invasive disease caused by Staphylococcus aureus	The CHMP denied eligibility to PRIME and adopted the critical summary report.
Endocrinology-Gynaecology-Fertility-Metabolism		
4.	**Givosiran,** Prevention of acute attacks of hepatic porphyria	The CHMP granted eligibility to PRIME and adopted the critical summary report.

G.3.2. List of procedures starting in February 2017 for March 2017 CHMP adoption of outcomes

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