20 October 2017
EMA/CHMP/479458/2017
Inspections, Human Medicines Pharmacovigilance and Committees Division

Committee for medicinal products for human use (CHMP)

Minutes of the meeting on 11-14 September 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).

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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) September 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 11-14 September 2017 (to be published post October 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.

The CHMP welcomed Jayne Crowe as new CHMP member and Peter Kiely as new CHMP alternate member from Ireland replacing David Lyons and Patrick Salmon. The CHMP also welcomed Simona Badoi as the new member from Romania replacing Nela Vilceanu and František Drafi as new member from Slovakia replacing Adriana Adameová.

1.2. Adoption of agenda

CHMP agenda for 11-14 September 2017

The CHMP adopted the agenda.

1.3. Adoption of the minutes

CHMP minutes for 17-20 July and August 2017 written procedure.

The CHMP adopted the CHMP minutes for 17-20 July 2017 and for the August written procedure.

The Minutes of the September 2017 CHMP ORGAM meeting held on 4 September 2017, together with all decisions taken at that meeting, were adopted.

2. Oral Explanations

2.1. Pre-authorisation procedure oral explanations

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

treatment of vulvovaginal atrophy

Scope: Oral explanation

Action: Oral explanation to be held on 13 September 2017 at time 16:00

List of Outstanding Issues adopted on 22.06.2017, 26.01.2017, 13.10.2016. List of Questions adopted on 26.05.2016.

An oral explanation was held on 13 September 2017 at time 16:00.

See 3.2

2.1.2. - ocrelizumab - EMEA/H/C/004043

treatment of multiple sclerosis

Scope: Oral explanation to be held on 13 September 2017 at time 09:00. Report from SAG Neurology.

Action: For adoption

List of Outstanding Issues adopted on 23.03.2017. List of Questions adopted on 15.09.2016.

Oral explanation was held on 13 September 2017 at time 09:00.

See 3.2

2.1.3. - sirukumab - EMEA/H/C/004165

treatment of rheumatoid arthritis

Scope: Oral explanation

Action: Oral explanation to be held on 14 September 2017 at time 09:00

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

An oral explanation was held on 14 September 2017 at time 09:00.

See 3.2

2.2. Re-examination procedure oral explanations

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

XBiotech Germany GmbH; treatment of metastatic colorectal cancer

Scope: Oral explanation to be held on 12 September 2017 at time 11:00

Action: For adoption

Opinion adopted on 18.05.2017

An oral explanation was held on 12 September 2017 at time 11:00.

See 3.5

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

AB Science; treatment of mastocytosis

Scope: Oral explanation to be held on 12 September 2017 at time 09:00, SAG report

Action: For adoption

Opinion adopted on 18.05.2017.

Oral explanation was held on 12 September 2017 at time 09:00.

See 3.5

2.3. Post-authorisation procedure oral explanations

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

Santhera Pharmaceuticals (Deutschland) GmbH

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

Scope: Oral explanation to be held on 13 September 2017 at time 14:00

Action: For adoption

Participation of patients' representatives

Request for Supplementary Information adopted on 18.05.2017, 23.02.2017, 15.09.2016.

An oral explanation was held on 13 September 2017 at time 14:00.

See 5.1

2.4. Referral procedure oral explanations

No items

3. Initial applications

3.1. Initial applications; Opinions

3.1.1. Cyltezo - adalimumab - EMEA/H/C/004319

Boehringer Ingelheim International GmbH; treatment of rheumatoid arthritis, axial spondyloarthritis, psoriasis, hidradenitis suppurativa (HS), Crohn's disease, ulcerative colitis and uveitis.

Scope: Opinion

Action: For adoption

Similar biological application (Article 10(4) of Directive No 2001/83/EC)

List of Outstanding Issues adopted on 20.07.2017. List of Questions adopted on 23.03.2017.

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.

The CHMP adopted the BWP report.

3.1.2. Elebrato Ellipta - fluticasone furoate / umeclidinium / vilanterol - EMEA/H/C/004781

GlaxoSmithKline Trading Services Limited; treatment of adult patients with chronic obstructive pulmonary disease (COPD)

Scope: Opinion

Action: For adoption

Fixed combination application (Article 10b of Directive No 2001/83/EC), Duplicate of Trelegy Ellipta

List of Outstanding Issues adopted on 20.07.2017. List of Questions adopted on 21.04.2017.

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.

3.1.3. Tremfya - guselkumab - EMEA/H/C/004271

Janssen-Cilag International N.V.; treatment of plaque psoriasis

Scope: Opinion

Action: For adoption

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

List of Outstanding Issues adopted on 20.07.2017. List of Questions adopted on 21.04.2017.

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 guselkumab 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 restricted medical prescription.

The summary of opinion was circulated for information.

The CHMP adopted the BWP report.

3.1.4. Imatinib Teva B.V. - imatinib - EMEA/H/C/004748

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

Scope: Opinion

Action: For adoption

Generic application (Article 10(1) of Directive No 2001/83/EC), Generic of Glivec

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.

The CHMP adopted the updated CHMP Assessment Report for Imatinib Teva B.V. on similarity.

3.1.5. Miglustat Gen.Orph - miglustat - EMEA/H/C/004366

Gen.Orph; treatment of Gaucher disease

Scope: Opinion

Action: For adoption

Generic application (Article 10(1) of Directive No 2001/83/EC), Generic of Zavesca

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

The CHMP adopted the CHMP Assessment Report for Miglustat Gen.Orph on similarity.

3.1.6. Nyxoid - naloxone - EMEA/H/C/004325

Mundipharma Corporation Limited; Nyxoid is intended for emergency use for known or suspected opioid overdose as manifested by respiratory and/or central nervous system depression

Scope: Opinion

Action: For adoption

Hybrid application (Article 10(3) of Directive No 2001/83/EC)

List of Outstanding Issues adopted on 20.07.2017. List of Questions adopted on 23.03.2017.

The CHMP discussed the post-authorisation study and whether a PAES or a PASS was considered more appropriate, and the CHMP agreed on a PAES.

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.

3.1.7. Ontruzant - trastuzumab - EMEA/H/C/004323

Samsung Bioepis UK Limited (SBUK); treatment of breast cancer and metastatic gastric cancer

Scope: Opinion

Action: For adoption

Similar biological application (Article 10(4) of Directive No 2001/83/EC)

List of Outstanding Issues adopted on 20.07.2017. List of Questions adopted on 26.01.2017.

The CHMP debated the apparent difference in terms of pathological complete response rate observed in the clinical efficacy trial between Ontruzant and Herceptin. Based on additional analysis, this observation was found to be likely the result of a temporary shift in some quality parameters of Herceptin impacting a number of batches used in the clinical efficacy/safety trial. Despite this apparent difference between Ontruzant and some batches of Herceptin used in this clinical study, the CHMP concluded that Ontruzant can be considered similar to the reference product Herceptin based on additional analyses conducted, pharmacodynamic knowledge of the product and considering all the evidence available from the comparative exercise.

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 CHMP noted the letter of recommendation dated 6 September 2017.

The summary of opinion was circulated for information.

The CHMP adopted the BWP report

3.1.8. Ritonavir Mylan - ritonavir - EMEA/H/C/004549

MYLAN S.A.S; treatment of HIV-1

Scope: Opinion

Action: For adoption

Generic application (Article 10(1) of Directive No 2001/83/EC), Generic of Norvir

List of Outstanding Issues adopted on 20.07.2017. List of Questions adopted on 21.04.2017.

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.

3.1.9. Tookad - padeliporfin - EMEA/H/C/004182

STEBA Biotech S.A; treatment of prostate cancer

Scope: Opinion

Action: For adoption

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

List of Outstanding Issues adopted on 20.07.2017, 21.04.2017, 15.12.2016. List of Questions adopted on 26.05.2016.

The Committee discussed the available clinical data and whether long term efficacy and safety data was required before concluding on the opinion. Some members expressed the need for long-term data. The Committee also discussed that the wording of the indication should adequately reflect the population which derive benefit.

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 majority (29 out of 31 votes) together with the CHMP assessment report and translation timetable.

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

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

The divergent position (Johann Lodewijk Hillege, Alexandre Moreau) was appended to the opinion.

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

The summary of opinion was circulated for information.

3.1.10. Trelegy Ellipta - fluticasone furoate / umeclidinium / vilanterol - EMEA/H/C/004363

GlaxoSmithKline Trading Services; treatment of adult patients with chronic obstructive pulmonary disease (COPD)

Scope: Opinion

Action: For adoption

Fixed combination application (Article 10b of Directive No 2001/83/EC)

List of Outstanding Issues adopted on 20.07.2017. List of Questions adopted on 21.04.2017.

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.

3.1.11. VeraSeal - human fibrinogen/human thrombin - EMEA/H/C/004446

Instituto Grifols, S.A.; treatment of haemostasis

Scope: Opinion

Action: For adoption

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

List of Questions adopted on 21.04.2017.

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.

The CHMP adopted the BWP report.

3.1.12. Zejula - niraparib - Orphan - EMEA/H/C/004249

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

Scope: Opinion

Action: For adoption

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

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

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 niraparib 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 restricted medical prescription.

The CHMP noted the letter of recommendation dated 14 September 2017.

The summary of opinion was circulated for information.

The CHMP adopted the assessment report on similarity for Zejula.

3.1.13. Zubsolv - buprenorphine / naloxone - EMEA/H/C/004407

Mundipharma Corporation Limited; treatment for opioid drug dependence

Scope: Opinion

Action: For adoption

Hybrid application (Article 10(3) of Directive No 2001/83/EC)

List of Outstanding Issues adopted on 20.07.2017. List of Questions adopted on 23.02.2017.

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 special and restricted 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. - anagrelide - EMEA/H/C/004585

reduction of elevated platelet counts in at risk essential thrombocythaemia patients

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 21.04.2017.

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.

3.2.2. - paclitaxel - Orphan - EMEA/H/C/004154

Oasmia Pharmaceutical AB; treatment of ovarian cancer

Scope: 2nd day 180 list of outstanding issue, letter from the applicant dated 8 September 2017 requesting an extension of clock stop to respond to the Day 180 list of outstanding issue.

Action: For adoption

List of Outstanding Issues adopted on 18.05.2017. List of Questions adopted on 23.06.2016.

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 for an extension to the clock stop to respond to the list of outstanding issues with a specific timetable.

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

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

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 23.02.2017.

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.

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

treatment of severe asthma with an eosinophilic phenotype

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 21.04.2017.

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.

3.2.5. - darunavir - EMEA/H/C/004273

treatment of HIV-1 infection

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 23.03.2017.

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.

3.2.6. - darunavir - EMEA/H/C/004891

treatment of HIV-1 infection

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.

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

treatment of breast cancer

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 21.04.2017.

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.

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

treatment of vulvovaginal atrophy

Scope: Oral explanation, Day 180 list of outstanding issue

Action: Oral explanation to be held on 13 September 2017 at time 16:00

List of Outstanding Issues adopted on 22.06.2017, 26.01.2017, 13.10.2016. List of Questions adopted on 26.05.2016.

An oral explanation took place on 13 September 2017 at time 16:00.

The Committee adopted a 4th list of outstanding issues with a specific timetable.

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

treatment of breast cancer, non-small cell lung cancer, renal cell cancer, advanced epithelial ovarian, fallopian tube, or primary peritoneal cancer, platinum-sensitive epithelial ovarian, fallopian tube or primary peritoneal cancer

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 21.04.2017.

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.

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

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

Scope: Day 180 list of outstanding issue, letter from the applicant dated 5 September 2017 requesting an extension of clock stop to respond to the Day 180 list of outstanding issue.

Action: For adoption

List of Questions adopted on 26.01.2017.

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.

The CHMP agreed to consult an Ad-hoc expert group. The list of questions to the Ad hoc expert group was adopted.

The CHMP adopted the BWP report.

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

Accelerated assessment

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

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 18.07.2017.

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.

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

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

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 21.04.2017.

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. - ocrelizumab - EMEA/H/C/004043

treatment of multiple sclerosis

Scope: Oral explanation, Day 180 list of outstanding issue. Report from SAG Neurology.

Action: For adoption

List of Outstanding Issues adopted on 23.03.2017. List of Questions adopted on 15.09.2016.

See 2.1

Oral explanation was held on 13 September 2017 at 09:00.

The CHMP noted the SAG report. Most of the experts considered the trial population to be an atypical PPMS cohort, consisting mainly of younger patients likely to have a more “active” disease. Therefore, they considered that the results from the trial will have to be cautiously interpreted and that extrapolation of these results to the whole PPMS population may be questionable. Limitations in the robustness of the efficacy data were mentioned. In particular, the differences between genders with respect to the results for the primary endpoint were noted. Similarly, the absence of significance in the results for patients aged over 45 years was considered as a concern.

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

The CHMP adopted the BWP report.

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

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

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 21.04.2017.

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.

3.2.15. - sirukumab - EMEA/H/C/004165

treatment of rheumatoid arthritis

Scope: Oral explanation, Day 180 list of outstanding issue

Action: Oral explanation to be held on 14 September 2017 at time 09:00

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

An oral explanation was held on 14 September 2017 at time 09:00.

The Committee adopted a 2nd list of outstanding issues.

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

The CHMP adopted the BWP report.

3.2.16. - d-biotin - EMEA/H/C/004153

treatment of progressive multiple sclerosis (primary or secondary)

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 15.12.2016.

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

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

The CHMP agreed to consult the SWP and the SmPC Advisory Group group.

3.2.17. - ciclosporin - EMEA/H/C/004229

for the treatment of moderate dry eye disease in adults

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 23.03.2017.

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.

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

Clovis Oncology UK Ltd; treatment of ovarian cancer

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 23.03.2017.

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 for rucaparib.

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

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

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 21.04.2017.

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.

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

3.3.1. - viable t-cells - Orphan - ATMP - EMEA/H/C/002397

Kiadis Pharma Netherlands B.V.; adjunctive treatment in haematopoietic stem cell transplantation (HSCT) for a malignant disease

Scope: Day 120 list of questions

Action: For information

The Committee discussed the issues identified in this application.

The Committee agreed to the recommendation and scientific discussion together with the list of questions as adopted by the CAT.

The CHMP adopted the BWP report.

3.3.2. - doxorubicin hydrochloride - EMEA/H/C/004110

treatment of breast and ovarian cancer

Scope: Day 120 list of questions, letter from the applicant dated 7 September 2017 requesting an extension of clock stop to respond to Day 120 list of questions, similarity assessment report

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 the request for an extension to the clock stop to respond to the list of questions with a specific timetable.

The CHMP adopted the CHMP assessment report on similarity.

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

treatment of HIV-1 infection

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.

3.3.4. - adalimumab - EMEA/H/C/004429

treatment of rheumatoid arthritis, psoriatic arthritis and ankylosing spondylitis

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.

3.3.5. - adalimumab - EMEA/H/C/004320

treatment of rheumatoid arthritis, psoriatic arthritis and ankylosing spondylitis

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.

3.3.6. - budesonide - Orphan - EMEA/H/C/004655

Accelerated assessment

Dr. Falk Pharma GmbH; treatment of eosinophilic esophagitis (EoE)

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.

3.3.7. - melatonin - PUMA - EMEA/H/C/004425

treatment of insomnia in children with Autism Spectrum Disorders and neurogenetic diseases

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 agreed to the request by the applicant for an extension to the clock stop to respond to the list of questions with a specific timetable.

3.3.8. - vestronidase alfa - Orphan - EMEA/H/C/004438

Ultragenyx Germany GmbH; indicated for the treatment of Mucopolysaccharidosis VII (MPS VII; Sly syndrome) for patients of all ages

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.

3.3.9. The CHMP adopted the BWP report. - pegfilgrastim - EMEA/H/C/003961

treatment of neutropenia

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.

3.3.10. - prasugrel - EMEA/H/C/004644

prevention of atherothrombotic events

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.

3.3.11. - infliximab - EMEA/H/C/004647

treatment of rheumatoid arthritis, Crohn's disease, ankylosing spondylitis, psoriatic arthritis, psoriasis and ulcerative colitis

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

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

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

treatment of haemophilia A

Scope: CHMP list of questions to BPWP/BWP

Action: For adoption

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

The CHMP adopted a list of questions to the BPWP/BWP.

3.4.2. - naldemedine - EMEA/H/C/004256

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

Scope: Letter from the applicant dated 8 September 2017 requesting an extension of clock stop to respond to Day 120 list of questions adopted on 20.07.2017

Action: For adoption

List of Questions adopted on 20.07.2017.

The CHMP agreed to the request by the applicant for an extension of clock stop to respond to Day 120 list of questions adopted on 20.07.2017

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

treatment of neutropenia

Scope: Letter from the applicant dated 23 August 2017 requesting an extension of clock stop to respond to Day 120 list of questions adopted on 23.03.2017

Action: For adoption

List of Questions adopted on 23.03.2017.

The CHMP did not agree to the request by the applicant for an additional extension of clock stop to respond to Day 120 list of questions adopted on 23.03.2017.

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

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

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

Scope: Opinion

Action: For adoption

List of experts for the SAG planned on 04.09.2017 adopted by written procedure on 30 August 2017.

Opinion adopted on 18.05.2017

The CHMP noted the SAG report. The SAG agreed to most grounds for the negative opinion adopted on 18 May 2017. Taking into account the co-primary and secondary analyses the experts considered that a clinically relevant effect has not been established. Concerning the efficacy of the product on lean body mass with concomitant (gluco)corticosteroid treatment the experts did not come to a conclusion due to the limited data. On the overall safety of the product the SAG did not express major concern but outlined some remaining uncertainties.

The members discussed the available data with focus on the benefit/risk of the product. It was noted that the applicant rejected the invitation for an oral explanation in front of the CHMP.

The CHMP adopted a negative opinion by consensus, recommending the refusal of the marketing authorisation application. The CHMP adopted the assessment report.

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

The refusal questions and answers document was circulated for information.

3.5.2. Fanaptum - iloperidone - EMEA/H/C/004149

Vanda Pharmaceuticals Ltd.; treatment of schizophrenia

Scope: Appointment of re-examination Rapporteurs

Action: For adoption

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

Opinion adopted on 20.07.2017, Oral explanation 17.05.2016, List of Outstanding Issues adopted on 18.05.2017, 23.02.2017. List of Questions adopted on 28.04.2016.

The CHMP appointed a re-examination Rapporteur and a re-examination Co-Rapporteur.

The CHMP adopted the re-examination timetable.

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

XBiotech Germany GmbH; treatment of metastatic colorectal cancer

Scope: Re-examination of Opinion /Oral explanation to be held on 12 September 2017 at time 11:00

Action: For adoption

Opinion adopted on 18.05.2017

See 2.2

An oral explanation was held on 12 September 2017 at time 11:00. The presentation focused on the clinical trial design, including the choice and clinical relevance of the endpoints. Furthermore the clinical trial results and analysis were explained.

The members further discussed the available efficacy data and whether it was considered sufficient to substantiate a positive benefit/risk ratio. The CHMP was also reminded of issues concerning the control strategy for the manufacture of the product.

After the oral explanation an orientation was sought on the benefit/risk of the product and a negative trend by consensus was observed.

The Committee was reassured that no commonly reported side effects appear to be directly linked to this medicine. However, insufficient safety data are available to properly assess its overall risks and the committee still had concerns about the medicine's benefits and manufacturing controls.

The CHMP therefore maintained its opinion that the benefits of this medicine did not outweigh its risks and recommended that it to be refused marketing authorisation.

The CHMP adopted a negative opinion by consensus, recommending the refusal of the marketing authorisation application. The CHMP adopted the assessment report.

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

The refusal questions and answers document was circulated for information.

The CHMP adopted the BWP report.

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

AB Science; treatment of mastocytosis

Scope: Oral explanation/Opinion, SAG report
Action: For adoption

List of experts for the SAG planned on 04.09.2017 adopted by written procedure on 30 August 2017.

Opinion adopted on 18.05.2017.

The CHMP noted the SAG report. The SAG outlined uncertainties on the efficacy and the safety taking into account deficiencies in the design, conduct, and analysis of the pivotal clinical trial. The experts also could not conclude on the acceptability of the safety database as doubts on the completeness of the source data were raised. The SAG highlighted the lack of long-term safety data. Overall, the experts seriously questioned the ability of the data from the pivotal study to support the applied indication, considering GCP findings.

An oral explanation was held on 12 September 2017 at time 09:00. During the oral explanation the company presented the answers to the relevance of efficacy results in the claim, including adequacy of pre-planned endpoints and clinical relevance of study results and safety questions.

The Committee further discussed the remaining uncertainties and GCP findings. The CHMP was concerned about the reliability of the study results because a routine GCP inspection at the study sites revealed serious failings in the way the study had been conducted. In addition, major changes were made to the study design while the study was ongoing, which made the results difficult to interpret. Finally, data on the safety of the medicine were limited and there were concerns regarding the medicine's side effects, including neutropenia and harmful effects on the skin and liver, which were of relevance particularly because the medicine was to be used long term.

Therefore, the CHMP was of the opinion that the benefits of Masipro did not outweigh its risks and recommended that it to be refused marketing authorisation.

The CHMP adopted a negative opinion by consensus, recommending the refusal of the marketing authorisation application. The CHMP adopted the assessment report.

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

The refusal question and answers document was circulated for information.

3.5.5. Onzeald - etirinotecan pegol - EMEA/H/C/003874

Nektar Therapeutics UK Limited; treatment of breast cancer with brain metastases

Scope: Appointment of re-examination Rapporteurs

Action: For adoption

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

Opinion adopted on 20.07.2017, Oral explanation 16.05.2017, List of Outstanding Issues adopted on 18.05.2017, 23.03.2017. List of Questions adopted on 10.11.2016.

The CHMP appointed a re-examination Rapporteur and a re-examination Co-Rapporteur.

The CHMP adopted the re-examination timetable.

3.6. Initial applications in the decision-making phase

No items

3.7. Withdrawals of initial marketing authorisation application

No items

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

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

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

Glaxo Group Ltd

Rapporteur: Kristina Dunder, PRAC Rapporteur: Ulla Wändel Liminga

Scope: “Extension of the marketing authorisation to include:
- a new strength: 200 mg
- a new pharmaceutical form: solution for injection
- a new route of administration: subcutaneous use
Type II variation: Update of sections 4.2, 4.8, 5.1 and 5.2 of the SmPC as a consequence of the data package submitted to support the new proposed solution for injection subcutaneous presentations. The Package Leaflet is updated accordingly. In addition, the Marketing authorisation holder (MAH) took the opportunity to bring the PI in line with the latest QRD template version 10.0 and to introduce some editorial changes.”

Action: For adoption

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

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.

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

Laboratoire HRA Pharma

Rapporteur: Paula Boudewina van Hennik

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

Action: For adoption

List of Questions adopted on 22.06.2017.

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.

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

Novartis Europharm Ltd

Rapporteur: Alexandre Moreau, PRAC Rapporteur: Ghania Chamouni

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

Action: For adoption

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

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.

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

Novartis Europharm Ltd

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

Scope: “Extension of Indication to include treatment of paediatric patients with newly diagnosed Philadelphia chromosome-positive chronic myelogenous leukemia in chronic phase (Ph+ CML-CP), or with Ph+ CML-CP resistant or intolerant to prior therapy including imatinib, based on results from two clinical studies in paediatric patients conducted in accordance with the approved Tasigna Paediatric Investigation Plan (PIP), the Phase I PK study CAMN107A2120 and the Phase II safety and efficacy study CAMN107A2203. An updated RMP version 18.0 was provided as part of the application.
Extension application to add a new strength of 50mg hard capsules.
In addition, the applicant proposes to merge the SmPCs for the 50 mg and 200 mg strengths.”

Action: For adoption

List of Outstanding Issues adopted on 20.07.2017. List of Questions adopted on 23.03.2017.

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 CHMP noted the letter of recommendation dated 14 September 2017.

The CHMP adopted the Similarity Assessment Report for Tasigna.

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

No items

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

AstraZeneca AB

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

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

Action: For adoption

The Committee discussed the issues identified in this application. The Committee noted the

issues relating to the indication.

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

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

Octapharma AB

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

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

Action: For adoption

The Committee noted the issues identified in this application.

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

4.3.3. Sprycel - dasatinib - EMEA/H/C/000709/X/0056/G

Bristol-Myers Squibb Pharma EEIG

Rapporteur: Sinan B. Sarac, Co-Rapporteur: Fátima Ventura, PRAC Rapporteur: Doris Stenver

Scope: "Extension application to introduce a new pharmaceutical form (powder for oral suspension) associated with a new strength (10 mg/ml)

2. C.I.6.a(type II) to include the treatment of children and adolescents aged 1 year to 18 years with Ph+ chronic phase CML for Sprycel. Sections 4.1, 4.2, 4.4, 4.5, 4.8, 5.1 and 5.2 of the SmPC are updated in order to add the new indication and its relevant posology, to add a warning on effects on growth and development in the paediatric population and to update the safety information. The Package Leaflet is updated in accordance."

Action: For adoption

The Committee discussed the issues identified in this application, which were related to Algae growth inhibition test.

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. Bydureon - exenatide - EMEA/H/C/002020/II/0045

AstraZeneca AB

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

Scope: "Extension of Indication to include treatment in combination with basal insulin for Bydureon; as a consequence, sections 4.1, 4.2, 4.4, 4.8 and 5.1 of the SmPC are updated based on the study D5553C00002 (Duration 7 study) which evaluated safety and efficacy of exenatide once weekly therapy added to titrated basal insulin in patients with type 2 diabetes who have inadequate glycemic control on basal insulin with or without metformin. The Package Leaflet is updated accordingly. In addition, the Marketing authorisation holder (MAH) took the opportunity to make minor corrections in sections 4.8 and 5.1 of the SmPC. Furthermore, the updated RMP version 26 has been submitted."

Action: For adoption

The Committee noted the issues identified in this application.

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

5.1.2. Firazyr - icatibant - Orphan - EMEA/H/C/000899/II/0034/G

Shire Orphan Therapies GmbH

Rapporteur: Kristina Dunder, Co-Rapporteur: Nithyanandan Nagercoil, PRAC Rapporteur: Qun-Ying Yue

Scope: Extension of indication to include adolescents and children aged 2 years and older, with C1-esterase-inhibitor deficiency, for the use of Firazyr for symptomatic treatment of acute attacks of hereditary angioedema; as a consequence, sections 4.1, 4.2, 4.4, 4.8, 4.9, 5.1, 5.2, 5.3 and 6.6 of the SmPC are updated. The Package Leaflet is updated in accordance. The possibility of caregiver/self-administration has also been introduced. In addition, the Marketing authorisation holder (MAH) took the opportunity to reflect the results of a juvenile toxicity study in SmPC section 5.3.

Update section 5.2 of the SmPC to update the effect of age (elderly), gender and race on

the pharmacokinetics of icatibant. The Package Leaflet is updated accordingly.

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

Action: For adoption

Request for Supplementary Information adopted on 18.05.2017, 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.

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

Merck Sharp & Dohme Limited

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

Scope: "Extension of Indication to include 1st line treatment of metastatic non-squamous non-small cell lung cancer (NSCLC) in combination with platinum-pemetrexed chemotherapy based on the results from study KEYNOTE-021 (cohort G); a Phase 1/2, open-label trial of pembrolizumab in combination with chemotherapy or immunotherapy in patients with locally advanced or metastatic NSCLC.
As a consequence sections 4.1, 4.2, 4.4, 4.8 and 5.1 of the SmPC have been updated and the Package Leaflet has been updated accordingly.
An updated RMP version 9.0 was provided as part of the application."

Action: For adoption

Request for Supplementary Information adopted on 22.06.2017.

The Committee discussed the issues identified in this application. The Committee discussed the available clinical data and agreed that further data was required to substantiate the extension of indication application.

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

5.1.4. Nplate - romiplostim - Orphan - EMEA/H/C/000942/II/0060/G

Amgen Europe B.V.

Rapporteur: Concepcion Prieto Yerro, Co-Rapporteur: Paula Boudewina van Hennik, PRAC Rapporteur: Eva A. Segovia

Scope: "C.I.6.a - Extension of Indication to include paediatric population for Nplate: to register Nplate for the use in the paediatric chronic immune (idiopathic) thrombocytopenic purpura (ITP) patients: 1 year of age and older.

As a consequence Product information has been updated accordingly.
The RMP version 18 has also been submitted.

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

B.II.e.5.c – To add a low-dose romiplostim 125 microgram vial presentation for powder for solution for injection (4 vials pack).

B.II.e.5.a.1 – To add a 1 vial pack size of a low-dose romiplostim 125 microgram presentation."

Action: For adoption

Request for Supplementary Information adopted on 23.03.2017.

The Committee discussed the issues identified in this application, mainly relating to the wording in different SmPC sections. Furthermore the Committee was informed about some dose discrepancy cases observed in the clinical study, which raised questions about possible medication errors. It was agreed to request further clarification from the MAH.

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

5.1.5. 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/Oral explanation

Participation of patients' representatives

Request for Supplementary Information adopted on 18.05.2017, 23.02.2017, 15.09.2016.

See 2.3

An oral explanation was held on 13 September 2017 at time 14:00

The presentation by the MAH focused on the clinical efficacy and safety data and proposed post-authorisation measures.

After the oral explanation the members discussed the clinical trial conduct and results with specific focus on the efficacy data. The Committee questioned the clinical relevance and validity of the trial endpoints as well as the trial integrity. It was discussed whether the efficacy has sufficiently been proven by available clinical data taking different approval types into account.

The CHMP was of the opinion that the study results provided by the company were insufficient to determine the benefit of Raxone in patients with Duchenne muscular dystrophy. Although a difference in peak expiratory flow (PEF) in favour of Raxone was observed, there was no clear improvement in other indicators of breathing function or in muscle strength, motor function or quality of life. The Committee also had some concerns about the way the study was conducted and analysed.

Therefore, the CHMP was of the opinion that the benefits of Raxone in patients with Duchenne muscular dystrophy did not outweigh its risks. Hence, the CHMP recommended that the change to the marketing authorisation be refused.

The CHMP adopted a negative opinion by majority (24 negative out of 30 votes) recommending the refusal of the variation. The CHMP adopted the assessment report.

The Norwegian Member was in agreement with the CHMP recommendation, the Icelandic Member was not.

The divergent position (Bart van der Schueren, Daniela Melchiorri, Ewa Balkowiec Iskra, Hrefna Gudmundsdottir, Jayne Crowe, John Joseph Borg, Koenraad Norga) was appended to the opinion.

The refusal questions and answers document was circulated for information.

Post-meeting note: The final opinion was adopted via written procedure on 27.09.2017.

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

Amgen Europe B.V.

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

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

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

Action: For adoption

The Committee discussed the issues identified in this application. The members were updated on the clinical trial results, which showed a significant reduction on the composite primary endpoint. It was noted that the treatment effect is different in Europe compared to the US and also differences regarding race. The Committee debated on the wording of the indication and whether it is in line with the study population. The CHMP agreed to request further clarification from the MAH.

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

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

Gilead Sciences International Limited

Rapporteur: Robert James Hemmings, Co-Rapporteur: Joseph Emmerich, PRAC Rapporteur: Julie Williams

Scope: “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”

Action: For adoption

Request for Supplementary Information adopted on 18.05.2017.

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.

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

Eli Lilly Nederland B.V.

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

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

Action: For adoption

The Committee discussed the issues identified in this application, relating to the wording of the indication. It was considered that more data was required to support the extension of indication.

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

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

Gilead Sciences International Limited

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

Scope: “Extension of Indication to include pre-exposure prophylaxis of HIV in adolescents

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

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

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

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

Action: For adoption

Request for Supplementary Information adopted on 18.05.2017.

The Committee discussed the issues identified in this application. The Committee noted that the MAH should discuss the feasibility of implementing a PrEP Registry in the countries known to be publicly providing PrEP, with the potential for other member states participating as use of PrEP expands.

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

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

Bristol-Myers Squibb Pharma EEIG

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

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

Action: For adoption

Request for Supplementary Information adopted on 22.06.2017.

The Committee discussed the issues identified in this application. The members discussed the available clinical data and concluded the need for more safety data in the paediatric population. The Committee looked at different options for data generation post-marketing and agreed to follow up with the MHA on this.

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

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

Pfizer Limited

Rapporteur: Kristina Dunder, PRAC Rapporteur: Ulla Wändel Liminga

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

Scope: Request by the applicant dated 8 September 2017 requesting an extension of clock stop to respond to the second Request for Supplementary Information adopted on 20.07.2017.

Action: For adoption

Request for Supplementary Information adopted on 20.07.2017

The CHMP agreed to the request by the applicant for an extension of clock stop to respond to the second Request for Supplementary Information adopted on 20.07.2017.

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

8.1.1. – fexinidazole - Art. 58 - H0002320

treatment of both, the first-stage (hemo-lymphatic) and second-stage (meningo-encephalitic) of human African trypanosomiasis due to T.b. gambiense in adults and children ≥ 6 years old and weighing 20 kg or more

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

Action: For adoption

The CHMP agreed to the request for accelerated assessment and adopted the briefing note and Rapporteurs' recommendation on the Request for Accelerated Assessment.

8.1.2. – cytarabine, daunorubicin - H0004282

treatment of adults with high-risk Acute Myeloid Leukaemia (AML) as defined by therapy-related AML (t-AML) or AML with myelodysplasia-related changes (AML-MRC).

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

Action: For adoption

The CHMP agreed to the request for accelerated assessment and adopted the briefing note and Rapporteurs' recommendation on the Request for Accelerated Assessment.

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

The CHMP noted the list of applications received.

8.2.2. Recommendation for PRIME eligibility

Action: For adoption

The CHMP adopted the recommendation for PRIME eligibility. The CHMP reviewed 4 recommendations for eligibility to PRIME: all 4 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. Aranesp - darbepoetin alfa - EMEA/H/C/000332/II/0143

Amgen Europe B.V.

Rapporteur: Martina Weise

Scope: CHMP request for PRAC advice on the "signal procedure (EMEA/H/C000332/SDA/090) and addresses the potential need for additional risk minimisation measures and an amendment of the RMP and Annex IID a formal request for PRAC advice is considered necessary for the assessment and conclusions on this variation procedure"

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.

9.1.2. Cotellic - cobimetinib -EMEA/H/C/003960

Roche Registration Limited

CHMP Rapporteur: Filip Josephson, PRAC Rapporteur: Sabine Straus

Scope: A non-interventional Study to investigate the Effectiveness, Safety and Utilisation of cobimetinib and vemurafenib in Patients with and without Brain Metastasis with BRAF V600 mutant melanoma under real world Conditions - (covenis)

Action: For discussion

The CHMP discussed the proposed post-authorisation study to address the safety concern 'safety and effectiveness in patients with CNS involvement' and proposed amendments to the protocol. The CHMP adopted a request for a revised protocol with a specific timetable.

9.1.3. Deltyba - delamanid - EMEA/H/C/002552/II/0021, Orphan

MAH: Otsuka Novel Products GmbH

Rapporteur: Greg Markey

Scope: "Update of sections 4.2, 4.4, 4.8, 4.9, 5.1 and 5.2 of the SmPC to reflect the results of the final study report of 242-09-213 (A Phase 3, Multicenter, Randomized, Double-blind, Placebo-controlled, Parallel Group Trial to Evaluate the Safety and Efficacy of Delamanid (OPC-67683) Administered Orally as 200 mg Total Daily Dose for Six Months in Patients With Pulmonary Sputum Culture-positive, Multidrug-resistant Tuberculosis), submitted to

fulfill SOB-01. The Package leaflet is updated accordingly."

Action: For discussion

The CHMP discussed the data from the final study report. The CHMP considered the outcome not conclusive in terms of demonstrating the clinical efficacy.

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

9.1.4. Ibrance - palbociclib - EMEA/H/C/003853

MAH: Pfizer Limited

Rapporteur: Filip Josephson, Co-Rapporteur: Alexandre Moreau, PRAC Rapporteur: Torbjorn Callreus

Scope: Discuss the rationale supporting the indication of Ibrance in combination with an aromatase inhibitor irrespective of the line of therapy

Action: For discussion

The members discussed the rationale supporting the current wording of the indication of Ibrance in combination with an aromatase inhibitor. The CHMP confirmed that extrapolation from first line to subsequent lines of therapy was the basis for the indication wording in combination with an AI. Extrapolation was based on the clinical data submitted in the dossier, including the results of PALOMA-2 which demonstrated the suitability of combining Ibrance with an AI, as well as the results of PALOMA-3 study where the use of Ibrance in combination with fulvestrant was studied in patients whose disease progressed after prior endocrine therapy, and which demonstrated that an add-on effect of Ibrance is also present in second-line.

9.1.5. Opdivo - nivolumab - EMEA/H/C/003985/II/0036/G

MAH: Bristol-Myers Squibb Pharma EEIG

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

Scope: "C.I.4 (Type II) - Update of sections 4.2, 5.1, 5.2 and 6.6 of the SmPC in order to introduce new dosing regimens and schedule.

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

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

Action: For discussion

The Committee discussed the issues identified in this application, mainly relating to the proposed changes to the posology, which would also apply to the already approved indications. Further clarification was considered necessary for the new dosing regimens.

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

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

MAH: Amgen Europe B.V.

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

Scope: List of experts at ad hoc expert group meeting and CHMP list of questions to the ad hoc expert group

Action: For adoption

Request for Supplementary Information adopted on 22.06.2017.

The CHMP agreed to the draft list of experts to the ad hoc expert group and adopted a list of questions to this group.

9.1.7. Neulasta – pegfilgrastim - EMEA/H/C/000420/0093/G

MAH: Amgen Europe B.V.

Rapporteur: Robert James Hemmings, PRAC Rapporteur: Patrick Batty

Scope: "B.IV.1.a.3 (type II) – To add a new device which may have a significant impact on the delivery of the product: the on-body injector (Onpro kit) to be used with Neulasta, 6mg solution for injection, pre-filled syringe
- B.II.e.5.c (type II) – To change the fill volume from 0.6 to 0.64 mL for Neulasta, 6 mg, solution for injection pre-filled syringe co-packed with the new on-body injector (Onpro kit)
In addition the MAH took the opportunity to introduce editorial changes to module 3.2.P.2.4 Container Closure System
Update of sections 3, 4.2, 5.1, 6.4, 6.5, 6.6 and 8 of the SmPC. The Labelling, Package Leaflet and the RMP (version 4.2) are updated accordingly. In addition the Marketing authorisation holder (MAH) took the opportunity to update the list of local representatives in the Package Leaflet, to include some editorial changes and correct some typos throughout the product information, and to bring the product information in line with the latest QRD template version 10."

Action: For discussion

Request for Supplementary Information adopted on 22.06.2017.

The Committee discussed the issues identified in this application. The members were updated on recent developments and additional data provided by the MAH on the on-body injector. The members also noted the discussion at the PRAC concerning the RMP.

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

9.1.8. Tarceva - erlotinib - EMEA/H/C/000618/II/0052

Roche Registration Limited

Rapporteur: Sinan B. Sarac

Scope: Letter from the applicant requesting extension of clock stop to respond to Request for supplementary information adopted on 20.07.2017

"Update of section 4.4 of the SmPC in order to include recommendations on Epidermal Growth Factor Receptor (EGFR) mutation status testing, to be in line with current technical and scientific progress.
In addition, the Marketing authorisation holder (MAH) took the opportunity to make minor editorial changes and to bring the PI in line with the latest QRD template version 10. Moreover, the MAH took the opportunity to make minor correction of section 4.2 of the SmPC. Furthermore, the Annex II has been corrected, as requested by the EMA, to include Educational Material as an additional risk minimisation measure, which has been already in place in the RMP."

Action: For adoption

Request for Supplementary Information adopted on 20.07.2017.

The CHMP agreed to the request by the applicant for an extension of clock stop to respond to Request for supplementary information adopted on 20.07.2017

9.1.9. Keytruda – Pembrolizumab – EMEA/H/C/0003820

Merck Sharp & Dohme Limited

Rapporteur: Daniela Melchiorri, Co-Rapporteur: Jan Mueller-Berghaus

Scope: Halted clinical trials for Keytruda in combination with dexamethasone and an immunomodulatory (IMID) agent (lenalidomide or pomalidomide) for the treatment of patients with multiple myeloma

Action: For information

The CHMP was updated on the hold of clinical trials KEYNOTE-183 and KEYNOTE-185. The members noted that an updated safety dataset for Keytruda was expected from the MAH. A request was made for a benefit/risk discussion of all authorised indications in light of these findings. Following the assessment of the dataset the rapporteurs will make a proposal whether a formal procedure for further assessment of the benefit/risk in the approved indication is necessary. Depending on the regulatory pathway, the assessment will be performed by the CHMP and/or PRAC.

10. Referral procedures

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

10.1.1. Zinbryta - Daclizumab - EMEA/ H/A-20/1456

Biogen Idec Ltd

Rapporteurs for the Article 20 referral: PRAC Rapporteur: Eva Segovia; PRAC Co-rapporteur: Marcia Sofia Sanches de Castro Lopes Silva,

Rapporteurs for Zinbryta: CHMP Rapporteur: Bruno Sepodes, CHMP Co-rapporteur: Greg Markey

Scope: Review of the benefit-risk balance following notification by the European Commission of a referral under Article 20 of Regulation (EC) No 726/2004 based on pharmacovigilance data

List of question to the SAG

Action: For adoption

The CHMP agreed to the list of questions to the SAG as adopted by the PRAC.

Call for additional experts for the SAG meeting: additional expertise required: experts in drug related hepatic injury.

Nominations should be sent by the 25th September 2017.

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

No items

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

No items

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

No items

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

10.5.1. Scandonest and associated names – mepivacaine - EMEA/H/A-30/1455

Septodont group of companies and associated companies

Rapporteur: Romaldas Maciulaitis, Co-rapporteur: Fatima Ventura,

Scope: Start of procedure and timetable

Action: For adoption

Harmonisation exercise for Scandonest and associated names. Summary of Product Characteristics and Module 3 harmonisation was triggered by the MAH.

The CHMP noted the start of procedure and adopted the procedure timetable.

Notification & submission: 25.08.2017

Start of procedure (CHMP): September 2017 CHMP

Rapporteur / co-rapporteur assessment report(s) circulated to CHMP: 28.09.2017

Comments: 02.10.2017

Updated Rapporteur / co-rapporteur assessment reports circulated to CHMP: 05.10.2017

CHMP list of questions / CHMP opinion: October 2017 CHMP

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

10.6.1. 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); efmoroctocog alfa – Elocta (CAP); moroctocog alfa – Refacto AF (CAP) octocog alfa – Advate (CAP), Helixate Nexgen (CAP), Iblias (CAP), Kogenate (CAP), Kovaltry (CAP); turoctocog alfa – Novoeight (CAP); simoctocog alfa – Nuwiq (CAP); susoctocog alfa – Obizur (CAP) - EMEA/H/A-31/1448

Baxter AG (Advate), Bayer Pharma AG (Helixate Nexgen, Iblias, Kogenate, Kovaltry), CSL Behring GmbH (Voncento), Novo Nordisk A/S (NovoEight), Octapharma AB (Nuwiq), Pfizer Limited (Refacto AF), Swedish Orphan Biovitrum AB (publ) (Elocta), Baxalta Innovations GmbH (Obizur), various

PRAC led referral - PRAC Rapporteur: Jan Neuhauser; PRAC Co-rapporteur: Ghania Chamouni

Scope: Opinion

Action: For adoption

Re-examination procedure under Article 32 of Directive 2001/83/EC of the 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, having considered the PRAC recommendation, adopted an opinion by consensus, recommending that the marketing authorisations for human coagulation factor VIII, efmoroctocog alfa, moroctocog alfa, octocog alfa, simoctocog alfa and turoctocog alfa should be varied.

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

10.6.2. Gadolinium-containing contrast agents (GdCA): gadobutrol (NAP); gadodiamide (NAP); gadopentetic acid (NAP); gadoteric acid (NAP); gadoteridol (NAP); gadoxetic acid (NAP) - EMEA/H/A-31/1097

Lead Rapporteur: Patrick Batty,

Scope: Annual cumulative reviews on NSF cases submission as a post-authorisation measure resulting from the 2010 Article 20 and Article 31 referral procedures for gadolinium-containing contrast agents, time table

Action: For adoption

The CHMP adopted the procedural timetable for assessment of the Annual cumulative reviews.

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

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

D&A Pharma

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

Scope: List of questions to the SAG, list of experts for the ad hoc expert group meeting

Action: For adoption

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

The CHMP adopted a list of questions to the SAG.

Post-meeting note: the draft list of experts for the ad hoc expert group meeting were adopted via written procedure on 29.09.2017.

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

No items

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

No items

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

No items

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

No items

11. Pharmacovigilance issue

11.1. Early Notification System

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

Action: For information

The CHMP noted the September 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

No items

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

Area of expertise of co-opted member

The mandate of co-opted member (expertise in Quality and safety of biological medicinal products, including biotechnology derived medicines, cell therapies and gene therapies) will expire on 13 November 2017.

Proposed timeline for appointment of the co-opted member:

Confirmation of areas of expertise: September 2017

Call for nomination for expert: September/October 2017

Election of co-opted member: November 2017

Action: For discussion

The CHMP noted the information. Further discussions will be held. Please send any proposals for areas of expertise **by 6th October 2017**.

CHMP/CAT joint membership

The Advanced Therapies Regulation ((EC) 1394/2007) requires that 5 members or co-opted members of the Committee for Medicinal Products for Human Use (CHMP) together with an alternate, either proposed by the Member state of the member or identified by the co-opted member, are appointed by the CHMP to the Committee for Advanced Therapies (CAT). The Member States, who are not represented through the members appointed by the CHMP, nominate then one member and alternate to the CAT.

Action: For discussion

The CHMP noted the information.

Improvement of Rapporteurship bidding process

Action: For information

The CHMP members were informed about a new 2-stage bidding process for Rapporteurships.

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 29 August – 1 September 2017

Action: For information

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

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

Action: For adoption

The CHMP adopted the list.

Comments on post consultation on 'Guideline on good pharmacovigilance practices (GVP), Module XV – Safety communication (Rev 1)' should be sent **by 18 September 2017**

Action: For information

The CHMP noted the information.

14.2.2. Committee for Advanced Therapies (CAT)

CAT draft minutes of meeting held on 6-8 September 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 11-12 July 2017

Action: For information

The CHMP noted the report.

14.2.4. Paediatric Committee (PDCO)

PIPs reaching D30 at September 2017 PDCO

Action: For information

The CHMP noted the information.

Report from the PDCO meeting held on 15-18 August 2017

Action: For information

The CHMP noted the information.

14.2.5. Committee for Orphan Medicinal Products (COMP)

Report from the COMP meeting held on 5-7 September 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 11-13 September 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)

Report from the SAWP meeting held on 29 August – 1 September 2017. Table of conclusions

Action: For information

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

14.3.2. The CHMP noted the report.Pharmacogenomics Working Party (PGWP)

Chair: Krishna Prasad/Markus Paulmichl

Election of PGWP Vice Chair, the term of the current Vice Chair ending September 2017

Action: For adoption

The CHMP elected Markus Paulmichl as PGWP Vice Chair for a second term of three years.

14.3.3. Pharmacokinetics Working Party (PKWP)

Chair: Jan Welink/Alfredo Garcia-Arieta

Election of PKWP Vice Chair, the term of the current Vice Chair ending September 2017

Action: For adoption

The CHMP elected Henrike Potthast as PKWP Vice Chair for a term of three years.

14.3.4. Infectious Disease Working Party (IDWP)

Chair: TBC

Election of IDWP Vice-Chair, the term of the current Vice Chair ended in July 2017

Action: For adoption

The CHMP elected María Jesús Fernández Cortizo as IDWP Vice Chair for a term of three years.

Call for nominations for Chair position: Nominations together with a brief resume in support of their candidature should be sent

Action: For information

The CHMP noted the call for nominations for the IDWP Chair.

14.3.5. Central Nervous System Working Party (CNSWP)

Chair: Karl Broich

Nomination of Petr Vrbata (CZ) as new observer to the CNSWP

Action: For adoption

The CHMP nominated Petr Vrbata (CZ) as new observer to the CNSWP.

14.3.6. Cardiovascular Working Party (CVSWP)

Chair: Pieter de Graeff/Kristina Dunder

Guideline on clinical investigation of medicinal products in the treatment or prevention of diabetes mellitus (CPMP/EWP/1080/00 Rev. 2)

Action: For adoption for 6-months public consultation

The CHMP decided to update further the guideline with regards to estimands concept in line with the ICH E9 (R1) addendum on estimands and sensitivity analysis in clinical trials to the guideline on statistical principles for clinical trials. Therefore the adoption of the guideline for public consultation postponed.

14.3.7. Oncology Working Party (ONCWP)

Chair: Pierre Demolis/Paolo Foggi

Guideline on the evaluation of anticancer medicinal products in man (EMA/CHMP/205/95 Rev.5)

Action: For adoption

The CHMP adopted the guideline. The purpose of the 5th revision of the main guideline is to address current changes in the therapeutic landscape that affect the requirements with regard to collection and reporting of safety data in order to inform the benefit-risk evaluation, including a need for more differentiated and detailed safety data presentation.

14.3.8. Biosimilar Medicinal Product Working Party (BMWP)

Chair: Elena Wolff-Holz/Martina Weise

Nomination of new member to the BMWP

Action: For adoption

The CHMP appointed Sean Barry as new member to BMWP.

Election of BMWP Vice-Chair, the term of the current Vice Chair ended in July 2017

Action: For adoption

The CHMP elected Niklas Ekman (FI) as Vice-Chair to BMWP for a term of three years.

14.3.9. Safety Working Party (SWP)

Chair: Jan Willem Van der Laan

Nomination of Bill Vestergaard (DK) and Camilla Ærtebjerg (DK) as observers to the SWP

Action: For adoption

The CHMP nominated Bill Vestergaard (DK) and Camilla Ærtebjerg (DK) as observers to the SWP

14.3.10. Extrapolation Working Group (EWG)

Reflection paper on the use of extrapolation in the development of medicines for paediatrics

CHMP: Robert James Hemmings

Action: For discussion

The CHMP noted the reflection paper. Comments should be sent by 25 September 2017. Further discussion and adoption is planned for the October CHMP.

14.3.11. Antimicrobial Advice Ad Hoc Expert Group (AMEG)

New mandate

Action: For discussion

Antimicrobial Advice Ad Hoc Expert Group (AMEG) provided a scientific advice on the impact of the use of antibiotics in animals on public health and animal health and measures to manage the possible risk to humans (2013-2016). In April 2017, the European Commission (EC) requested EMA to revise the AMEG categorisation of antimicrobials. In addition there is a need to further elaborate on the proposed early hazard categorisation. The group will be

composed of members of the CVMP and its Antimicrobials Working Party (AWP), CHMP and its Infectious Disease Working Party (IDWP) and experts from ECDC, EFSA, JIACRA, EURL-AR.

14.4. Cooperation within the EU regulatory network

None

14.5. Cooperation with International Regulators

None

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

None

14.7. CHMP work plan

14.7.1. CHMP 2018 Work Plan: proposed list of topics

Action: For information

The CHMP noted the proposed list of topics for 2018 and agreed to it.

14.8. Planning and reporting

14.8.1. New marketing authorisation applications for 2017 with and without appointed rapporteurs

Action: For information

The CHMP noted the new marketing authorisation applications for 2017 with and without appointed rapporteurs.

14.9. Others

None

15. Any other business

15.1. AOB topic

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

CHMP Rapporteur: Harald Enzmann

Scope: Overview of comments of the revised guideline

Action: For information

The revised guideline was adopted at the July 2017 plenary.

The CHMP noted the overview of comments. The comments will be published.

15.1.2. Preparedness of the system and capacity increase

Action: For discussion

The CHMP noted the update and next steps.

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 11 – 14 September 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	
Emilia Mavrokordatou	Member	Cyprus	No interests declared	
Ondřej Slanař	Member	Czech Republic	No interests declared	
Tomas Boran	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	

Name	Role	Member State or affiliation	Outcome restriction following evaluation of e-DoI	Topics on agenda for which restrictions apply
Alexandre Moreau	Member	France	No interests declared	
Joseph Emmerich	Alternate	France	No interests declared	
Harald Enzmann	Member (Vice-Chair)	Germany	No interests declared	
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	
Jayne Crowe	Member	Ireland	No interests declared	
Peter Kiely	Alternate	Ireland	No interests declared	
Daniela Melchiorri	Member	Italy	No restrictions applicable to this meeting	
Luca Pani	Alternate	Italy	No restrictions applicable to this meeting	
Juris Pokrotnieks	Member	Latvia	No participation in final deliberations and voting on:	3.3.6. - budesonide - Orphan - EMEA/H/C/004655
Natalja Karpova	Alternate	Latvia	No interests declared	
Romaldas Mačiulaitis	Member	Lithuania	No participation in final deliberations and voting on:	2.1.2. - ocrelizumab - EMEA/H/C/004043 3.1.7. - ocrelizumab - EMEA/H/C/004043 9.1.2. Cotellic - cobimetinib - EMEA/H/C/003960 9.1.9. Tarceva - erlotinib - EMEA/H/C/000618/II/0052
John Joseph Borg	Member	Malta	No interests	

Name	Role	Member State or affiliation	Outcome restriction following evaluation of e-DoI	Topics on agenda for which restrictions apply
			declared	
Johann Lodewijk Hillege	Member	Netherlands	No interests declared	
Paula Boudewina van Hennik	Alternate	Netherlands	No interests declared	
Svein Rune Andersen	Member	Norway	No interests declared	
Bjorg Bolstad	Alternate	Norway	No restrictions applicable to this meeting	
Ewa Balkowiec Iskra	Member	Poland	No interests declared	
Aldona Paluchowska	Alternate	Poland	No interests declared	
Bruno Sepodes	Member - via Adobe	Portugal	No interests declared	
Fatima Ventura	Alternate	Portugal	No participation in final deliberations and voting on:	3.2.13. - d-biotin - EMEA/H/C/004153
Simona Badoi	Member	Romania	No interests declared	
Frantisek Drafi	Member	Slovakia	No interests declared	
Eva Malikova	Alternate	Slovakia	No interests declared	
Nevenka Trsinar Brodt	Alternate	Slovenia	No interests declared	
Concepcion Prieto Yerro	Member	Spain	No interests declared	
Jorge Camarero Jiménez	Member	Spain	No participation in final deliberations and voting on:	2.1.2. - ocrelizumab - EMEA/H/C/004043 3.1.7. - ocrelizumab - EMEA/H/C/004043 9.1.2. Cotellic - cobimetinib - EMEA/H/C/003960 9.1.9. Tarceva - erlotinib - EMEA/H/C/000618/II/0052
Kristina Dunder	Member	Sweden	No interests declared	
Filip Josephson	Alternate	Sweden	No interests declared	
Greg Markey	Member	United	No interests	

Name	Role	Member State or affiliation	Outcome restriction following evaluation of e-DoI	Topics on agenda for which restrictions apply
		Kingdom	declared	
Nithyanandan Nagercoil	Alternate	United Kingdom	No restrictions applicable to this meeting	
Koenraad Norga	Co-opted member	Belgium	No participation in final deliberations and voting on:	3.1.2. Elebrato Ellipta - fluticasone furoate / umeclidinium / vilanterol - EMEA/H/C/004781 3.1.10. Trelegy Ellipta - fluticasone furoate / umeclidinium / vilanterol - EMEA/H/C/004363 3.2.19. - human herpesvirus 3 - EMEA/H/C/004336 4.1.1. Benlysta - belimumab - EMEA/H/C/002015/X/0046/G
Jan Mueller-Berghaus	Co-opted member	Germany	No interests declared	
Jean-Louis Robert	Co-opted member	Luxembourg	No interests declared	
Sol Ruiz	Co-opted member	Spain	No interests declared	
Robert James Hemmings	Co-opted member	United Kingdom	No restrictions applicable to this meeting	
Philip Lange Moller	Expert - in person*	Denmark	No interests declared	
Trine Jensen	Expert - in person*	Denmark	No restrictions applicable to this meeting	
Sabine Mayrhofer	Expert - in person*	Germany	No interests declared	
Thomas Lang	Expert - by phone*	Austria	No interests declared	
Marleen Laloup	Expert - by Adobe	Belgium	No restrictions applicable to this meeting	
Eskild Colding-Jorgensen	Expert - by phone*	Denmark	No restrictions applicable to this meeting	
Pauline Dayani	Expert - by phone*	France	No interests declared	
Serge Bakchine	Expert - by phone*	France	No interests declared	

Name	Role	Member State or affiliation	Outcome restriction following evaluation of e-DoI	Topics on agenda for which restrictions apply
Martina Schussler-Lenz	Expert - by phone*	Germany	No interests declared	
Franco Granella	Expert - by phone*	Italy	No restrictions applicable to this meeting	
Maura O'Donovan	Expert - by phone*	Ireland	No interests declared	
Menno van der Elst	Expert - by phone*	Netherlands	No interests declared	
Didier Meulendijks	Expert - by phone*	Netherlands	No restrictions applicable to this meeting	
Joao Freire	Expert - by phone*	Portugal	No restrictions applicable to this meeting	
Maria Escudero Galindo	Expert - in person*	Spain	No restrictions applicable to this meeting	
Rolf Gedeborg	Expert - by phone*	Sweden	No interests declared	
Jonas Bergh	Expert - by phone*	Sweden	Indirect interests declared	
Sabine Lenton	Expert - by phone*	United Kingdom	No interests declared	
Cecilia Chisholm	Expert - by phone*	United Kingdom	No interests declared	
Patients' Representative		Patient observer	No interests declared	
Patients' Representative	via telephone*	Patient observer	No interests declared	
Olga Kholmanskikh	Expert - by Adobe	Belgium	No interests declared	
Diederica Claeys	Expert - by Adobe	Belgium	No interests declared	
Janet Schriever	Expert - by Adobe	Germany	No interests declared	
A representative from the European Commission attended the meeting				

Name	Role	Member State or affiliation	Outcome restriction following evaluation of e-DoI	Topics on agenda for which restrictions apply
Meeting run with support from relevant EMA staff				

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)*

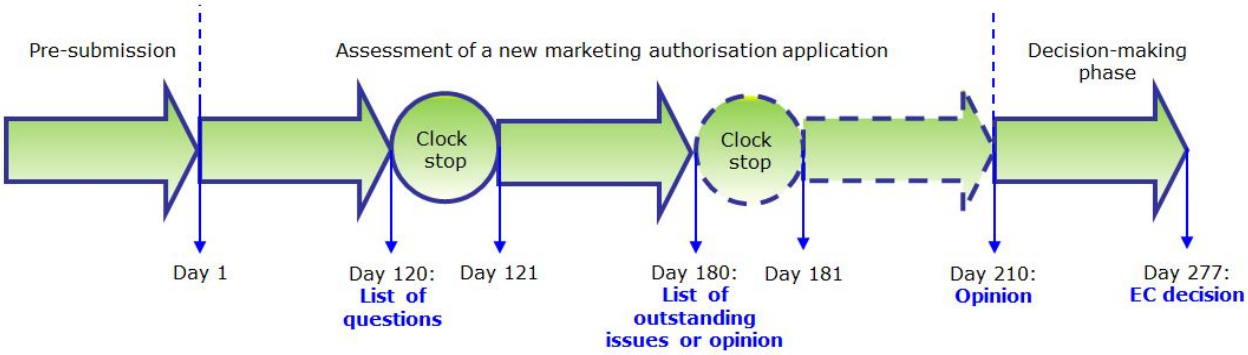
The items listed in this section are those for which marketing authorisation holders (MAHs) or applicants have been invited to the CHMP plenary meeting to address questions raised by the Committee. Oral explanations normally relate to on-going applications (section 3, 4 and 5) or referral procedures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

20 October 2017
EMA/CHMP/479459/2017[1]

Annex to 11-14 September 2017 CHMP Minutes

Pre submission and post authorisations issues

[1] Correction in section B.4.

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

A. PRE SUBMISSION ISSUES

A.1. ELIGIBILITY REQUESTS

Report on Eligibility to Centralised Procedure for September 2017: **For adoption**	Adopted.

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

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

A.3. PRE-SUBMISSION ISSUES FOR INFORMATION

Information related to pre-submission of initial applications cannot be released at the present time as these contain commercially confidential information.

B. POST-AUTHORISATION PROCEDURES OUTCOMES

B.1. Annual re-assessment outcomes

B.1.1. Annual reassessment for products authorised under exceptional circumstances

Firdapse - amifampridine - EMEA/H/C/001032/S/0049, Orphan MAH: BioMarin Europe Ltd, Rapporteur: Greg Markey, PRAC Rapporteur: Julie Williams Request for Supplementary Information adopted on 22.06.2017.	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.

B.2. RENEWALS OF MARKETING AUTHORISATIONS OUTCOMES

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

NexoBrid - concentrate of proteolytic enzymes enriched in bromelain - EMEA/H/C/002246/R/0031, Orphan MAH: MediWound Germany GmbH, Rapporteur: Harald Enzmann, Co-Rapporteur: Robert James Hemmings, PRAC Rapporteur: Valerie Strassmann Request for Supplementary Information adopted on 22.06.2017.	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 Opinion.

B.2.2. Renewals of Marketing Authorisations for unlimited validity

Adasuve - loxapine - EMEA/H/C/002400/R/0024 MAH: Ferrer Internacional s.a., Rapporteur: Johann Lodewijk Hillege, Co-Rapporteur: Daniela Melchiorri, PRAC Rapporteur: Sabine Straus Request for Supplementary Information adopted on 14.09.2017.	Request for Supplementary Information adopted
Glubrava - pioglitazone / metformin hydrochloride - EMEA/H/C/000893/R/0054 MAH: Takeda Pharma A/S, Informed Consent of Competact, Rapporteur: Peter Kiely, Co-Rapporteur: Concepcion Prieto Yerro, PRAC Rapporteur: Almath Spooner Request for Supplementary Information adopted on 22.06.2017.	Positive Opinion adopted by consensus together with the CHMP assessment report. 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.
Jetrea - ocriplasmin - EMEA/H/C/002381/R/0033 MAH: ThromboGenics NV, Rapporteur: Greg Markey, Co-Rapporteur: Kristina Dunder, PRAC Rapporteur: Julie Williams Request for Supplementary Information adopted on 14.09.2017.	Request for Supplementary Information adopted
Pradaxa - dabigatran etexilate - EMEA/H/C/000829/R/0105 MAH: Boehringer Ingelheim International GmbH, Rapporteur: Hanne Lomholt Larsen, Co-Rapporteur: Joseph Emmerich, PRAC Rapporteur: Doris Stenver Request for Supplementary Information adopted on 14.09.2017.	Request for Supplementary Information adopted
Selincro - nalmefene - EMEA/H/C/002583/R/0022 MAH: H. Lundbeck A/S, Rapporteur: Harald Enzmann, Co-Rapporteur: Peter Kiely, PRAC Rapporteur: Martin Huber, Procedure Manager: Viktor Vlcek, EPL: Lorenzo Guizzaro	Positive Opinion adopted by consensus together with the CHMP assessment report. 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

Adcetris - brentuximab vedotin - EMEA/H/C/002455/R/0051, Orphan MAH: Takeda Pharma A/S, Rapporteur: Paula Boudewina van Hennik, PRAC Rapporteur:	Positive Opinion adopted by consensus together with the CHMP assessment report and translation timetable.

Sabine Straus Request for Supplementary Information adopted on 20.07.2017.	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.
Ocaliva - obeticholic acid - EMEA/H/C/004093/R/0002, Orphan MAH: Intercept Pharma Ltd, Rapporteur: Jorge Camarero Jiménez, Co-Rapporteur: Daniela Melchiorri, PRAC Rapporteur: Menno van der Elst Request for Supplementary Information adopted on 14.09.2017.	Request for Supplementary Information adopted
Venclyxto - venetoclax - EMEA/H/C/004106/R/0005, Orphan MAH: AbbVie Limited, Rapporteur: Filip Josephson, PRAC Rapporteur: Patrick Batty Request for Supplementary Information adopted on 20.07.2017.	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 29 August – 1 September 2017 PRAC:	
Signal of dystonia **Pramipexole - Mirapexin - pramipexole - EMEA/H/C/000134** Boehringer Ingelheim International Rapporteur: Hanne Lomholt Larsen, Co-Rapporteur: Outi Mäki-Ikola, PRAC Rapporteur: Doris Stenver, **Sifrol - pramipexole - EMEA/H/C/000133** Boehringer Ingelheim International Rapporteur: Hanne Lomholt Larsen, Co-Rapporteur: Outi Mäki-Ikola, PRAC Rapporteur: Doris Stenver, P **Oprymea - pramipexole - EMEA/H/C/00941** KRKA, d.d., Novo mesto	Adopted.

Rapporteur: Hanne Lomholt Larsen, PRAC Rapporteur: Doris Stenver, **Pramipexole Teva - pramipexole - EMEA/H/C/00940** Teva B.V. Rapporteur: Filip Josephson, Co-Rapporteur: Hanne Lomholt Larsen, PRAC Rapporteur: Qun-Ying Yue, **Pramipexole Accord - pramipexole - EMEA/H/C/00291** Accord Healthcare Limited Rapporteur: Svein Rune Andersen, Co-Rapporteur: Hanne Lomholt Larsen, PRAC Rapporteur: Doris Stenver,	
PSUR procedures for which PRAC adopted a recommendation for variation of the terms of the MA at its 29 August – 1 September 2017 PRAC meeting:	
Pyramax - pyronaridine / artesunate - EMEA/H/W/002319/PSUV/0016 (without RMP) MAH: Shin Poong Pharmaceutical Co., Ltd., PRAC Rapporteur: Caroline Laborde, “16 August 2016 – 15 February 2017 Update of Annex II.C Other conditions and requirements of the scientific opinion holder to reflect the revised PSUR submission frequency of 1 year.”	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 Article 58 CHMP Scientific Opinion for the above mentioned medicinal product, concerning the following change: Update of Annex II.C Other conditions and requirements of the scientific opinion holder to reflect the revised PSUR submission frequency of 1 year. The Icelandic and the Norwegian CHMP members agree with the above-mentioned recommendation of the CHMP.
EMEA/H/C/PSUSA/00000057/201612 (adalimumab (except for biosimilars)) CAPS: **Humira** (EMEA/H/C/000481) (adalimumab), MAH: AbbVie Limited, Rapporteur: Kristina Dunder, PRAC Rapporteur: Ulla Wändel Liminga, “DLP 31/12/2016, 3 years”	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 2 of the PL to add a clarifying statement that allergic reactions with Humira in rare cases can be life-threatening.” The Icelandic and the Norwegian CHMP members agree with the above-mentioned recommendation of the CHMP.

EMEA/H/C/PSUSA/00000985/201701 (dexamethasone (centrally authorised product indicated in uveitis and macular oedema)) CAPS: **Ozurdex** (EMEA/H/C/001140) (dexamethasone), MAH: Allergan Pharmaceuticals Ireland, Rapporteur: Greg Markey, PRAC Rapporteur: Julie Williams, "28 Jan 2016 – 27 Jan 2017"	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 the SmPC sections 4.2 and 4.8 to reflect that patients receiving more than 2 injections experience more adverse reactions. The Icelandic and the Norwegian CHMP members agree with the above-mentioned recommendation of the CHMP
EMEA/H/C/PSUSA/00001152/201611 (docetaxel) CAPS: **Docetaxel Winthrop** (EMEA/H/C/000808) (docetaxel), MAH: Aventis Pharma S.A., Rapporteur: Alexandre Moreau **Taxotere** (EMEA/H/C/000073) (docetaxel), MAH: Aventis Pharma S.A., Rapporteur: Alexandre Moreau NAPS: **Taxotan** - MEDICOPHARM AG PRAC Rapporteur: Ghania Chamouni, "01-Dec-2013 – 30-Nov-2016"	The CHMP, having considered in accordance with Article 28 of Regulation (EC) No 726/2004 and Article 107g(3) of Directive 2001/83/EC the PSUR on the basis of the PRAC recommendation and the PRAC assessment report as appended together with the detailed explanation of the scientific grounds for the differences with the PRAC recommendation, recommends by consensus the variation to the terms of the marketing authorisation(s) for the medicinal products containing the above referred active substance(s), concerning the following change(s): Update of section 4.8 of the SmPC to add the adverse reaction injection site recall reaction with a frequency unknown. Update of sections 4.4 and 4.8 of the SmPC to add a warning on hypersensitivity reactions to docetaxel in patients that have had previous hypersensitivity reaction to paclitaxel and to add this adverse drug reaction with a frequency unknown. Update of sections 4.4, 4.5 and 4.7 of the SmPC to update the information related to the risk of potential effects of alcohol and interactions with other medicinal products. 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/00002875/201701 (gimeracil / oteracil monopotassium / tegafur)	The CHMP, having considered in accordance with Article 28 of Regulation (EC) No 726/2004 the

CAPS: **Teysuno** (EMEA/H/C/001242) (tegafur / gimeracil / oteracil), MAH: Nordic Group B.V., Rapporteur: Paula Boudewina van Hennik, PRAC Rapporteur: Sabine Straus, “25 Jan 2016 – 24 Jan 2017”	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 products, concerning the following changes: Update of sections 4.4 and 4.8 of the SmPC to add Hepatitis B reactivation as an adverse reaction with a frequency of rare/ very rare .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/00002973/201612 (tipranavir) CAPS: **Aptivus** (EMEA/H/C/000631) (tipranavir), MAH: Boehringer Ingelheim International GmbH, Rapporteur: Joseph Emmerich, PRAC Rapporteur: Caroline Laborde, “01 January 2014 to 3 I December 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 for the above mentioned medicinal product, concerning the following change: Update of section 4.3 and 4.5 of the SmPC to add the contraindication to use concomitantly lurasidone. 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/00010275/201701 (peginterferon beta-1A) CAPS: **Plegridy** (EMEA/H/C/002827) (peginterferon beta-1a), MAH: Biogen Idec Ltd, Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Julie Williams, “19 Jul 2016 to 18 Jan 2017”	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 together with the detailed explanation of the scientific grounds for the differences with the PRAC recommendation, 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 add “alopecia” with a frequency “common”. 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/00010303/201701 (idelalisib)	The CHMP, having considered in accordance with Article 28 of Regulation (EC) No 726/2004 the

CAPS: **Zydelig** (EMEA/H/C/003843) (idelalisib), MAH: Gilead Sciences International Limited, Rapporteur: Filip Josephson, PRAC Rapporteur: Patrick Batty, “23 Jul 2016 - 22 Jan 2017”	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): To update section 4.4 of SmPC with hepatotoxicity data and to update section 5.1 of SmPC with drug induced lymphocytosis, section 4.8 has been updated with hepatocellular injury as common adverse reaction and lymphocytosis as very common adverse reaction. The Icelandic and the Norwegian CHMP members agree with the above-mentioned recommendation of the CHMP.
EMEA/H/C/PSUSA/00010363/201701 (dasabuvir) CAPS: **Exviera** (EMEA/H/C/003837) (dasabuvir), MAH: AbbVie Limited, Rapporteur: Filip Josephson, PRAC Rapporteur: Dolores Montero Corominas, “15 July 2016 – 14 January 2017”	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 for the above mentioned medicinal product, concerning the following changes: Update of section 4.4 of the SmPC to add a warning on depression, suicidal ideation and suicide attempt. 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/00010367/201701 (ombitasvir / paritaprevir / ritonavir) CAPS: **Viekirax** (EMEA/H/C/003839) (ombitasvir / paritaprevir / ritonavir), MAH: AbbVie Limited, Rapporteur: Filip Josephson, PRAC Rapporteur: Dolores Montero Corominas, “15 July 2016 – 14 January 2017”	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 for the above mentioned medicinal product, concerning the following change: Update of section 4.4 of the SmPC to add a warning on depression, suicidal ideation and suicide attempt. 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/00010448/201701 (carfilzomib) CAPS: **Kyprolis** (EMEA/H/C/003790) (carfilzomib),	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,

MAH: Amgen Europe B.V., Rapporteur: Jorge Camarero Jiménez, PRAC Rapporteur: Nikica Mirošević Skvrce, "20 July 2016 to 19 January 2017"	recommends by consensus, the variation to the terms of the marketing authorisation for the above mentioned medicinal product, concerning the following changes: Update of section 4.8 of the SmPC to add the adverse reaction tinnitus with a frequency common. The Package leaflet is updated accordingly. In addition the Marketing Authorisation Holder took the opportunity to make some small formatting changes throughout the Product Information and to amend the contact details of the Slovak local representative in the Package Leaflet. The Icelandic and the Norwegian CHMP members agree with the above-mentioned recommendation of the CHMP.
EMEA/H/C/PSUSA/00010094/201702 (florbetaben (18f)) CAPS: **Neuraceq** (EMEA/H/C/002553) (florbetaben (18F)), MAH: Piramal Imaging Limited, Rapporteur: Concepcion Prieto Yerro, PRAC Rapporteur: Patrick Batty, "21 August 2016 - 20 February 2017"	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 change the frequency of the adverse reaction 'Injection site irritation' from common to uncommon. The Package leaflet is updated accordingly. The Icelandic and the Norwegian CHMP members agree with the above-mentioned recommendation of the CHMP.

B.4. EPARs / WPARs

Bavencio - avelumab - EMEA/H/C/004338 Applicant: Merck Serono Europe Limited, treatment of adult patients with metastatic Merkel cell carcinoma (MCC New active substance (Article 8(3) of Directive No 2001/83/EC)	adopted.
Dupixent - dupilumab - EMEA/H/C/004390 Applicant: sanofi-aventis groupe, treatment of moderate-to-severe atopic dermatitis, New active substance (Article 8(3) of Directive No 2001/83/EC)	adopted.
Entecavir Mylan - entecavir - EMEA/H/C/004377 Applicant: Mylan S.A.S, treatment of chronic hepatitis B virus infection, Generic, Generic of	adopted.

Baraclude Generic application (Article 10(1) of Directive No 2001/83/EC)	
Fulphila - pegfilgrastim - EMEA/H/C/004262 Applicant: Mylan S.A.S, treatment of neutropenia Similar biological application (Article 10(4) of Directive No 2001/83/EC) **WPAR**	adopted.
Infinia - alpha-1-antitrypsin - EMEA/H/C/003934, Orphan Applicant: Kamada BioPharma Limited at Fieldfisher LLP, treatment and maintenance therapy of adult patients with congenital deficiency of alpha-1 antitrypsin and lung disease with clinical evidence of emphysema and airway obstruction (FEV1/SVC<70%), Known active substance (Article 8(3) of Directive No 2001/83/EC) **WPAR**	adopted.
Ogivri - trastuzumab - EMEA/H/C/004346 Applicant: Mylan S.A.S, treatment of metastatic and early breast cancer and metastatic gastric cancer (MGC), Similar biological application (Article 10(4) of Directive No 2001/83/EC) **WPAR**	adopted.
Samsca - tolvaptan - EMEA/H/C/000980 Applicant: Otsuka Pharmaceutical Europe Ltd, treatment of adult patients with hyponatraemia secondary to syndrome of inappropriate antidiuretic hormone secretion (SIADH) New active substance (Article 8(3) of Directive No 2001/83/EC)	adopted.
Xermelo - telotristat ethyl - EMEA/H/C/003937, Orphan Applicant: Ipsen Pharma, treatment of carcinoid syndrome, New active substance (Article 8(3) of Directive No 2001/83/EC)	adopted.
Tigecycline Accord - tigecycline - EMEA/H/C/004419 Applicant: Accord Healthcare Ltd; 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	adopted.

not suitable	
Generic application (Article 10(1) of Directive No 2001/83/EC), Generic of Tygacil **WPAR**	

B.5. TYPE II VARIATION, WORKSHARING PROCEDURE OUTCOMES

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

B.5.1. CHMP assessed procedures scope: Pharmaceutical aspects

Advate - octocog alfa - EMEA/H/C/000520/II/0085 MAH: Baxter AG, Rapporteur: Jan Mueller-Berghaus Opinion adopted on 14.09.2017. Request for Supplementary Information adopted on 20.07.2017.	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
BeneFIX - nonacog alfa - EMEA/H/C/000139/II/0146 MAH: Pfizer Limited, Rapporteur: Jan Mueller-Berghaus Request for Supplementary Information adopted on 14.09.2017.	Request for Supplementary Information adopted with a specific timetable.
Benepali - etanercept - EMEA/H/C/004007/II/0026 MAH: Samsung Bioepis UK Limited, Rapporteur: Andrea Laslop Request for Supplementary Information adopted on 14.09.2017.	Request for Supplementary Information adopted with a specific timetable.
Betaferon - interferon beta-1b - EMEA/H/C/000081/II/0114 MAH: Bayer AG, Rapporteur: Greg Markey Opinion adopted on 14.09.2017. Request for Supplementary Information adopted on 15.06.2017.	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Cosentyx - secukinumab - EMEA/H/C/003729/II/0026 MAH: Novartis Europharm Ltd, Rapporteur: Tuomo Lapveteläinen Opinion adopted on 14.09.2017.	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Cystadane - betaine anhydrous - EMEA/H/C/000678/II/0029 MAH: Orphan Europe SARL, Rapporteur: Harald	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP

Enzmann Opinion adopted on 14.09.2017.	recommendation.
Darunavir Mylan - darunavir - EMEA/H/C/004068/II/0001/G MAH: Mylan S.A.S, Generic, Generic of Prezista, Rapporteur: John Joseph Borg Request for Supplementary Information adopted on 14.09.2017, 05.05.2017.	Request for Supplementary Information adopted with a specific timetable.
Deltyba - delamanid - EMEA/H/C/002552/II/0020/G, Orphan MAH: Otsuka Novel Products GmbH, Rapporteur: Greg Markey Opinion adopted on 14.09.2017.	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Elocta - efmoroctocog alfa - EMEA/H/C/003964/II/0016/G MAH: Swedish Orphan Biovitrum AB (publ), Rapporteur: Jan Mueller-Berghaus Request for Supplementary Information adopted on 14.09.2017.	Request for Supplementary Information adopted with a specific timetable.
Extavia - interferon beta-1b - EMEA/H/C/000933/II/0084 MAH: Novartis Europharm Ltd, Informed Consent of Betaferon, Rapporteur: Greg Markey Opinion adopted on 14.09.2017. Request for Supplementary Information adopted on 15.06.2017.	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Foscan - temoporfin - EMEA/H/C/000318/II/0042 MAH: biolitec Pharma Ltd, Rapporteur: Paula Boudewina van Hennik Request for Supplementary Information adopted on 14.09.2017.	Request for Supplementary Information adopted with a specific timetable.
Ganfort - bimatoprost / timolol - EMEA/H/C/000668/II/0027/G MAH: Allergan Pharmaceuticals Ireland, Rapporteur: Hanne Lomholt Larsen Request for Supplementary Information adopted on 14.09.2017, 09.06.2017.	Request for Supplementary Information adopted with a specific timetable.
Hizentra - human normal immunoglobulin - EMEA/H/C/002127/II/0074/G MAH: CSL Behring GmbH, Rapporteur: Jan Mueller-Berghaus Opinion adopted on 14.09.2017. Request for Supplementary Information adopted on 22.06.2017, 23.02.2017.	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Hizentra - human normal immunoglobulin -	Request for Supplementary Information adopted

EMEA/H/C/002127/II/0086 MAH: CSL Behring GmbH, Rapporteur: Jan Mueller-Berghaus Request for Supplementary Information adopted on 14.09.2017.	with a specific timetable.
Imvanex - modified vaccinia ankara virus - EMEA/H/C/002596/II/0027 MAH: Bavarian Nordic A/S, Rapporteur: Greg Markey Opinion adopted on 14.09.2017. Request for Supplementary Information adopted on 22.06.2017.	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Kadcyla - trastuzumab emtansine - EMEA/H/C/002389/II/0034 MAH: Roche Registration Limited, Rapporteur: Sinan B. Sarac Opinion adopted on 14.09.2017. Request for Supplementary Information adopted on 29.06.2017.	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Keytruda - pembrolizumab - EMEA/H/C/003820/II/0030 MAH: Merck Sharp & Dohme Limited, Rapporteur: Daniela Melchiorri Opinion adopted on 14.09.2017. Request for Supplementary Information adopted on 06.07.2017.	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Keytruda - pembrolizumab - EMEA/H/C/003820/II/0031/G MAH: Merck Sharp & Dohme Limited, Rapporteur: Daniela Melchiorri Opinion adopted on 14.09.2017. Request for Supplementary Information adopted on 13.07.2017.	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Myozyme - alglucosidase alfa - EMEA/H/C/000636/II/0063/G MAH: Genzyme Europe BV, Rapporteur: Alexandre Moreau Opinion adopted on 14.09.2017. Request for Supplementary Information adopted on 20.07.2017.	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Opdivo - nivolumab - EMEA/H/C/003985/II/0037/G MAH: Bristol-Myers Squibb Pharma EEIG, Rapporteur: Jorge Camarero Jiménez Request for Supplementary Information adopted on 14.09.2017.	Request for Supplementary Information adopted with a specific timetable.
Pergoveris - follitropin alfa / lutropin alfa -	Positive Opinion adopted by consensus on

EMEA/H/C/000714/II/0052/G MAH: Merck Serono Europe Limited, Rapporteur: Nithyanandan Nagercoil Opinion adopted on 14.09.2017.	14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Perjeta - pertuzumab - EMEA/H/C/002547/II/0030 MAH: Roche Registration Limited, Rapporteur: Sinan B. Sarac Opinion adopted on 14.09.2017. Request for Supplementary Information adopted on 29.06.2017.	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Praluent - alirocumab - EMEA/H/C/003882/II/0021/G MAH: sanofi-aventis groupe, Rapporteur: Johann Lodewijk Hillege Opinion adopted on 14.09.2017. Request for Supplementary Information adopted on 18.05.2017.	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Soliris - eculizumab - EMEA/H/C/000791/II/0100, Orphan MAH: Alexion Europe SAS, Rapporteur: Jorge Camarero Jiménez Opinion adopted on 14.09.2017.	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Suliqua - insulin glargine / lixisenatide - EMEA/H/C/004243/II/0003/G MAH: sanofi-aventis groupe, Rapporteur: Kristina Dunder Opinion adopted on 14.09.2017.	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Uptravi - selexipag - EMEA/H/C/003774/II/0010 MAH: Actelion Registration Limited, Rapporteur: Martina Weise Opinion adopted on 14.09.2017.	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Voncento - human coagulation factor VIII / human von willebrand factor - EMEA/H/C/002493/II/0030/G MAH: CSL Behring GmbH, Rapporteur: Paula Boudewina van Hennik Request for Supplementary Information adopted on 14.09.2017.	Request for Supplementary Information adopted with a specific timetable.
WS1084/G Ganfort-EMEA/H/C/000668/WS1084/0028/G Lumigan-EMEA/H/C/000391/WS1084/0053/G MAH: Allergan Pharmaceuticals Ireland, Lead Rapporteur: Hanne Lomholt Larsen	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

Opinion adopted on 14.09.2017.	
WS1177/G **Neulasta-EMEA/H/C/000420/WS1177/0097/G** **Ristempa - (SRD) - EMEA/H/C/003910/WS1177/0012/G** MAH: Amgen Europe B.V., Lead Rapporteur: Robert James Hemmings Request for Supplementary Information adopted on 14.09.2017.	Request for Supplementary Information adopted with a specific timetable.
WS1226 **Humalog-EMEA/H/C/000088/WS1226/0158** **Liprolog-EMEA/H/C/000393/WS1226/0121** MAH: Eli Lilly Nederland B.V., Lead Rapporteur: Robert James Hemmings Opinion adopted on 14.09.2017.	Positive Opinion adopted by consensus on 14.09.2017. 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

Aranesp - darbepoetin alfa - EMEA/H/C/000332/II/0143 MAH: Amgen Europe B.V., Rapporteur: Martina Weise, "Update of section of section 4.8 the SmPC in order to add a warning on injection site bruise and haemorrhage with frequency unknown and to provide additional instructions on the use of the device in the PL following signal procedure EMEA/H/C000332/SDA/090 on cases of incorrect device use / device malfunction (EU/1/01/185/045 - EU/1/01/185/068). Provide results of further research on the effectiveness of training tools and a discussion on adequate training methods and the need for additional risk minimisation methods including, as appropriate, a proposal for an educational package including key elements for inclusion in Annex IID/RMP. Provide a detailed review of the type of device malfunctions and their outcomes, including a discussion on the need of further risk minimisation regarding this issue. Comments were received from two member states supporting the Rapporteur's conclusions." Opinion adopted on 14.09.2017.	See 9.1 in main agenda. Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Cinryze - C1-esterase inhibitor, human - EMEA/H/C/001207/II/0048 MAH: Shire Services BVBA, Rapporteur: Jan	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP

Mueller-Berghaus, “To replace the term Unit (U) by International Unit (IU) for the product potency in the product information in line with World Health Organization (WHO) international C1-inhibitor concentrate standard (08/256) as well as in compliance with ICH guidelines (Q6B, 2.2.1) analytical considerations, reference standards and reference materials and also to be consistent with Module 3 of the product dossier.” Opinion adopted on 14.09.2017. Request for Supplementary Information adopted on 18.05.2017, 26.01.2017.	recommendation.
Deltyba - delamanid - EMEA/H/C/002552/II/0021, Orphan MAH: Otsuka Novel Products GmbH, Rapporteur: Greg Markey, “Update of sections 4.2, 4.4, 4.8, 4.9, 5.1 and 5.2 of the SmPC to reflect the results of the final study report of 242-09-213 (A Phase 3, Multicenter, Randomized, Double-blind, Placebo-controlled, Parallel Group Trial to Evaluate the Safety and Efficacy of Delamanid (OPC-67683) Administered Orally as 200 mg Total Daily Dose for Six Months in Patients With Pulmonary Sputum Culture-positive, Multidrug-resistant Tuberculosis), submitted to fulfill SOB-01. The Package leaflet is updated accordingly.” Request for Supplementary Information adopted on 14.09.2017.	Request for Supplementary Information adopted with a specific timetable.
Elonva - corifollitropin alfa - EMEA/H/C/001106/II/0034 MAH: Merck Sharp & Dohme Limited, Rapporteur: Paula Boudewina van Hennik, “Update of section 4.5 of the SmPC to add information pertaining to potential hCG cross-reactivity resulting in a false positive pregnancy test. The Package Leaflet has been updated accordingly. In addition, the MAH is taking the opportunity to implement changes in the annexes in line with the QRD templates (versions 9.1 and 10) and to propose combined versions of the SmPCs and Package Leaflets for the different strengths.” Opinion adopted on 14.09.2017. Request for Supplementary Information adopted on 20.07.2017, 18.05.2017.	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Emend - aprepitant - EMEA/H/C/000527/II/0055	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP

MAH: Merck Sharp & Dohme Limited, Rapporteur: Filip Josephson, “Update of sections 4.2 of the SmPC in order to replace the nomogram for the paediatric formulation provided in ml/kg with purely weight-based dosing instructions (in mg/kg) This is based on data that were already submitted as part of the paediatric application X/49. The Package Leaflet is updated accordingly. In addition, the Marketing authorisation holder (MAH) took the opportunity to bring the PI in line with the latest QRD template version 10.0.” Opinion adopted on 14.09.2017. Request for Supplementary Information adopted on 20.07.2017.	Members were in agreement with the CHMP recommendation.
Epclusa - sofosbuvir / velpatasvir - EMEA/H/C/004210/II/0012 MAH: Gilead Sciences International Limited, Rapporteur: Filip Josephson, “Update of section 5.3 of the SmPC in order to add non-clinical safety findings based on a 6-month carcinogenicity study conducted with velpatasvir in transgenic mice” Opinion adopted on 14.09.2017.	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Eperzan - albiglutide - EMEA/H/C/002735/II/0031 MAH: GlaxoSmithKline Trading Services Limited, Rapporteur: Kristina Dunder, “Update of section 4.8 of the SmPC in order to include angioedema under the description of “Allergic reactions”. The Package Leaflet is updated accordingly. In addition, the Marketing authorisation holder took the opportunity to implement minor editorial amendments throughout the Product Information.” Opinion adopted on 14.09.2017. Request for Supplementary Information adopted on 22.06.2017, 21.04.2017.	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Esbriet - pirfenidone - EMEA/H/C/002154/II/0043, Orphan MAH: Roche Registration Limited, Rapporteur: Greg Markey, “Update of sections 4.2 and 5.2 of the SmPC in order to update the existing safety information with revised recommendations for patients with moderate renal impairment based on the totality of data from clinical studies; the Package Leaflet is updated accordingly.” Request for Supplementary Information adopted on 14.09.2017, 29.06.2017.	Request for Supplementary Information adopted with a specific timetable.

Eviplera - emtricitabine / rilpivirine / tenofovir disoproxil - EMEA/H/C/002312/II/0082 MAH: Gilead Sciences International Limited, Rapporteur: Johann Lodewijk Hillege, "Update of section 4.5 of the SmPC with Drug-Drug Interaction information for Eviplera based on the results from Study TMC435-TiDP16-C114; this is a Phase I, 2-panel, open-label, randomized, cross-over trial in healthy subjects to investigate the pharmacokinetic interaction between TMC435 and antiretroviral agents, TMC278 and tenofovir disoproxil fumarate (TDF), at steady-state. The Package Leaflet is updated accordingly. In addition, the Marketing authorisation holder (MAH) took the opportunity to introduce minor administrative changes in the SmPC and to update the list of local representatives in the Package Leaflet for Estonia, Latvia and Lithuania. Minor linguistic amendments (MLAs) have been implemented to the translations of the product information annexes: CS, DE, ES, FR, IS, IT, NL, NO, PT, SE and SK." Opinion adopted on 14.09.2017. Request for Supplementary Information adopted on 15.06.2017.	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Forsteo - teriparatide - EMEA/H/C/000425/II/0046 MAH: Eli Lilly Nederland B.V., Rapporteur: Greg Markey, "Update of section 5.1 of the SmPC of the SmPC based on the results of study B3D-EW-GHDW (VERO), a phase 4 multi-centre, prospective, randomized, parallel, double-blind, double-dummy, active controlled study comparing the effect of teriparatide for injection versus risedronate on the incidence of fractures and low bone mass. In addition, the Marketing authorisation holder (MAH) took the opportunity to correct the formatting throughout the Product Information and to bring Annex II in line with the latest QRD template version 10." Request for Supplementary Information adopted on 14.09.2017, 20.07.2017.	Request for Supplementary Information adopted with a specific timetable.
Glivec - imatinib - EMEA/H/C/000406/II/0108 MAH: Novartis Europharm Ltd, Rapporteur: Jorge Camarero Jiménez, "Update of SmPC section 4.4	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

based on the final CSR for study STI571A2405; the International Study for Chronic Myeloid Leukaemia (CML) in childhood and adolescents (I-CML-Ped Study). The provision of the study report addresses the post-authorisation measure MEA 162.8." Opinion adopted on 14.09.2017. Request for Supplementary Information adopted on 22.06.2017, 21.04.2017.	
Imnovid - pomalidomide - EMEA/H/C/002682/II/0025, Orphan MAH: Celgene Europe Limited, Rapporteur: Robert James Hemmings, "Submission of a biomarker analysis report based on the clinical study CC-4047-MM-010 following a recommendation from the CHMP at the time of the initial authorisation." Opinion adopted on 14.09.2017. Request for Supplementary Information adopted on 06.07.2017.	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Pradaxa - dabigatran etexilate - EMEA/H/C/000829/II/0103 MAH: Boehringer Ingelheim International GmbH, Rapporteur: Hanne Lomholt Larsen, "Update of sections 4.2, 4.4 and 5.1 to reflect the final study results of the phase IV study 1160.204 (The RE-CIRCUIT Trial), " A Randomised Evaluation of dabigatran etexilate Compared to warfar/n inpulmonaRy vein ablation: assessment of an uninterrupted periproCedUralant/coagulation sTrategy"" Opinion adopted on 14.09.2017. Request for Supplementary Information adopted on 20.07.2017.	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
ReFacto AF - moroctocog alfa - EMEA/H/C/000232/II/0140 MAH: Pfizer Limited, Rapporteur: Hanne Lomholt Larsen, "Submission of the report 'The Immunogenicity of ReFacto AF in UK PUPs Who Started Treatment from 2010' prepared by the United Kingdom Haemophilia Centre Doctors' Organisation (UKHCDO). This report is being submitted in the context of a post-approval commitment, MEA 115.1 ('The MAH commits to submit the CSR for "A Postauthorization Safety Surveillance Registry or ReFacto AF in Previously Untreated Patients (PUPs) in Usual Care Settings – study number 4435" and to initiate the registry'), as supporting	Request for Supplementary Information adopted with a specific timetable.

evidence of the ongoing safety evaluation of ReFacto AF in PUPs with haemophilia A and with a specific focus on the development of inhibitors." Request for Supplementary Information adopted on 14.09.2017.	
Revatio - sildenafil - EMEA/H/C/000638/II/0077 MAH: Pfizer Limited, Rapporteur: Johann Lodewijk Hillege, "Update of section 4.6 of the SmPC in order to revise the statement concerning the detection of sildenafil and its active metabolite in human mik and the potential for impact on the breastfed infact. The Package Leaflet and Labelling are updated accordingly. In addition, the Marketing authorisation holder (MAH) took the opportunity to bring the PI in line with the latest QRD template version 10.0." Request for Supplementary Information adopted on 14.09.2017.	Request for Supplementary Information adopted with a specific timetable.
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." Opinion adopted on 14.09.2017. Request for Supplementary Information adopted on 18.05.2017, 23.02.2017.	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Stelara - ustekinumab - EMEA/H/C/000958/II/0058 MAH: Janssen-Cilag International NV, Rapporteur: Greg Markey, "Update of section 4.8 of the SmPC in order to include Lower Respiratory Tract Infection as an Adverse Drug Reaction based on a comprehensive evaluation of safety information from the STELARA clinical studies database and post-marketing database, as well as available literature. The Package Leaflet is updated accordingly." Opinion adopted on 14.09.2017.	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Strensiq - asfotase alfa - EMEA/H/C/003794/II/0019/G, Orphan MAH: Alexion Europe SAS, Rapporteur: Greg Markey, "Update of section 5.1 of the SmPC in	Request for Supplementary Information adopted with a specific timetable.

order to update information following final results from studies ENB-006-09 [A Randomized, Open-Label, Multicenter, Multinational, Dose-Ranging, Historical Control Study of the Safety,Efficacy, Pharmacokinetics, and Pharmacodynamics of Asfotase Alfa (Human Recombinant Tissue-Nonspecific Alkaline Phosphatase Fusion Protein) in Children with Hypophosphatasia (HPP)] (and its extension ENB-008-10 [Extension Study of Protocol ENB-006-09 Evaluating the Long-Term Safety and Efficacy of Asfotase Alfa (Human Recombinant Tissue-Nonspecific Alkaline Phosphatase Fusion Protein) in Children with Hypophosphatasia (HPP)]) and ENB-009-10 [A Randomized, Open-Label, Multicenter, Multinational, Dose-Ranging, Concurrent Control Study of the Safety, Efficacy, and Pharmacokinetics of ENB-0040 (Human Recombinant Tissue-Nonspecific Alkaline Phosphatase Fusion Protein) in Adolescents and Adults with Hypophosphatasia (HPP)] listed as an obligation in the Annex II (ANX002). In addition, the Marketing authorisation holder (MAH) took the opportunity to propose editorial changes for section 4.5 to better clarify the information provided." Request for Supplementary Information adopted on 14.09.2017.	
Stribild - elvitegravir / cobicistat / emtricitabine / tenofovir disoproxil - EMEA/H/C/002574/II/0083 MAH: Gilead Sciences International Limited, Rapporteur: Robert James Hemmings, "Update of sections 4.5 of the SmPC in order to add drug-drug interaction data from Study GS-US-292-1316; this is a Phase 1, Open-Label, Fixed Sequence Study Evaluating the Pharmacokinetics and Drug Interaction Potential Between Elvitegravir/Cobicistat/Emtricitabine/Tenofovir Alafenamide Single-Table Regimen and Sertraline in Healthy Subjects. In addition, the Marketing authorisation holder (MAH) took the opportunity make administrative amendments to section 4.8 of the SmPC." Opinion adopted on 14.09.2017.	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Tecfidera - dimethyl fumarate - EMEA/H/C/002601/II/0041	Request for Supplementary Information adopted with a specific timetable.

MAH: Biogen Idec Ltd, Rapporteur: Martina Weise, “Update of sections 4.4 and 4.8 of the SmPC in order to add anaphylactic reaction as a warning and as an adverse reaction with unknown frequency, based on post-marketing experience. The Package Leaflet is updated accordingly. In addition, the Biogen Idec Ltd took the opportunity to bring the PI in line with the latest QRD template version 10.” Request for Supplementary Information adopted on 14.09.2017.	
Tecfidera - dimethyl fumarate - EMEA/H/C/002601/II/0042 MAH: Biogen Idec Ltd, Rapporteur: Martina Weise, “Update of sections 4.2, 4.8, 5.1 and 5.2 of the SmPC in order to update the safety and efficacy information in the paediatric population based on the clinical study results from study 109MS202, listed as a category 3 study in the RMP; this is an open-label, multicentre, multidose study designed to assess the effect of Tecfidera on magnetic resonance imaging lesions and pharmacokinetics, safety and tolerability in paediatric population with relapsing-remitting multiple sclerosis. There are no updates proposed in the package leaflet or RMP.” Request for Supplementary Information adopted on 14.09.2017.	Request for Supplementary Information adopted with a specific timetable.
Tecfidera - dimethyl fumarate - EMEA/H/C/002601/II/0043/G MAH: Biogen Idec Ltd, Rapporteur: Martina Weise, “Submission of non-clinical study report for study PD-15-73: haemotoxicity study of dimethyl fumarate (DMF) and monomethyl fumarate (MMF) on T lymphocyte colony forming progenitor stem cells (T CFC) and T-cells derived from mononuclear cells (MNCs) of bone marrow and peripheral blood of humans and Cynomolgus monkeys. This submission is linked to a category 3 study in the RMP. Submission of non-clinical study report for study P00012-15-05: 3-Month repeated-dose oral (nasogastric) toxicity and toxicokinetic study of dimethyl fumarate (DMF) and hydroxyurea (HU) in cynomolgus monkeys. This submission is linked to a category 3 study in the RMP.”	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

Opinion adopted on 14.09.2017.	
Tecfidera - dimethyl fumarate - EMEA/H/C/002601/II/0044 MAH: Biogen Idec Ltd, Rapporteur: Martina Weise, "Submission of the final report from study 109MS407: an interventional PASS listed as a category 4 study in the RMP: a multicentre, open-label, single-arm study to evaluate gastrointestinal tolerability in subjects with relapsing-remitting multiple sclerosis receiving dimethyl fumarate (TOLERATE)." Opinion adopted on 14.09.2017.	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Toviaz - fesoterodine - EMEA/H/C/000723/II/0049 MAH: Pfizer Limited, Rapporteur: Concepcion Prieto Yerro, "Update of the SmPC sections 4.6 and 5.3 with revised information from reproductive toxicity studies in mice. In addition, the MAH took the opportunity to bring the PI in line with the latest QRD template version 10.0." Opinion adopted on 14.09.2017.	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Translarna - ataluren - EMEA/H/C/002720/II/0036, Orphan MAH: PTC Therapeutics International Limited, Rapporteur: Johann Lodewijk Hillege, "Update of section 4.5 of the SmPC in order to include information regarding the effects of ataluren on the pharmacokinetics of sensitive probe substrate of organic anion transporter 3 (OAT3)) following results from study PTC124-GD-037-HV (MEA015). In addition, the Marketing authorisation holder (MAH) took the opportunity to introduce some editorial changes in the PI." Opinion adopted on 14.09.2017.	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Trevicta - paliperidone - EMEA/H/C/004066/II/0011 MAH: Janssen-Cilag International NV, Informed Consent of Xeplion, Rapporteur: Kristina Dunder, "Update of section 4.8 of the SmPC in order to update the safety information after assessment of study R092670-SCA-3004 (A Randomized, Double-Blind, Placebo-Controlled, Parallel-Group Study of Paliperidone Palmitate Evaluating Time to Relapse in Subjects With Schizoaffective Disorder). The Package Leaflet has been updated accordingly." Opinion adopted on 14.09.2017.	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Truvada - emtricitabine / tenofovir	Positive Opinion adopted by consensus on

disoproxil - EMEA/H/C/000594/II/0138/G MAH: Gilead Sciences International Limited, Rapporteur: Greg Markey, "Submission of the final report from studies GS-US-276-0101 and GS-US-276-0105, listed as a category 3 studies in the RMP. GS-US-276-0101 - This is a A Prospective, Observational Study of Pregnancy Outcomes among Women Exposed to Truvada for PrEP Indication Nested in the Antiretroviral Pregnancy Registry GS-US-276-0105 – This is a A Prospective, Observational, Drug Utilization Study of Subjects Taking Truvada for Pre-exposure Prophylaxis in the USA." Opinion adopted on 14.09.2017. Request for Supplementary Information adopted on 09.06.2017.	14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Vargatef - nintedanib - EMEA/H/C/002569/II/0017 MAH: Boehringer Ingelheim International GmbH, Rapporteur: Sinan B. Sarac, "Update of section 4.8 of the SmPC in order to add 'weight decreased' as a new adverse drug reaction based on a safety review of clinical trials and post-marketing data. The Package Leaflet is updated accordingly. In addition, the Marketing authorisation holder (MAH) took the opportunity to implement a minor correction in the English product information, minor corrections to the Croatian, Danish, Dutch and Finnish translations and to bring section 4 of the Package Leaflet in line with QRD template version 10." Opinion adopted on 14.09.2017.	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Vargatef - nintedanib - EMEA/H/C/002569/II/0018 MAH: Boehringer Ingelheim International GmbH, Rapporteur: Sinan B. Sarac, "Update of section 4.4 of the SmPC to amend the current warning on hepatic function to include that drug liver induced injury was associated with nintendanib administration, to include low body weight, Asian origin, female sex and age as factors of increased risk of liver enzymes elevations, update of section 4.8 of the SmPC to add 'drug-induced liver injury' (DILI) as new ADR with an 'uncommon' frequency and update of section 5.2 of the SmPC to amend the current information related to the mean exposure to nintedanib by race, based on a	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

review of clinical trials and post-marketing data on DILI and on the exposure safety relationship between nintedanib plasma exposure and liver enzyme elevations, as requested by the PRAC as part of PSUSA/00010318/201611. The Package Leaflet is updated accordingly. In addition, the MAH took the opportunity to make some minor changes to section 4.4 and 4.8 of the SmPC." Opinion adopted on 14.09.2017.	
Venclyxto - venetoclax - EMEA/H/C/004106/II/0003, Orphan MAH: AbbVie Limited, Rapporteur: Filip Josephson, "Submission of the final report from study R&D 16/1398: Assessment of Cytochrome P450 mRNA Induction by A-1195425 in Cultured Human Hepatocytes to evaluate CYP1A2 and CYP2B6 induction response which is included as a Post Authorisation Measure (Category 3) in RMP 2.0." Opinion adopted on 14.09.2017. Request for Supplementary Information adopted on 15.06.2017.	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Xeljanz - tofacitinib - EMEA/H/C/004214/II/0003 MAH: Pfizer Limited, Rapporteur: Robert James Hemmings, "Submission of 2 transported inhibition studies evaluating tofacitinib for its potential to inhibit organic anion transporter (OAT) 1, OAT3 and to interact with Human MRP2 Efflux (ABC) Transporter in fulfilment of the Recommendation dated 26 January 2017." Request for Supplementary Information adopted on 14.09.2017.	Request for Supplementary Information adopted with a specific timetable.
Xeplion - paliperidone - EMEA/H/C/002105/II/0035 MAH: Janssen-Cilag International NV, Rapporteur: Kristina Dunder, "Update of section 4.2 of the SmPC in order to add a dosage conversion table to provide guidance for healthcare professionals when switching patients from paliperidone ER tablets to paliperidone palmitate long acting injection (PP1M). The Package Leaflet is updated accordingly." Request for Supplementary Information adopted on 14.09.2017, 20.07.2017.	Request for Supplementary Information adopted with a specific timetable.
Xgeva - denosumab - EMEA/H/C/002173/II/0054 MAH: Amgen Europe B.V., Rapporteur: Kristina	Positive Opinion adopted by consensus on 01.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP

Dunder, PRAC Rapporteur: Ulla Wändel Liminga, "Submission of an updated RMP version 25 in order to remove cataracts from the list of potential risks associated with denosumab therapy based on the results of study 20080560 (Multicentre, randomized, double blind, placebo-controlled study in men with nonmetastatic prostate cancer receiving androgen deprivation therapy cataract development and progression study using a slit-lamp-based evaluation system (Lens Opacities Classification System III (LOCS III))." Opinion adopted on 01.09.2017. Request for Supplementary Information adopted on 09.06.2017.	recommendation.
Xultophy - insulin degludec / liraglutide - EMEA/H/C/002647/II/0021 MAH: Novo Nordisk A/S, Rapporteur: Kristina Dunder, "Update of section 5.1 of the SmPC in order to reflect data for transfer from insulin glargine U100 to Xultophy as compared to a basal-bolus regimen. The update is based on data from the clinical trial NN9068-4185: "A clinical trial comparing efficacy and safety of insulin degludec/liraglutide (IDegLira) versus basal-bolus therapy in subjects with type 2 diabetes mellitus". The MAH has taken the opportunity to make minor editorial and formatting changes throughout the Annexes." Opinion adopted on 14.09.2017.	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Xyrem - sodium oxybate - EMEA/H/C/000593/II/0067/G MAH: UCB Pharma Limited, Rapporteur: Bruno Sepodes, "Update of section 4.8 of the SmPC in order to add the adverse reactions "increased libido" and "seborrhea" with an unknown frequency. Update of section 4.6 of the SmPC in order to amend the information about breast-feeding. The Package Leaflet is updated accordingly." Opinion adopted on 14.09.2017.	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
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,	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

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." Opinion adopted on 14.09.2017. Request for Supplementary Information adopted on 22.06.2017, 21.04.2017.	
Zyclara - imiquimod - EMEA/H/C/002387/II/0013 MAH: Meda AB, Rapporteur: Nithyanandan Nagercoil, "Update of sections 4.2, 4.4 and 5.1 of the SmPC in order to add data on the clinical experience gained with study X-03016-3284 (LEIDA 2) and a meta-analysis of X-03016-3271 and X-03016-3284. The MAH took the opportunity to update the details of local representatives in the PIL." Request for Supplementary Information adopted on 14.09.2017, 22.06.2017.	Request for Supplementary Information adopted with a specific timetable.
Zykadia - ceritinib - EMEA/H/C/003819/II/0016 MAH: Novartis Europharm Ltd, Rapporteur: Jorge Camarero Jiménez, "Update of sections 4.2, 4.4, 4.8 and 5.2 of the SmPC in order to include amendments to the posology in hepatically impaired patients and update the safety information, respectively. The updates are based on the results from the hepatic function Study CLDK378A2110 which evaluated the PK, safety and tolerability of a single oral dose of ceritinib in subjects with varying degrees of impaired hepatic function and results from physiology-based pharmacokinetic (PBPK) modeling at steady-state. Submission of the Report for Study A2110 fulfils MEA 001 for Zykadia." Request for Supplementary Information adopted on 14.09.2017.	Request for Supplementary Information adopted with a specific timetable.
WS1137 Lyrica-EMEA/H/C/000546/WS1137/0087 Pregabalin Pfizer-EMEA/H/C/003880/WS1137/0017 MAH: Pfizer Limited, Lead Rapporteur: Johann Lodewijk Hillege, "Update of sections 4.8 and 5.1 of the SmPC in order to reflect final results from	Request for Supplementary Information adopted with a specific timetable.

paediatric study A0081041: “A Double-Blind, Placebo-Controlled, Parallel-Group, Multicenter Study of the Efficacy and Safety of Pregabalin as Adjunctive Therapy in Children 4-16 Years of Age with Partial Onset Seizures”.” Request for Supplementary Information adopted on 14.09.2017, 08.05.2017.	
WS1203/G **Docetaxel** **Winthrop-EMEA/H/C/000808/WS1203/0053/G** **Taxotere-EMEA/H/C/000073/WS1203/0128/G** MAH: Aventis Pharma S.A., Lead Rapporteur: Alexandre Moreau, “Update of sections 4.4 and 4.8 of the SmPC to add information about ventricular arrhythmia including ventricular tachycardia based on review of the MAH’s global pharmacovigilance database and scientific literature. The Package Leaflet is updated accordingly. Update of section 4.8 of the SmPC on the 10-year follow-up data for studies TAX316 and GEICAM 9805 studies in order to clarify the persisting events in the follow-up periods.” Opinion adopted on 14.09.2017.	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1205 **Descovy-EMEA/H/C/004094/WS1205/0020** **Genvoya-EMEA/H/C/004042/WS1205/0034** **Odefsey-EMEA/H/C/004156/WS1205/0018** MAH: Gilead Sciences International Limited, Lead Rapporteur: Robert James Hemmings, “Update of section 4.5 of the SmPC in order to provide the final study report for the in vitro study AD-120-2045; this is a non-clinical study on the Effect of Xanthine Oxidase Inhibitors on Metabolism of Tenofovir alafenamide fumarate in Primary Human Hepatocytes. This study is listed in their respective Risk Management Plans (RMPs) as an additional pharmacovigilance activity (Category 3) (Genvoya: MEA 006; Descovy: MEA 004; Odefsey: MEA 007). The requested worksharing procedure proposed amendments to the Summary of Product	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

Characteristics." Opinion adopted on 14.09.2017.	
WS1218 **Brimica Genuair-EMEA/H/C/003969/WS1218/0015** **Duaklir Genuair-EMEA/H/C/003745/WS1218/0015** MAH: AstraZeneca AB, Lead Rapporteur: Nithyanandan Nagercoil, "Update of section 5.1 of the SmPC in order to update information following results from study M-40464-33 (A Multiple Dose, Randomised, Double-Blind, Placebo Controlled, Parallel Clinical Trial to Assess the Effect of Aclidinium Bromide/Formoterol Fumarate Fixed-Dose Combination on Lung Hyperinflation, Exercise Capacity and Physical Activity in Patients with Moderate to Severe Chronic Obstructive Pulmonary Disease (COPD))" Opinion adopted on 14.09.2017.	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1219 **Brimica Genuair-EMEA/H/C/003969/WS1219/0014** **Duaklir Genuair-EMEA/H/C/003745/WS1219/0014** MAH: AstraZeneca AB, Lead Rapporteur: Nithyanandan Nagercoil, "Update of section 5.2 of the SmPC in order to update information based on results from study KRP-AB1102F-302 [KRP-AB1102F Phase II Clinical Pharmacology Study - An Investigation into the Pharmacokinetics upon Repeated Administration of KRP-AB1102F to COPD Patients as Subjects]. In addition, the Worksharing applicant (WSA) took the opportunity to update footnotes of the table in section 4.8 as requested during PSUR procedure EMEA/H/C/PSUSA/00010307/201511 and to amend annex II following request from procedure EMEA/H/C/PSA/S/0017." Request for Supplementary Information adopted on 14.09.2017.	Request for Supplementary Information adopted with a specific timetable.
WS1225/G **Exviera-EMEA/H/C/003837/WS1225/0031/G** **Viekirax-EMEA/H/C/003839/WS1225/003**	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

5/G MAH: AbbVie Limited, Lead Rapporteur: Filip Josephson, “Submission of the final reports for two phase IIIb studies (studies M14-226 and M15-461) listed as category 3 studies in the RMP. These are open-label studies evaluating the safety and efficacy of ombitasvir/paritaprevir/ritonavir and dasabuvir with or without ribavirin in hepatitis C virus infected patients with several renal impairment or end-stage renal disease with or without compensated cirrhosis.” Opinion adopted on 14.09.2017.	
WS1234/G **Genvoya-EMEA/H/C/004042/WS1234/0036/G** **Stribild-EMEA/H/C/002574/WS1234/0084/G** **Tybost-EMEA/H/C/002572/WS1234/0038/G** MAH: Gilead Sciences International Limited, Lead Rapporteur: Robert James Hemmings, “Update of sections 4.4 and 4.5 of the SmPC based on data from the following Pharamacology Studies (GS-US-216-1008 and GS-US-216-4032). Study GS-US-216-1008 is a Phase 1, randomized, fixed-sequence, open-label, single and multiple-dose, multiple-cohort, single-center study that evaluated the drug interaction potential between darunavir (DRV)+COBI, atazanavir (ATV)+COBI, or Genvoya and the 3 hydroxy-3-methylglutaryl-coenzyme A (HMG CoA) reductase inhibitors rosuvastatin and/or atorvastatin. Study GS-US-216-4032 is an open-label, single -center, multiple-cohort, fixed_sequence, Phase 1 study that evaluated the effect of DRV+COBI or ATV+COBI on the pharmacokinetic (PK) of a representative hormonal contraceptive medication, drospirenone/ethinyl estradiol. The Package Leaflet is updated accordingly. In addition, the Worksharing applicant (WSA) took the opportunity to make administrative changes to the PI of all three products and update the list of local representatives for Estonia, Latvia and Lithuania for Tybost and Stribild. Minor linguistic amendments were made to the Product Information.”	Request for Supplementary Information adopted with a specific timetable.

Request for Supplementary Information adopted on 14.09.2017.	

B.5.3. CHMP-PRAC assessed procedures

Abasaglar - insulin glargine - EMEA/H/C/002835/II/0014 MAH: Eli Lilly Regional Operations GmbH, Rapporteur: Robert James Hemmings, PRAC Rapporteur: Carmela Macchiarulo, "Submission of the final report from study I4L-MC-ABER(ABER). This is a Prospective, Randomized, Open-Label Comparison of a Long-Acting Basal Insulin Analog LY2963016 to LANTUS® in Adult Patients with Type 2 Diabetes Mellitus: the ELEMENT 5 Study. This study was conducted in non European countries. This study replaces the cancelled studies that were planned to be conducted in China and other countries and that were described in the RMP. An updated RMP version 1.6 is submitted accordingly." Opinion adopted on 01.09.2017. Request for Supplementary Information adopted on 09.06.2017.	Positive Opinion adopted by consensus on 01.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Adcetris - brentuximab vedotin - EMEA/H/C/002455/II/0049, Orphan MAH: Takeda Pharma A/S, Rapporteur: Paula Boudewina van Hennik, PRAC Rapporteur: Sabine Straus, "Update of sections 4.2, 4.8, 5.1 and 5.2 of the SmPC with data from study C25002; a phase 1/2 study of brentuximab vedotIn (SGN-35) in paediatric patients with relapsed or refractory systemic anaplastic large cell lymphoma or hodgkin lymphoma (listed in the agreed PIP covering the conditions of Hodgkin lymphoma and anaplastic large cell lymphoma for ADCETRIS (EMEA-000980-PIP01-10-M04)). An updated RMP version 11.0 was provided as part of the application." Request for Supplementary Information adopted on 01.09.2017.	Request for Supplementary Information adopted with a specific timetable.
Adenuric - febuxostat - EMEA/H/C/000777/II/0047 MAH: Menarini International Operations Luxembourg S.A., Rapporteur: Andrea Laslop, PRAC Rapporteur: Jan Neuhauser, "Update of sections 4.4 and 4.5 of the SmPC in order to reflect the results of preclinical study	Request for Supplementary Information adopted with a specific timetable.

MRPO-2015-PKM-005 “Pharmacokinetic of azathioprine in the rat after one-week daily oral treatment at three different dosages and with the concomitant oral administration of febuxostat or allopurinol" and clinical study REP-POPPK-MRP-2015-PKM-005 “Population Pharmacokinetic analysis from study titled Pharmacokinetic of azathioprine in the rat after one-week daily oral treatment at three different dosages and with the concomitant oral administration of febuxostat or allopurinol”, investigating the drug-drug interaction with azathioprine when co-administered with febuxostat. The RMP version 6.0 has also been submitted. In addition, the MAH took the opportunity to correct the typing errors and to bring the PI in line with the latest QRD template version 10.” Request for Supplementary Information adopted on 14.09.2017.	
Cabometyx - cabozantinib - EMEA/H/C/004163/II/0002/G MAH: Ipsen Pharma, Rapporteur: Robert James Hemmings, PRAC Rapporteur: Sabine Straus, “1) C.I.4 (type II) Update of section 5.1 of the SmPC to reflect the final study results from clinical study XL184-308: A Phase 3, Randomized, Controlled Study of Cabozantinib (XL184) vs Everolimus in Subjects with Metastatic Renal Cell Carcinoma that has Progressed after Prior VEGFR Tyrosine Kinase Inhibitor Therapy, to fulfil the condition to the marketing authorisation listed as a PAES in the Annex II. The RMP version 2.0 has also been submitted. 2) C.I.4 (type II) Update of section 5.3 of the SmPC to reflect the final study results from non-clinical study XL184-NC-036: 104-Week Oral Gavage Carcinogenicity and Toxicokinetic Study with Cabozantinib (XL184) in Rats. The RMP version 2.0 has also been submitted. 3) C.I.3.z (type IB) Update of section 4.5 of the SmPC to implement the wording agreed by the PRAC following the outcome of the PSUR procedure EMEA/H/C/PSUSA/10180/201603.” Request for Supplementary Information adopted	Request for Supplementary Information adopted with a specific timetable.

on 14.09.2017.	
Cerdelga - eliglustat - EMEA/H/C/003724/II/0013, Orphan MAH: Genzyme Europe BV, Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Dolores Montero Corominas, "Update of section 4.8. of the SmPC in order to amend the safety information based on the analysis of Adverse Events from the following clinical trials: GZGD00304 (Phase 2), GZGD02507 (ENGAGE), GZGD02607 (ENCORE) and GZGD03109 (EDGE) to address post-authorisation MEA011.1 which is included in the current approved Risk Management Plan. Update of the labelling in order to reflect the instructions on use for the sleeve of the intermediate packaging of the single blister. The RMP version 4.0 has also been submitted." Opinion adopted on 01.09.2017.	Positive Opinion adopted by consensus on 01.09.2017. 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/0085 MAH: GlaxoSmithkline Biologicals SA, Rapporteur: Bart Van der Schueren, Co-Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Jean-Michel Dogné, "Submission of Study EPI-HPV-069, a meta-analysis assessing the risk of three autoimmune diseases following vaccination with Cervarix: autoimmune thyroiditis (AIT), Guillain-Barre Syndrome (GBS) and Inflammatory Bowel Disease (IBD). The EPI-HPV-069 study is a post-licensure commitment to the EMA (PASS register number EUPAS13332). As part of this submission, an updated RMP (version 18) is provided, including changes related to the EPI-HPV-069 meta-analysis submitted and minor updates related to other studies." Request for Supplementary Information adopted on 14.09.2017, 18.05.2017, 15.12.2016.	Request for Supplementary Information adopted with a specific timetable.
Cimzia - certolizumab pegol - EMEA/H/C/001037/II/0060 MAH: UCB Pharma S.A., Rapporteur: Kristina Dunder, PRAC Rapporteur: Ulla Wändel Liminga, "Update of section 4.6 of the SmPC in order to update the information on pregnancy and lactation based on two pharmacokinetics studies	Request for Supplementary Information adopted with a specific timetable.

evaluation the transfer of Cimzia into breastmilk and via the placenta (UP0016 and UP0017). The Package Leaflet is updated accordingly. The RMP version 12 has also been submitted." Request for Supplementary Information adopted on 14.09.2017, 22.06.2017.	
Eperzan - albiglutide - EMEA/H/C/002735/II/0033 MAH: GlaxoSmithKline Trading Services Limited, Rapporteur: Kristina Dunder, PRAC Rapporteur: Julie Williams, "Update of the Package Leaflet in order to amend the layout and content of the Instructions for Use (IFU). In addition, the RMP version 8 has also been submitted to implement additional pharmacovigilance and risk minimisation activities addressing the safety concern of "medication errors/device issue potentially leading to lack of efficacy or inadequate diabetes control"." Opinion adopted on 01.09.2017.	Positive Opinion adopted by consensus on 01.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Ibrance - palbociclib - EMEA/H/C/003853/II/0007 MAH: Pfizer Limited, Rapporteur: Filip Josephson, PRAC Rapporteur: Doris Stenver, "Update of sections 4.2, 4.4 and 5.2 of the SmPC to reflect the results of studies A5481013 and A5481014. The mentioned studies provide information of the impact of hepatic impairment (Study A5481013) on the PK of a single oral dose of 75 mg palbociclib and the impact of renal impairment (Study A5481014) on the PK of a single oral dose of 125 mg palbociclib both administered under fed conditions to subjects with varying degrees of hepatic function or renal function. The RMP (version 1.4) is proposed to be amended to reflect the completion of these studies." Request for Supplementary Information adopted on 14.09.2017.	Request for Supplementary Information adopted with a specific timetable.
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	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

wording and advices." Opinion adopted on 14.09.2017. Request for Supplementary Information adopted on 22.06.2017, 23.02.2017.	
Mozobil - plerixafor - EMEA/H/C/001030/II/0032, Orphan MAH: Genzyme Europe BV, Rapporteur: Paula Boudewina van Hennik, PRAC Rapporteur: Sabine Straus, "Update of sections 4.2 and 5.2 of the SmPC in order to reflect the results of the completed study MSC12830 (MOZ11809), entitled "A Phase 4, Multicenter, Randomized, Comparator Trial Evaluating the Standard Weight-Based Dose (0.24 mg/kg) Compared to a Fixed Dose (20 mg) of Plerixafor Injection in Combination with G-CSF to Mobilize and Collect ≥5 x 106 CD34+ cells/kg in ≤4 Days and to Evaluate the Difference in Total Systemic Exposure in Patients with Non-Hodgkin's Lymphoma Weighing ≤70 kg" listed as a category 3 study in the RMP. The Package Leaflet is updated accordingly. The RMP version 9.0 has also been submitted. In addition, the Marketing authorisation holder (MAH) took the opportunity to update the list of local representatives in the Package Leaflet." Opinion adopted on 01.09.2017. Request for Supplementary Information adopted on 06.07.2017.	Positive Opinion adopted by consensus on 01.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Mysimba - naltrexone hydrochloride / bupropion hydrochloride - EMEA/H/C/003687/II/0017 MAH: Orexigen Therapeutics Ireland Limited, Rapporteur: Hanne Lomholt Larsen, PRAC Rapporteur: Martin Huber, "Submission of the final report from phase I study NaltrexBuprop-1001 (TQT) to evaluate the potential effect of Naltrexone and Bupropion extended-release combination on cardiac repolarization in healthy subjects and updated RMP to include study NaltrexBuprop-1001 but also studies recently completed (NB-CVOT, NaltrexBuprop-4001, NaltrexBuprop-1004 and NB-404). The MAH also took the opportunity to include throughout the RMP references to the PASS protocols currently under discussion at the PRAC." Opinion adopted on 14.09.2017.	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

Request for Supplementary Information adopted on 22.06.2017.	
Neulasta - pegfilgrastim - EMEA/H/C/000420/II/0093/G MAH: Amgen Europe B.V., Rapporteur: Robert James Hemmings, PRAC Rapporteur: Patrick Batty, Procedure Manager: Margaux Philippe, EPL: Silvy Da Rocha Dias, "- B.IV.1.a.3 (type II) – To add a new device which may have a significant impact on the delivery of the product: the on-body injector (Onpro kit) to be used with Neulasta, 6mg solution for injection, pre-filled syringe - B.II.e.5.c (type II) – To change the fill volume from 0.6 to 0.64 mL for Neulasta, 6 mg, solution for injection pre-filled syringe co-packed with the new on-body injector (Onpro kit) In addition the MAH took the opportunity to introduce editorial changes to module 3.2.P.2.4 Container Closure System Update of sections 3, 4.2, 5.1, 6.4, 6.5, 6.6 and 8 of the SmPC. The Labelling, Package Leaflet and the RMP (version 4.2) are updated accordingly. In addition the Marketing authorisation holder (MAH) took the opportunity to update the list of local representatives in the Package Leaflet, to include some editorial changes and correct some typos throughout the product information, and to bring the product information in line with the latest QRD template version 10." Request for Supplementary Information adopted on 14.09.2017, 22.06.2017.	See 9.1 in main agenda.
Nulojix - belatacept - EMEA/H/C/002098/II/0045 MAH: Bristol-Myers Squibb Pharma EEIG, Rapporteur: Filip Josephson, PRAC Rapporteur: Ulla Wändel Liminga, "Update of sections 4.4 and 4.8 of the SmPC in order to add a warning and update the safety information on the risk of venous thrombosis of the renal allograft when anti-thymocyte globulin (ATG) and belatacept are coadministered (at the same or nearly the same time) in patients with other predisposing risk factors for thrombosis. The update is based on a review of the potential increased risk for allograft thrombosis with belatacept given in close temporal relation to Thymoglobulin, as requested during assessment of PSUR 8	Positive Opinion adopted by consensus on 01.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

(EMEA/H/C/PSUSA/00000311/201606). In addition, the MAH took the opportunity update section 6.6 “Special precautions for disposal and other handling” of the SmPC and the “Information for healthcare professionals (HCPs)”in the Package Leaflet (PL) with additional safety instructions for the co-administration of Belatacept. Submission of this variation application fulfils LEG 021 for Nulojix. Consistently with the above, RMP version 14 has also been submitted, including addition of the potential risk of venous thrombosis of the allograft when ATG and belatacept are coadministered in patients with other predisposing risk factors for thrombosis and a number of administrative changes.” Opinion adopted on 01.09.2017.	
Nuwiq - simoctocog alfa - EMEA/H/C/002813/II/0017/G MAH: Octapharma AB, Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Ulla Wändel Liminga, “C.I.4: Update of sections 4.2, 4.8 and 5.1 of the SmPC to reflect avaialble data from Previously Untreated Patients (PUP) from GENA-05 (interim report) study. The Package Leaflet is updated accordingly. The RMP version 8.0 has also been submitted. In addition, the Marketing authorisation holder (MAH) took the opportunity to update the Product Information throughout to bring it in line with the Core summary of product characteristics for human plasma-derived and recombinant coagulation factor VIII products (EMA/CHMP/BPWP/1619/1999 rev. 2) and with the QRD version 10 (affects labelling only). Moreover the MAH is combining the SmPC for all strengths and updating Annex A with detailed information on the packaging. C.I.11: Submission of an updated RMP version 8.0 in order to align the content in a single harmonised worldwide version for simoctocog alfa (rFVIII).” Opinion adopted on 14.09.2017. Request for Supplementary Information adopted on 22.06.2017, 21.04.2017.	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Olumiant - baricitinib - EMEA/H/C/004085/II/0002	Request for Supplementary Information adopted

MAH: Eli Lilly Nederland B.V., Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Patrick Batty, “Update of sections 4.5 and 5.2 of the SmPC, based on the final study report of in vitro study to investigate the inhibitory effect of baricitinib on the organic anion transporter 2 (OAT2) in fulfilment of PAM (MEA 001). The updated RMP version 3.0 has been submitted as part of this application.” Request for Supplementary Information adopted on 14.09.2017.	with a specific timetable.
Opdivo - nivolumab - EMEA/H/C/003985/II/0032 MAH: Bristol-Myers Squibb Pharma EEIG, Rapporteur: Jorge Camarero Jiménez, PRAC Rapporteur: Brigitte Keller-Stanislawski, “Update of sections 4.1, 4.2, 4.4, 4.8, 5.1 and 5.2 of the SmPC in order to update the statement on outcome benefit, to add administration guidance, to update the safety information and updated overall survival data based on final results from study CA209067 (listed as an imposed PAES in the Annex II). Study CA209067 is an interventional, randomized, double-blind study of nivolumab monotherapy or nivolumab combined with ipilimumab versus ipilimumab monotherapy in adult subjects with previously untreated, unresectable or metastatic Stage III or Stage IV melanoma. The Package Leaflet is updated accordingly. The RMP version 7.6 has also been submitted. This submission fulfils ANX 016 and Annex II is updated accordingly. In addition, the Marketing authorisation holder (MAH) took the opportunity to make other changes to the Annex II conditions to reflect the fact that ANX/005 has been fulfilled, i.e. the initial ANX 005 commitment has been removed and was replaced by the new ANX 005.1 and ANX005.2 commitments. Moreover, the MAH took the opportunity to introduce minor editorial and formatting revisions in the PI.” Opinion adopted on 14.09.2017. Request for Supplementary Information adopted on 18.05.2017.	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Opdivo - nivolumab - EMEA/H/C/003985/II/0036/G	See 9.1 in main agenda.

MAH: Bristol-Myers Squibb Pharma EEIG, Co-Rapporteur: Paula Boudewina van Hennik, PRAC Rapporteur: Brigitte Keller-Stanislawski, Procedure Manager: Elisa Pedone, EPL: Silvy Da Rocha Dias, "C.I.4 (Type II) - Update of sections 4.2, 5.1, 5.2 and 6.6 of the SmPC in order to introduce new dosing regimens and schedule. C.I.4 (Type II) - Update of sections 4.2, 5.1, 5.2 and 6.6 of the SmPC in order to introduce change the infusion time from 60 minutes to 30 minutes These changes are based on interim results from study CA209153; this is a phase IIIb/IV safety trial of nivolumab in subjects with advanced or metastatic non-small cell Lung cancer who have progressed during or after receiving at least one prior systemic regimen; The Package Leaflet is updated accordingly. The RMP version 10.0 has also been submitted." Request for Supplementary Information adopted on 14.09.2017.	
Opdivo - nivolumab - EMEA/H/C/003985/II/0038 MAH: Bristol-Myers Squibb Pharma EEIG, Rapporteur: Jorge Camarero Jiménez, PRAC Rapporteur: Brigitte Keller-Stanislawski, "Update of section 4.8 of the SmPC with longer follow-up for subjects proceeding to allogeneic transplant following nivolumab treatment, of section 5.1 of the SmPC with efficacy data from longer follow-up based on final results from study CA209205 listed as a PAES in the Annex II; this is a Phase 2, non-comparative, multi-cohort, single-arm, open-label study of nivolumab (BMS-936558) in cHL subjects after failure of ASCT Annex II is updated to remove the commitment. Version 7.5 of the RMP has been submitted." Request for Supplementary Information adopted on 14.09.2017.	Request for Supplementary Information adopted with a specific timetable.
Orkambi - lumacaftor / ivacaftor - EMEA/H/C/003954/II/0021 MAH: Vertex Pharmaceuticals (Europe) Ltd., Rapporteur: Nithyanandan Nagercoil, PRAC Rapporteur: Almath Spooner, "Update of section 4.8 of the SmPC in order to add information on respiratory events based on final results from study Study VX14-809-106 (Study 106), a Phase 3b, open-label study to evaluate safety and tolerability of lumacaftor and ivacaftor	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

combination therapy in subjects 12 years and older with Cystic Fibrosis and advanced lung disease, homozygous for the F508del-CFTR Mutation. Efficacy was evaluated as a secondary objective. This study report is being submitted to fulfil MEA 002. An updated RMP (version 3.4) has also been submitted." Opinion adopted on 14.09.2017. Request for Supplementary Information adopted on 22.06.2017.	
Praxbind - idarucizumab - EMEA/H/C/003986/II/0007 MAH: Boehringer Ingelheim International GmbH, Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Menno van der Elst, "Update of sections 4.2, 4.8, 5.1 and 5.2 of the SmPC in order to reflect the final results from a study 1321.3 titled "A Phase III, case series clinical study of the reversal of the anticoagulant effects of dabigatran by intravenous administration of 5.0 g idarucizumab (BI 655075) in patients treated with dabigatran etexilate who have uncontrolled bleeding or require emergency surgery or procedures. RE-VERSE-AD (A study of the RE-VERSal Effects of Idarucizumab on Active Dabigatran) trial" listed as a category 3 study in the RMP (MEA 001). The RMP version 3.0 has also been submitted. In addition, the Marketing authorisation holder took the opportunity to update the immunogenicity section in 5.1 of SmPC and to bring the PI in line with the latest QRD template version 10." Request for Supplementary Information adopted on 14.09.2017, 20.07.2017.	Request for Supplementary Information adopted with a specific timetable.
Prolia - denosumab - EMEA/H/C/001120/II/0069 MAH: Amgen Europe B.V., Rapporteur: Kristina Dunder, PRAC Rapporteur: Ulla Wändel Liminga, "Update of section 4.8 of the SmPC and PL in order to remove cataracts from the list of adverse reaction associated with denosumab therapy based on final data from study 20080560, a category 3 study in the RMP (multicentre, randomized, double blind, placebo-controlled study in men with non-metastatic prostate cancer receiving androgen deprivation therapy cataract development and progression study using a	Positive Opinion adopted by consensus on 01.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

slit-lamp-based evaluation system (Lens Opacities Classification System III (LOCS III).) In addition, the RMP has been updated to remove the important potential risk 'cataract in men with prostate cancer receiving androgen deprivation therapy'." Opinion adopted on 01.09.2017. Request for Supplementary Information adopted on 09.06.2017.	
Remicade - infliximab - EMEA/H/C/000240/II/0204 MAH: Janssen Biologics B.V., Rapporteur: Kristina Dunder, PRAC Rapporteur: Ulla Wändel Liminga, "Submission of the final registry report from the C0168T71 study (a review and analysis of birth outcomes from Swedish, Danish and Finish medical birth registers) and an evaluation of pregnancy data from multiple sources. Section 4.6 of the SmPC, relevant section of the PL and the RMP version 13.2 has been updated to reflect the study results. The MAH has also taken the opportunity to bring the product in line with the QRD template and update the local representative section of the PL." Request for Supplementary Information adopted on 14.09.2017, 22.06.2017, 23.03.2017.	Request for Supplementary Information adopted with a specific timetable.
Reyataz - atazanavir / atazanavir sulfate - EMEA/H/C/000494/II/0111 MAH: Bristol-Myers Squibb Pharma EEIG, Rapporteur: Joseph Emmerich, PRAC Rapporteur: Caroline Laborde, "Update of sections 4.4 and 4.8 of the SmPC to add a warning on chronic kidney disease observed in HIV infected patients during treatment with atazanavir (with or without ritonavir). This update is based on a review of the MAH safety database, a cohort study of patients with laboratory values from a large US administrative claims database and a review of published scientific literature. The Package Leaflet is updated accordingly. The RMP version 12.0 has also been submitted." Opinion adopted on 01.09.2017. Request for Supplementary Information adopted on 09.06.2017.	Positive Opinion adopted by consensus on 01.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Soliris - eculizumab - EMEA/H/C/000791/II/0098, Orphan MAH: Alexion Europe SAS, Rapporteur: Jorge Camarero Jiménez, PRAC Rapporteur: Eva A.	Request for Supplementary Information adopted with a specific timetable.

Segovia, “Update of sections 4.6 and 5.3 of the SmPC in order to update the safety information related to pregnancy, lactation and fertility following the review of data in PSUR 13 and 14. Annex II and the Package Leaflet are updated accordingly. The RMP version 17 has also been submitted with updated information on pregnancy and lactation and fertility.” Request for Supplementary Information adopted on 14.09.2017.	
Spedra - avanafil - EMEA/H/C/002581/II/0027/G MAH: Menarini International Operations Luxembourg S.A., Rapporteur: Concepcion Prieto Yerro, PRAC Rapporteur: Dolores Montero Corominas, “Update of section 4.4. to reflect the results of clinical study TA-402 “A Double-Blind, Randomized, Placebo-Controlled, Single-Dose, Parallel Study to Assess the Effects of Avanafil on Multiple Parameters of Vision, including, but Not Limited to Visual Acuity, Intraocular Pressure, Pupillometry, and Color Vision Discrimination, in Healthy Male Subjects). Update of section 4.6. of the SmPC in order to reflect the results of clinical study TA-401 “A Randomized, Double-Blind, Placebo-Controlled, Parallel Group, Multicenter Clinical Trial of the Effect of Avanafil on Spermatogenesis in Healthy Adult Males and Adult Males with Mild Erectile Dysfunction”. The Package Leaflet is updated accordingly. The RMP version 5.1has also been submitted. In addition, the MAH took the opportunity to make an editorial correction on the approved SmPc by adding the missing adverse reaction epistaxis from the tabulated list of adverse reactions reported in section 4.8. Additionally, the MAH took the opportunity of this variation to align the information included in Section 3 “How to take Spedra” in the Package Leaflet to section 4.2 “Posology” in the SmPC. Some additional minor amendments, due to translation mistakes are proposed for the French Product Information.” Request for Supplementary Information adopted on 14.09.2017.	Request for Supplementary Information adopted with a specific timetable.

Tresiba - insulin degludec - EMEA/H/C/002498/II/0028 MAH: Novo Nordisk A/S, Rapporteur: Kristina Dunder, PRAC Rapporteur: Qun-Ying Yue, "Update of section 5.1 of the SmPC based on new clinical data from a cardiovascular outcome trial EX1250-4080 (DEVOTE) conducted for Tresiba. DEVOTE was a randomised, double-blind and event-driven clinical trial with a median duration of 2 years comparing the cardiovascular safety of Tresiba versus insulin glargine (100 units/mL) in patients with type 2 diabetes mellitus at high risk of cardiovascular events. The RMP version 8 has also been submitted, with updates consequent to the data in support of the application." Request for Supplementary Information adopted on 01.09.2017.	Request for Supplementary Information adopted with a specific timetable.
Wakix - pitolisant - EMEA/H/C/002616/II/0004/G, Orphan MAH: BIOPROJET PHARMA, Rapporteur: Joseph Emmerich, PRAC Rapporteur: Kirsti Villikka, "Update of sections 4.2, 4.5 and 5.2 of the SmPC based on the final CSR of study P15-02 (to assess the mass balance recovery, metabolite profile and metabolite identification of 14C-pitolisant at steady state conditions, in healthy CYP2D6 phenotyped subjects), P14-07 (to evaluate pharmacokinetic interaction of pitolisant with sodium oxybate and modafinil in healthy male volunteers) and P15-15 (to evaluate pharmacokinetic interaction of pitolisant with CYP3A4 substrates (midazolam), CYP2B6 substrates (bupropion), UGT2B7 inhibotors (probenecide)) in fulfilment of PAM (MEA 02, 03 and 04). The Package Leaflet is updated accordingly. In addition, the Marketing authorisation holder (MAH) took the opportunity to make minor editorial change in section 4.6 and 4.8 of the SmPC. Moreover, updated RMP version 5.2 has been agreed as part of this procedure." Opinion adopted on 14.09.2017. Request for Supplementary Information adopted on 22.06.2017, 23.03.2017.	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Xarelto - rivaroxaban - EMEA/H/C/000944/II/0052/G MAH: Bayer AG, Rapporteur: Kristina Dunder, PRAC Rapporteur: Qun-Ying Yue, "Group of variations consisting of:	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

1) C.1.4. To add the authorised indications "Treatment of deep vein thrombosis (DVT) and pulmonary embolism (PE), and prevention of recurrent DVT and PE in adults" to Xarelto 10 mg based on Einstein Choice trial (A randomised phase III clinical study to evaluate efficacy and safety of Reduced-dosed rivaroxaban and standard-dosed rivaroxaban versus ASA in the long-term prevention of recurrent symptomatic venous thromboembolism in patients with symptomatic deep-vein thrombosis and/or pulmonary embolism) in section 4.1 of the SmPC 10 mg.

Consequently:

- Changes in sections 4.2, 4.8 and 5.1 for Xarelto 10mg, 15mg and 20 mg are made in order to update the posology, efficacy and safety information.
- Annex III is updated to include Xarelto 10 mg into Patient alert card to support management of bleeding when the 10 mg is treated for long-term prevention of recurrent VTE
- RMP (version 10.2) is updated

2) B.II.e.5.a.1- to add a new pack size of 14 film coated tablets in blister (PP/alu) for Xarelto 10 mg

3) B.II.e.5.a.1- to add a new pack size of 28 film coated tablets in blister (PP/alu) for Xarelto 10 mg

4) B.II.e.5.a.1- to add a new pack size of 98 film coated tablets in blister (PP/alu) for Xarelto 10 mg

5) B.II.e.1.b.1 to change immediate packaging of the finished product for 10 mg film coated tablets to introduce HDPE bottle with screw cap including new presentation (pack containing 100 film coated tablets for 10 mg strength)

6) C.1.4 To add information on interactions with SSRIs and SNRIs in section 4.5 and a related warning in section 4.4 of the SmPC based on post-hoc analyses to investigate bleeding risk for rivaroxaban in patients with and without use of SSRI or SNRIs from the pivotal studies.

In addition, MedDRA terminology is updated in the adverse drug reactions table in section 4.8 of the SmPC

7) C.1.11.z To delete from the summary of safety concerns: "Patients undergoing major orthopaedic surgery other than elective hip or knee replacement surgery" and "Remedial pro-coagulant therapy for excessive

haemorrhage". Part II - Modules SVIII: Summary of the safety concerns, Part III, Section 1 Safety Concerns and overview of planned pharmacovigilance action were amended accordingly. In addition, Part II, Safety Specification, module SIV, Populations not studied in clinical trials: "Patients undergoing major orthopaedic surgery other than elective hip or knee replacement surgery" and "Remedial pro-coagulant therapy for excessive haemorrhage" was updated. Opinion adopted on 14.09.2017. Request for Supplementary Information adopted on 20.07.2017, 18.05.2017.	
Yervoy - ipilimumab - EMEA/H/C/002213/II/0042 MAH: Bristol-Myers Squibb Pharma EEIG, Rapporteur: Paula Boudewina van Hennik, PRAC Rapporteur: Sabine Straus, "Update of sections 4.4, 4.8 and 5.1 of the SmPC to reflect the final results of study CA184-169, a randomized double-blind phase III study of ipilimumab administered at 3 mg/kg versus at 10 mg/kg in subjects previously treated or untreated with unresectable or metastatic melanoma, in order to fulfil ANX 014.1. The MAH also provided with this variation application efficacy and safety data from study CA184-169 in two subgroups: female ≥ 50 years of age and with brain metastases in order to fulfil MEA 015.1. Annex II.D and the RMP (version 14.0) are updated accordingly. In addition the Marketing authorisation holder (MAH) took the opportunity to update the list of local representatives in the Package Leaflet, to include some editorial changes and correct some typos throughout the product information, and to bring the product information in line with the latest QRD template version 10." Request for Supplementary Information adopted on 01.09.2017, 05.05.2017.	Request for Supplementary Information adopted with a specific timetable.
Yervoy - ipilimumab - EMEA/H/C/002213/II/0047/G MAH: Bristol-Myers Squibb Pharma EEIG, Rapporteur: Paula Boudewina van Hennik, PRAC Rapporteur: Sabine Straus, "Update of section 4.4 to revised the current warning on concurrent administration with vemurafenib to enhance awareness on the potential of hypersensitivity reactions when ipilimumab is used sequentially with vemurafenib as requested by the PRAC	Positive Opinion adopted by consensus on 01.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

following the assessment of PSUSA/00009200/201603. Update of sections 4.8 of the SmPC to amend the frequency of the adverse drug reaction 'Vogt-Konyanagi-Haranda syndrome' from 'not know' to 'very rare'. The RMP (version 16) has been updated accordingly.]In addition, the Marketing authorisation holder (MAH) took the opportunity to implement some editorial changes to sections 4.2 and 4.4 of the SmPC to update the dose modification information for hepatotoxicity management guidelines in line with National Cancer Institute (NCI) Common Terminology Criteria for Adverse Events (CTCAE) recommendations (version 4)." Opinion adopted on 01.09.2017. Request for Supplementary Information adopted on 09.06.2017.	
WS1026 **Rasilez-EMEA/H/C/000780/WS1026/0110** **Rasilez** **HCT-EMEA/H/C/000964/WS1026/0080** MAH: Noden Pharma DAC, Lead Rapporteur: Daniela Melchiorri, Lead PRAC Rapporteur: Carmela Macchiarulo, "Update of section 5.1 of the SmPC in order to reflect the results of study SPP100F2301 (ATMOSPHERE) a multicenter, randomized, doubleblind, parallel group, active-controlled study to evaluate the efficacy and safety of both aliskiren monotherapy and aliskiren/enalapril combination therapy compared to enalapril monotherapy, on morbidity and mortality in patients with chronic heart failure (NYHA Class II - IV). The RMP (v 13) has also been updated to reflect the study results." Request for Supplementary Information adopted on 14.09.2017, 22.06.2017, 21.04.2017, 15.12.2016.	Request for Supplementary Information adopted with a specific timetable.
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	Request for Supplementary Information adopted with a specific timetable.

safety information on QTc prolongation based on the final results from study AI266959; this is an interventional study to determine the concentration-electrocardiographic effects of efavirenz in healthy subjects enriched for cyp2b6 polymorphisms; the Package Leaflet is updated accordingly. The RMP version 8 has also been submitted. C.I.4 (Type II) – Update of sections 4.4 and 4.8 to add catatonia as a Psychiatric symptom following an assessment of catatonia cases reported in the literature and via the United States (US) Food and Drug Administration Adverse Event Reporting System (FAERS)." Request for Supplementary Information adopted on 01.09.2017, 06.07.2017, 06.04.2017.	
WS1158/G **Humalog-EMEA/H/C/000088/WS1158/0154/G** **Liprolog-EMEA/H/C/000393/WS1158/0117/G** MAH: Eli Lilly Nederland B.V., Lead Rapporteur: Robert James Hemmings, Lead PRAC Rapporteur: Julie Williams"Type II (B.IV.1.c): to add a pre-filled pen: the Humalog and Liprolog 100 U/ml Junior KwikPen. The Junior KwikPen can administer insulin in half unit increments and contains the insulin lispro 3ml cartridge that is already approved for use. The pack contains 5 pre-filled pens. Type IAin (B.II.e.5.a.1): to add a new pack size of 10 (2x5) pre-filled pens (multipack) for the Humalog and Liprolog 100 U/ml Junior KwikPen. This presentation contains the insulin lispro 3ml cartridge that is already approved for use. Type II (C.I.z): Update of sections 4.2 and 4.4 of the SmPC of the already authorised 100 U/ml Humalog and Liprolog presentations to indicate in section 4.2 that it can be used in paediatric population instead of in section 4.4, where the text that states that the product should only be used in children in preference to soluble insulin when a fast action of insulin might be beneficial, is deleted. The Package leaflet is updated accordingly." Opinion adopted on 14.09.2017. Request for Supplementary Information adopted on 09.06.2017, 05.05.2017.	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

WS1168 **Azilect-EMEA/H/C/000574/WS1168/0077** **Rasagiline** **ratiopharm-EMEA/H/C/003957/WS1168/0010** MAH: Teva B.V., Lead Rapporteur: Bruno Sepodes, Lead PRAC Rapporteur: Ana Sofia Diniz Martins, "Update of sections 4.4, 4.7 and 4.8 to include a new warning on excessive daytime sleepiness and sudden sleep onset episodes, update of section 4.9 to remove 'dysphoria' as a symptom reported following overdose of rasagiline based on a CCDS update. The Package Leaflet is updated accordingly. The RMP version 2.0 has also been submitted. In addition, the Worksharing applicant (WSA) took the opportunity to make editorial changes throughout the PI, to correct the invented name for Rasagiline Ratiopharm in the Czech annexes and to bring the PI in line with the latest QRD template version 10." Opinion adopted on 01.09.2017.	Positive Opinion adopted by consensus on 01.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1180 **Corlentor-EMEA/H/C/000598/WS1180/0047** **Ivabradine** **Anpharm-EMEA/H/C/004187/WS1180/0006** **Procoralan-EMEA/H/C/000597/WS1180/0046** MAH: Les Laboratoires Servier, Lead Rapporteur: Johann Lodewijk Hillege, Lead PRAC Rapporteur: Menno van der Elst, "Update to the section 4.8 of the SmPC with new ADRs: Ventricular tachycardia, Ventricular fibrillation and Torsade de pointes. The PL is updated accordingly. The RMP version 6 has also been submitted. In addition the MAH took the opportunity to align the PI with the latest QRD template 10.0." Request for Supplementary Information adopted on 01.09.2017.	Request for Supplementary Information adopted with a specific timetable.
WS1182 **Amgevita-EMEA/H/C/004212/WS1182/0001** **SOLYMBIC-EMEA/H/C/004373/WS1182/0001** MAH: Amgen Europe B.V., Lead Rapporteur: Kristina Dunder, Lead PRAC Rapporteur: Ulla Wändel Liminga, "Submission of the final report from study/studies 20130258, an open-label,	Positive Opinion adopted by consensus on 01.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

single-arm extension study to evaluate the long-term safety and efficacy of ABP 501 in subjects with moderate to severe rheumatoid arthritis, listed as a category 3 study in the RMP (MEA002). No changes of the PI are proposed; the RMP is updated accordingly (version 2.0)." Opinion adopted on 01.09.2017. Request for Supplementary Information adopted on 06.07.2017.	
WS1211 **Januvia-EMEA/H/C/000722/WS1211/0059** **Ristaben-EMEA/H/C/001234/WS1211/0051** **TESAVEL-EMEA/H/C/000910/WS1211/0059** **Xelevia-EMEA/H/C/000762/WS1211/0063** MAH: Merck Sharp & Dohme Limited, Lead Rapporteur: Johann Lodewijk Hillege, Lead PRAC Rapporteur: Menno van der Elst, "Update of sections 4.2, 4.4 and 5.2 of the SmPC in order to modify the information on dosing, an existing warning and administration instructions, respectively for use of sitagliptin in patients with type 2 diabetes mellitus and renal impairment. Consequently, the RMP version 8 has also been updated accordingly. In addition, the Worksharing applicant (WSA) took the opportunity to update the list of local representatives in the Package Leaflet for Tesavel and to bring the PI in line with the latest QRD template version 10. Minor editorial changes are also introduced in the Product Information." Request for Supplementary Information adopted on 14.09.2017.	Request for Supplementary Information adopted with a specific timetable.
WS1212/G **Efficib-EMEA/H/C/000896/WS1212/0085/G** **Janumet-EMEA/H/C/000861/WS1212/0085/G** **Ristfor-EMEA/H/C/001235/WS1212/0072/G** **Velmetia-EMEA/H/C/000862/WS1212/0088/G** MAH: Merck Sharp & Dohme Limited, Lead Rapporteur: Johann Lodewijk Hillege, Lead PRAC Rapporteur: Menno van der Elst, "Update of sections 4.2, and 5.2 of the SmPC in order to modify the information on dosing, and	Request for Supplementary Information adopted with a specific timetable.

administration instructions respectively for use of sitagliptin/metformin in patients with type 2 diabetes mellitus and moderate renal impairment. Consequently, the RMP version 8 has also been updated accordingly. Section 4.5 of the SmPC is also updated to include information on the concominant use of ranolazine, vandetanib, dolutegravir and cimetidine. In addition, the Worksharing applicant (WSA) took the opportunity to update the list of local representatives in the Package Leaflet for Efficib and to bring the PI in line with the latest QRD template version 10. Minor editorial changes are also introduced in the Product Information." Request for Supplementary Information adopted on 14.09.2017.	

B.5.4. PRAC assessed procedures

PRAC Led **Eliquis - apixaban - EMEA/H/C/002148/II/0043** MAH: Bristol-Myers Squibb / Pfizer EEIG, Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Menno van der Elst, PRAC-CHMP liaison: Johann Lodewijk Hillege, "Submission of the final report from study (CV185-365) listed as a category 3 study in the RMP. This is a post authorisation safety study which evaluates the effectiveness of Eliquis (apixaban) risk minimisation tools in the European Economic Area countries. A RMP (version 17.0) has also been submitted to reflect the completion of the study CV185-365." Opinion adopted on 01.09.2017.	Positive Opinion adopted by consensus on 01.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
PRAC Led **Invokana - canagliflozin - EMEA/H/C/002649/II/0030** MAH: Janssen-Cilag International NV, Rapporteur: Martina Weise, PRAC Rapporteur: Valerie Strassmann, PRAC-CHMP liaison: Martina Weise, "Submission of an updated RMP version 7.1 in order to include prior commitments made to PRAC during the PSUR/LEG procedural review of pancreatitis cases and the Article 20 referral procedure reviewing lower limb amputation in relation to the use of SGLT-2 inhibitors. In addition, the updated RMP reflects labelling	Positive Opinion adopted by consensus on 01.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

changes that resulted from a variation to add information regarding fatal DKA cases to the existing DKA warning and the Article 31 procedure reviewing metformin-containing medicines." Opinion adopted on 01.09.2017.	
PRAC Led **Plenadren - hydrocortisone - EMEA/H/C/002185/II/0024, Orphan** MAH: Shire Services BVBA, Rapporteur: Kristina Dunder, PRAC Rapporteur: Qun-Ying Yue, PRAC-CHMP liaison: Kristina Dunder, "Submission of an updated RMP (version 3.1) in order to submit protocol amendments of SHP 617-400 (EU-AIR) study – A European multicentre, multi-country, post-authorisation, observation study (registry) of patients with chronic adrenal insufficiency (category 3). Additionally, the opportunity is being taken to implement a change agreed by the PRAC/CHMP as part of the assessment of MEA 005.3 in July 2016 and remove from the RMP reference to study SHP617-404 (SWE-DUS), a Category 3 study to monitor off-label use of Plenadren to evaluate physician prescribing patterns." Request for Supplementary Information adopted on 01.09.2017, 05.05.2017.	Request for Supplementary Information adopted with a specific timetable.
PRAC Led **Viread - tenofovir disoproxil - EMEA/H/C/000419/II/0182** MAH: Gilead Sciences International Limited, Rapporteur: Joseph Emmerich, PRAC Rapporteur: Caroline Laborde, PRAC-CHMP liaison: Joseph Emmerich, "Submission of the final report from Study GX-US-174-0172, listed as a category 3 study in the RMP. This is a 5-year observational (non-interventional) renal safety registry conducted to provide further safety data in HBV-infected patients with decompensated liver disease." Opinion adopted on 01.09.2017.	Positive Opinion adopted by consensus on 01.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
PRAC Led **Vokanamet - canagliflozin / metformin - EMEA/H/C/002656/II/0031** MAH: Janssen-Cilag International NV, Rapporteur: Martina Weise, PRAC Rapporteur: Menno van der Elst, PRAC-CHMP liaison: Johann Lodewijk Hillege, "Submission of an updated RMP version 7.1 in order to include prior commitments	Positive Opinion adopted by consensus on 01.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

made to PRAC during the PSUR/LEG procedural review of pancreatitis cases and the Article 20 referral procedure reviewing lower limb amputation in relation to the use of SGLT-2 inhibitors. In addition, the updated RMP reflects labelling changes that resulted from a variation to add information regarding fatal DKA cases to the existing DKA warning and the Article 31 procedure reviewing metformin-containing medicines." Opinion adopted on 01.09.2017.	
PRAC Led **Xeplion - paliperidone - EMEA/H/C/002105/II/0031** MAH: Janssen-Cilag International NV, Rapporteur: Kristina Dunder, PRAC Rapporteur: Qun-Ying Yue, PRAC-CHMP liaison: Filip Josephson, "Submission of the final study report of the "Post-Authorization Safety Study Using European Union Databases to Assess the Risk of Cardiovascular and Cerebrovascular Adverse Events in Elderly Patients Treated with Paliperidone Palmitate, Paliperidone Prolonged-Release, and Other Antipsychotics"." Opinion adopted on 14.09.2017. Request for Supplementary Information adopted on 22.06.2017, 23.02.2017.	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
PRAC Led **Xyrem - sodium oxybate - EMEA/H/C/000593/II/0066** MAH: UCB Pharma Limited, Rapporteur: Bruno Sepodes, PRAC Rapporteur: Ana Sofia Diniz Martins, PRAC-CHMP liaison: Bruno Sepodes, "Submission of the final report from study (C00302) listed as a category 3 study in the RMP. This is a post marketing non-interventional surveillance pharmacoepidemiology study (PMSS) to evaluate long-term safety, tolerability and compliance in administration of Xyrem (sodium oxybate) oral solution in patients who receive treatment with this medication in regular clinical practice. In addition, the MAH submitted a revised risk management plan version 8.0." Opinion adopted on 01.09.2017. Request for Supplementary Information adopted on 09.06.2017.	Positive Opinion adopted by consensus on 01.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
PRAC Led **Yervoy - ipilimumab - EMEA/H/C/002213/II/0049**	Positive Opinion adopted by consensus on 01.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP

MAH: Bristol-Myers Squibb Pharma EEIG, Rapporteur: Paula Boudewina van Hennik, PRAC Rapporteur: Sabine Straus, PRAC-CHMP liaison: Johann Lodewijk Hillege, “Submission of an updated RMP (version 17.1) in order to amend the study objectives and milestones for two studies: - study CA184332, a multi-site retrospective observational study of US patients with unresectable or metastatic melanoma receiving ipilimumab (Yervoy) as first line therapy in a community setting, a category 3 study in the RMP (MEA 029): to submit the final study report with 2-years of follow-up - study CA184338, a multi-site retrospective observational study of US patients with unresectable or metastatic melanoma receiving ipilimumab (Yervoy) as first line therapy, a category 3 study in the RMP (MEA 030): to submit the final study report with 4-years of follow-up.” Opinion adopted on 01.09.2017. Request for Supplementary Information adopted on 06.07.2017.	recommendation.
PRAC Led **WS1188** **Humalog-EMEA/H/C/000088/WS1188/0157** **Liprolog-EMEA/H/C/000393/WS1188/0120** MAH: Eli Lilly Nederland B.V., Lead Rapporteur: Robert James Hemmings, Lead PRAC Rapporteur: Julie Williams, PRAC-CHMP liaison: Robert James Hemmings, “Submission of the final report of a non-interventional post-authorisation safety study EUPAS 13422. This study is aimed to evaluate the impact of additional risk minimisation measures on healthcare professionals and on patients’ understanding and their behaviour regarding the risk of hypoglycaemia and/or hyperglycaemia due to medication errors associated with administration of Humalog 200 U/ml KwikPen.” Opinion adopted on 01.09.2017. Request for Supplementary Information adopted on 06.07.2017.	Positive Opinion adopted by consensus on 01.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
PRAC Led **WS1197** **Actraphane-EMEA/H/C/000427/WS1197/0072** **Actrapid-EMEA/H/C/000424/WS1197/006**	Request for Supplementary Information adopted with a specific timetable.

6 **Insulatard-EMEA/H/C/000441/WS1197/0069** **Mixtard-EMEA/H/C/000428/WS1197/0073** **Protaphane-EMEA/H/C/000442/WS1197/0068** MAH: Novo Nordisk A/S, Lead Rapporteur: Hanne Lomholt Larsen, Lead PRAC Rapporteur: Doris Stenver, PRAC-CHMP liaison: Sinan B. Sarac, "Submission of an updated RMP version 3.0 according to GVP Module V, in order to remove three important potential risks (immunogenicity, allergic reactions and lack of efficacy) related to the new NN729 manufacturing process from the RMP, remove hypoglycaemia and anaphylactic reactions, remove peripheral neuropathy, refraction disorders, lipodystrophy, urticaria, rash, oedema and diabetic retinopathy and remove missing information concerning special populations. No changes are proposed to the product information." Request for Supplementary Information adopted on 01.09.2017.	

B.5.5. CHMP-CAT assessed procedures

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

WS0935/G **Filgrastim** **Hexal-EMEA/H/C/000918/WS0935/0035/G** **Zarzio-EMEA/H/C/000917/WS0935/0036/G** MAH: Sandoz GmbH, Lead Rapporteur: Greg Markey Opinion adopted on 14.09.2017.	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1172 **Infanrix hexa-EMEA/H/C/000296/WS1172/0221** MAH: GlaxoSmithkline Biologicals SA, Lead Rapporteur: Bart Van der Schueren	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

Opinion adopted on 14.09.2017.	
WS1184 **Eucreas-EMEA/H/C/000807/WS1184/0063** **Icandra-EMEA/H/C/001050/WS1184/0064** **Zomarist-EMEA/H/C/001049/WS1184/0064** MAH: Novartis Europharm Ltd, Lead Rapporteur: Kristina Dunder Opinion adopted on 14.09.2017.	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1185/G **Hexacima-EMEA/H/C/002702/WS1185/0065/G** **Hexaxim-EMEA/H/W/002495/WS1185/0071/G** **Hexyon-EMEA/H/C/002796/WS1185/0069/G** MAH: Sanofi Pasteur Europe, Duplicate, Duplicate of Hexacima, Lead Rapporteur: Jan Mueller-Berghaus Request for Supplementary Information adopted on 14.09.2017.	Request for Supplementary Information adopted with a specific timetable.
WS1187/G **Kalydeco-EMEA/H/C/002494/WS1187/0061/G** **Orkambi-EMEA/H/C/003954/WS1187/0022/G** MAH: Vertex Pharmaceuticals (Europe) Ltd., Lead Rapporteur: Nithyanandan Nagercoil Opinion adopted on 14.09.2017.	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1192 **Hexacima-EMEA/H/C/002702/WS1192/0066** **Hexaxim-EMEA/H/W/002495/WS1192/0072** **Hexyon-EMEA/H/C/002796/WS1192/0070** MAH: Sanofi Pasteur SA, Lead Rapporteur: Kristina Dunder Opinion adopted on 14.09.2017. Request for Supplementary Information adopted on 20.07.2017.	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1194 **Infanrix hexa-EMEA/H/C/000296/WS1194/0222** MAH: GlaxoSmithkline Biologicals SA, Lead Rapporteur: Bart Van der Schueren	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

Opinion adopted on 14.09.2017.	
WS1196/G **Ebymect-EMEA/H/C/004162/WS1196/0023/G** **Xigduo-EMEA/H/C/002672/WS1196/0034/G** MAH: AstraZeneca AB, Lead Rapporteur: Kristina Dunder Opinion adopted on 14.09.2017.	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1201/G **Glyxambi-EMEA/H/C/003833/WS1201/0009/G** **Jentadueto-EMEA/H/C/002279/WS1201/0041/G** **Trajenta-EMEA/H/C/002110/WS1201/0031/G** MAH: Boehringer Ingelheim International GmbH, Lead Rapporteur: Johann Lodewijk Hillege Opinion adopted on 14.09.2017.	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1202/G **Efficib-EMEA/H/C/000896/WS1202/0084/G** **Janumet-EMEA/H/C/000861/WS1202/0084/G** **Januvia-EMEA/H/C/000722/WS1202/0058/G** **Ristaben-EMEA/H/C/001234/WS1202/0050/G** **Ristfor-EMEA/H/C/001235/WS1202/0071/G** **TESAVEL-EMEA/H/C/000910/WS1202/0058/G** **Velmetia-EMEA/H/C/000862/WS1202/0087/G** **Xelevia-EMEA/H/C/000762/WS1202/0062/G** MAH: Merck Sharp & Dohme Limited, Lead Rapporteur: Johann Lodewijk Hillege Request for Supplementary Information adopted on 14.09.2017.	Request for Supplementary Information adopted with a specific timetable.
WS1204/G **Herceptin-EMEA/H/C/000278/WS1204/0134/G** **Kadcyla-EMEA/H/C/002389/WS1204/0037/G** MAH: Roche Registration Limited, Lead Rapporteur: Jan Mueller-Berghaus Opinion adopted on 14.09.2017.	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

WS1213 **Lyrica-EMEA/H/C/000546/WS1213/0090** **Pregabalin** **Pfizer-EMEA/H/C/003880/WS1213/0020** MAH: Pfizer Limited, Lead Rapporteur: Johann Lodewijk Hillege Opinion adopted on 14.09.2017.	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1214 **Aflunov-EMEA/H/C/002094/WS1214/0039** **Foclivia-EMEA/H/C/001208/WS1214/0033** MAH: Seqirus S.r.l, Lead Rapporteur: Daniela Melchiorri Request for Supplementary Information adopted on 14.09.2017.	Request for Supplementary Information adopted with a specific timetable.
WS1216 **IntronA-EMEA/H/C/000281/WS1216/0112** **PegIntron-EMEA/H/C/000280/WS1216/0131** **ViraferonPeg-EMEA/H/C/000329/WS1216/0124** MAH: Merck Sharp & Dohme Limited, Lead Rapporteur: Koenraad Norga Opinion adopted on 14.09.2017.	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1224 **Relvar Ellipta-EMEA/H/C/002673/WS1224/0031** **Revinty Ellipta-EMEA/H/C/002745/WS1224/0027** MAH: Glaxo Group Ltd, Lead Rapporteur: Concepcion Prieto Yerro Opinion adopted on 14.09.2017.	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1235/G **Incresync-EMEA/H/C/002178/WS1235/0020/G** **Vipdomet-EMEA/H/C/002654/WS1235/0022/G** **Vipidia-EMEA/H/C/002182/WS1235/0017/G** MAH: Takeda Pharma A/S, Lead Rapporteur: Johann Lodewijk Hillege Opinion adopted on 14.09.2017.	Positive Opinion adopted by consensus on 14.09.2017. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

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

Advate - octocog alfa -	The MAH withdrew the procedure on 24.08.2017.

EMEA/H/C/000520/II/0087/G MAH: Baxter AG, Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Brigitte Keller-Stanislawski Withdrawal request submitted on 24.08.2017.	
MACI - matrix applied characterised autologous cultured chondrocytes - EMEA/H/C/002522/II/0015/G, ATMP MAH: Vericel Denmark ApS, Rapporteur: Christiane Niederlaender Withdrawal request submitted on 05.09.2017.	The MAH withdrew the procedure on 05.09.2017.
Sebivo - telbivudine - EMEA/H/C/000713/II/0047 MAH: Novartis Europharm Ltd, Rapporteur: Joseph Emmerich, PRAC Rapporteur: Caroline Laborde Withdrawal request submitted on 01.09.2017.	The MAH withdrew the procedure on 01.09.2017.
WS1179 **Invega-EMEA/H/C/000746/WS1179/0055** **Trevicta-EMEA/H/C/004066/WS1179/0010** **Xeplion-EMEA/H/C/002105/WS1179/0034** MAH: Janssen-Cilag International NV, Lead Rapporteur: Kristina Dunder, "Update of section 4.6 (Fertility, pregnancy and lactation) of the SmPC in order to add new information concerning a retrospective observational cohort study with risperidone and risk of congenital malformations. Nationally approved products are also affected by this variation." Request for Supplementary Information adopted on 20.07.2017. Withdrawal request submitted on 05.09.2017.	The MAH withdrew the procedure on 05.09.2017.

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

Tarceva - erlotinib - EMEA/H/C/000618/II/0052 MAH: Roche Registration Limited, Rapporteur: Sinan B. Sarac, "Update of section 4.4 of the SmPC in order to include recommendations on Epidermal Growth Factor Receptor (EGFR) mutation status testing, to be in line with current technical and scientific progress. In addition, the Marketing authorisation holder (MAH) took the opportunity to make minor editorial changes and to bring the PI in line with the latest QRD template version 10. Moreover, the MAH took the opportunity to make minor	CHMP adopted clock stop extension

correction of section 4.2 of the SmPC.
Furthermore, the Annex II has been corrected, as requested by the EMA, to include Educational Material as an additional risk minimisation measure, which has been already in place in the RMP."
Request for Supplementary Information adopted on 20.07.2017.

B.6. START OF THE PROCEDURES TIMETABLES FOR INFORMATION

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

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

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

- trastuzumab - EMEA/H/C/002575
, treatment of metastatic and early breast cancer and metastatic gastric cancer (MGC)
List of Questions adopted on 23.02.2017.

- andexanet alfa - EMEA/H/C/004108
, treatment of direct or indirect factor Xa(FXa) inhibitor when reversal of anticoagulation is needed
List of Questions adopted on 15.12.2016.

- trastuzumab - EMEA/H/C/004361
, treatment of metastatic breast cancer, early breast cancer, metastatic gastric cancer
List of Questions adopted on 20.07.2017.

- binimetinib - EMEA/H/C/004052
, treatment of unresectable or metastatic melanoma
Treatment of unresectable melanoma, with NRA Q61 mutation.
List of Questions adopted on 26.01.2017.

- lumacaftor / ivacaftor - EMEA/H/C/003954/X/0020
List of Questions adopted on 20.07.2017.
Request for 1 year of market protection for a new indication (Article 14(11) of Regulation (EC) 726/2004)

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

List of Questions adopted on 22.06.2017.

- ertugliflozin - EMEA/H/C/004315
, type 2 diabetes mellitus
List of Questions adopted on 22.06.2017.

- ertugliflozin / sitagliptin -
EMEA/H/C/004313
, type 2 diabetes mellitus
List of Questions adopted on 22.06.2017.

- recombinant human albumin solution -
EMEA/H/D/004693
, human assisted reproductive techniques including in-vitro fertilisation procedures
List of Questions adopted on 18.05.2017.

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

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

Caprelsa - vandetanib -
EMEA/H/C/002315/R/0027
MAH: Genzyme Europe BV, Rapporteur: Alexandre Moreau, PRAC Rapporteur: Ghania Chamouni,

Cometriq - cabozantinib -
EMEA/H/C/002640/R/0027, Orphan
MAH: Ipsen Pharma, Rapporteur: Paula Boudewina van Hennik, Co-Rapporteur: Bjorg Bolstad, PRAC Rapporteur: Sabine Straus,

Sirturo - bedaquiline -
EMEA/H/C/002614/R/0024, Orphan
MAH: Janssen-Cilag International NV, Rapporteur: Filip Josephson, PRAC Rapporteur: Qun-Ying Yue,

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

B.6.8. CHMP assessed procedures scope: Pharmaceutical aspects

Atripla - efavirenz / emtricitabine / tenofovir disoproxil -
EMEA/H/C/000797/II/0125/G
MAH: Bristol-Myers Squibb and Gilead Sciences

Ltd., Rapporteur: Martina Weise

Cerezyme - imiglucerase -
EMEA/H/C/000157/II/0105
MAH: Genzyme Europe BV, Rapporteur: Johann Lodewijk Hillege

Daptomycin Hospira - daptomycin -
EMEA/H/C/004310/II/0003
MAH: Hospira UK Limited, Generic, Generic of Cubicin, Rapporteur: Kolbeinn Gudmundsson,

Elaprase - idursulfase -
EMEA/H/C/000700/II/0071/G
MAH: Shire Human Genetic Therapies AB, Rapporteur: Greg Markey,

Elonva - corifollitropin alfa -
EMEA/H/C/001106/II/0037/G
MAH: Merck Sharp & Dohme Limited, Rapporteur: Paula Boudewina van Hennik,

Erbitux - cetuximab -
EMEA/H/C/000558/II/0078/G
MAH: Merck KGaA, Rapporteur: Filip Josephson,

Natpar - parathyroid hormone -
EMEA/H/C/003861/II/0004/G, Orphan
MAH: Shire Pharmaceuticals Ireland Ltd, Rapporteur: Bart Van der Schueren,

Opatanol - olopatadine -
EMEA/H/C/000407/II/0035/G
MAH: Novartis Europharm Ltd, Rapporteur: Peter Kiely

Privigen - human normal immunoglobulin -
EMEA/H/C/000831/II/0126
MAH: CSL Behring GmbH, Rapporteur: Jan Mueller-Berghaus

Strensiq - asfotase alfa -
EMEA/H/C/003794/II/0023, Orphan
MAH: Alexion Europe SAS, Rapporteur: Greg Markey

Trulicity - dulaglutide -
EMEA/H/C/002825/II/0021
MAH: Eli Lilly Nederland B.V., Rapporteur: Greg Markey

Vaniqa - eflornithine -
EMEA/H/C/000325/II/0051
MAH: Almirall S.A, Rapporteur: Peter Kiely,

Vyndaqel - tafamidis -

EMEA/H/C/002294/II/0041/G, Orphan
MAH: Pfizer Limited, Rapporteur: Joseph Emmerich

Xadago - safinamide - EMEA/H/C/002396/II/0020
MAH: Zambon S.p.A., Rapporteur: Johann Lodewijk Hillege

WS1206/G
Exelon-EMEA/H/C/000169/WS1206/0114/G
Prometax-EMEA/H/C/000255/WS1206/0114/G
MAH: Novartis Europharm Ltd, Lead Rapporteur: Alexandre Moreau

WS1254/G
Hirobriz Breezhaler-EMEA/H/C/001211/WS1254/0042/G
Onbrez Breezhaler-EMEA/H/C/001114/WS1254/0041/G
Oslif Breezhaler-EMEA/H/C/001210/WS1254/0041/G
Ultibro Breezhaler-EMEA/H/C/002679/WS1254/0017/G
Ulunar Breezhaler-EMEA/H/C/003875/WS1254/0017/G
Xoterna Breezhaler-EMEA/H/C/003755/WS1254/0020/G
MAH: Novartis Europharm Ltd, Lead Rapporteur: Hanne Lomholt Larsen

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

Abilify - aripiprazole - EMEA/H/C/000471/II/0127
MAH: Otsuka Pharmaceutical Europe Ltd, Rapporteur: Bruno Sepodes,"Update of sections 4.4 and 4.8 of the SmPC with further information about the risk of impulse control disorders, and section 4.8 of the SmPC to include the new ADRs 'impulse control disorders', 'binge eating' and 'compulsive shopping' and to delete the ADR 'hyperglycaemia'. The Package Leaflet has been updated accordingly. Further, the MAH has

implemented minor editorial changes in section 6.1 of the SmPC, section 6 of the Package leaflet and module 3.2.P.1 to include lactose as one of the components of the excipient vanilla flavour. In addition, the MAH takes the opportunity to align the annexes with the product information of Abilify Maintena and the latest QRD template."

Abilify Maintena - aripiprazole - EMEA/H/C/002755/II/0023

MAH: Otsuka Pharmaceutical Europe Ltd, Rapporteur: Bruno Sepodes"Update of sections 4.4 and 4.8 of the SmPC with further information about the risk of impulse control disorders, and section 4.8 of the SmPC to include the new ADRs 'impulse control disorders', 'binge eating' and 'compulsive shopping'. The Package Leaflet has been updated accordingly. In addition, the MAH took the opportunity to add a unique identifier (barcode) in the labelling, to implement minor editorial changes and align the annexes with the latest QRD template."

Eliquis - apixaban - EMEA/H/C/002148/II/0047

MAH: Bristol-Myers Squibb / Pfizer EEIG, Rapporteur: Johann Lodewijk Hillege"Update of section 4.5 of the SmPC to include clarithromycin as one of the active substances which are not considered strong inhibitors of both CYP3A4 and P-gp and which are expected to increase apixaban plasma concentration to a lesser extent based on the final results from study CV185547. The final study report of study CV185547 (an open-label, non-randomised, single-sequence, crossover study in healthy subjects to determine the effect of multiple-dose clarithromycin on the single-dose pharmacokinetics of apixaban) is also submitted. In addition, the MAH took the opportunity to make some corrections in the SmPC and to update the labelling in line with the latest QRD template version 10.0."

Elonva - corifollitropin alfa - EMEA/H/C/001106/II/0038

MAH: Merck Sharp & Dohme Limited, Rapporteur: Paula Boudewina van Hennik, "Update of section 5.1 of the SmPC to include updated information regarding congenital malformations reported in infants born after a frozen_thawed embryo transfer (FTET) cycle."

Enbrel - etanercept -

EMEA/H/C/000262/II/0213

MAH: Pfizer Limited, Rapporteur: Robert James Hemmings,"Update of section 4.8 of the SmPC to update the frequency category of 7 ADRs currently listed and to split one ADR into 2, following a re-analyse of the frequencies of all listed ADRs as proposed by the MAH as part of Enbrel LEG 0168. For the ADR 'interstitial lung disease and 'autoimmune hepatitis' the description of these ADRs has also been amended as a consequence. The Marketing authorisation holder (MAH) also took the opportunity to reformat the ADR listing in section 4.8 of the SmPC. Section 4.4 of the SmPC and the Package Leaflet are updated accordingly. In addition, the MAH took the opportunity to combine the 25 mg and 50 mg pre-filled syringe (PFS) SmPCs and Package Leaflets."

Gazyvaro - obinutuzumab - EMEA/H/C/002799/II/0020, Orphan

MAH: Roche Registration Limited, Rapporteur: Sinan B. Sarac, "Update of section 4.4 of the SmPC in order to update the safety information on delayed hypersensitivity reactions based on drug safety report (DSR) number 1072198. In addition, the Marketing authorisation holder (MAH) took the opportunity to introduce editorial changes to the SmPC and package leaflet."

Praluent - alirocumab - EMEA/H/C/003882/II/0029

MAH: sanofi-aventis groupe, Rapporteur: Johann Lodewijk Hillege "Submission of the final report from study R727-CL-1032 (study title: A Phase 2, Open-Label Extension of Study R727-CL-1003 to Evaluate the Long-Term Safety and Efficacy of REGN727 Administered by Subcutaneous Injection in Patients with Heterozygous Familial Hypercholesterolemia), listed as a category 3 study in the RMP (MEA013)."

Spinraza - nusinersen - EMEA/H/C/004312/II/0002/G, Orphan

MAH: Biogen Idec Ltd, Rapporteur: Bruno Sepodes, "Update of sections 4.8 and 5.1 of the SmPC to reflect efficacy and immunogenicity data from the final clinical study reports for study CS4 and CS12 and the final update to the CS2-12 longitudinal analysis."

Stivarga - regorafenib - EMEA/H/C/002573/II/0024/G

MAH: Bayer AG, Rapporteur: Paula Boudewina van Hennik,“Submission of final results from non-clinical PK studies and physiologically-based pharmacokinetic (PBPK) modelling:

- report from a study investigating the substrate characteristics and the inhibitory potential of major human plasma metabolites towards OATP1B1 and OATP1B3.

- report from a study investigating the hepatobiliary disposition of regorafenib and its metabolites in human hepatocytes, and the inhibitory potential of regorafenib and metabolites M-2 and M-5 towards BSEP.

- report from the study 16671 using physiologically-based pharmacokinetic (PBPK) modeling investigating CYP3A4, UGT1A9, P-gp-inhibition.”

Toujeo - insulin glargine - EMEA/H/C/000309/II/0100

MAH: Sanofi-Aventis Deutschland GmbH, Duplicate, Duplicate of Lantus, Rapporteur: Johann Lodewijk Hillege,“Update of sections 4.2, 4.4 and 6.6 of the SmPC in order to add a warning on the risk for medication error associated with pre-filled pens and cartridges presentations following the evaluation of a signal (EPITT 18893) .The Package Leaflet is updated accordingly.”

Triumeq - dolutegravir / abacavir / lamivudine - EMEA/H/C/002754/II/0047

MAH: ViiV Healthcare UK Limited, Rapporteur: Kristina Dunder,“Update of section 4.5 of the SmPC with new transporter data available for abacavir and lamivudine. In addition, the MAH took the opportunity to implement some minor editorial changes in the SmPC.”

Visudyne - verteporfin - EMEA/H/C/000305/II/0095

MAH: Novartis Europharm Ltd, Rapporteur: Alexandre Moreau“Update of sections 4.4 and 4.8 of the SmPC in order to add a warning to update the safety information to reflect current knowledge about the product based on new data from spontaneous reports on localised (skin) necrosis at the injection site following extravasation; 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 and to bring the PI in line with the latest QRD template version 10."

Xalkori - crizotinib - EMEA/H/C/002489/II/0051

MAH: Pfizer Limited, Rapporteur: Alexandre Moreau, "Update of section 4.5 and 5.2 of the SmPC based on the results from the crizotinib-itraconazole drug-drug interaction (DDI) substudy of Study A8081001 (to determine the effect of the coadministration of a strong cytochrome P450 (CYP) 3A inhibitior, itraconazole, on the multiple-dose plasma pharmacokinetic of crizotinib) and the assessment of potential DDIs between crizotinib and weak and moderate CYP3A inhibitors. The labelling is also updated in line with the QRD template."

elboraf - vemurafenib - EMEA/H/C/002409/II/0043

MAH: Roche Registration Limited, Rapporteur: Filip Josephson, "Update of section 4.8 of the SmPC in order to update the safety information following results from pooled safety analysis of the final results from pivotal phase II (NP22657 BRIM-2) and pivotal phase III (NO25026 BRIM-3) trials. The Package Leaflet is updated accordingly. In addition, the Marketing authorisation holder (MAH) took the opportunity to review the SmPC and Package Leaflet in order to improve clarity and consistency across sections."

WS1251
Eviplera-EMEA/H/C/002312/WS1251/0086
Odefsey-EMEA/H/C/004156/WS1251/0019

MAH: Gilead Sciences International Limited, Lead Rapporteur: Johann Lodewijk HillegeUpdates to the Summary of Product Characteristics (SmPC) sections 4.2, 4.4 , 4.6, 5.1 and 5.2 for Eviplera and Odefsey with data from Study TMC114HIV3015 , a Category 4 additional pharmacovigilance activity in the pharmacovigilance plan for both the Eviplera and Odefsey. This is a single-arm, open-label study to assess the pharmacokinetics of Darunavir and Ritonavir, Darunavir and Cobicistat, Etravirine, and Rilpivirine in HIV-1

infected pregnant women results for the Rilpivirine arm. The Labelling and Package Leaflet are updated accordingly.

In addition, the Worksharing Applicant (WSA) has taken the opportunity to introduce some minor administrative amendments and to implement some minor linguistic amendments (MLAs) to the translations of the product information annexes."

WS1267
Docetaxel
Winthrop-EMEA/H/C/000808/WS1267/0054
Taxotere-EMEA/H/C/000073/WS1267/0129
MAH: Aventis Pharma S.A., Lead Rapporteur: Alexandre Moreau"Update of sections 4.4 and 4.8 of the SmPC in order to add a warning of enterocolitis in patients with neutropenia and to update the safety information on enterocolitis to reflect fatal outcomes based on the review of the MAH global pharmacovigilance data base, worldwide scientific literature and main pharmacovigilance textbooks; the Package Leaflet is updated accordingly.

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

B.6.10. CHMP-PRAC assessed procedures

Defitelio - defibrotide - EMEA/H/C/002393/II/0027, Orphan
MAH: Gentium S.r.l., Rapporteur: Nithyanandan Nagercoil, PRAC Rapporteur: Julie Williams, "Submission of an updated RMP version 4.0 in order to re-classify the imposed non-interventional PASS listed as a category 2 study in the RMP (specific obligation) to a study listed as a category 3 in the RMP (required additional pharmacovigilance activities). This study is an observational registry (DF-VOD2013-03-REG) which aims to record safety and outcome data in patients diagnosed with severe VOD following HSCT treated or not with defitelio. The Annex II of the product information is updated accordingly."

Vemlidy - tenofovir alafenamide - EMEA/H/C/004169/II/0004

MAH: Gilead Sciences International Limited, Rapporteur: Robert James Hemmings, PRAC Rapporteur: Amelia Cupelli,"Update of sections 4.8 and 5.1 of the Vemlidy SmPC in order to provide 96 week data from Studies GS-US-320-0108 and GS-US-320-0110, listed as category 3 studies in the RMP;

GS-US-320-0108 is an ongoing Phase 3, randomized, double-blind, non-inferiority study evaluating the safety and efficacy of Vemlidy 25 mg compared with tenofovir disoproxil fumarate 300 mg in HBeAg-negative subjects with Chronic hepatitis B.

GS-US-320-0110 is a an ongoing Phase 3, randomized, double-blind, noninferiority study evaluating the safety and efficacy of Vemlidy versus tenofovir disoproxil fumarate for the treatment of HBeAg-positive subjects with chronic hepatitis B; the Package Leaflet is updated accordingly.

The RMP version 2.0 has also been submitted.

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

B.6.11. PRAC assessed procedures

PRAC Led

Multaq - dronedarone - EMEA/H/C/001043/II/0039/G

MAH: sanofi-aventis groupe, Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Menno van der Elst, PRAC-CHMP liaison: Johann Lodewijk Hillege "C.I.13: Submission of the final report from study DRONE_C_05917 listed as a category 3 study in the RMP. This is a non-interventional epidemiological study aimed for the surveillance of serious liver injuries/diseases (SLD) with the use of dronedarone using multiple databases in the US, including the addendum on surveillance of interstitial lung disease (ILD). The RMP version 11.0 has also been submitted.

C.I.13: Submission of the final report from study DRONE_C_05911 listed as a category 3 study in the RMP. This is a non-interventional epidemiological study aimed to study the concomitant use of dronedarone and digoxin (or statins) and the risk of digitalis intoxication (or

rhabdomyolysis and myopathy). The RMP version 11.0 has also been submitted."

PRAC Led

WS1221

Brimica

Genuair-EMEA/H/C/003969/WS1221/0017

Duaklir

Genuair-EMEA/H/C/003745/WS1221/0017

MAH: AstraZeneca AB, Lead Rapporteur: Nithyanandan Nagercoil, Lead PRAC Rapporteur: Julie Williams, PRAC-CHMP liaison: Robert James Hemmings"To provide an updated RMP, version 3, to promote "Hypersensitivity (anaphylactic responses, angioedema, and urticaria)" from Important Potential Risk to Important Identified Risk, remove "Use in non-Caucasian patients" as Missing Information (with the completion of clinical studies in Asian patients), and include milestones and due dates for a cardiovascular PASS (D6560R00004) and a drug utilisation study (DUS2: D6560R00002)."

PRAC Led

WS1261

Enbrel-EMEA/H/C/000262/WS1261/0212

LIFMIOR-EMEA/H/C/004167/WS1261/0010

AH: Pfizer Limited, Lead Rapporteur: Robert James Hemmings, Lead PRAC Rapporteur: Patrick Batty, PRAC-CHMP liaison: Robert James Hemmings"Submission of the final report for the Anti-Rheumatic Treatment in Sweden Registry-Etanercept Cohort Study listed as a category 3 study in the RMP. This non-interventional PASS aimed at providing an assessment of a number of pre-specified safety outcomes for Enbrel as used in the treatment of RA in Sweden, using data from the ARTIS system, in total and from 2006."

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

WS1215
Infanrix
hexa-EMEA/H/C/000296/WS1215/0224
MAH: GlaxoSmithkline Biologicals SA, Lead
Rapporteur: Bart Van der Schueren

WS1223
Ambirix-EMEA/H/C/000426/WS1223/008
6
Fendrix-EMEA/H/C/000550/WS1223/005
9
Infanrix
hexa-EMEA/H/C/000296/WS1223/0226
Twinrix
Adult-EMEA/H/C/000112/WS1223/0120
Twinrix
Paediatric-EMEA/H/C/000129/WS1223/0
121
MAH: GlaxoSmithkline Biologicals SA, Lead
Rapporteur: Bart Van der Schueren

WS1227
Infanrix
hexa-EMEA/H/C/000296/WS1227/0225
MAH: GlaxoSmithkline Biologicals SA, Lead
Rapporteur: Bart Van der Schueren

WS1230
Lixiana-EMEA/H/C/002629/WS1230/0014
Roteas-EMEA/H/C/004339/WS1230/0002
MAH: Daiichi Sankyo Europe GmbH, Lead
Rapporteur: Concepcion Prieto Yerro

WS1239/G
Infanrix
hexa-EMEA/H/C/000296/WS1239/0227/
G
MAH: GlaxoSmithkline Biologicals SA, Lead
Rapporteur: Bart Van der Schueren

WS1240/G
Ambirix-EMEA/H/C/000426/WS1240/008
7/G
Twinrix

Adult-EMEA/H/C/000112/WS1240/0121/G
Twinrix
Paediatric-EMEA/H/C/000129/WS1240/0122/G
MAH: GlaxoSmithkline Biologicals SA, Lead Rapporteur: Robert James Hemmings

WS1253
Iblias-EMEA/H/C/004147/WS1253/0009
Kovaltry-EMEA/H/C/003825/WS1253/0012
MAH: Bayer AG, Lead Rapporteur: Kristina Dunder

WS1260
Blitzima-EMEA/H/C/004723/WS1260/0003
Ritemvia-EMEA/H/C/004725/WS1260/0003
Rituzena-EMEA/H/C/004724/WS1260/0004
MAH: Celltrion Healthcare Hungary Kft., Duplicate, Duplicate of Truxima, Lead Rapporteur: Sol Ruiz

Hexacima-EMEA/H/C/002702/WS1231/0069
Hexaxim-EMEA/H/W/002495/WS1231/0074
Hexyon-EMEA/H/C/002796/WS1231/0073
MAH: Sanofi Pasteur SA, Lead Rapporteur: Jan Mueller-Berghaus

B.7. DOCUMENTS TABLED IN MMD AFTER THE CHMP PLENARY

B.7.1. Yearly Line listing for Type I and II variations

B.7.2. Monthly Line listing for Type I variations

B.7.3. Opinion on Marketing Authorisation transfer (MMD only)

B.7.4. Notifications in accordance with Article 61(3) of Council Directive 2001/83/EC (MMD only)

B.7.5. Request for supplementary information relating to Notification of Type I variation (MMD only)

B.7.6. Notifications of Type I Variations (MMD only)

C. Annex C - Post-Authorisation Measures (PAMs), (Line listing of Post authorisation measures with a description of the PAM. Procedures starting in that given month with assessment timetabled)

D. Annex D - Post-Authorisation Measures (PAMs), (Details on PAMs including description and conclusion, for adoption by CHMP in that given month, or finalised ones with PRAC recommendation and no adoption by CHMP needed)

E. Annex E - EMEA CERTIFICATION OF PLASMA MASTER FILES

Information related to plasma master files cannot be released at the present time as these contain commercially confidential information.

E.1. PMF Certification Dossiers:

E.1.1. Annual Update

E.1.2. Variations:

E.1.3. Initial PMF Certification:

E.2. Time Tables – starting & ongoing procedures: For information

PMF timetables starting and ongoing procedures	Tabled in MMD and sent by post mail (folder E).

F. ANNEX F - Decision of the Granting of a Fee Reduction/Fee Waiver

Draft Executive Decision on the granting of a fee reduction: **For adoption**	Adopted.

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

Executive Decision on the granting of a fee reduction: EMA/610080/2017 **For adoption**	Adopted.

G. ANNEX G

G.1. Final Scientific Advice (Reports and Scientific Advice letters):

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

G.2. Ongoing procedures

G.3. PRIME

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

G.3.1. List of procedures concluding at 11-14 September 2017 CHMP plenary:

Gastroenterology-Hepatology		
1.	Treatment of eosinophilic oesophagitis	The CHMP denied eligibility to PRIME and adopted the critical summary report.
Cardiovascular Diseases		
2.	Increase in fistula survival and use for haemodialysis	The CHMP denied eligibility to PRIME and adopted the critical summary report.
Vaccines		
3.	Prevention against Influenza	The CHMP denied eligibility to PRIME and adopted the critical summary report.
Oncology		
4.	Treatment of stage IV Oesophageal Carcinoma	The CHMP denied eligibility to PRIME and adopted the critical summary report.

G.3.2. List of procedures starting in September 2017 for October 2017 CHMP adoption of outcomes

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