05 January 2021
EMA/CHMP/708154/2020
Human Medicines Division

Committee for medicinal products for human use (CHMP)

Minutes for the meeting on 12-15 October 2020

Chair: Harald Enzmann – Vice-Chair: Bruno Sepodes

Disclaimers

Some of the information contained in these minutes 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, these minutes are a working document primarily designed for CHMP members and the work the Committee undertakes.

Note on access to documents

Some documents mentioned in the minutes 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

The participants had no objection to hold the meeting remotely.

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 the restricted involvement of some meeting participants in upcoming discussions as included in the pre-meeting list of participants and restrictions. See October 2020 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 12 – 15 October 2020.

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.

The members were informed that due to a vacant CHMP co-opted member position the total number of members eligible to vote is reduced to 31 and the majority is 16.

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. 16 or more members were present remotely). All decisions, recommendations and advice were agreed by consensus, unless otherwise specified.

1.2. Adoption of agenda

CHMP agenda for 12-15 October 2020.

The CHMP adopted the agenda.

1.3. Adoption of the minutes

CHMP minutes for 20-23 July 2020 adopted via written procedure on 07.10.2020.

The CHMP noted the July minutes which were adopted via written procedure.

2. Oral Explanations

2.1. Pre-authorisation procedure oral explanations

2.1.1. selinexor - Orphan - EMEA/H/C/005127

Karyopharm Europe GmbH; treatment of patients with Relapsed Refractory Multiple Myeloma (RRMM).

Scope: Possible oral explanation

Action: Oral explanation to be held on Tuesday, 13 October 2020 at 11:00

List of Outstanding Issues adopted on 30.01.2020, 19.09.2019. List of Questions adopted on 24.04.2019.

The CHMP agreed that an oral explanation was not needed at this time.

See 3.2.

2.2. Re-examination procedure oral explanations

No items

2.3. Post-authorisation procedure oral explanations

2.3.1. Opdivo - nivolumab - EMEA/H/C/003985/II/0080

Bristol-Myers Squibb Pharma EEIG

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

Scope: "Extension of indication to include treatment of adult patients with unresectable advanced, recurrent or metastatic oesophageal squamous cell carcinoma (OSCC) after prior fluoropyrimidine- and platinum-based chemotherapy. As a consequence, sections 4.1, 4.2, 4.4, 4.8 and 5.1 of the SmPC are updated. The Package Leaflet is updated in accordance. Version 16.0 of the RMP has also been submitted."

List of experts for the SAG Oncology meeting held on 07.10.2020 adopted via written procedure on 07 October 2020

SAG report

Possible oral explanation

Action: Oral explanation to be held on Wednesday, 14 October 2020 at 16:00

Request for Supplementary Information adopted on 23.07.2020, 30.04.2020.

The CHMP noted the SAG report.

An oral explanation was held on Wednesday 14 October 2020. The presentation by the applicant focused on the clinical data in different subgroups in support of the application.

See 5.1

2.4. Referral procedure oral explanations

No items

3. Initial applications

3.1. Initial applications; Opinions

3.1.1. Trixeo Aerosphere - budesonide / glycopyrronium / formoterol fumarate dihydrate - EMEA/H/C/004983

AstraZeneca AB; as a maintenance treatment in adult patients with moderate to very severe 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 30.01.2020. List of Questions adopted on 26.04.2019.

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 CHMP noted the letter of recommendation dated 08.10.2020.

The summary of opinion was circulated for information.

3.1.2. Fintepla - fenfluramine - Orphan - EMEA/H/C/003933

Zogenix ROI Limited; treatment of seizures associated with Dravet syndrome in children aged 2 years to 17 years and adults.

Scope: Opinion

Action: For adoption

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

List of Outstanding Issues adopted on 17.09.2020, 25.06.2020, 26.03.2020. List of Questions adopted on 27.06.2019.

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

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

The divergent position (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.

The CHMP adopted the similarity assessment report.

3.1.3. Tecartus - autologous peripheral blood T cells CD4 and CD8 selected and CD3 and CD28 activated transduced with retroviral vector expressing anti-CD19 CD28/CD3-zeta chimeric antigen receptor and cultured - Orphan - ATMP - EMEA/H/C/005102

Kite Pharma EU B.V.; treatment of adult patients with relapsed or refractory Mantle Cell Lymphoma (MCL).

Scope: Opinion

Action: For adoption

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

List of Outstanding Issues adopted on 11.09.2020. List of Questions adopted on 20.05.2020.

The CHMP was updated on discussions at the CAT.

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

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

Furthermore, the CHMP considered that Autologous anti-CD19-transduced CD3+ cells 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 06.10.2020.

The summary of opinion was circulated for information.

The CHMP adopted the similarity assessment.

3.1.4. Lenalidomide Mylan - lenalidomide - EMEA/H/C/005306

Mylan Ireland Limited; treatment of multiple myeloma.

Scope: Opinion

Action: For adoption

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

List of Outstanding Issues adopted on 17.09.2020, 25.06.2020. List of Questions adopted on 30.01.2020.

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 similarity assessment report.

3.1.5. Leqvio - inclisiran - EMEA/H/C/005333

Novartis Europharm Limited; treatment for primary hypercholesterolaemia or mixed dyslipidaemia.

Scope: Opinion

Action: For adoption

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

List of Outstanding Issues adopted on 17.09.2020. List of Questions adopted on 28.05.2020.

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 inclisiran is a new active substance, as claimed by the applicant.

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

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

The CHMP noted the letter of recommendation dated 14.10.2020.

The summary of opinion was circulated for information.

3.1.6. Libmeldy - autologous CD34+ cell enriched population that contains hematopoietic stem and progenitor cells transduced ex vivo using a lentiviral vector encoding the human arylsulfatase A gene - Orphan - ATMP - EMEA/H/C/005321

Orchard Therapeutics (Netherlands) BV; treatment of Metachromatic Leukodystrophy (MLD).

Scope: Opinion

Action: For adoption

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

List of Outstanding Issues adopted on 11.09.2020. List of Questions adopted on 20.03.2020.

The CHMP was updated on discussions at the CAT.

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

The Committee, based on the draft opinion prepared by the CAT, 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 autologous CD34+ cell enriched population that contains hematopoietic stem and progenitor cells transduced ex vivo using a lentiviral vector encoding the human arylsulfatase A gene 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.

3.1.7. Oxlumo - lumasiran - Orphan - EMEA/H/C/005040

Alnylam Netherlands B.V.; primary hyperoxaluria type 1 (PH1)

Scope: Opinion

Action: For adoption

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

List of Outstanding Issues adopted on 15.09.2020. List of Questions adopted on 21.07.2020.

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 lumasiran 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 14.10.2020.

The summary of opinion was circulated for information.

3.1.8. Palforzia - Defatted powder of *Arachis hypogaea L.*, semen (peanuts) - EMEA/H/C/004917

Aimmune Therapeutics Ireland Limited; desensitisation of children and adolescents to peanut allergy.

Scope: Opinion

Action: For adoption

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

List of Outstanding Issues adopted on 17.09.2020, 25.06.2020. List of Questions adopted on 14.11.2019.

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 defatted powder of *Arachis hypogaea* L., semen (peanuts) is not a new active substance, as claimed by the applicant.

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

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

The CHMP noted the letter of recommendation dated 12.10.2020.

The summary of opinion was circulated for information.

3.1.9. Rekambys - rilpivirine - EMEA/H/C/005060

Janssen-Cilag International N.V.; treatment of Human Immunodeficiency Virus type 1 (HIV-1).

Scope: Opinion

Action: For adoption

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

List of Outstanding Issues adopted on 23.07.2020. List of Questions adopted on 12.12.2019.

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.

3.1.10. Vocabria - cabotegravir - EMEA/H/C/004976

ViiV Healthcare B.V.; treatment of Human Immunodeficiency Virus type 1 (HIV-1).

Scope: Opinion

Action: For adoption

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

List of Outstanding Issues adopted on 23.07.2020. List of Questions adopted on 12.12.2019.

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 cabotegravir 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.

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

3.2.1. abiraterone acetate - EMEA/H/C/005408

treatment of metastatic prostate cancer.

Scope: List of outstanding issues

Action: For adoption

List of Questions adopted on 30.01.2020.

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. bevacizumab - EMEA/H/C/005286

Treatment of metastatic carcinoma of the colon or rectum, metastatic breast cancer and recurrence of platinum-sensitive epithelial ovarian, fallopian tube or primary peritoneal cancer;
First-line treatment of patients with unresectable advanced, metastatic or recurrent non-small cell lung cancer;
First-line treatment of patients with advanced and/or metastatic renal cell cancer.

Scope: List of outstanding issues

Action: For adoption

List of Questions adopted on 28.05.2020.

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.

3.2.3. remimazolam - EMEA/H/C/005246

indicated for procedural sedation.

Scope: List of outstanding issues

Action: For adoption

List of Questions adopted on 30.04.2020.

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. salmeterol xinafoate / fluticasone propionate - EMEA/H/C/005591

treatment of asthma.

Scope: List of outstanding issues

Action: For adoption

List of Questions adopted on 27.02.2020.

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. ioflupane (^{123}I) - EMEA/H/C/005135

indicated for detecting loss of functional dopaminergic neuron terminals in the striatum.

Scope: List of outstanding issues

Action: For adoption

List of Questions adopted on 14.11.2019.

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. risperidone - EMEA/H/C/005406

treatment of schizophrenia.

Scope: List of outstanding issues

Action: For adoption

List of Questions adopted on 28.05.2020.

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. hepatitis B surface antigen - EMEA/H/C/005063

prevention of hepatitis B virus infection.

Scope: List of outstanding issues

Action: For adoption

List of Questions adopted on 25.07.2019.

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. fedratinib - Orphan - EMEA/H/C/005026

Celgene Europe BV; treatment of primary myelofibrosis, post polycythaemia vera myelofibrosis or post essential thrombocythaemia myelofibrosis.

Scope: List of outstanding issues

Action: For adoption

List of Questions adopted on 28.05.2020.

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.

3.2.9. azathioprine - EMEA/H/C/005055

indicated for the prophylaxis of transplant rejection, used as an immunosuppressant antimetabolite, indicated in patients who are intolerant to glucocorticosteroids, and chronic inflammatory bowel disease (IBD) (Crohn’s disease or ulcerative colitis), relapsing multiple sclerosis, generalised myasthenia gravis.

Scope: List of outstanding issues

Action: For adoption

List of Questions adopted on 30.04.2020.

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

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

3.2.10. insulin aspart - EMEA/H/C/004965

treatment of diabetes mellitus.

Scope: List of outstanding issues

Action: For adoption

List of Questions adopted on 27.02.2020.

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.

3.2.11. lenalidomide - EMEA/H/C/005348

treatment of multiple myeloma.

Scope: List of outstanding issues

Action: For adoption

List of Questions adopted on 28.05.2020.

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. lenalidomide - EMEA/H/C/005734

treatment of multiple myeloma and Follicular lymphoma.

Scope: List of outstanding issues

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.

3.2.13. lenalidomide - EMEA/H/C/005729

treatment of multiple myeloma, myelodysplastic syndromes and follicular lymphoma.

Scope: List of outstanding issues

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.

3.2.14. moxetumomab pasudotox - Orphan - EMEA/H/C/005322

AstraZeneca AB; relapsed or refractory Hairy Cell Leukaemia (HCL) after receiving at least

two prior systemic therapies.

Scope: List of outstanding issues

Action: For adoption

List of Questions adopted on 30.04.2020.

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.

3.2.15. selinexor - Orphan - EMEA/H/C/005127

Karyopharm Europe GmbH; treatment of patients with Relapsed Refractory Multiple Myeloma (RRMM).

Scope: List of outstanding issues

Action: Oral explanation to be held on Tuesday, 13 October 2020 at 11:00

List of Outstanding Issues adopted on 30.01.2020, 19.09.2019. List of Questions adopted on 24.04.2019.

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

See 2.1.

The CHMP agreed that an oral explanation was not needed at this time.

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

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

3.2.16. ofatumumab - EMEA/H/C/005410

treatment of relapsing forms of multiple sclerosis.

Scope: List of outstanding issues

Action: For adoption

List of Questions adopted on 28.05.2020.

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

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

3.2.17. bevacizumab - EMEA/H/C/005556

Treatment of metastatic carcinoma of the colon or rectum, metastatic breast cancer and recurrence of platinum-sensitive epithelial ovarian, fallopian tube or primary peritoneal cancer;
First-line treatment of patients with unresectable advanced, metastatic or recurrent non-small cell lung cancer;
First line treatment of patients with advanced and/or metastatic renal cell cancer.

Scope: List of outstanding issues

Action: For adoption

List of Questions adopted on 28.05.2020.

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

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

3.2.18. pertuzumab / trastuzumab - EMEA/H/C/005386

treatment of early breast cancer, metastatic breast cancer.

Scope: List of outstanding issues

Action: For adoption

List of Questions adopted on 28.05.2020.

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

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

3.2.19. fostemsavir - EMEA/H/C/005011

indicated, in combination with other antiretrovirals, for the treatment of adults with multidrug resistant HIV-1 infection for whom it is otherwise not possible to construct a suppressive anti-viral regimen due to resistance, intolerance or safety considerations.

Scope: List of outstanding issues

Action: For adoption

List of Questions adopted on 28.04.2020.

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.

3.2.20. salmeterol xinafoate / fluticasone propionate - EMEA/H/C/004881

treatment of asthma.

Scope: List of outstanding issues

Action: For adoption

List of Questions adopted on 27.02.2020.

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.

3.2.21. selpercatinib - EMEA/H/C/005375

indicated for the treatment of adults with advanced RET fusion-positive Non-Small Cell Lung Cancer (NSCLC) who require systemic therapy; advanced RET fusion-positive thyroid cancer who require systemic therapy and who have progressed following prior treatment. As monotherapy is indicated for the treatment of adults and adolescents 12 years and older with advanced RET-mutant Medullary Thyroid Cancer (MTC) who require systemic therapy.

Scope: List of outstanding issues

Action: For adoption

List of Questions adopted on 28.05.2020.

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.

3.2.22. potassium - Orphan - EMEA/H/C/005407

Advicenne S.A.; treatment of distal Renal Tubular Acidosis (dRTA) in patients aged 6 months and older.

Scope: List of outstanding issues

Action: For adoption

List of Questions adopted on 26.03.2020.

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.

3.2.23. sunitinib - EMEA/H/C/005419

treatment of Gastrointestinal Stromal Tumour (GIST) and Metastatic Renal Cell Carcinoma (MRCC) and pancreatic Neuro-Endocrine Tumours (pNET).

Scope: List of outstanding issues

Action: For adoption

List of Questions adopted on 27.02.2020.

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.

3.2.24. tucatinib - EMEA/H/C/005263

treatment of metastatic breast cancer or brain metastases.

Scope: List of outstanding issues

Action: For adoption

List of Questions adopted on 28.05.2020.

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

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

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

3.3.1. zanubrutinib - Orphan - EMEA/H/C/004978

BeiGene Ireland Ltd; treatment of Waldenström's Macroglobulinaemia (WM).

Scope: List of questions

Action: For adoption

The Committee discussed the issues identified in this application.

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

3.3.2. dexamethasone phosphate - EMEA/H/C/005740

indicated for cerebral oedema, post-traumatic shock-lung syndrome, asthma, skin diseases, autoimmune diseases, rheumatoid arthritis, prophylaxis and treatment of post-operative or cytostatic-induced vomiting, treatment of COVID-19, eye inflammation and infection.

Scope: 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.

3.3.3. eflornithine / sulindac - Orphan - EMEA/H/C/005043

Cancer Prevention Pharma (Ireland) Limited; treatment of adults patients with Familial Adenomatous Polyposis (FAP).

Scope: 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. setmelanotide - Orphan - EMEA/H/C/005089

TMC Pharma (EU) Limited; treatment of obesity and the control of hunger associated with deficiencies in the leptin-melanocortin pathway.

Scope: 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.

3.3.5. vericiguat - EMEA/H/C/005319

treatment of symptomatic chronic heart failure.

Scope: List of questions

Action: For adoption

The Committee discussed the issues identified in this application.

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

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

3.4.1. abiraterone acetate - EMEA/H/C/005368

treatment of metastatic castration resistant prostate cancer.

Scope: Letter from the applicant requesting an extension of clock-stop to respond to the list of questions adopted in July 2020 – adopted via written procedure on 08 October 2020.

Action: For information

List of Questions adopted on 23.07.2020.

The CHMP noted the agreement of the request by the applicant for an extension of clock-stop to respond to the list of questions adopted in July 2020, which was adopted via written procedure.

3.4.2. duvelisib - Orphan - EMEA/H/C/005381

Verastem Europe GmbH; treatment of adult patients with relapsed or refractory Chronic Lymphocytic Leukaemia (CLL) or Small Lymphocytic Lymphoma (SLL) and relapsed or refractory Follicular Lymphoma (FL).

Scope: Letter from the applicant dated 28 September 2020 requesting an extension of clock-stop to respond to the list of outstanding issues adopted in September 2020.

Action: For adoption

List of Outstanding Issues adopted 17.09.2020. List of Questions adopted on 30.04.2020.

The CHMP agreed to the request by the applicant for an extension of clock-stop to respond to the list of outstanding issues adopted in September 2020.

3.4.3. dasatinib - EMEA/H/C/005446

treatment of leukaemia.

Scope: Letter from the applicant dated 07 October 2020 requesting an extension of clock-stop to respond to the list of outstanding issues adopted in June 2020.

Action: For adoption

List of Outstanding Issues adopted on 25.06.2020. List of Questions adopted on 17.10.2019.

The CHMP agreed to the request by the applicant for an extension of clock-stop to respond to the list of outstanding issues adopted in June 2020.

3.4.4. dasatinib - EMEA/H/C/005317

treatment of leukaemia.

Scope: Letter from the applicant dated 07 October 2020 requesting an extension of clock-stop to respond to the list of outstanding issues adopted in June 2020.

Action: For adoption

List of Outstanding Issues adopted on 25.06.2020. List of Questions adopted on 17.10.2019.

The CHMP agreed to the request by the applicant for an extension of clock-stop to respond to the list of outstanding issues adopted in June 2020.

3.4.5. lonafarnib - Orphan - EMEA/H/C/005271

EigerBio Europe Limited; treatment of Hutchinson-Gilford progeria syndrome and progeroid laminopathies.

Scope: Letter from the applicant dated 24 September 2020 requesting an extension of clock-stop to respond to the list of questions adopted in July 2020 – adopted via written procedure on 29.09.2020.

Action: For information

List of questions adopted on 23.07.2020.

The CHMP noted the agreement of the request by the applicant for an extension of clock-stop to respond to the list of questions adopted in July 2020, which was adopted via written procedure.

3.4.6. somapacitan - Orphan - EMEA/H/C/005030

Novo Nordisk A/S; indicated for the replacement of endogenous GH with growth hormone deficiency (AGHD).

Scope: List of experts for the ad-hoc expert meeting scheduled on 29 October 2020

Action: For adoption

List of Outstanding Issues adopted on 17.09.2020, 25.06.2020. List of Questions adopted

on 30.01.2020.

The CHMP adopted the list of experts for the ad-hoc expert meeting scheduled on 29 October 2020.

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

3.5.1. Gamifant - emapalumab - Orphan - EMEA/H/C/004386

Swedish Orphan Biovitrum AB; treatment of paediatric patients with primary Haemophagocytic Lymphohistiocytosis (HLH).

Scope: Draft list of experts for the ad-hoc expert meeting scheduled on 30 October 2020

Action: For adoption

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

Opinion adopted on 23.07.2020. List of Outstanding Issues adopted on 25.06.2020, 27.06.2019. List of Questions adopted on 13.12.2018.

The CHMP adopted the list of experts for the ad-hoc expert meeting scheduled on 30 October 2020.

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

3.7.1. Tibsovo - ivosidenib - Orphan - EMEA/H/C/005056

Agios Netherlands B.V.; treatment of adult patients (≥ 18 years old) with relapsed or refractory acute myeloid leukaemia (AML) with an isocitrate dehydrogenase-1 (IDH1) R132 mutation.

Scope: Withdrawal of initial marketing authorisation application

Action: For information

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

List of Outstanding Issues adopted on 25.06.2020, 27.02.2020. List of Questions adopted on 29.05.2019.

The CHMP noted the withdrawal of initial marketing authorisation application.

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

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

4.1.1. Hulio - adalimumab - EMEA/H/C/004429/X/0016

Mylan S.A.S

Rapporteur: Christophe Focke, PRAC Rapporteur: Ulla Wändel Liminga

Scope: "Extension application to add a new strength of 20 mg solution for injection. The RMP (version 3.1) is updated in accordance."

Action: For adoption

List of Questions adopted on 23.07.2020.

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. Plegridy - peginterferon beta-1a - EMEA/H/C/002827/X/0056

Biogen Netherlands B.V.

Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Ulla Wändel Liminga

Scope: "Extension application to introduce a new route of administration (intramuscular use) for the 125 µg solution for injection."

Action: For adoption

List of Outstanding Issues adopted on 17.09.2020. List of Questions adopted on 25.06.2020.

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 recommendations dated 08.10.2020.

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

4.2.1. Nuceiva - botulinum toxin type A - EMEA/H/C/004587/X/0005

Evolus Pharma Limited

Rapporteur: Peter Kiely, PRAC Rapporteur: Adam Przybylkowski

Scope: "Extension application to add a new strength of 50 U for botulinum toxin type A for powder for solution for injection in vial (glass), for intramuscular administration."

Action: For adoption

List of Questions adopted on 28.05.2020.

The Committee discussed the issues identified in this application, mainly relating to quality aspects and the RMP.

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

4.2.2. Pradaxa - dabigatran etexilate - EMEA/H/C/000829/X/0122/G

Boehringer Ingelheim International GmbH

Rapporteur: Kirstine Moll Harboe, Co-Rapporteur: Jean-Michel Race, PRAC Rapporteur: Anette Kirstine Stark

Scope: "Extension application to add two new pharmaceutical forms for Pradaxa (coated granules (20 mg, 30 mg, 40 mg, 50 mg, 110 mg, 150 mg) and powder and solvent for oral solution (6.25 mg/ml)), grouped with:
-A type II variation (C.I.6.a) - Extension of indication to include new indication for Pradaxa 75 mg, 110 mg, 150 mg capsules based on the paediatric trials 1160.106 and 1160.108. As a consequence, sections 4.1, 4.2, 4.3, 4.4, 4.5, 4.7, 4.8, 4.9, 5.1, 5.2 and 5.3 of the SmPC are updated. The package leaflet and labelling are updated in accordance. In addition, the marketing authorisation holder took the opportunity to update the list of local representatives in the package leaflet. The RMP version 37.0 has also been submitted.
-Type IB (B.I.b.1.c)
-Type IA (B.I.b.1.b)
-Type IB (B.I.b.1.d)
-Type IA (B.I.b.2.a)
-Type IA (B.I.b.1.d)
-Type IA (B.I.d.1.a.1)
-Type IA (B.II.d.1.a)
-Type IB (B.II.d.1.d)
-Type IA (B.II.d.2.a)
-Type IA (B.II.c.1.c)"

Action: For adoption

List of Outstanding Issues adopted on 23.07.2020. List of Questions adopted on 27.02.2020.

The Committee discussed the issues identified in this application, mainly concerning quality and clinical aspects.

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

4.2.3. Tivicay - dolutegravir - EMEA/H/C/002753/X/0058/G

ViiV Healthcare B.V.

Rapporteur: Filip Josephson, PRAC Rapporteur: Martin Huber

Scope: "Extension application to add a new pharmaceutical form associated with new strength (5 mg dispersible tablet). The new presentation is indicated for the treatment of HIV infected children from 4 weeks of life and weighing at least 3 kilograms.
- Type II variation (C.I.4) to update the currently approved product information, labelling and package leaflet for the existing film-coated tablets (10 mg, 25 mg and 50 mg) for children 6 years and older and weighing at least 15 kg. The application comprises PK, safety, and efficacy data from the Phase I/II study (P1093) and PK and safety data from relevant sub-studies nested within the phase II/III Study ODYSSEY (PENTA 20).
In addition, the applicant took the opportunity to amend section 4.1 of SmPC, the indication for the approved Tivicay film-coated tablets to clarify that children should be "aged at least 6 years" as the current approved indication is inclusive of those aged 6 years. The RMP (version 16) is updated in accordance."

Action: For adoption

List of Questions adopted on 28.05.2020.

The Committee discussed the issues identified in this application, mainly concerning clinical aspects.

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

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

4.3.1. Buvidal - buprenorphine - EMEA/H/C/004651/X/0008/G

Camurus AB

Rapporteur: Peter Kiely, PRAC Rapporteur: Tiphaine Vaillant

Scope: "Line extension to add a new strength of 160 mg for prolonged-release solution for injection pharmaceutical form. In addition the MAH took the opportunity to align the PI to the latest QRD template and to implement new text regarding the content of ethanol in accordance with the EMA document "Information for the package leaflet regarding ethanol used as an excipient in medicinal products for human use" (EMA/CHMP/43486/2018) in the package leaflet. Variations included:
A.4 -
A.5.b -"

Action: For adoption

The Committee discussed the issues identified in this application, mainly concerning clinical aspects, including pharmacokinetics.

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

4.3.2. Xeljanz - tofacitinib - EMEA/H/C/004214/X/0024/G

Pfizer Europe MA EEIG

Rapporteur: Armando Genazzani, PRAC Rapporteur: Liana Gross-Martirosyan

Scope: "Extension application to introduce a new pharmaceutical form (oral solution, 1 mg/ml) grouped with a type II variation (C.I.6.a) to add a new indication (treatment of active polyarticular course Juvenile Idiopathic Arthritis (pJIA) in patients 2 years of age and older). The RMP (version 12.1) is updated in accordance. Additionally, the marketing authorisation holder took the opportunity to align the product information with the latest QRD template."

Action: For adoption

The Committee discussed the issues identified in this application, mainly concerning quality issues and the clinical part.

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. Benlysta - belimumab - EMEA/H/C/002015/II/0080

GlaxoSmithKline (Ireland) Limited

Rapporteur: Kristina Dunder, PRAC Rapporteur: Ulla Wändel Liminga

Scope: "Extension of indication to include treatment of lupus nephritis for belimumab. As a consequence, sections 4.1, 4.2, 4.4, 4.8, 5.1 and 5.2 of the SmPC are updated. The package leaflet is updated in accordance. Version 38 of the RMP has also been submitted."

Action: For adoption

The Committee discussed the issues identified in this application, mainly concerning clinical efficacy aspects.

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

5.1.2. BiResp Spiromax - budesonide / formoterol - EMEA/H/C/003890/II/0033/G

Teva Pharma B.V.

Rapporteur: John Joseph Borg, Co-Rapporteur: Peter Kiely, PRAC Rapporteur: Hans Christian Siersted

Scope: "Type II variation – C.I.6.a. - Extension of indication to include adolescents (12 years and older) for the regular treatment of asthma, where use of a combination (inhaled corticosteroid and long-acting β2 adrenoceptor agonist) is appropriate: in patients not adequately controlled with inhaled corticosteroids and "as needed" inhaled short-acting β2 adrenoceptor agonists; or in patients already adequately controlled on both inhaled corticosteroids and long-acting β2 adrenoceptor agonists. The proposed extension to the indication is based upon data from the literature. As a consequence, sections 4.1, 4.2, 5.1 and 5.2 of the SmPC have been updated and the Package Leaflet has been updated accordingly. In addition, the MAH took the opportunity to make an administrative update to the Greek, Islandic, Irish and Maltese local representatives phone numbers in the Package Leaflet.
Furthermore, the PI is brought in line with the latest QRD template version 10.1.
An updated RMP version 3.0 was submitted as part of the application.
Type IB variation - C.I.z other – updates of sections 4.2, 5.1 and 5.2 of the SmPC to update the information on paediatric data and section 4.4 of the SmPC to remove the warning regarding the risk of growth retardation in children and the guidance on how to address this risk as agreed during the assessment of the duplicate Budesonide/Formoterol Teva Pharma B.V. (EMEA/H/C/004882), which was approved in Jan 2020."

Action: For adoption

The Committee discussed the issues identified in this application, mainly concerning clinical efficacy and safety aspects.

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

5.1.3. Blincyto - blinatumomab - Orphan - EMEA/H/C/003731/II/0030

Amgen Europe B.V.

Rapporteur: Alexandre Moreau, Co-Rapporteur: Armando Genazzani, PRAC Rapporteur: Eva Jirsová

Scope: "To modify the approved therapeutic indication to include the treatment of Philadelphia chromosome positive CD19 positive B-cell precursor acute lymphoblastic leukaemia (ALL) in adult and paediatric patients with relapsed or refractory ALL and adult patients in first or second complete remission with minimal residual disease (MRD) greater

than or equal to 0.1%. As a consequence, sections 4.1, 4.2, 4.4, 4.8, 5.1 and 5.2 of the SmPC and the PL are updated accordingly. The updated RMP version 10.0 has also been submitted."

Action: For adoption

Request for Supplementary Information adopted on 25.06.2020, 14.11.2019.

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

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

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

The summary of opinion was circulated for information.

The CHMP adopted the similarity assessment report.

5.1.4. DuoResp Spiromax - budesonide / formoterol - EMEA/H/C/002348/II/0033/G

Teva Pharma B.V.

Rapporteur: John Joseph Borg, Co-Rapporteur: Peter Kiely, PRAC Rapporteur: Hans Christian Siersted

Scope: "- Type II variation – C.I.6.a. - Extension of indication to include adolescents (12 years and older) for the regular treatment of asthma, where use of a combination (inhaled corticosteroid and long-acting β2 adrenoceptor agonist) is appropriate: in patients not adequately controlled with inhaled corticosteroids and "as needed" inhaled short-acting β2 adrenoceptor agonists; or in patients already adequately controlled on both inhaled corticosteroids and long-acting β2 adrenoceptor agonists. The proposed extension to the indication is based upon data from the literature. As a consequence, sections 4.1, 4.2, 5.1 and 5.2 of the SmPC have been updated and the package leaflet has been updated accordingly. In addition, the marketing authorisation holder took the opportunity to make an administrative update to the Greek, Islandic, Irish and Maltese local representatives phone numbers in the package leaflet. Furthermore, the product information is brought in line with the latest QRD template version 10.1.
An updated RMP version 3.0 was submitted as part of the application.
- Type IB variation - C.I.z other – updates of sections 4.2, 5.1 and 5.2 of the SmPC to update the information on paediatric data and section 4.4 of the SmPC to remove the warning regarding the risk of growth retardation in children and the guidance on how to address this risk as agreed during the assessment of the duplicate Budesonide/Formoterol Teva Pharma B.V. (EMEA/H/C/004882), which was approved in January 2020."

Action: For adoption

The Committee discussed the issues identified in this application, mainly concerning clinical efficacy and safety aspects.

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

5.1.5. Dupixent - dupilumab - EMEA/H/C/004390/II/0027

sanofi-aventis groupe

Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Kimmo Jaakkola

Scope: "Extension of indication to include atopic dermatitis patients from 6 years to 11 years. Consequently, the sections 4.1, 4.2, 4.8, 5.1 and 5.2 are updated. The package leaflet is updated accordingly."

Action: For adoption

Request for Supplementary Information adopted on 23.07.2020, 30.04.2020.

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

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

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

The summary of opinion was circulated for information.

5.1.6. Epidyolex - cannabidiol - Orphan - EMEA/H/C/004675/II/0005

GW Pharma (International) B.V.

Rapporteur: Kirstine Moll Harboe, Co-Rapporteur: Ondřej Slanař, PRAC Rapporteur: Ana Sofia Diniz Martins

Scope: "Extension of indication for use as adjunctive therapy of seizures associated with Tuberous Sclerosis Complex (TSC) for patients 1 year of age and older. As a consequence, sections 4.1, 4.2, 4.4, 4.5, 4.8, 5.1, 5.2 and 5.3 of the SmPC are updated. The package leaflet is updated accordingly. The updated RMP version 1.1 has been submitted.
The marketing authorisation holder also took the opportunity to correct typographic errors and implement editorial updates as well as implement the updated ethanol statement in compliance with the EC Guideline on Excipient." Request for 1 year of market protection for a new indication (Article 14(11) of Regulation (EC) 726/2004)

Action: For adoption

Request for Supplementary Information adopted on 23.07.2020.

The Committee discussed the issues identified in this application, mainly in relation to the clinical part and the request for 1 year of market protection.

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

5.1.7. Humira - adalimumab - EMEA/H/C/000481/II/0198

AbbVie Deutschland GmbH & Co. KG

Rapporteur: Kristina Dunder, PRAC Rapporteur: Ulla Wändel Liminga

Scope: "Extension of indication to include treatment of moderately to severely active

ulcerative colitis in paediatric patients for Humira. As a consequence, sections 4.1, 4.2, 5.1 and 5.2 of the SmPC for 40 mg/0.8 mL, 40 mg/0.4 mL and 80 mg/0.8 mL presentations are updated. Furthermore, sections 5.1 and 5.2 of the SmPC for 20 mg/0.2 mL are updated. The package leaflet is updated in accordance. Version 15.0 of the RMP has also been submitted."

Action: For adoption

Request for Supplementary Information adopted on 23.07.2020.

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

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

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

The summary of opinion was circulated for information.

5.1.8. Keytruda - pembrolizumab - EMEA/H/C/003820/II/0091

Merck Sharp & Dohme B.V.

Rapporteur: Armando Genazzani, Co-Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Menno van der Elst

Scope: "C.I.6 - Extension of indication to include first-line treatment of unresectable or Metastatic Microsatellite Instability-High (MSI H) or Mismatch Repair Deficient (dMMR) colorectal cancer in adults for Keytruda based on the results from KEYNOTE-177 (an international, randomised, open-label phase 3 trial of pembrolizumab versus chemotherapy in MSI-H or dMMR Stage IV Colorectal Carcinoma). As a consequence, sections 4.1, 4.2 and 5.1 of the SmPC are updated. The package leaflet is updated in accordance. In addition, a minor correction has been made in section 4.4, "Immune related endocrinopathies" subsection. Version 29.1 of the RMP has also been submitted."

Action: For adoption

The Committee discussed the issues identified in this application, mainly concerning clinical efficacy aspects.

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

5.1.9. Opdivo - nivolumab - EMEA/H/C/003985/II/0080

Bristol-Myers Squibb Pharma EEIG

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

Scope: "Extension of indication to include treatment of adult patients with unresectable advanced, recurrent or metastatic oesophageal squamous cell carcinoma (OSCC) after prior fluoropyrimidine- and platinum-based chemotherapy. As a consequence, sections 4.1, 4.2, 4.4, 4.8 and 5.1 of the SmPC are updated. The Package Leaflet is updated in accordance. Version 16.0 of the RMP has also been submitted."

Draft list of experts for the SAG Oncology meeting held on 07.10.2020 adopted via written procedure on 07 October 2020

SAG report

Possible oral explanation

Action: For adoption

Request for Supplementary Information adopted on 23.07.2020, 30.04.2020.

See 2.3

The CHMP noted the SAG report.

An oral explanation was held on Wednesday 14 October 2020. The presentation by the applicant focused on the clinical data in different subgroups in support of the application.

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

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

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

The summary of opinion was circulated for information.

5.1.10. Recarbrio - imipenem / cilastatin / relebactam - EMEA/H/C/004808/II/0001

Merck Sharp & Dohme B.V.

Rapporteur: Filip Josephson, PRAC Rapporteur: Adam Przybylkowski

Scope: "Extension of indication to include the treatment of hospital-acquired pneumonia (HAP) including ventilator-associated pneumonia (VAP), with or without concurrent bacteraemia in adults for Recarbrio; as a consequence, sections 4.1, 4.2, 4.4, 4.8, 5.1, 5.2 and 6.6 of the SmPC are updated. The Package Leaflet is updated in accordance. Furthermore, the Marketing authorisation holder (MAH) made editorial corrections and brought the PI in line with the latest QRD template version 10.1.
Version 2.0 of the RMP is approved with this variation. The variation leads to amendments to the Summary of Product Characteristics, Annex II and Package Leaflet and to the Risk Management Plan (RMP)."

Action: For adoption

Request for Supplementary Information adopted on 23.07.2020.

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

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

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

The summary of opinion was circulated for information.

5.1.11. Saxenda - liraglutide - EMEA/H/C/003780/II/0026

Novo Nordisk A/S

Rapporteur: Johann Lodewijk Hillege, Co-Rapporteur: Kirstine Moll Harboe, PRAC Rapporteur: Menno van der Elst

Scope: "Extension of indication to include treatment as an adjunct to a healthy nutrition and physical activity counselling for weight management in adolescent patients from the age of 12 years and above with body weight above 60 kg and obesity (BMI corresponding to ≥30 kg/m^2 for adults), based on study NN8022-4180 that evaluated the efficacy of liraglutide 3.0 mg in adolescents aged 12 to less than 18 years with obesity. As a consequence, sections 4.1, 4.2, 4.8, 5.1 and 5.2 of the SmPC are being updated and the package leaflet is updated in accordance. The application relates to paediatric studies submitted according to Article 46 of the paediatric regulation. The application included an updated RMP version 32.0."

Action: For adoption

Request for Supplementary Information adopted on 28.05.2020.

The Committee discussed the issues identified in this application, mainly in relation to clinical efficacy.

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

5.1.12. Siklos - hydroxycarbamide - EMEA/H/C/000689/II/0047

Addmedica S.A.S.

Rapporteur: Karin Janssen van Doorn

Scope: "Extension of indication to include treatment of severe chronic anemia (haemoglobin level < 6 g/dL or < 7 g/dL with poor clinical or functional tolerance) in adults, adolescents and children older than 2 years suffering from sickle cell syndrome for Siklos. As a consequence, sections 4.1, 4.2 and 5.1 of the SmPC are updated. The package leaflet is updated in accordance."

Action: For adoption

The Committee discussed the issues identified in this application, mainly concerning clinical efficacy aspects.

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

5.1.13. Spravato - esketamine - EMEA/H/C/004535/II/0001/G

Janssen-Cilag International N.V.

Rapporteur: Martina Weise, Co-Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Kirsti Villikka

Scope: "- C.I.6(a): Extension of indication to include a new indication for Spravato for the rapid reduction of depressive symptoms in adult patients with a moderate to severe depressive episode of MDD who have current suicidal ideation with intent.

As a consequence, sections 4.1, 4.2, 4.4, 4.8 and 5.1 the SmPC are updated. The RMP version 2.1 has also been submitted.
- B.II.e.5.a.2: Addition of a new pack size corresponding to 4 weeks of treatment in the new indication. The package leaflet and labelling are updated in accordance. In addition, the marketing authorisation holder took the opportunity to clarify the wording in Annex II.D."

Action: For adoption

Request for Supplementary Information adopted on 30.04.2020.

The Committee discussed the issues identified in this application, mainly concerning clinical efficacy and safety aspects.

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

5.1.14. Tremfya - guselkumab - EMEA/H/C/004271/II/0017

Janssen-Cilag International N.V.

Rapporteur: Agnes Gyurasics, Co-Rapporteur: Peter Kiely, PRAC Rapporteur: Brigitte Keller-Stanislawski

Scope: "Addition of a new indication for the treatment of active psoriatic arthritis in adult patients who have had an inadequate response or who have been intolerant to a prior Disease-Modifying Antirheumatic Drug (DMARD) therapy. Consequently sections 4.1, 4.2 4.5 , 4.8, 5.1 and 5.2 of the SmPC are updated. The package leaflet is updated accordingly. Additionally minor QRD changes are introduced in annex II."

Action: For adoption

Request for Supplementary Information adopted on 25.06.2020, 30.01.2020.

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

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

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

The summary of opinion was circulated for information.

5.1.15. Venclyxto - venetoclax - EMEA/H/C/004106/II/0030

AbbVie Deutschland GmbH & Co. KG

Rapporteur: Filip Josephson, Co-Rapporteur: Paula Boudewina van Hennik, PRAC Rapporteur: Eva Jirsová

Scope: "Extension of indication for Venclyxto (venetoclax) in combination with Hypomethylating Agents (HMAs) or Low Dose Cytarabine (LDAC) for the treatment of adult patients with newly-diagnosed acute myeloid leukaemia (AML) who are ineligible for intensive chemotherapy. As a consequence, sections 4.2, 4.3, 4.4, 4.5, 4.7, 4.8, 5.1 and 5.2 of the SmPC are updated. The package leaflet and RMP version 6.1 are also 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, mainly concerning the clinical efficacy.

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

5.1.16. WS1737
Edistride - dapagliflozin - EMEA/H/C/004161/WS1737/0034
Forxiga - dapagliflozin - EMEA/H/C/002322/WS1737/0053

AstraZeneca AB

Lead Rapporteur: Kristina Dunder, Lead Co-Rapporteur: Martina Weise, PRAC Rapporteur: Annika Folin

Scope: "Update of sections 4.1, 4.2, 4.4, 4.8 and 5.1 of the SmPC for Edistride and Forxiga to add a new indication for the treatment of symptomatic heart failure with reduced ejection fraction in adults. The Package Leaflet and Labelling are updated in accordance.
The RMP version 18 has also been submitted.
Furthermore, the PI is brought in line with the latest QRD template version 10.1, as well as editorial change (addition of SI unit for blood glucose).
The worksharing procedure requested amendments to the Summary of Product Characteristics, Labelling and Package Leaflet and to the Risk Management Plan (RMP)."
Request for 1 year of market protection for a new indication (Article 14(11) of Regulation (EC) 726/2004)

Action: For adoption

Request for Supplementary Information adopted on 17.09.2020, 25.06.2020, 27.02.2020.

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

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

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

The summary of opinion was circulated for information.

5.1.17. WS1782
Lacosamide UCB - lacosamide - EMEA/H/C/005243/WS1782/0006
Vimpat - lacosamide - EMEA/H/C/000863/WS1782/0088

UCB Pharma S.A.

Lead Rapporteur: Filip Josephson, Lead Co-Rapporteur: Armando Genazzani, PRAC Rapporteur: Ulla Wändel Liminga

Scope: "Extension of indication to include the treatment as adjunctive therapy of primary generalised tonic-clonic seizures in adults, adolescents and children from 4 years of age with idiopathic generalised epilepsy for Lacosamide UCB and Vimpat. Consequently sections 4.1, 4.2, 4.8 and 5.1 of the SmPC are updated. The Package Leaflet is updated in accordance. The RMP version 15.0 has also been submitted.

Furthermore, the PI is brought in line with the latest QRD template version 10.1. The MAH also takes the opportunity to align the PI of Lacosamide UCB with the PI of Vimpat."

Action: For adoption

Request for Supplementary Information adopted on 23.07.2020, 30.04.2020.

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

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

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

The summary of opinion was circulated for information.

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

No items

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. adrenalin - H0005584

adrenalin should be used for the emergency treatment of allergic reactions, including anaphylaxis.

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

Action: For adoption

The CHMP did not agree to the request for accelerated assessment and adopted the briefing note and Rapporteurs' recommendation on the Request for Accelerated Assessment.

8.1.2. finerenone - H0005200

finerenone is indicated for the treatment of patients with kidney disease and type 2 diabetes to reduce the risk of cardiovascular mortality and morbidity and to reduce the rate of progression of renal disease.

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

Action: For adoption

The CHMP did not agree to the request for accelerated assessment and adopted the briefing note and Rapporteurs' recommendation on the Request for Accelerated Assessment.

8.1.3. voxelotor - H0004869

treatment of haemolytic anaemia associated with sickle cell disease in adults and paediatric patient 12 years and older.

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

Action: For adoption

The CHMP did not agree 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 information.

8.2.2. Recommendation for PRIME eligibility

Action: For adoption

The CHMP adopted the recommendation for PRIME eligibility. The CHMP reviewed 8 recommendations for eligibility to PRIME: 4 were accepted and 3 were denied and 1 is currently under consideration.

The individual outcomes are listed in PRIME Monthly Report on EMA website.

9. Post-authorisation issues

9.1. Post-authorisation issues

9.1.1. Desloratadine ratiopharm - desloratadine - EMEA/H/C/002404/II/0023/G

ratiopharm GmbH

Rapporteur: Christophe Focke, PRAC Rapporteur: Laurence de Fays

Scope: "C.I.5.b - Change in the legal status of desloratadine ratiopharm from 'medicinal product subject to medical prescription' to 'medicinal product not subject to medical prescription' in view of the safety profile of desloratadine ratiopharm. As a consequence, sections 4.1, 4.2, 4.3, 4.4, 4.5, 4.6, 4.8 and 5.1 of the SmPC are updated. The annexe II, package leaflet and labelling are updated accordingly. Furthermore, the product information is brought in line with the latest QRD template (version 10.1) and the list of local representatives in the package leaflet is updated. The RMP is updated to version 1.2. As a consequence of the variation, the pack sizes of 40, 50, 60, 90 and 100 tables are deleted (EU/1/11/746/007-011);
C.I.6.b - To delete the therapeutic indication in adolescents aged 12 years and older for the relief of symptoms associated with allergic rhinitis and urticaria. Section 4.1 of the SmPC and section 1 of the package leaflet are updated accordingly."

Action: For adoption

Request for Supplementary Information adopted on 23.07.2020, 26.03.2020.

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. Forxiga/Edistride (dapagliflozin) – EMEA/H/C/002322 / EMEA/H/C/004161/WS1844

Astra Zeneca AB

Rapporteur: Kristina Dunder

Scope: MAH's intention to re-classify the imposed category 1 PASS in the T1DM indication

Action: For adoption

Request for Supplementary Information adopted on 23.07.2020.

The Committee discussed the issues identified in this application, concerning the provided clinical data.

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

9.1.3. Kanuma - sebelipase alfa - EMEA/H/C/004004/II/0026/G, Orphan

Alexion Europe SAS

Rapporteur: Karin Janssen van Doorn, PRAC Rapporteur: Ulla Wändel Liminga

Scope: "Grouping consisting of the following variations:
- Update of sections 4.2, 4.4, 4.8 and 5.1 of the Summary of Product Characteristics (SmPC) in order to update the clinical information based on the pooled safety and efficacy analysis of already submitted studies (Studies LAL-CL04, LAL-CL03, LAL-CL06, LAL-CL08 and LAL-CL02) and updated population PK analyses in children and adults. The Package Leaflet has been amended accordingly. The ATC code has been added to the SmPC. The RMP version 4 has also been submitted. Annex II is also updated to remove the obligation related to the provision of study LAL-CL08.
- Submission of the final report from study LAL-EA01 An open-label study with sebelipase alfa (1 mg/kg every other week for up to 78 weeks or until drug commercialization) in United States (US) patients who did not otherwise qualify for an active sebelipase alfa trial (expanded access protocol)"

Action: For adoption

Request for Supplementary Information adopted on 23.07.2020, 26.03.2020.

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.

9.1.4. Ondexxya - andexanet alfa - EMEA/H/C/004108/II/0011

Portola Netherlands B.V.

Rapporteur: Jan Mueller-Berghaus

Scope: "C.I.4 Update of sections 4.4 and 4.5 of the SmPC in order to add a new warning on

use of heparin after administration of andexanet based on spontaneous reports, medical literature, reports, clinical trials and in vitro data. The package leaflet is updated accordingly. Additional editorial changes were proposed in Annex I and IIIB."

DHPC and communication plan

Action: For adoption

Request for Supplementary Information adopted on 25.06.2020.

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.

9.1.5. Polivy - polatuzumab vedotin - EMEA/H/C/004870/R/0003, Orphan

Roche Registration GmbH

Rapporteur: Alexandre Moreau, Co-Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Annika Folin

Scope: Renewal of conditional marketing authorisations

Action: For adoption

Request for Supplementary Information adopted on 17.09.2020.

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.

9.1.6. Tecfidera - dimethyl fumarate - EMEA/H/C/002601/II/0063

Biogen Netherlands B.V.

Rapporteur: Martina Weise, PRAC Rapporteur: Martin Huber

Scope: "Update of sections 4.4 and 4.8 of the SmPC to reflect PML in the setting of mild lymphopenia based on data submitted in the ongoing PSUSA/00010143/201903. The package leaflet is updated accordingly. Additionally, the product information has been updated in line with QRD template (version 10.1)."

Action: For adoption

Request for Supplementary Information adopted on 17.09.2020, 28.05.2020, 30.01.2020, 19.09.2019.

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 agreed to the DHPC and communication plan.

9.1.7. WS1820 Iscover-EMEA/H/C/000175/WS1820/0142, Plavix-EMEA/H/C/000174/WS1820/0140

sanofi-aventis groupe

Lead Rapporteur: Bruno Sepodes, Lead PRAC Rapporteur: Marcia Sofia Sanches de Castro Lopes Silva

Scope: "Update of section 4.2 of the SmPC in order to add 600 mg as an alternative loading dose to the existing 300 mg to be used at initiation of treatment in the indication 'Secondary prevention of atherothrombotic events in adult patients suffering from acute coronary syndrome'. This update is based on a bibliographic review of published studies. The package leaflet is updated accordingly. The RMP version 2.0 has also been submitted."

Action: For discussion

Request for Supplementary Information adopted on 25.06.2020.

The Committee discussed the issues identified in this application, concerning non-clinical and clinical aspects.

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

9.1.8. NexoBrid - concentrate of proteolytic enzymes enriched in bromelain - EMEA/H/C/002246/II/0049, Orphan

MediWound Germany GmbH,

Rapporteur: Janet Koenig

Scope: "Update of sections 4.2, 4.4, 4.8, 4.9, 5.1 and 5.2 of the SmPC based on interim results from study MW2010-03-02 (DETECT) listed as an obligation in the Annex II; this is a multicenter, multinational, randomized, controlled, assessor blinded, three-arm study performed in subjects with thermal burns, to evaluate the efficacy and safety of NexoBrid compared to Gel Vehicle and compared to Standard of Care. The submitted Clinical Study Report includes data of Acute Phase and 12M Follow Up Period. The MAH also took the opportunity to submit integrated pooled analyses of efficacy and safety data obtained from phase 2 and phase 3 clinical studies to provide a comprehensive assessment of the safety and efficacy of NexoBrid. The Package Leaflet is updated accordingly."

Action: For adoption

The Committee discussed the issues identified in this application, mainly relating to clinical

aspects.

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

9.1.9. Nitrosamine testing of metformin medicines

Action: For discussion

The CHMP agreed that all marketing authorisation holders of metformin-containing medicines are contacted to request to implement interim measure of testing before release of the medicines for the presence of nitrosamines before they are released onto the market. This is a precautionary step to ensure patient safety while ongoing investigations on these medicines are being finalised. The request is in line with this year's Article 5(3) review (EMEA/H/A-5(3)/1490).

10. Referral procedures

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

No items

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

No items

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

10.6.1. Angiotensin-II-receptor antagonists (sartans) containing a tetrazole group - EMEA/H/A-31/1471

MAHs: various

Overall Rapporteur: Martina Weise, Co-Rapporteurs:
Candesartan: Kristina Dunder, Irbesartan: Concepcion Prieto Yerro, Losartan: Johann Lodewijk Hillege, Olmesartan: Milena Stain, Valsartan: Armando Genazzani

Scope: Impact of the Article 5(3) referral on nitrosamines in human medicinal products on the referral under article 31 of Directive 2001/83/EC for sartans medicinal products containing a tetrazole ring (opinion adopted in January 2019), following a request from the European Commission, draft timetable for adoption via written procedure

Action: For information

The CHMP noted the draft timetable adopted via written procedure.

Request from EC for a CHMP Opinion: 29.07.2020

(Joint) Assessment Report circulated to CHMP: 24.10.2020

Comments: 30.10.2020

Updated (joint) assessment report circulated to CHMP: 05.11.2020

CHMP opinion: November 2020 CHMP

10.6.2. Esmya (CAP); NAP - Ulipristal acetate - EMEA/H/A-31/1496

MAH(s): Gedeon Richter Plc.; various

Referral PRAC Rapporteur: Annika Folin, Referral PRAC Co-Rapporteur: Menno van der Elst, CHMP Rapporteurs (Esmya): Kristina Dunder, CHMP Co-Rapporteur (Esmya): Paula Boudewina van Hennik

Scope: CHMP Opinion

Action: For adoption

Review of the benefit-risk balance following notification by the European Commission of a referral under Article 31 of Directive 2001/83/EC, based on pharmacovigilance data; PRAC recommendation.

The CHMP noted the PRAC recommendation to revoke the marketing authorisation for Esmya.

The CHMP discussed the available data and considered that the benefit/risk profile may be favourable in a restricted indication.

The CHMP adopted a list of questions to the MAHs with a specific timetable.

List of questions adopted by CHMP: 15.10.2020

Responses to list of questions by MAH(s): 22.10.2020

Joint Assessment Report from CHMP rapporteurs: 29.10.2020

Comments from CHMP: 05.11.2020

Oral Explanation and/or CHMP opinion: November 2020 CHMP

10.6.3. Gadolinium-containing contrast agents (GdCA): gadobenic acid (NAP), gadoteridol (NAP) - EMEA/H/A-31/1097

Applicant: Bracco Imaging SpA

Lead Rapporteur: Johann Lodewijk Hillege (NL)

Scope: Annual cumulative reviews on NSF cases submission as a post-authorisation measure resulting from the 2010 Article 31 referral procedure for gadolinium-containing contrast agents.

Action: For adoption

The CHMP adopted the annual cumulative reviews with the conclusion that the post authorisation measure is fulfilled and no further NSF cases need to be submitted.

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

No items

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

No items

10.9. Disagreement between Member States on Type II variation– Arbitration procedure initiated by MAH under Article 6(13) 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

October 2020 Early Notification System on envisaged CHMP/CMDh outcome accompanied by communication to the general public.

Action: For information

The CHMP noted the information.

12. Inspections

12.1. GMP inspections

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

12.2. GCP inspections

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

12.3. Pharmacovigilance inspections

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

12.4. GLP inspections

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

No items

13.2. Innovation Task Force briefing meetings

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

14.1.1. Update on CHMP Co-opted Members

Call for nomination of CHMP co-opted member in light of the end of the mandate of Jan Mueller-Berghaus as co-opted member on 13 November 2020.

Agreed areas of expertise: Quality, safety and efficacy of biological medicinal products, including advanced therapies and with specific emphasis on vaccines.

Nominations should be sent.

Discussion of area of expertise in light of Koenraad Norga's resignation as CHMP co-opted member as of 30 September 2020.

Proposals should be sent.

Action: For information

The CHMP noted the information.

14.2. Coordination with EMA Scientific Committees

14.2.1. Pharmacovigilance Risk Assessment Committee (PRAC)

Summary of recommendations and advice of PRAC meeting held on 27-30 September 2020

Action: For information

The CHMP noted the Summary of recommendations and advice.

List of Union Reference Dates and frequency of submission of Periodic Safety Update Reports (EURD list) for October 2020

Action: For adoption

The CHMP adopted the EURD list.

14.2.2. Paediatric Committee (PDCO)

PIPs reaching D30 at October 2020 PDCO

Action: For information

The CHMP noted the information.

Report from the PDCO meeting held on 12-15 October 2020

Action: For information

The CHMP noted the information.

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

14.3.1. Biologics Working Party (BWP)

Chair: Sol Ruiz/Nanna Aaby Kruse

Reports from BWP October 2020 meeting to CHMP for adoption:

- 27 reports on products in scientific advice and protocol assistance
- 9 reports on products in pre-authorisation procedures
- 2 reports on products in plasma master file

Action: For adoption

The CHMP adopted the BWP reports.

14.3.2. Name Review Group (NRG)

Table of Decisions of the NRG meeting held on 15-16 September 2020

Action: For adoption

The CHMP adopted the Table of Decisions.

14.3.3. Scientific Advice Working Party (SAWP)

Chair: Anja Schiel

Report from the SAWP meeting held on 28 September - 1 October 2020. Table of conclusions

Action: For information

The CHMP noted the report.

Scientific advice letters:

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

14.3.4. Safety Working Party (SWP)

Chair(s): Jan Willem Van der Laan/Susanne Brendler-Schwaab

SWP position on the issue related to the MeNP limit in Rifampcin

Action: For adoption

The CHMP adopted the SWP position.

CMDh question to SWP - "Diethanolamine" and "coconut oil diethanolamine condensate" excipients

Action: For adoption

The CHMP endorsed the questions to SWP.

14.4. Cooperation within the EU regulatory network

No items

14.5. Cooperation with International Regulators

No items

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

No items

14.7. CHMP work plan

No items

14.8. Planning and reporting

No items

14.9. Others

No items

15. Any other business

15.1. AOB topic

15.1.1. Update on COVID-19

Action: For information

The CHMP noted the update.

15.1.2. Update on Brexit

Action: For information

The members noted the update.

15.1.3. Clarification on authorised indication of melanoma products

CHMP letter to third party on clarification regarding the authorised indication of melanoma products

Action: For discussion

The CHMP adopted the response letter.

15.1.4. Covid-19 mRNA vaccine (nucleoside-modified) - H0005735

Active immunisation against COVID-19 disease.

Scope: Start of rolling review, draft timetable

Action: For adoption

The CHMP adopted the timetable.

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 12-15 October 2020 CHMP meeting.

Name	Role	Member State or affiliation	Outcome restriction following evaluation of e-DoI	Topics on agenda for which restrictions apply
Harald Enzmann	Chair	Germany	No interests declared	
Andrea Laslop	Member	Austria	No interests declared	
Milena Stain	Alternate	Austria	No interests declared	
Christophe Focke	Member	Belgium	No restrictions applicable to this meeting	
Karin Janssen van Doorn	Alternate	Belgium	No interests declared	
Margareta Bego	Member	Croatia	No interests declared	
Selma Arapovic Dzakula	Alternate	Croatia	No interests declared	
Emilia Mavrokordatou	Alternate	Cyprus	No participation in final deliberations and voting on	covid-19 mRNA vaccine (nucleoside-modified)
Ondřej Slanař	Member	Czech Republic	No interests declared	
Tomas Radimersky	Alternate	Czech Republic	No interests declared	
Sinan B. Sarac	Member	Denmark	No interests declared	
Kirstine Moll Harboe	Alternate	Denmark	No interests declared	
Alar Irs	Member	Estonia	No restrictions applicable to this meeting	
Edward Laane	Alternate	Estonia	No restrictions applicable to this meeting	
Outi Mäki-Ikola	Member	Finland	No restrictions applicable to this meeting	
Alexandre Moreau	Member	France	No interests declared	
Jean-Michel Race	Alternate	France	No interests declared	
Martina Weise	Member	Germany	No restrictions applicable to this meeting	
Janet Koenig	Alternate	Germany	No interests declared	

Name	Role	Member State or affiliation	Outcome restriction following evaluation of e-DoI	Topics on agenda for which restrictions apply
Konstantinos Markopoulos	Member	Greece	No interests declared	
Eleftheria Nikolaidi	Alternate	Greece	No interests declared	
Agnes Gyurasics	Alternate	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	
Armando Genazzani	Member	Italy	No interests declared	
Elita Poplavska	Member	Latvia	No interests declared	
Simona Stankeviciute	Alternate	Lithuania	No interests declared	
Martine Trauffler	Member	Luxembourg	No interests declared	
John Joseph Borg	Member	Malta	No interests declared	
Johann Lodewijk Hillege	Member	Netherlands	No interests declared	
Paula Boudewina van Hennik	Alternate	Netherlands	No interests declared	
Bjorg Bolstad	Member	Norway	No restrictions applicable to this meeting	
Ingrid Wang	Alternate	Norway	No interests declared	
Ewa Balkowiec Iskra	Member	Poland	No interests declared	
Bruno Sepodes	Member (Vice-Chair)	Portugal	No interests declared	
Fatima Ventura	Alternate	Portugal	No restrictions applicable to this meeting	
Simona Badoi	Member	Romania	No interests declared	
Dana Gabriela Marin	Alternate	Romania	No interests declared	
Francisek Drafi	Member	Slovakia	No interests declared	
Dorota Distlerova	Alternate	Slovakia	No participation in discussions, final deliberations and voting on	Plegridy - EMEA/H/C/002827/X/0056 Tecfidera - EMEA/H/C/002601/II/0063
Nevenka Trsinar Brodt	Alternate	Slovenia	No interests declared	
Maria Concepcion Prieto Yerro	Member	Spain	No interests declared	

Name	Role	Member State or affiliation	Outcome restriction following evaluation of e-DoI	Topics on agenda for which restrictions apply
Blanca Garcia-Ochoa	Alternate	Spain	No interests declared	
Kristina Dunder	Member	Sweden	No interests declared	
Filip Josephson	Alternate	Sweden	No interests declared	
Christian Gartner	Co-opted member	Austria	No restrictions applicable to this meeting	
Jan Mueller-Berghaus	Co-opted member	Germany	No interests declared	
Blanka Hirschlerova	Co-opted member	Czech Republic	No interests declared	
Sol Ruiz	Co-opted member	Spain	No interests declared	
Eskild Colding-Jorgensen	Expert - via telephone*	Denmark	No restrictions applicable to this meeting	
Menno van der Elst	Expert - via telephone*	Netherlands	No interests declared	
Eduard Dolusic	Expert - via Adobe*	Belgium	No interests declared	
Edwige Haelterman	Expert - via Adobe*	Belgium	No interests declared	
Flora Musuamba	Expert - via Adobe*	Belgium	No interests declared	
Helene Van de Ven	Expert - via Adobe*	Belgium	No interests declared	
Miranda Vroenhove	Expert - via Adobe*	Belgium	No interests declared	
Roel van Loock	Expert - via Adobe*	Belgium	No interests declared	
Tim Leest	Expert - via Adobe*	Belgium	No interests declared	
Violette Dirix	Expert - via Adobe*	Belgium	No interests declared	
Eva Jirsova	Expert - via Adobe*	Czech Republic	No interests declared	
Aaron Sosa	Expert - via Adobe*	Denmark	No restrictions applicable to this meeting	
Ebru Gulsun Karakoc Madsen	Expert - via Adobe*	Denmark	No restrictions applicable to this meeting	
Mette Linnert Jensen	Expert - via Adobe*	Denmark	No interests declared	
Mogens Westergaard	Expert - via Adobe*	Denmark	No interests declared	
Nanna Borup Johansen	Expert - via Adobe*	Denmark	No interests declared	
Ole Weis Bjerrum	Expert - via Adobe*	Denmark	No restrictions applicable to this meeting	

Name	Role	Member State or affiliation	Outcome restriction following evaluation of e-DoI	Topics on agenda for which restrictions apply
Elina Asikanius	Expert - via Adobe*	Finland	No restrictions applicable to this meeting	
Johanna Lahteenvuo	Expert - via Adobe*	Finland	No interests declared	
Karri Penttila	Expert - via Adobe*	Finland	No interests declared	
Claudia Reichmann	Expert - via Adobe*	Germany	No interests declared	
Elmer Schabel	Expert - via Adobe*	Germany	No interests declared	
Gregorios Aislaitner	Expert - via Adobe*	Germany	No interests declared	
Heiko Preusser	Expert - via Adobe*	Germany	No interests declared	
Hilke Zander	Expert - via Adobe*	Germany	No interests declared	
Marion Haberkamp	Expert - via Adobe*	Germany	No interests declared	
Miriam Fuerst-Wilmes	Expert - via Adobe*	Germany	No interests declared	
Agnieszka Przybyszewska	Expert - via Adobe*	Ireland	No interests declared	
Caoimhin Concannon	Expert - via Adobe*	Ireland	No interests declared	
Catherine Byrne	Expert - via Adobe*	Ireland	No interests declared	
Cathy Byrne	Expert - via Adobe*	Ireland	No interests declared	
Elma O`Reilly	Expert - via Adobe*	Ireland	No interests declared	
Larissa Higgins	Expert - via Adobe*	Ireland	No interests declared	
Mair Powell	Expert - via Adobe*	Ireland	No interests declared	
Antonella Isgro	Expert - via Adobe*	Italy	No interests declared	
Cristina Migali	Expert - via Adobe*	Italy	No interests declared	
Danila Renzo	Expert - via Adobe*	Italy	No interests declared	
Sara Galluzzo	Expert - via Adobe*	Italy	No interests declared	
Svetlana Lorenzano	Expert - via Adobe*	Italy	No restrictions applicable to this meeting	
Valeria Zoccano	Expert - via Adobe*	Italy	No interests declared	
Yanica Cassar	Expert - via Adobe*	Malta	No interests declared	
Adrianus van Gompel	Expert - via Adobe*	Netherlands	No interests declared	
Frank Holtkamp	Expert - via Adobe*	Netherlands	No interests declared	

Name	Role	Member State or affiliation	Outcome restriction following evaluation of e-DoI	Topics on agenda for which restrictions apply
Hinke Johanna van der Woude	Expert - via Adobe*	Netherlands	No interests declared	
Illiana Meurs	Expert - via Adobe*	Netherlands	No interests declared	
Jorn Mulder	Expert - via Adobe*	Netherlands	No interests declared	
Loes den Otter	Expert - via Adobe*	Netherlands	No interests declared	
Martijn van Gils	Expert - via Adobe*	Netherlands	No interests declared	
Quirine Fillekes	Expert - via Adobe*	Netherlands	No interests declared	
Richard Ijzerman	Expert - via Adobe*	Netherlands	no part in discussions, final deliberations and voting as appropriate	WS1737: Edistride - dapagliflozin - EMEA/H/C/004161/WS1737/0034 Forxiga - dapagliflozin - EMEA/H/C/002322/WS1737/0053
Taina Mattila	Expert - via Adobe*	Netherlands	No interests declared	
Anja Schiel	Expert - via Adobe*	Norway	No interests declared	
Jens-Uwe Bleich	Expert - via Adobe*	Norway	No interests declared	
Veronika Mihalcova	Expert - via Adobe*	Slovakia	No interests declared	
Jonas Bergh	Expert - via Adobe*	Sweden	No restrictions applicable to this meeting	
Meeting run with the help of EMA staff				

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

17. Explanatory notes

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

Oral explanations *(section 2)*

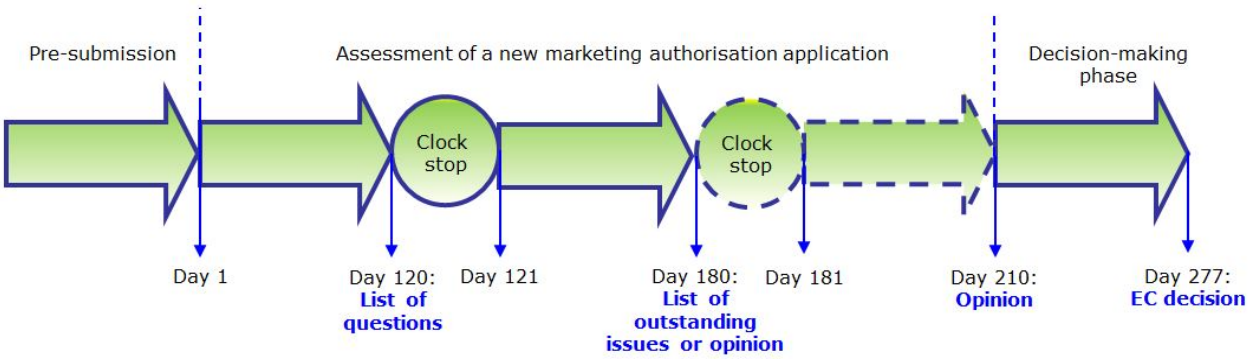
The items listed in this section are those for which marketing authorisation holders (MAHs) or applicants have been invited to the CHMP plenary meeting to address questions raised by the Committee. Oral explanations normally relate to on-going applications (section 3, 4 and 5) or referral procedures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

Initial applications *(section 3)*

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

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

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



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

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

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

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

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

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

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

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

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

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

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

Re-examination procedures *(section5.3)*

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

Withdrawal of application *(section 3.7)*

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

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

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

Pre-submission issues *(section 8)*

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

Post-authorisation issues *(section 9)*

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

Referral procedures *(section 10)*

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

Pharmacovigilance issues *(section 11)*

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

Inspections Issues *(section 12)*

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

Innovation task force *(section 13)*

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

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

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

Satellite groups / other committees *(section 14.2)*

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

Invented name issues *(section 14.3)*

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

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

EUROPEAN MEDICINES AGENCY
SCIENCE MEDICINES HEALTH

05 January 2021
EMA/CHMP/708237/2020

Annex to 12-15 October 2020 CHMP Minutes

Pre-submission and post-authorisations issues

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A. PRE-SUBMISSION ISSUES

A.1. ELIGIBILITY REQUESTS

Report on Eligibility to Centralised Procedure for October 2020: **For adoption**	Adopted.

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

Final Outcome of Rapporteurship allocation for October 2020: **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

Qarziba - dinutuximab beta - EMEA/H/C/003918/S/0022, Orphan EUSA Pharma (Netherlands) B.V., Rapporteur: Paula Boudewina van Hennik, PRAC Rapporteur: Brigitte Keller-Stanislawski	Positive Opinion adopted by consensus together with the CHMP assessment report. The Marketing Authorisation remains under exceptional circumstances. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP opinion.

B.2. RENEWALS OF MARKETING AUTHORISATIONS OUTCOMES

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

Neofordex - dexamethasone - EMEA/H/C/004071/R/0016 Laboratoires CTRS, Rapporteur: Ondřej Slanař, Co-Rapporteur: Armando Genazzani, PRAC Rapporteur: Tiphaine Vaillant	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

Alprolix - eftrenonacog alfa - EMEA/H/C/004142/R/0032, Orphan Swedish Orphan Biovitrum AB (publ), Rapporteur: Andrea Laslop, Co-Rapporteur: Maria Concepcion Prieto Yerro, PRAC Rapporteur: Brigitte Keller-Stanislawski Request for Supplementary Information adopted on 15.10.2020.	Request for supplementary information adopted with a specific timetable.
Coagadex - human coagulation factor X - EMEA/H/C/003855/R/0031, Orphan BPL Bioproducts Laboratory GmbH, Rapporteur: Andrea Laslop, Co-Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Menno van der Elst Request for Supplementary Information adopted on 15.10.2020.	Request for supplementary information adopted with a specific timetable.
Descovy - emtricitabine / tenofovir alafenamide - EMEA/H/C/004094/R/0051 Gilead Sciences Ireland UC, Rapporteur: Bruno Sepodes, Co-Rapporteur: Jean-Michel Race, PRAC Rapporteur: Ana Sofia Diniz Martins Request for Supplementary Information adopted on 15.10.2020.	Request for supplementary information adopted with a specific timetable.
Empliciti - elotuzumab - EMEA/H/C/003967/R/0024 Bristol-Myers Squibb Pharma EEIG, Rapporteur: Paula Boudewina van Hennik, Co-Rapporteur: Armando Genazzani, PRAC Rapporteur: Brigitte Keller-Stanislawski	Positive Opinion adopted by consensus together with the CHMP assessment report and translation timetable. Based on the review of the available information, the CHMP was of the opinion that the renewal of the marketing authorisation can be granted with unlimited validity. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP Opinion.
Jevtana - cabazitaxel - EMEA/H/C/002018/R/0042 sanofi-aventis groupe, Rapporteur: Alexandre Moreau, Co-Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Tiphaine Vaillant Request for Supplementary Information adopted on 23.07.2020.	Positive Opinion adopted by consensus together with the CHMP assessment report and translation timetable. Based on the review of the available information, the CHMP was of the opinion that the renewal of the marketing authorisation can be granted with unlimited validity. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP Opinion.
Lonsurf - trifluridine / tipiracil -	Positive Opinion adopted by consensus together

EMEA/H/C/003897/R/0020 Les Laboratoires Servier, Rapporteur: Paula Boudewina van Hennik, Co-Rapporteur: Blanca Garcia-Ochoa, PRAC Rapporteur: Annika Folin	with the CHMP assessment report and translation timetable. Based on the review of the available information, the CHMP was of the opinion that the renewal of the marketing authorisation can be granted with unlimited validity. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP Opinion.
Taltz - ixekizumab - EMEA/H/C/003943/R/0039 Eli Lilly Nederland B.V., Rapporteur: Kristina Dunder, Co-Rapporteur: Peter Kiely, PRAC Rapporteur: Brigitte Keller-Stanislawski	Positive Opinion adopted by consensus together with the CHMP assessment report and translation timetable. Based on the review of the available information, the CHMP was of the opinion that the renewal of the marketing authorisation can be granted with unlimited validity. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP Opinion.
Uptravi - selexipag - EMEA/H/C/003774/R/0030 Janssen-Cilag International N.V., Rapporteur: Martina Weise, Co-Rapporteur: Maria Concepcion Prieto Yerro, PRAC Rapporteur: Adrien Inoubli	Positive Opinion adopted by consensus together with the CHMP assessment report and translation timetable. Based on the review of the available information, the CHMP was of the opinion that the renewal of the marketing authorisation can be granted with unlimited validity. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP Opinion.
Wakix - pitolisant - EMEA/H/C/002616/R/0024, Orphan Bioprojet Pharma, Rapporteur: Alexandre Moreau, Co-Rapporteur: Outi Mäki-Ikola, PRAC Rapporteur: Kirsti Villikka	Positive Opinion adopted by consensus together with the CHMP assessment report and translation timetable. Based on the review of the available information, the CHMP was of the opinion that the renewal of the marketing authorisation can be granted with unlimited validity. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP Opinion.

B.2.3. Renewals of Conditional Marketing Authorisations

Caprelsa - vandetanib - EMEA/H/C/002315/R/0046 Genzyme Europe BV, Rapporteur: Alexandre Moreau, Co-Rapporteur: Paula Boudewina van Hennik, PRAC Rapporteur: Tiphaine Vaillant	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.
CRYSVITA - burosumab - EMEA/H/C/004275/R/0019, Orphan Kyowa Kirin Holdings B.V., Rapporteur: Kristina Dunder, Co-Rapporteur: Jayne Crowe, PRAC Rapporteur: Brigitte Keller-Stanislawski	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.
Holoclar - ex vivo expanded autologous human corneal epithelial cells containing stem cells - EMEA/H/C/002450/R/0032, Orphan, ATMP Holostem Terapie Avanzate s.r.l., Rapporteur: Egbert Flory, Co-Rapporteur: Paolo Gasparini, CHMP coordinators: Jan Mueller-Berghaus and Armando Genazzani, PRAC Rapporteur: Rhea Fitzgerald	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.
Polivy - polatuzumab vedotin - EMEA/H/C/004870/R/0003, Orphan Roche Registration GmbH, Rapporteur: Alexandre Moreau, Co-Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Annika Folin Request for Supplementary Information adopted on 17.09.2020.	See agenda 9.1 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 28 September 2020 – 01 October 2020 PRAC:**	

Signal of post-partum haemorrhage BRINTELLIX - vortioxetine; amitriptyline; bupropion; citalopram; escitalopram; fluoxetine; mirtazapine; paroxetine; sertraline; trazodone; venlafaxine Rapporteurs: various Brintellix: Rapporteur: Karin Janssen van Doorn Co-Rapporteur: Martina Weise PRAC recommendation on a variation / Monitor in PSUR **Action:** For adoption	Adopted.
Signal of Sjögren's Syndrome KEYTRUDA – pembrolizumab Rapporteur: Armando Genazzani Co-Rapporteur: Jan Mueller-Berghaus PRAC recommendation on a variation **Action:** For adoption	Adopted.
PSUR procedures for which PRAC adopted a recommendation for variation of the terms of the MA at its October 2020 meeting:	
EMEA/H/C/PSUSA/00001393/202002 (fingolimod) CAPS: **Gilenya** (EMEA/H/C/002202) (fingolimod), Novartis Europharm Limited, Rapporteur: Alexandre Moreau, PRAC Rapporteur: Tiphaine Vaillant, "Period Covered From: 01/03/2019 To: 28/02/2020"	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 4.4 of the SmPC, subsections Infections and Liver injury as well as section 4.8 to add Hepatobiliary disorders with a frequency not known: acute hepatic failure. In addition, the PL sections 2 and 4 should be updated accordingly. The Icelandic and the Norwegian CHMP members agree with the above-mentioned recommendation of the CHMP. The CHMP also agreed to the DHPC and communication plan.
EMEA/H/C/PSUSA/00001729/202001 (imiquimod) CAPS: **Aldara** (EMEA/H/C/000179) (imiquimod), Meda	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

AB, Rapporteur: Ewa Balkowiec Iskra **Zyclara** (EMEA/H/C/002387) (imiquimod), Meda AB, Rapporteur: Ewa Balkowiec Iskra NAPS: **NAPs** - EU PRAC Rapporteur: Adam Przybylkowski, "Period Covered From: 26/01/2017 To: 26/01/2020"	and the PRAC assessment report as appended, 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 sections 2 and 4 of the Package Leaflet to reflect the risk of exacerbation of autoimmune disorders. The Icelandic and the Norwegian CHMP members agree with the above-mentioned recommendation of the CHMP.
EMEA/H/C/PSUSA/00010015/202002 (ruxolitinib) CAPS: **Jakavi** (EMEA/H/C/002464) (ruxolitinib), Novartis Europharm Limited, Rapporteur: Filip Josephson, PRAC Rapporteur: Annika Folin, "Period Covered From: 23/02/2019 To: 22/02/2020"	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): The product information should be updated adding adverse drug reactions 'pancytopenia' (frequency - common) and 'HBV reactivation' (frequency not known) and revising the warning on HBV reactivation in sections 4.8 and 4.4 of the SmPC, respectively. The Icelandic and the Norwegian CHMP members agree with the above-mentioned recommendation of the CHMP.
EMEA/H/C/PSUSA/00010039/202002 (brentuximab vedotin) CAPS: **Adcetris** (EMEA/H/C/002455) (brentuximab vedotin), Takeda Pharma A/S, Rapporteur: Paula Boudewina van Hennik, PRAC Rapporteur: Menno van der Elst, "Period Covered From: 17/02/2019 To: 17/02/2020"	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, concerning the following change: Update of section 4.8 of the SmPC to add 'infusion site extravasation' (frequency unknown) for brentuximab vedotin monotherapy. The package leaflet should be updated accordingly. The Icelandic and the Norwegian CHMP members agree with the above-mentioned recommendation of the CHMP.
EMEA/H/C/PSUSA/00010073/202003 (bosutinib) CAPS: **Bosulif** (EMEA/H/C/002373) (bosutinib), Pfizer	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

Europe MA EEIG, Rapporteur: Janet Koenig, PRAC Rapporteur: Martin Huber, "Period Covered From: 04/03/2019 To: 03/03/2020"	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 sections 4.4 and 4.8 of the SmPC to add the adverse reaction "photosensitivity reaction" with a frequency common and a warning on the risk of photosensitivity. 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/00010368/202003 (oritavancin) CAPS: **Orbactiv** (EMEA/H/C/003785) (oritavancin), Menarini International Operations Luxembourg S.A., Rapporteur: Janet Koenig, PRAC Rapporteur: Adam Przybylkowski, "Period Covered From: 20/03/2019 To: 19/03/2020"	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 sections 4.4 and 4.8 of the SmPC to add information about possible adverse reactions is recommended. 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

EXPAREL liposomal - bupivacaine - EMEA/H/C/004586 PACIRA IRELAND LIMITED, indicated for prolonged acute pain management and reduction in need for opioids in adults compared to immediate-release bupivacaine., Known active substance (Article 8(3) of Directive No 2001/83/EC)	For information only. Comments can be sent to the PL in case necessary.
MenQuadfi - meningococcal group A, C, W135 and Y conjugate vaccine - EMEA/H/C/005084, Article 28 Sanofi Pasteur, immunization against Neisseria meningitidis serogroups A, C, W-135 and Y, Known active substance (Article 8(3) of Directive No 2001/83/EC)	For information only. Comments can be sent to the PL in case necessary.
Nyvepria - pegfilgrastim - EMEA/H/C/005085 Pfizer Europe MA EEIG, treatment of neutropenia, Similar biological application	For information only. Comments can be sent to the PL in case necessary.

(Article 10(4) of Directive No 2001/83/EC)	
Obiltoxaximab SFL - obiltoxaximab - EMEA/H/C/005169, Orphan SFL Pharmaceuticals Deutschland GmbH, treatment of inhalational anthrax due to Bacillus anthracis, New active substance (Article 8(3) of Directive No 2001/83/EC)	For information only. Comments can be sent to the PL in case necessary.
PHELINUN - melphalan - EMEA/H/C/005173 ADIENNE S.r.l., High-dose used alone or in combination with other cytotoxic drugs and/or total body irradiation is indicated in the treatment of: - multiple myeloma, - malignant lymphoma (Hodgkin, non-Hodgkin lymphoma), - acute lymphoblastic and myeloblastic leukemia, - childhood neuroblastoma, - ovarian adenocarcinoma, - mammary adenocarcinoma. In combination with other cytotoxic drugs and/or total body irradiation, in adult and paediatric population, is indicated as conditioning treatment prior to allogeneic haematopoietic stem cell transplantation (HSCT) in haematological diseases., Hybrid application (Article 10(3) of Directive No 2001/83/EC)	For information only. Comments can be sent to the PL in case necessary.
Rivaroxaban Accord - rivaroxaban - EMEA/H/C/005279 Accord Healthcare S.L.U., prevention of atherothrombotic events, Generic, Generic of Xarelto, Generic application (Article 10(1) of Directive No 2001/83/EC)	For information only. Comments can be sent to the PL in case necessary.
Supemtek - influenza quadrivalent vaccine (rDNA) - EMEA/H/C/005159 Sanofi Pasteur, prevention of influenza disease, New active substance (Article 8(3) of Directive No 2001/83/EC)	For information only. Comments can be sent to the PL in case necessary.

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

ADYNOVI - rurioctocog alfa pegol -	Positive Opinion adopted by consensus on

EMEA/H/C/004195/II/0015/G Baxalta Innovations GmbH, Rapporteur: Andrea Laslop Opinion adopted on 24.09.2020. Request for Supplementary Information adopted on 23.07.2020.	24.09.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Bemfola - follitropin alfa - EMEA/H/C/002615/II/0027 Gedeon Richter Plc., Rapporteur: Paula Boudewina van Hennik Opinion adopted on 15.10.2020. Request for Supplementary Information adopted on 17.09.2020.	Positive Opinion adopted by consensus on 15.10.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Bexsero - meningococcal group B vaccine (recombinant, component, adsorbed) - EMEA/H/C/002333/II/0094 GSK Vaccines S.r.l, Rapporteur: Kristina Dunder Opinion adopted on 15.10.2020.	Positive Opinion adopted by consensus on 15.10.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Caprelsa - vandetanib - EMEA/H/C/002315/II/0044/G Genzyme Europe BV, Rapporteur: Alexandre Moreau Opinion adopted on 01.10.2020. Request for Supplementary Information adopted on 11.06.2020.	Positive Opinion adopted by consensus on 01.10.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Desloratadine ratiopharm - desloratadine - EMEA/H/C/002404/II/0025/G ratiopharm GmbH, Generic, Generic of Aerius, Rapporteur: Christophe Focke Request for Supplementary Information adopted on 15.10.2020.	Request for supplementary information adopted with a specific timetable.
Dupixent - dupilumab - EMEA/H/C/004390/II/0031/G sanofi-aventis groupe, Rapporteur: Jan Mueller-Berghaus Opinion adopted on 01.10.2020. Request for Supplementary Information adopted on 16.07.2020.	Positive Opinion adopted by consensus on 01.10.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Evenity - romosozumab - EMEA/H/C/004465/II/0005 UCB Pharma S.A., Rapporteur: Kristina Dunder Opinion adopted on 15.10.2020.	Positive Opinion adopted by consensus on 15.10.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Flixabi - infliximab - EMEA/H/C/004020/II/0062 Samsung Bioepis NL B.V., Rapporteur: Jan Mueller-Berghaus	Positive Opinion adopted by consensus on 24.09.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

Opinion adopted on 24.09.2020.	
Flucelvax Tetra - influenza vaccine surface antigen inactivated prepared in cell cultures - EMEA/H/C/004814/II/0017 Seqirus Netherlands B.V., Rapporteur: Sol Ruiz Request for Supplementary Information adopted on 08.10.2020.	Request for supplementary information adopted with a specific timetable.
Herzuma - trastuzumab - EMEA/H/C/002575/II/0032/G Celltrion Healthcare Hungary Kft., Rapporteur: Jan Mueller-Berghaus Request for Supplementary Information adopted on 01.10.2020.	Request for supplementary information adopted with a specific timetable.
Hizentra - human normal immunoglobulin - EMEA/H/C/002127/II/0119/G CSL Behring GmbH, Rapporteur: Jan Mueller-Berghaus Request for Supplementary Information adopted on 15.10.2020.	Request for supplementary information adopted with a specific timetable.
IDELVION - albutrepenonacog alfa - EMEA/H/C/003955/II/0041/G, Orphan CSL Behring GmbH, Rapporteur: Jan Mueller-Berghaus Request for Supplementary Information adopted on 15.10.2020, 16.07.2020.	Request for supplementary information adopted with a specific timetable.
IMVANEX - smallpox vaccine (live modified vaccinia virus Ankara) - EMEA/H/C/002596/II/0049 Bavarian Nordic A/S, Rapporteur: Jan Mueller-Berghaus Opinion adopted on 01.10.2020. Request for Supplementary Information adopted on 23.07.2020.	Positive Opinion adopted by consensus on 01.10.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
IMVANEX - smallpox vaccine (live modified vaccinia virus Ankara) - EMEA/H/C/002596/II/0055 Bavarian Nordic A/S, Rapporteur: Jan Mueller-Berghaus Request for Supplementary Information adopted on 08.10.2020.	Request for supplementary information adopted with a specific timetable.
Lucentis - ranibizumab - EMEA/H/C/000715/II/0088 Novartis Europharm Limited, Rapporteur: Kristina Dunder Opinion adopted on 15.10.2020.	Positive Opinion adopted by consensus on 15.10.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Mosquirix - plasmodium falciparum and	Positive Opinion adopted by consensus on

hepatitis B vaccine (recombinant, adjuvanted) - EMEA/H/W/002300/II/0047 GlaxoSmithkline Biologicals SA, Rapporteur: Jan Mueller-Berghaus Opinion adopted on 24.09.2020.	24.09.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
NexoBrid - concentrate of proteolytic enzymes enriched in bromelain - EMEA/H/C/002246/II/0050/G, Orphan MediWound Germany GmbH, Rapporteur: Janet Koenig Opinion adopted on 15.10.2020.	Positive Opinion adopted by consensus on 15.10.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Nulojix - belatacept - EMEA/H/C/002098/II/0069 Bristol-Myers Squibb Pharma EEIG, Rapporteur: Filip Josephson Opinion adopted on 24.09.2020.	Positive Opinion adopted by consensus on 24.09.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Nulojix - belatacept - EMEA/H/C/002098/II/0071 Bristol-Myers Squibb Pharma EEIG, Rapporteur: Filip Josephson Request for Supplementary Information adopted on 15.10.2020.	Request for supplementary information adopted with a specific timetable.
Orencia - abatacept - EMEA/H/C/000701/II/0137/G Bristol-Myers Squibb Pharma EEIG, Rapporteur: Outi Mäki-Ikola Opinion adopted on 24.09.2020. Request for Supplementary Information adopted on 17.04.2020.	Positive Opinion adopted by consensus on 24.09.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Orencia - abatacept - EMEA/H/C/000701/II/0140/G Bristol-Myers Squibb Pharma EEIG, Rapporteur: Outi Mäki-Ikola Opinion adopted on 24.09.2020. Request for Supplementary Information adopted on 09.07.2020.	Positive Opinion adopted by consensus on 24.09.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Pemetrexed Hospira - pemetrexed - EMEA/H/C/003970/II/0024 Pfizer Europe MA EEIG, Generic, Generic of Alimta, Rapporteur: Alar Irs Opinion adopted on 24.09.2020.	Positive Opinion adopted by consensus on 24.09.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Polivy - polatuzumab vedotin - EMEA/H/C/004870/II/0002/G, Orphan Roche Registration GmbH, Rapporteur: Alexandre Moreau Opinion adopted on 15.10.2020.	Positive Opinion adopted by consensus on 15.10.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

Request for Supplementary Information adopted on 10.09.2020.	
Posaconazole Accord - posaconazole - EMEA/H/C/005005/II/0002 Accord Healthcare S.L.U., Generic, Generic of Noxafil, Rapporteur: Kolbeinn Gudmundsson Opinion adopted on 08.10.2020. Request for Supplementary Information adopted on 03.09.2020.	Positive Opinion adopted by consensus on 08.10.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Praluent - alirocumab - EMEA/H/C/003882/II/0058/G sanofi-aventis groupe, Rapporteur: Johann Lodewijk Hillege Request for Supplementary Information adopted on 15.10.2020.	Request for supplementary information adopted with a specific timetable.
Privigen - human normal immunoglobulin - EMEA/H/C/000831/II/0163 CSL Behring GmbH, Rapporteur: Jan Mueller-Berghaus Opinion adopted on 24.09.2020. Request for Supplementary Information adopted on 23.07.2020.	Positive Opinion adopted by consensus on 24.09.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Reblozyl - luspatercept - EMEA/H/C/004444/II/0001/G, Orphan Celgene Europe BV, Rapporteur: Milena Stain Request for Supplementary Information adopted on 15.10.2020.	Request for supplementary information adopted with a specific timetable.
Riluzole Zentiva - riluzole - EMEA/H/C/002622/II/0027 Zentiva, k.s., Rapporteur: Kirstine Moll Harboe Request for Supplementary Information adopted on 24.09.2020.	Request for supplementary information adopted with a specific timetable.
Ritonavir Mylan - ritonavir - EMEA/H/C/004549/II/0007/G Mylan S.A.S, Generic, Generic of Norvir, Rapporteur: John Joseph Borg Request for Supplementary Information adopted on 01.10.2020, 16.07.2020, 17.04.2020.	Request for supplementary information adopted with a specific timetable.
Rixubis - nonacog gamma - EMEA/H/C/003771/II/0035/G Baxalta Innovations GmbH, Rapporteur: Andrea Laslop Request for Supplementary Information adopted on 08.10.2020.	Request for supplementary information adopted with a specific timetable.
Ruconest - conestat alfa - EMEA/H/C/001223/II/0056	Positive Opinion adopted by consensus on 01.10.2020. The Icelandic and Norwegian CHMP

Pharming Group N.V, Rapporteur: Andrea Laslop Opinion adopted on 01.10.2020.	Members were in agreement with the CHMP recommendation.
Simulect - basiliximab - EMEA/H/C/000207/II/0107 Novartis Europharm Limited, Rapporteur: Jan Mueller-Berghaus Request for Supplementary Information adopted on 15.10.2020.	Request for supplementary information adopted with a specific timetable.
Skyrizi - risankizumab - EMEA/H/C/004759/II/0010/G AbbVie Deutschland GmbH & Co. KG, Rapporteur: Peter Kiely Opinion adopted on 15.10.2020. Request for Supplementary Information adopted on 23.07.2020.	Positive Opinion adopted by consensus on 15.10.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Trecondi - treosulfan - EMEA/H/C/004751/II/0004/G medac Gesellschaft fur klinische Spezialpraparate mbH, Rapporteur: Fátima Ventura Opinion adopted on 24.09.2020. Request for Supplementary Information adopted on 23.07.2020.	Positive Opinion adopted by consensus on 24.09.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Verzenios - abemaciclib - EMEA/H/C/004302/II/0012/G Eli Lilly Nederland B.V., Rapporteur: Filip Josephson Opinion adopted on 15.10.2020.	Positive Opinion adopted by consensus on 15.10.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
VITRAKVI - larotrectinib - EMEA/H/C/004919/II/0010/G Bayer AG, Rapporteur: Filip Josephson Request for Supplementary Information adopted on 15.10.2020.	Request for supplementary information adopted with a specific timetable.
Xenical - orlistat - EMEA/H/C/000154/II/0083 CHEPLAPHARM Arzneimittel GmbH, Rapporteur: Jean-Michel Race Request for Supplementary Information adopted on 15.10.2020.	Request for supplementary information adopted with a specific timetable.
Yervoy - ipilimumab - EMEA/H/C/002213/II/0083/G Bristol-Myers Squibb Pharma EEIG, Rapporteur: Paula Boudewina van Hennik Opinion adopted on 15.10.2020.	Positive Opinion adopted by consensus on 15.10.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Zavicefta - ceftazidime / avibactam - EMEA/H/C/004027/II/0023/G	Request for supplementary information adopted with a specific timetable.

Pfizer Ireland Pharmaceuticals, Rapporteur: Bjorg Bolstad Request for Supplementary Information adopted on 01.10.2020.	
WS1863/G **Incruse Ellipta-EMEA/H/C/002809/ WS1863/0030/G** **Rolufta Ellipta-EMEA/H/C/004654/ WS1863/0015/G** GlaxoSmithKline (Ireland) Limited, Lead Rapporteur: Maria Concepcion Prieto Yerro Opinion adopted on 24.09.2020.	Positive Opinion adopted by consensus on 24.09.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1904/G **Hexacima-EMEA/H/C/002702/WS1904/ 0105/G** **Hexaxim-EMEA/H/W/002495/WS1904/ 0110/G** **Hexyon-EMEA/H/C/002796/WS1904/ 0109/G** Sanofi Pasteur Europe, Duplicate, Duplicate of Hexacima, Lead Rapporteur: Jan Mueller-Berghaus Opinion adopted on 15.10.2020.	Positive Opinion adopted by consensus on 15.10.2020. 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

AUBAGIO - teriflunomide - EMEA/H/C/002514/II/0029 sanofi-aventis groupe, Rapporteur: Martina Weise, "To update section 4.4 of the SmPC to add information on cases of drug-induced liver injury (DILI) observed in the post-marketing setting and section 4.8 of the SmPC to add the adverse event DILI under the frequency unknown. The package leaflet is updated accordingly." Request for Supplementary Information adopted on 24.09.2020, 14.05.2020.	Request for supplementary information adopted with a specific timetable.
AUBAGIO - teriflunomide - EMEA/H/C/002514/II/0032 sanofi-aventis groupe, Rapporteur: Martina Weise, "C.I.4. Update of section 4.4 of the SmPC in order to update information on the liver monitoring schedule and the use of concomitant potentially hepatotoxic drugs based on evidence from diverse clinical and post-marketing sources including results from three studies, namely TENERE/EFC10891 (Phase 3 multi-center, randomized, double-blind, open-	Request for supplementary information adopted with a specific timetable.

label (for IFN β -1a), parallel-group study), Teri-PRO/LPS13567 study (Phase 4, multicenter, prospective, single-arm, open-label study) and TERIKIDS/EFC11759 (Phase 3 multicenter, randomized, double-blind, placebo-controlled, parallel-group study in patients with 10 to 17 years of age) together with post-marketing data including real-world data from two European National Disease registries (The Danish Multiple Sclerosis Registry and Belgian Treatment in Multiple Sclerosis, or BELTRIMS registry) and one US-based database of electronic health records (Optum Humedica Database) and post-marketing experience included in the Sanofi Global pharmacovigilance database." Request for Supplementary Information adopted on 24.09.2020.	
Busilvex - busulfan - EMEA/H/C/000472/II/0031 Pierre Fabre Medicament, Rapporteur: Blanca Garcia-Ochoa, "Update of section 4.5 of the SmPC regarding the interaction with deferasirox and iron chelating agents. The patient leaflet is updated accordingly. Update of the SmPC section 5.2 with minor changes in the paediatric population PK parameters. In addition, the MAH took the opportunity to clarify statement on incompatibilities in sections 6.2 and 6.6 and to expand the incompatibility of the polycarbonates syringes with Busilvex to the incompatibility of any infusion components containing polycarbonate with Busilvex. This change has been reflected on the subsection "Instructions for use" of the section 2 "recommendations for safe handling" in the preparation guide of the Package Leaflet. In addition, QRD-related changes have been implemented in annex II." Opinion adopted on 01.10.2020. Request for Supplementary Information adopted on 25.06.2020.	Positive Opinion adopted by consensus on 01.10.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Cresemba - isavuconazole - EMEA/H/C/002734/II/0031, Orphan Basilea Pharmaceutica Deutschland GmbH, Rapporteur: Johann Lodewijk Hillege, "Update of section 4.2 of SmPC to clarify the instructions on the start of therapy, according to current treatment guidelines for aspergillus and mucor diseases. Additionally, correction of an oversight is carried out in SmPC section 4.8 and PL	Request for supplementary information adopted with a specific timetable.

section 4 of Cresemba 100 mg capsules, to reinstate "odema peripheral" as "uncommon" adverse event, which was unintentionally omitted from the PI at the time of the initial MAA. Re-instatement of text about the potential interaction between isavuconazole and protease inhibitors other than indinavir and lopinavir/ritonavir, wrongly deleted from the interactions table in SmPC section 4.5 and PL section 2 as part of a previous procedure, is also carried out." Request for Supplementary Information adopted on 15.10.2020.	
DaTSCAN - ioflupane (123I) - EMEA/H/C/000266/II/0059 GE Healthcare B.V., Rapporteur: Alexandre Moreau, "C.I.4, Update of sections 4.2 and 5.1 of the SmPC in order to describe the possibility of quantitative reading in line with current scientific knowledge." Request for Supplementary Information adopted on 15.10.2020, 16.07.2020.	Request for supplementary information adopted with a specific timetable.
Enbrel - etanercept - EMEA/H/C/000262/II/0234 Pfizer Europe MA EEIG, Rapporteur: Maria Concepcion Prieto Yerro, "Update of section 5.1 of the SmPC in order to update the efficacy and safety information based on the final results from study (B1801381); this is a multicenter open-label study which evaluated withdrawal and retreatment of etanercept in subjects with non-radiographic axial spondyloarthritis who achieved an adequate response following 24 weeks of treatment. In addition, the MAH took the opportunity to make some editorial changes and bring the PI in line with the latest QRD template version 10.1." Request for Supplementary Information adopted on 15.10.2020.	Request for supplementary information adopted with a specific timetable.
Ervebo - recombinant vesicular stomatitis virus - Zaire ebolavirus vaccine (live) - EMEA/H/C/004554/II/0007/G Merck Sharp & Dohme B.V., Rapporteur: Christophe Focke, "C.I.4: To update section 5.1 of the SmPC with the description and final results from study V920-018; this is a phase 3 open-label trial conducted in Guinea to evaluate the safety and immunogenicity of Ervebo in vaccinated frontline workers 18 years of age	Request for supplementary information adopted with a specific timetable.

and older that was implemented as Part B of the Phase 3 ring vaccination study V920-010. With this submission, REC 20 is fulfilled. C.I.4: To update section 5.1 of the SmPC based on the result of the final study reports on the Correlate of Protection. With this submission, REC 16 is fulfilled. C.I.4: To update section 5.1 of the SmPC, based on results from the integrated summary of immunogenicity (ISI). With this submission, RECs 15 and 22 are fulfilled. C.I.13 - Submission of Non-Human Primates (NHP) Correlate of Protection analysis report (non-clinical report). Analysis is based upon previous submitted NHP studies which are already part of the dossier. The MAH takes the opportunity to implement changes in the Package Leaflet following the assessment of the User Acceptance Test, procedure EMEA/H/C/004554/REC/011. With the implementation of these changes to the PL, the MAH fulfils REC011. In addition, a minor editorial change has been included in section 4.4 of the SmPC and section 2 of the patient leaflet." Request for Supplementary Information adopted on 15.10.2020.	
Eylea - aflibercept - EMEA/H/C/002392/II/0064 Bayer AG, Rapporteur: Alexandre Moreau, "C.1.4 to update section 5.1 of the SmPC based on the ALTAIR Study with additional long-term efficacy information on patients with wet AMD." Request for Supplementary Information adopted on 08.10.2020.	Request for supplementary information adopted with a specific timetable.
Eylea - aflibercept - EMEA/H/C/002392/II/0065 Bayer AG, Rapporteur: Alexandre Moreau, "C.I.4, Update of section 4.2 to modify the posology in wet AMD and of 5.1 to reflect the underlying data." Request for Supplementary Information adopted on 15.10.2020.	Request for supplementary information adopted with a specific timetable.
Fluad Tetra - influenza vaccine (surface antigen, inactivated, adjuvanted) - EMEA/H/C/004993/II/0003 Seqirus Netherlands B.V., Rapporteur: Sol Ruiz, "Update of section 5.1 of the SmPC in order to introduce effectiveness information of Fluad	Request for supplementary information adopted with a specific timetable.

(trivalent formulation) in the product information of Fluad Tetra based on the results from three retrospective observational studies "CORE 17-18 and 18-19", "HEOR 17-18" and "HEOR 18-19") and a randomised clinical trial "LTCF 16-17".” Request for Supplementary Information adopted on 15.10.2020.	
Jevtana - cabazitaxel - EMEA/H/C/002018/II/0043/G sanofi-aventis groupe, Rapporteur: Alexandre Moreau, “Update of sections 4.8 and 5.1 of the SmPC with new clinical data from CARD study - a randomized, multicenter, Phase 4 study comparing cabazitaxel at 25 mg/m2 every 3 weeks in combination with prednisone versus alternate AR-targeted agent (abiraterone or enzalutamide) for the treatment of mCRPC patients previously treated with docetaxel and who failed a prior AR-targeted agent. Section 4.4 of the SmPC is also updated in accordance with the updated annex to the European Commission guideline on ‘Excipients in the labelling and package leaflet of medicinal products for human use’ (SANTE-2017-11668) regarding ethanol used as an excipient. The Package Leaflet is updated accordingly.” Request for Supplementary Information adopted on 15.10.2020.	Request for supplementary information adopted with a specific timetable.
Lyumjev - insulin lispro - EMEA/H/C/005037/II/0005 Eli Lilly Nederland B.V., Rapporteur: Outi Mäki-Ikola, “Update of sections 4.2 and 5.2 of the SmPC in order to update pharmacokinetic information based on final results from study I8B-MC-ITSA (submitted in accordance with Article 46 of regulation (EC) No 1901/2006). This study was conducted to evaluate the pharmacokinetics and glucodynamics of Lyumjev compared to Humalog in children, adolescents, and adults with Type 1 Diabetes Mellitus.” Request for Supplementary Information adopted on 15.10.2020.	Request for supplementary information adopted with a specific timetable.
Nerlynx - neratinib - EMEA/H/C/004030/II/0015 Pierre Fabre Medicament, Rapporteur: Bruno Sepodes, “Update of section 5.1 of the SmPC in order to include final OS results from study	Positive Opinion adopted by consensus on 15.10.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

3144A2-3004-WW, a randomised, double-blind, placebo-controlled trial of neratinib after trastuzumab in women with early-stage HER-2/neu overexpressed/amplified breast cancer. The MAH has also taken the opportunity to make some corrections in the SmPC and package leaflet." Opinion adopted on 15.10.2020. Request for Supplementary Information adopted on 03.09.2020.	
NexoBrid - concentrate of proteolytic enzymes enriched in bromelain - EMEA/H/C/002246/II/0049, Orphan MediWound Germany GmbH, Rapporteur: Janet Koenig, "Update of sections 4.2, 4.4, 4.8, 4.9, 5.1 and 5.2 of the SmPC based on interim results from study MW2010-03-02 (DETECT) listed as an obligation in the Annex II; this is a multicenter, multinational, randomized, controlled, assessor blinded, three-arm study performed in subjects with thermal burns, to evaluate the efficacy and safety of NexoBrid compared to Gel Vehicle and compared to Standard of Care. The submitted Clinical Study Report includes data of Acute Phase and 12M Follow Up Period. The MAH also took the opportunity to submit integrated pooled analyses of efficacy and safety data obtained from phase 2 and phase 3 clinical studies to provide a comprehensive assessment of the safety and efficacy of NexoBrid. The Package Leaflet is updated accordingly." Request for Supplementary Information adopted on 15.10.2020.	See agenda 9.1 Request for supplementary information adopted with a specific timetable.
Ondexxya - andexanet alfa - EMEA/H/C/004108/II/0011 Portola Netherlands B.V., Rapporteur: Jan Mueller-Berghaus, "C.I.4 Update of sections 4.4 and 4.5 of the SmPC in order to add a new warning on use of heparin after administration of andexanet based on spontaneous reports, medical literature reports, clinical trials and In vitro data. The Package Leaflet is updated accordingly. Additional editorial changes were proposed in Annex I and IIIB." Opinion adopted on 15.10.2020. Request for Supplementary Information adopted on 25.06.2020.	Positive Opinion adopted by consensus on 15.10.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

Ovitrelle - choriogonadotropin alfa - EMEA/H/C/000320/II/0081 Merck Europe B.V., Rapporteur: Paula Boudewina van Hennik, "Changes in sections 4.1, 4.2, 4.4 and 4.6 of the SmPC in order to update the terminology, in 4.3 to amend existing contraindications and in 4.8 to delete certain adverse drug reactions (ADRs) and add gastrointestinal ADRs with frequency common, with the aim to align the Product Information with similar text provided for other gonadotropin products. 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 10.1 and performed minor linguistic changes. The requested variation proposed amendments to the Summary of Product Characteristics, Labelling and Package Leaflet." Opinion adopted on 15.10.2020. Request for Supplementary Information adopted on 16.07.2020.	Positive Opinion adopted by consensus on 15.10.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Prolia - denosumab - EMEA/H/C/001120/II/0085/G Amgen Europe B.V., Rapporteur: Kristina Dunder, "Updates to SmPC section 4.8 adding the adverse reactions "hypersensitivity vasculitis" with a frequency category of very rare and "drug reaction with eosinophilia and systemic symptoms (DRESS) syndrome" with a frequency category of not known, and section 4.4 to introduce QRD traceability statement. The package leaflet has been updated accordingly." Opinion adopted on 24.09.2020.	Positive Opinion adopted by consensus on 24.09.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Ranexa - ranolazine - EMEA/H/C/000805/II/0063 Menarini International Operations Luxembourg S.A., Rapporteur: Kristina Dunder, "C.I.4 Update of section 4.8 of the SmPC in order to add "myoclonus" to the list of adverse drug reactions (ADRs) with frequency "rare" based on post-marketing data and update to section 4.9 based on review of the data regarding events of overdose. The Package Leaflet was updated accordingly. In addition, the MAH took the opportunity to update the list of local representatives in the Package Leaflet, to bring the PI in line with the latest QRD template	Positive Opinion adopted by consensus on 08.10.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

version 10.1, to correct linguistic mistakes in the SmPC and in some national translations of the Product Information." Opinion adopted on 08.10.2020.	
Rekovelle - follitropin delta - EMEA/H/C/003994/II/0023 Ferring Pharmaceuticals A/S, Rapporteur: Jean-Michel Race, "Update of section 5.1 of the SmPC in order to add information on dose equivalence factor to follitropin alfa based on clinical data from literature. The MAH took the opportunity to amend section 4.4. of the SmPC with traceability information and to bring the PI in line with the latest QRD template version 10.1." Request for Supplementary Information adopted on 15.10.2020.	Request for supplementary information adopted with a specific timetable.
Shingrix - herpes zoster vaccine (recombinant, adjuvanted) - EMEA/H/C/004336/II/0036 GlaxoSmithkline Biologicals SA, Rapporteur: Christophe Focke, "Submission of the final report from study Zoster-056, in order to fulfil the post-authorisation measure MEA/FSR 006. This is a cross-vaccination study in subjects who previously received placebo in studies Zoster-006 and Zoster-022." Request for Supplementary Information adopted on 01.10.2020.	Request for supplementary information adopted with a specific timetable.
Strensiq - asfotase alfa - EMEA/H/C/003794/II/0047, Orphan Alexion Europe SAS, Rapporteur: Armando Genazzani, "Update of section 5.1 of the SmPC in order to remove the Paediatric Investigation Plan (PIP) compliance statement as per Article 28(3) of Regulation (EC) No 1901/2006, following submission of the results and reports of all the PIP measures, including results of the Extrapolation Study AXN100107PIP ("Extrapolation of Efficacy to Asfotase Alfa Treatment in Paediatric Patients Ages 6 months to <3 years with Juvenile-Onset Hypophosphatasia")." Opinion adopted on 15.10.2020.	Positive Opinion adopted by consensus on 15.10.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Taltz - ixekizumab - EMEA/H/C/003943/II/0038/G Eli Lilly Nederland B.V., Rapporteur: Kristina Dunder, "Clinical studies in adult plaque psoriasis: Type II- C.I.4 -Update of section 5.1 of the	Request for supplementary information adopted with a specific timetable.

SmPC regarding long-term data in the treatment of psoriasis: RHAZ, RHBA, RHBC (3 studies of the "UNCOVER" series) were the pivotal registration studies, with response data up to 60 weeks already included in the SmPC. The current update relates to the extension data available, covering a total of 5 years. Section 5.1 of the SmPC has also been updated with information from study RHCR (known as "IXORA-R") which is a 24-week head-to-head comparison of Taltz vs guselkumab. Clinical studies in adult psoriatic arthritis: Type II- C.I.4 -Update of section 5.1 of the SmPC regarding long-term data in the treatment of psoriatic arthritis: RHAP and RHBE (also known as SPIRIT-P1 and P2) were pivotal registration studies, with response data at 24 weeks and up to 52 weeks (SPIRIT-P1) already included in the SmPC. The current update relates to extension data available covering a total of 3 years. Section 5.1 of the SmPC has also been updated with longer-term data from study RHCF ("SPIRIT-H2H" Taltz vs adalimumab). Response data of up to 24 weeks are already in the SmPC and the addition of 52 week data are being proposed." Request for Supplementary Information adopted on 15.10.2020.	
Tecentriq - atezolizumab - EMEA/H/C/004143/II/0030 Roche Registration GmbH, Rapporteur: Sinan B. Sarac, "Update of section 4.8 of the SmPC to reflect the outcome of anti-drug antibody (ADA) analyses conducted across studies POPLAR, OAK, IMpower 150, IMpower 130, IMPower 131, IMpower 132, IMvigor 211, IMmotion 151, IMpower 133 and IMpassion 130, further to the CHMP recommendation." Request for Supplementary Information adopted on 08.10.2020, 23.04.2020, 05.12.2019.	Request for supplementary information adopted with a specific timetable.
Tivicay - dolutegravir - EMEA/H/C/002753/II/0064 ViiV Healthcare B.V., Rapporteur: Filip Josephson, "Update of section 5.1 in order to add long-term efficacy and safety data, following the week 96 results from studies 204861 (GEMINI-1) and 205543 (GEMINI-2), listed as specific Category 3 studies in the RMP. These are two identical ongoing pivotal, randomised, double-blind, parallel group, 148-	Positive Opinion adopted by consensus on 15.10.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

week, phase III studies to evaluate the efficacy, safety and tolerability of dolutegravir plus lamivudine compared to dolutegravir plus tenofovir/emtricitabine in HIV-1-infected treatment naïve patients. In addition, the MAH took the opportunity to introduce minor editorial changes in the Product Information." Opinion adopted on 15.10.2020.	
Veltassa - patiromer - EMEA/H/C/004180/II/0018 Vifor Fresenius Medical Care Renal Pharma France, Rapporteur: Jayne Crowe, "Update of section 5.1 of the SmPC in order to update efficacy information based on final results from Study RLY5016-207; this is a randomised, double-blind, placebo-controlled, parallel group study of patiromer to enable concomitant spironolactone treatment in patients with resistant hypertension and CKD. Editorial changes are also made to section 6.4 of the SmPC. The PI is also brought to the latest QRD version 10.1." Opinion adopted on 15.10.2020. Request for Supplementary Information adopted on 18.06.2020.	Positive Opinion adopted by consensus on 15.10.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Vemlidy - tenofovir alafenamide - EMEA/H/C/004169/II/0023 Gilead Sciences Ireland UC, Rapporteur: Janet Koenig, "Update of sections 4.4, 4.8, 5.1 and 5.2 of the SmPC based on interim analysis data at Week 24 from study GS-US-320-4035. This is a Phase 2, open-label study evaluating the efficacy and safety of tenofovir alafenamide (TAF) in virologically suppressed chronic hepatitis B subjects with renal and/or hepatic impairment who switch from tenofovir disoproxil fumarate and/or other oral antiviral agents to TAF 25 mg once daily. Supportive safety and pharmacokinetic data are also provided from Study GS-US-292-1825 (phase 3b trial in which Elvitegravir/cobicistat/ emtricitabine/tenofovir alafenamide an fixed-dose TAF containing combination indicated for the treatment of HIV-1 infection, was evaluated in HIV-1 infected subjects with end stage renal disease). Study GS-US-320-4035 is included in the RMP version 4.2 under additional PhV activities." Opinion adopted on 24.09.2020. Request for Supplementary Information adopted	Positive Opinion adopted by consensus on 24.09.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

on 23.07.2020, 07.05.2020, 06.02.2020.	
Votrient - pazopanib - EMEA/H/C/001141/II/0059 Novartis Europharm Limited, Rapporteur: Sinan B. Sarac, "To update sections 4.2, 4.8 and 5.1 of the SmPC to update the safety information based on results from studies 2012-001306-20 (ADVL0815 / PZP114411) and study 2013-003595-12 (ADVL1322 / VEG116731 / PZP034X2203) listed in the agreed PIP; these are a phase 1 clinical trial of single-agent pazopanib in children with a relapsed or refractory solid (including CNS) tumour, and a therapeutic-exploratory (phase 2) clinical trials of single-agent pazopanib in children (including adolescents) and young adults with a refractory tumour." Request for Supplementary Information adopted on 15.10.2020.	Request for supplementary information adopted with a specific timetable.
VPRIV - velaglucerase alfa - EMEA/H/C/001249/II/0048, Orphan Shire Pharmaceuticals Ireland Limited, Rapporteur: Martina Weise, "Update of section 4.4 of the SmPC to include information on 1 additional patient with IgG anti-velaglucerase antibodies with neutralizing activity reported during extension Study HGT-GCB-044, and to include vomiting as an infusion-related reaction that has been reported in post-marketing experience. Further, the MAH is updating the instructions in sections 4.2 and 6.6 of the SmPC to state that a 0.2 µm filter and a 0.22 µm filter are both considered acceptable when administering the product. In addition, the MAH took the opportunity to implement some minor editorial changes in SmPC section 5.1 and a clarification that paediatric patients included in the studies were 4 years of age and older. The Package Leaflet is updated accordingly." Request for Supplementary Information adopted on 15.10.2020.	Request for supplementary information adopted with a specific timetable.
XOSPATA - gilteritinib - EMEA/H/C/004752/II/0001, Orphan Astellas Pharma Europe B.V., Rapporteur: Bjorg Bolstad, "Submission of a pooled analysis report from studies 2215-CL-0101 (Phase 1/2), 2215-CL-0102 (Phase 1), 2215-CL-0301, 2215-CL-9100 (phase 3) listed as "Other forms of routine pharmacovigilance activities in section III.1 of	Request for supplementary information adopted with a specific timetable.

the RMP". This is a pooled analysis to characterise gilteritinib-related differentiation syndrome, specifically incidence, observed signs and symptoms, duration and response to intervention based on patient-level data from on-going trials in patients with acute myeloid leukemia." Request for Supplementary Information adopted on 08.10.2020.	
WS1822 **Relvar Ellipta-EMEA/H/C/002673/ WS1822/0045** **Revinty Ellipta-EMEA/H/C/002745/ WS1822/0043** GlaxoSmithKline (Ireland) Limited, Lead Rapporteur: Maria Concepcion Prieto Yerro, "Update of section 5.1 of the Relvar/Revinty SmPC to include safety information based on results from Therapeutic Index study 203162. This study compared the therapeutic index of fluticasone furoate (FF) and other inhaled corticosteroid (ICS) agents using the efficacy marker of adenosine5'-monophosphate (AMP) challenge and the systemic exposure marker of cortisol suppression. The results provide new information that will help prescribers to understand the relative potency for efficacy and systemic activity of the ICS component of Relvar/Revinty, fluticasone furoate (FF), compared to other ICS drug molecules. In addition, GSK has taken the opportunity to add text related to SUMMIT data to section 5.1 of the high strength label (184/22 mcg) for Relvar/Revinty Ellipta. The text was agreed in procedure EMEA/H/C/XXXX/WS/0992 finalised on 21st April 2017, however the change was mistakenly not implemented to the SmPC during this procedure. Additionally, minor corrections are introduced in the PL." Request for Supplementary Information adopted on 15.10.2020, 25.06.2020.	Request for supplementary information adopted with a specific timetable.
WS1886/G **Aprovel-EMEA/H/C/000141/WS1886/ 0181/G** **CoAprovel-EMEA/H/C/000222/WS1886/ 0199/G** **Karvea-EMEA/H/C/000142/WS1886/ 0183/G** **Karvezide-EMEA/H/C/000221/WS1886/**	Request for supplementary information adopted with a specific timetable.

0199/G sanofi-aventis groupe, Lead Rapporteur: Maria Concepcion Prieto Yerro, "Group of variations consisting of: C.I.4 - Update of sections 4.4 and 4.8 of the SmPC to add information on hypoglycaemia based on a review of available data including the MAH pharmacovigilance data base and a literature review. The Package leaflet is updated accordingly. C.I.4 - Update of 4.4 and 4.5 of the SmPC to add information on a drug -drug interaction with irbesartan and repaglinide based on a review of the available data including the MAH database and a literature review. The package leaflet is updated accordingly. In addition, the MAH took the opportunity to implement the updated annex to the European Commission guideline on Excipients in the labelling and package leaflet of medicinal products for human use' to update the excipient sodium." Request for Supplementary Information adopted on 08.10.2020.	
WS1898 **Efficib-EMEA/H/C/000896/WS1898/0095** **Janumet-EMEA/H/C/000861/WS1898/0095** **Ristfor-EMEA/H/C/001235/WS1898/0082** **Velmetia-EMEA/H/C/000862/WS1898/0098** Merck Sharp & Dohme B.V., Lead Rapporteur: Johann Lodewijk Hillege, "Update of the SmPC sections 4.2, 4.8, 5.1 and 5.2, to include data from the pooled analysis of paediatric studies P170 (sitagliptin/metformin) and P289 (sitagliptin/metformin extended release). The package leaflet is revised accordingly, and update of the product information is performed to comply with QRD Version 10.1." Opinion adopted on 24.09.2020.	Positive Opinion adopted by consensus on 24.09.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

B.5.3. CHMP-PRAC assessed procedures

Desloratadine ratiopharm - desloratadine - EMEA/H/C/002404/II/0023/G ratiopharm GmbH, Generic, Generic of Aerius, Rapporteur: Christophe Focke, PRAC Rapporteur: Laurence de Fays, "C.I.5.b - Change in the legal status of desloratadine ratiopharm from 'medicinal product subject to	See agenda 9.1 Positive Opinion adopted by consensus on 15.10.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

medical prescription' to 'medicinal product not subject to medical prescription'. As a consequence, sections 4.1, 4.2, 4.3, 4.4, 4.5, 4.6, 4.8 and 5.1 of the SmPC are updated. The annex II, package leaflet and labelling are updated accordingly. Furthermore, the product information is brought in line with the latest QRD template (version 10.1) and the list of local representatives in the package leaflet is updated. The RMP is updated to version 1.2. As a consequence of the variation, the pack sizes of 40, 50, 60, 90 and 100 tables are deleted (EU/1/11/746/007-011). C.I.6.b - To delete the therapeutic indication in adolescents aged 12 years and older for the relief of symptoms associated with allergic rhinitis and urticaria. Section 4.1 of the SmPC and section 1 of the PL are updated accordingly." Opinion adopted on 15.10.2020. Request for Supplementary Information adopted on 23.07.2020, 26.03.2020.	
Erleada - apalutamide - EMEA/H/C/004452/II/0008 Janssen-Cilag International N.V., Rapporteur: Blanca Garcia-Ochoa, PRAC Rapporteur: Tiphaine Vaillant, "Update of sections 4.4, 4.8 and 5.1 of the SmPC in order to update efficacy and safety information based on final results from study ARN-509-003 (SPARTAN) listed as a PAES in Annex II; this is a multicenter, randomised, double-blind, placebo-controlled, phase III study of ARN-509 in men with non-metastatic (M0) castration-resistant prostate cancer; the package leaflet and Annex II are updated accordingly. The RMP version 3.1 has also been submitted. In addition, the MAH took the opportunity to update the list of local representatives in the Package leaflet." Request for Supplementary Information adopted on 01.10.2020.	Request for supplementary information adopted with a specific timetable.
Imfinzi - durvalumab - EMEA/H/C/004771/II/0023 AstraZeneca AB, Rapporteur: Sinan B. Sarac, PRAC Rapporteur: David Olsen, "Update of sections 4.2 and 5.1 of the SmPC in order to introduce a new posology regimen of 1500 mg every 4 weeks (Q4W) for the approved indication of the treatment of patients with locally advanced, unresectable non-small cell	Request for supplementary information adopted with a specific timetable.

lung cancer (NSCLC) whose tumours express PD-L1 on ≥ 1% of tumour cells and whose disease has not progressed following platinum based chemoradiation therapy. The RMP version 4.1 has also been submitted." Request for Supplementary Information adopted on 15.10.2020.	
Kanuma - sebelipase alfa - EMEA/H/C/004004/II/0026/G, Orphan Alexion Europe SAS, Rapporteur: Karin Janssen van Doorn, PRAC Rapporteur: Ulla Wändel Liminga, "Grouping consisting of the following variations: - Update of sections 4.2, 4.4, 4.8, 5.1, 5.2 and 6.6 of the Summary of Product Characteristics (SmPC) in order to update the clinical information based on the pooled safety and efficacy analysis of already submitted studies (Studies LAL-CL04, LAL-CL03, LAL-CL06, LAL-CL08 and LAL-CL02) and updated population pharmacokinetics analyses in children and adults. The Package Leaflet has been amended accordingly. The RMP version 4.1 has also been submitted. Annex II is also updated to remove the specific obligation related to the provision of study LAL-CL08. - Submission of the final report from study LAL-EA01 An open-label study with sebelipase alfa (1 mg/kg every other week for up to 78 weeks or until drug commercialization) in United States (US) patients who did not otherwise qualify for an active sebelipase alfa trial (expanded access protocol)" Opinion adopted on 15.10.2020. Request for Supplementary Information adopted on 23.07.2020, 26.03.2020.	See agenda 9.1 Positive Opinion adopted by consensus on 15.10.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Myalepta - metreleptin - EMEA/H/C/004218/II/0012, Orphan Amryt Pharmaceuticals DAC, Rapporteur: Karin Janssen van Doorn, PRAC Rapporteur: Adam Przybylkowski, "Update of section 4.4 of the SmPC in order to add a new warning on the risk of autoimmune disease following exposure to metreleptin; the Package Leaflet and the key elements to be included in the Guide/training material for healthcare professionals are updated accordingly. The RMP version 2.0 has also been submitted." Opinion adopted on 01.10.2020. Request for Supplementary Information adopted	Positive Opinion adopted by consensus on 01.10.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

on 09.07.2020.	
NUBEQA - darolutamide - EMEA/H/C/004790/II/0002 Bayer AG, Rapporteur: Alexandre Moreau, PRAC Rapporteur: Jan Neuhauser, "Update of section 5.1 of the SmPC in order to update efficacy information based on the final CSR from study 17772 (ARAMIS) listed as a PAES in the Annex II; this is a multinational, randomised, double-blind, placebo-controlled, phase III efficacy and safety study of darolutamide in men with high-risk non-metastatic castration-resistant prostate cancer; the Annex II is updated accordingly. The RMP version 1.1 is accepted." Opinion adopted on 08.10.2020. Request for Supplementary Information adopted on 03.09.2020.	Positive Opinion adopted by consensus on 08.10.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Ocrevus - ocrelizumab - EMEA/H/C/004043/II/0020 Roche Registration GmbH, Rapporteur: Kirstine Moll Harboe, PRAC Rapporteur: Brigitte Keller-Stanislawski, "Submission of the final report for study 17-1133 listed as a Category 3 study in the RMP (MEA 006). This is a study to assess the effects of ocrelizumab on embryo-fetal and pre- and postnatal development in cynomolgus monkeys. The RMP ver. 5.0 has also been submitted." Request for Supplementary Information adopted on 01.10.2020.	Request for supplementary information adopted with a specific timetable.
OFEV - nintedanib - EMEA/H/C/003821/II/0038 Boehringer Ingelheim International GmbH, Rapporteur: Peter Kiely, PRAC Rapporteur: Nikica Mirošević Skvrce, "Update of sections 4.5, 4.6 and 5.2 of the SmPC in order to add drug-drug interaction information with the oral contraceptive Microgynon, a combination of ethynilestradiol and levonorgestrel based on final results from clinical study №1199-0340. This was a phase I, open-label, 2-period cross-over, fixed-sequence design trial, investigated the effect of multiple oral doses of nintedanib on the single dose kinetics of a combination of ethinylestradiol and levonorgestrel (Microgynon). The Package Leaflet is updated accordingly. The RMP version 10 has also been submitted." Request for Supplementary Information adopted	Request for supplementary information adopted with a specific timetable.

on 01.10.2020.	
Oncaspar - pegaspargase - EMEA/H/C/003789/II/0036/G Les Laboratoires Servier, Rapporteur: Alexandre Moreau, PRAC Rapporteur: Annika Folin, "Group of 2 Type II variations to submit the study results of study 12-266 A(12) an open label single arm phase II trial evaluating the efficacy and toxicity of treatment regimens including Oncaspar in adults (aged 18-60) with newly diagnosed Philadelphia chromosome-negative acute lymphoblastic leukaemia and study CAALL-F01 a prospective multicentre cohort study evaluating Oncaspar used in the first-line treatment of children and adolescents with ALL along with multi-agent chemotherapy. Consequently, Annex II is proposed to be updated to remove both PAES. Additionally, update of the product information to remove the need for additional monitoring and to implement editorial changes. The RMP (version 4.1) is updated accordingly." Request for Supplementary Information adopted on 01.10.2020.	Request for supplementary information adopted with a specific timetable.
Privigen - human normal immunoglobulin - EMEA/H/C/000831/II/0161/G CSL Behring GmbH, Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Brigitte Keller-Stanislawski, "C.I.4: Update of sections 4.4, 4.8 and 5.1 of the SmPC in order to amend an existing warning on haemolytic anaemia and to update safety information based on final results from study IgPro10_5003 listed as a category 3 study in the RMP; this is an observational hospital-based cohort study in the US to evaluate Privigen use and haemolytic anaemia in adults and children and the Privigen safety profile in children with CIDP; C.I.4: Update of sections 4.8 and 5.1 of the SmPC in order to update to the list of adverse drug reactions based on final results from study IgPro10_3004; this is a Prospective Open-Label Single-Arm Study of the Pharmacokinetics and Safety of Intravenous IgPro10 in Japanese Subjects with Primary Immunodeficiency The Package Leaflet is updated accordingly. The RMP version 8.0 has also been submitted. In addition, the MAH took the opportunity to align the SmPC (sections 4.1, 4.2, 4.3, 4.4, 4.5 and 4.8) with the EU Core SmPC for IVIG, to update	Positive Opinion adopted by consensus on 01.10.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

the local representative for Bulgaria and Slovenia in the Package Leaflet and to bring the PI in line with the latest QRD template version 10.1." Opinion adopted on 01.10.2020. Request for Supplementary Information adopted on 09.07.2020.	
SCENESSE - afamelanotide - EMEA/H/C/002548/II/0033, Orphan Clinuvel Europe Limited, Rapporteur: Janet Koenig, PRAC Rapporteur: Martin Huber, "Update of section 4.8 of the SmPC to revise the frequencies of adverse drug reactions (ADRs) based on safety reports and to add new ADRs based on post-marketing spontaneous reports as requested during Scenesse Renewal procedure (EMEA/H/C/002548/R/0026); the Package Leaflet is updated accordingly. The revised RMP version 9.0 (in line with rev 2 of the template) is acceptable. In addition, the MAH took the opportunity to introduce minor editorial changes in section 2 of the SmPC and Annex IIIA." Opinion adopted on 01.10.2020. Request for Supplementary Information adopted on 14.05.2020.	Positive Opinion adopted by consensus on 01.10.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Stelara - ustekinumab - EMEA/H/C/000958/II/0081/G Janssen-Cilag International NV, Rapporteur: Jayne Crowe, PRAC Rapporteur: Rhea Fitzgerald, "Update of section 4.2 of Stelara SmPC solution for injection presentations in order to change posology recommendations for patients with ulcerative colitis, and 5.1 of Stelara SmPC to update efficacy information based on 2-year results from study 3001 listed as a category 3 study in the RMP; this is a Phase 3, randomized, double blind, placebo controlled, parallel-group, multicenter protocol to evaluate the safety and efficacy of ustekinumab induction and maintenance therapy in subjects with moderately to severely active ulcerative colitis. Update of section 5.1 of the SmPC in order to update efficacy information based on 5-year results from study 3003 listed as a category 3 study in the RMP; this is a Phase 3, randomized, double blind, placebo controlled, parallel-group, multicenter trial to evaluate the safety and efficacy of ustekinumab maintenance therapy in	Request for supplementary information adopted with a specific timetable.

adult patients with moderately to severely active Crohn's disease. The RMP version 18.1 has also been submitted." Request for Supplementary Information adopted on 15.10.2020.	
TECFIDERA - dimethyl fumarate - EMEA/H/C/002601/II/0063 Biogen Netherlands B.V., Rapporteur: Martina Weise, PRAC Rapporteur: Martin Huber, "Update of sections 4.3, 4.4 and 4.8 of the SmPC to reflect new available information on Progressive Multifocal Leukoencephalopathy (PML) risk monitoring based on a cumulative review of PML cases in the setting of mild lymphopenia. The Package Leaflet is updated accordingly. Additionally, the Product Information has been updated in line with QRD template (version 10.1)." Opinion adopted on 15.10.2020. Request for Supplementary Information adopted on 17.09.2020, 28.05.2020, 30.01.2020, 19.09.2019.	See agenda 9.1 Positive Opinion adopted by consensus on 15.10.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Tevagrastim - filgrastim - EMEA/H/C/000827/II/0077 TEVA GmbH, Duplicate, Duplicate of Biograstim (SRD), Rapporteur: Outi Mäki-Ikola, PRAC Rapporteur: Kirsti Villikka, "Submission of a variation to update the RMP to remove the additional pharmacovigilance activity "Cooperation with SCNIR (Severe Chronic Neutropenia International Registry) and analysis of corresponding Ratiograstim/Tevagrastim-SCNIR data"." Opinion adopted on 01.10.2020.	Positive Opinion adopted by consensus on 01.10.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Trumenba - meningococcal group B vaccine (recombinant, adsorbed) - EMEA/H/C/004051/II/0027/G Pfizer Europe MA EEIG, Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Jean-Michel Dogné, "C.I.11.b- To update the RMP for Trumemba to version 4.0 to provide a revised protocol outline for study B1971060 in immunocompromised individuals: Although the study was originally designed to evaluate 3 doses of Trumenba administered on a 0-, 2-, and 6-month schedule, the MAH is now proposing a 2-dose regimen administered on 0- and 6-month schedule. C.I.11.b- To submit the protocol outline for the	Positive Opinion adopted by consensus on 01.10.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

co-administration study (C3511006). The MAH is proposing that the commitment to conduct a co-administration study with Trumenba may be met by a study of the MAH's candidate pentavalent meningococcal vaccine (which contains Trumenba) co-administered with MMR and PnC vaccines." Opinion adopted on 01.10.2020. Request for Supplementary Information adopted on 09.07.2020.	
Vizimpro - dacomitinib - EMEA/H/C/004779/II/0003/G Pfizer Europe MA EEIG, Rapporteur: Blanca Garcia-Ochoa, PRAC Rapporteur: Menno van der Elst, "Update of sections 4.2 and 5.2 of the SmPC in order to revise the dosing recommendation for patients with hepatic impairment and include relevant pharmacokinetics data based on results of study A7471058, evaluating the effect of severe hepatic impairment on the plasma PK, safety and tolerability after a single dose of dacomitinib. As a consequence, the MAH is proposing to remove the missing information "Safety in Patient with Severe Hepatic Impairment" from the list of safety concerns in the RMP. In addition, the MAH took the opportunity to bring the PI in line with the latest QRD template version 10.1. The MAH has also taken the opportunity to update the EU RMP to include PASS Study A7471064 "Single Arm Study to Evaluate the Safety of Dacomitinib for the First-Line Treatment of Participants in India with Metastatic NSCLC with Epidermal Growth Factor Receptor (EGFR)-Activating Mutations" as a "Category 3 required additional pharmacovigilance activity". A revised RMP v1.1 (clean and tracked) has been submitted." Request for Supplementary Information adopted on 15.10.2020.	Request for supplementary information adopted with a specific timetable.
WS1792/G **Hexacima-EMEA/H/C/002702/WS1792/0099/G** **Hexaxim-EMEA/H/W/002495/WS1792/0104/G** **Hexyon-EMEA/H/C/002796/WS1792/0103/G** Sanofi Pasteur, Lead Rapporteur: Jan Mueller-Berghaus, Lead PRAC Rapporteur: Brigitte	Positive Opinion adopted by consensus on 24.09.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

Keller-Stanislawski, "C.I.4 (type II) - Update of sections 4.4 and 5.1 of the SmPC in order to revise the warning regarding preterm infants and to add new information on immunogenicity in preterm infants and in infants born from women vaccinated during pregnancy, based on the final results from study A3L00053-EXT; this is an observational cohort study conducted with DTaP-IPV-HB-PRP~T vaccine, aimed to describe the concentrations of IgG against the different antigens. The RMP version 12.0 has been submitted and updated accordingly, following revision 2 with consequential update to the safety concerns. C.I.z (type IB) - Update of sections 2 and 4.4 of the SmPC in order to add warning for excipients with known effect: phenylalanine, potassium and sodium, according to the European guideline "Excipients in the labelling and package leaflet of medicinal products for human use SANTE-2017-11668". The package leaflet is updated accordingly. In addition, the MAH/SOH took the opportunity to introduce editorial changes in sections 4.2, 4.4, 4.5 and 4.8 of the SmPC and to update the list of local representatives in the Package Leaflet." Opinion adopted on 24.09.2020. Request for Supplementary Information adopted on 11.06.2020.	
WS1820 **Iscover-EMEA/H/C/000175/WS1820/0142** **Plavix-EMEA/H/C/000174/WS1820/0140** sanofi-aventis groupe, Lead Rapporteur: Bruno Sepodes, Lead PRAC Rapporteur: Marcia Sofia Sanches de Castro Lopes Silva, "Update of section 4.2 of the SmPC in order to add 600 mg as an alternative loading dose to the existing 300 mg to be used at initiation of treatment in the indication "Secondary prevention of atherothrombotic events in adult patients suffering from acute coronary syndrome ". This update is based on a bibliographic review of published studies. The Package Leaflet is updated accordingly. The RMP version 2.0 has also been submitted." Request for Supplementary Information adopted on 15.10.2020, 25.06.2020.	See agenda 9.1 Request for supplementary information adopted with a specific timetable.

WS1844 **Edistride-EMEA/H/C/004161/WS1844/0039** **Forxiga-EMEA/H/C/002322/WS1844/0057** AstraZeneca AB, Lead Rapporteur: Kristina Dunder, Lead PRAC Rapporteur: Annika Folin, "Re-categorisation of the Forxiga/Edistride PASS (D169C00011): Retrospective Cohort Study on the Risk of Diabetic Ketoacidosis (DKA) to determine the effectiveness of additional risk minimisation measures in place for DKA by assessing the impact of the risk minimisation measures on the risk of DKA in T1DM patients who are treated with dapagliflozin in Europe, from a category 1 to category 3 PASS in the RMP Version 20 and removal of the following Annex IID of the PI: Obligation to conduct the following Non-interventional PASS: In order to estimate the incidence of DKA in T1DM dapagliflozin users following implementation of RMMs in Europe, the MAH should conduct and submit the results from an observational cohort study using existing data sources in European countries where dapagliflozin will be launched for T1DM" Request for Supplementary Information adopted on 15.10.2020, 23.07.2020.	See agenda 9.1 Request for supplementary information adopted with a specific timetable.
WS1850 **Anoro Ellipta-EMEA/H/C/002751/WS1850/0030** **Laventair Ellipta-EMEA/H/C/003754/WS1850/0033** GlaxoSmithKline (Ireland) Limited, Lead Rapporteur: Peter Kiely, Lead PRAC Rapporteur: Ilaria Baldelli, "To update the RMP with the completion of study WWE117397 "A Postauthorization safety Electronic Medical Records database retrospective cohort study of new users of inhaled UMEC/VI or new users of inhaled UMEC in the primary care setting." As part of the assessment of EMEA/H/C/WS1761 the MAH was requested to update the RMP. In addition, the MAH has amendment the RMP with the study 201038 "A Post authorisation Safety Observational Cohort Study to quantify the incidence of selected cardiovascular and cerebrovascular events in COPD patients using inhaled UMEC/VI combination or inhaled UMEC versus Tiotropium" as approved during	Positive Opinion adopted by consensus on 15.10.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

procedure EMEA/H/C/PSA/S/0032.3." Opinion adopted on 15.10.2020. Request for Supplementary Information adopted on 23.07.2020.	

B.5.4. PRAC assessed procedures

PRAC Led **Aclasta - zoledronic acid - EMEA/H/C/000595/II/0076** Novartis Europharm Limited, Rapporteur: Kristina Dunder, PRAC Rapporteur: Ulla Wändel Liminga, PRAC-CHMP liaison: Kristina Dunder, "Provision of an updated RMP version 13.0. Proposed changes: 1. Alignment with requirements of GVP Module V Rev.2 with consequential removal/reclassification of a number of important potential risks; 2. Consequential removal of education material for renal risk (renal dysfunction and use in patients with severe renal impairment); 3. Removal of 'post-dose symptoms' from the list of important identified risks (following the assessment of LEG 037 & variation II/74/G); 4. Update of the targeted questionnaire related to the ONJ risk (following the assessment of LEG 035); 5. Inclusion of the completed 5-year registry study ZOL446H2422 (following the assessment of variation II-69 & MEA 017.1). The additional risk minimisation measures in the Annex II of the product information are proposed to be updated accordingly." Request for Supplementary Information adopted on 01.10.2020.	Request for supplementary information adopted with a specific timetable.
PRAC Led **Circadin - melatonin - EMEA/H/C/000695/II/0061** RAD Neurim Pharmaceuticals EEC SARL, Rapporteur: Bruno Sepodes, PRAC Rapporteur: Ana Sofia Diniz Martins, PRAC-CHMP liaison: Bruno Sepodes, "Risk Management Plan update to remove the following risks from the list of potential risks: "Drug interaction with levothyroxine" "Panic Attacks", "Potential interaction with warfarin", "Sperm motility decreased/Spermatozoa morphology abnormal" and "Withdrawal"." Request for Supplementary Information adopted	Request for supplementary information adopted with a specific timetable.

on 01.10.2020.	
PRAC Led **EMEND - aprepitant - EMEA/H/C/000527/II/0063** Merck Sharp & Dohme B.V., Rapporteur: Filip Josephson, PRAC Rapporteur: Annika Folin, PRAC-CHMP liaison: Filip Josephson, "update of the RMP to version 5.1 to remove all the safety concerns (important identified risks, important potential risks and missing information) and information related to both 40 mg and 165 mg capsules strengths and the Postoperative Nausea and Vomiting indication (PONV), as well as to update data in the post-authorisation exposure (Part II: Module SV) and epidemiology (Part II: Module SI) sections." Opinion adopted on 01.10.2020.	Positive Opinion adopted by consensus on 01.10.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
PRAC Led **Esbriet - pirfenidone - EMEA/H/C/002154/II/0066/G, Orphan** Roche Registration GmbH, Rapporteur: Peter Kiely, PRAC Rapporteur: Rhea Fitzgerald, PRAC-CHMP liaison: Peter Kiely, "Update of sections 4.4 and 4.8 of the SmPC in order to amend an existing warning on drug-induced liver injury (DILI) subsequent to the latest PSUSA (EMEA/H/C/PSUSA/00002435/201902) and post-authorisation measure (EMEA/H/C/2154/LEG/015). The annex II and package leaflet (PL) are updated accordingly. The RMP version 10.2 has also been updated. In addition, the MAH took the opportunity to add in the PL information about sodium content in line with excipients guideline (EMA/CHMP/302620/2017) and to correct formatting, punctuation and spelling mistakes in the product information. Update of sections 4.4 and 4.8 of the SmPC in order to add a warning on hyponatraemia and to add hyponatraemia with a frequency 'uncommon' to the list adverse reactions subsequent to the latest PSUSA (EMEA/H/C/PSUSA/00002435/201902) and post-authorisation measure (EMEA/H/C/2154/LEG/015). The PL is updated accordingly." Opinion adopted on 01.10.2020. Request for Supplementary Information adopted	Positive Opinion adopted by consensus on 01.10.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

on 11.06.2020.	
PRAC Led **Imbruvica - ibrutinib - EMEA/H/C/003791/II/0061, Orphan** Janssen-Cilag International NV, Rapporteur: Filip Josephson, PRAC Rapporteur: Nikica Mirošević Skvrce, PRAC-CHMP liaison: Selma Arapovic Dzakula, "Update of the RMP introducing changes to safety concerns following the assessment of the renewal R/0049. The MAH is taking this opportunity to include additional changes related to two post-authorisation measures; postponement of the completion date of cat3 study PCI-32765MCL3002 of ibrutinib in combination with BR versus BR alone and removal of study 54179060CLL1017 on DDI as assessed in II/0058." Opinion adopted on 01.10.2020. Request for Supplementary Information adopted on 11.06.2020.	Positive Opinion adopted by consensus on 01.10.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
PRAC Led **Ivemend - fosaprepitant - EMEA/H/C/000743/II/0043** Merck Sharp & Dohme B.V., Rapporteur: Filip Josephson, PRAC Rapporteur: Annika Folin, PRAC-CHMP liaison: Filip Josephson, "update of the RMP for IVEMEND to version 5.1 to remove all the safety concerns (important identified risks, important potential risks and missing information), as well as to update data in the post-authorisation exposure (Part II: Module SV) and epidemiology (Part II: Module SI) sections." Opinion adopted on 01.10.2020.	Positive Opinion adopted by consensus on 01.10.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
PRAC Led **Keppra - levetiracetam - EMEA/H/C/000277/II/0189** UCB Pharma S.A., Rapporteur: Karin Janssen van Doorn, PRAC Rapporteur: Laurence de Fays, PRAC-CHMP liaison: Karin Janssen van Doorn, "Submission of the final report of the PASS EUPAS26595 'Comparing the incidence of acute renal failure in patients with epilepsy exposed to levetiracetam versus other antiepileptics drugs'." Opinion adopted on 01.10.2020.	Positive Opinion adopted by consensus on 01.10.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
PRAC Led **Mysimba - naltrexone hydrochloride /**	Positive Opinion adopted by consensus on 01.10.2020. The Icelandic and Norwegian CHMP

bupropion hydrochloride - EMEA/H/C/003687/II/0044/G Orexigen Therapeutics Ireland Limited, Rapporteur: Kirstine Moll Harboe, Co-Rapporteur: Andrea Laslop, PRAC Rapporteur: Martin Huber, PRAC-CHMP liaison: Janet Koenig, "Update of product information resulting from PRAC Assessment Report request in PSUSA/00010366/201909: - Introduction of a warning concerning the interaction between Naltrexone/Bupropion and Digoxin in SmPC section 4.5 and related PL section. - Update of SmPC section 4.8 and related PL section on drug-induced lupus erythematosus with Naltrexone/Bupropion and its individual substances." Opinion adopted on 01.10.2020.	Members were in agreement with the CHMP recommendation.
PRAC Led **NovoEight - turoctocog alfa - EMEA/H/C/002719/II/0035** Novo Nordisk A/S, Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Brigitte Keller-Stanislawski, PRAC-CHMP liaison: Jan Mueller-Berghaus, "Update of section 5.1 of the SmPC to include the results of the completed study PASS Guardian 5 NN70083553. The RMP has been updated accordingly." Opinion adopted on 01.10.2020.	Positive Opinion adopted by consensus on 01.10.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
PRAC Led **Ratiograstim - filgrastim - EMEA/H/C/000825/II/0069** ratiopharm GmbH, Rapporteur: Outi Mäki-Ikola, PRAC Rapporteur: Kirsti Villikka, PRAC-CHMP liaison: Outi Mäki-Ikola, "Submission of an updated RMP version 10.0 in order to remove the additional pharmacovigilance activity "Cooperation with SCNIR (Severe Chronic Neutropenia International Registry) and analysis of corresponding Ratiograstim/Tevagrastim-SCNIR data"." Opinion adopted on 01.10.2020.	Positive Opinion adopted by consensus on 01.10.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
PRAC Led **Tecentriq - atezolizumab - EMEA/H/C/004143/II/0048** Roche Registration GmbH, Rapporteur: Sinan B. Sarac, PRAC Rapporteur: Marcia Sofia Sanches de Castro Lopes Silva, PRAC-CHMP liaison: Bruno Sepodes, "Submission of the results of	Request for supplementary information adopted with a specific timetable.

study WO41486 evaluating the effectiveness of the HCP brochure designed to mitigate important immune-related risks in patients receiving atezolizumab in the European Union. As a consequence, the MAH is updating section 4.4 of the SmPC, Annex II.D and the RMP. In addition, the MAH is proposing a delay in the due date for the submission of the CSR for IMvigor210 ." Request for Supplementary Information adopted on 01.10.2020.	
PRAC Led **Xadago - safinamide - EMEA/H/C/002396/II/0035** Zambon S.p.A., Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Rhea Fitzgerald, PRAC-CHMP liaison: Peter Kiely, "C.I.13 - Results of a Drug utilisation study (DUS) and changes to RMP." Opinion adopted on 01.10.2020. Request for Supplementary Information adopted on 14.05.2020.	Positive Opinion adopted by consensus on 01.10.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
PRAC Led **Xarelto - rivaroxaban - EMEA/H/C/000944/II/0080** Bayer AG, Rapporteur: Kristina Dunder, PRAC Rapporteur: Ulla Wändel Liminga, PRAC-CHMP liaison: Kristina Dunder, "Submission of the final report from a Survey on Prescribers' Guide/Patient Alert listed as a category 3 study in the RMP." Opinion adopted on 01.10.2020.	Positive Opinion adopted by consensus on 01.10.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
PRAC Led **WS1919 Lyrica-EMEA/H/C/000546/WS1919/0109 Pregabalin Pfizer-EMEA/H/C/003880/WS1919/0038** Upjohn EESV, Lead Rapporteur: Johann Lodewijk Hillege, Lead PRAC Rapporteur: Liana Gross-Martirosyan, PRAC-CHMP liaison: Johann Lodewijk Hillege, "Update to the Risk Management Plan to include results of recently completed PASS studies." Request for Supplementary Information adopted on 01.10.2020.	Request for supplementary information adopted with a specific timetable.

B.5.5. CHMP-CAT assessed procedures

Kymriah - tisagenlecleucel -	Request for supplementary information adopted

EMEA/H/C/004090/II/0026/G, Orphan, ATMP Novartis Europharm Limited, Rapporteur: Rune Kjeken, CHMP Coordinator: Ingrid Wang Request for Supplementary Information adopted on 09.10.2020.	with a specific timetable.
Kymriah - tisagenlecleucel - EMEA/H/C/004090/II/0027, Orphan, ATMP Novartis Europharm Limited, Rapporteur: Rune Kjeken, CHMP Coordinator: Ingrid Wang Request for Supplementary Information adopted on 09.10.2020.	Request for supplementary information adopted with a specific timetable.
Zolgensma - onasemnogene abeparvovec - EMEA/H/C/004750/II/0003/G, Orphan, ATMP AveXis EU Limited, Rapporteur: Johannes Hendrikus Ovelgonne, CHMP Coordinator: Johann Lodewijk Hillege Opinion adopted on 15.10.2020, 09.10.2020. Request for Supplementary Information adopted on 11.09.2020.	Positive Opinion adopted by consensus on 15.10.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Zolgensma - onasemnogene abeparvovec - EMEA/H/C/004750/II/0006, Orphan, ATMP AveXis EU Limited, Rapporteur: Johannes Hendrikus Ovelgonne, CHMP Coordinator: Johann Lodewijk Hillege Request for Supplementary Information adopted on 09.10.2020.	Request for supplementary information adopted with a specific timetable.

B.5.6. CHMP-PRAC-CAT assessed procedures

Yescarta - axicabtagene ciloleucel - EMEA/H/C/004480/II/0028, Orphan, ATMP Kite Pharma EU B.V., Rapporteur: Jan Mueller-Berghaus, CHMP coordinator: Jan Mueller-Berghaus, PRAC Rapporteur: Anette Kirstine Stark, "To update SmPC sections; 4.4 on CRS grading and neurologic adverse reactions; 4.8 on safety profile summary; 5.1 on follow-up analysis; to update the safety information based on updates from study KTE-C19-101, entitled "A Phase 1/2 Multicenter Study Evaluating the Safety and Efficacy of KTE-C19 in Subjects with Refractory Aggressive Non-Hodgkin Lymphoma (ZUMA-1)", the pivotal study for Yescarta. The updates include the Phase 2 safety management	Request for supplementary information adopted with a specific timetable.

ZUMA-1 Cohort 4, which was intended to assess the impact of earlier interventions (tocilizumab and/or corticosteroids, in addition to prophylactic levetiracetam) on the rate and severity of CRS and neurologic events; and data from a 36-month analysis from ZUMA-1 Cohorts 1 and 2. The updated RMP version 3.1 has also been submitted." Request for Supplementary Information adopted on 09.10.2020.	

B.5.7. PRAC assessed ATMP procedures

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

WS1855/G **Bretaris Genuair-EMEA/H/C/002706/ WS1855/0044/G** **Eklira Genuair-EMEA/H/C/002211/ WS1855/0044/G** AstraZeneca AB, Lead Rapporteur: Ewa Balkowiec Iskra Opinion adopted on 24.09.2020. Request for Supplementary Information adopted on 23.07.2020.	Positive Opinion adopted by consensus on 24.09.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1856/G **Brimica Genuair-EMEA/H/C/003969/ WS1856/0030/G** **Duaklir Genuair-EMEA/H/C/003745/ WS1856/0030/G** AstraZeneca AB, Lead Rapporteur: Ewa Balkowiec Iskra Opinion adopted on 24.09.2020. Request for Supplementary Information adopted on 23.07.2020.	Positive Opinion adopted by consensus on 24.09.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1896 **Keppra-EMEA/H/C/000277/WS1896/0190** UCB Pharma S.A., Lead Rapporteur: Karin Janssen van Doorn, "To update section 4.4, Special warnings and precautions for use, to add a warning "Electrocardiogram QT interval prolongation" and section 4.8 of the SmPC, Undesirable effects to add ADR "Electrocardiogram QT prolonged" following outcome of LEG 088.2. Sections 2 and 4 of the Package Leaflet were updated accordingly." Opinion adopted on 15.10.2020.	Positive Opinion adopted by consensus on 15.10.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

WS1901 **Actraphane-EMEA/H/C/000427/WS1901/0086** **Actrapid-EMEA/H/C/000424/WS1901/0079** **Fiasp-EMEA/H/C/004046/WS1901/0024** **Insulatard-EMEA/H/C/000441/WS1901/0084** **Levemir-EMEA/H/C/000528/WS1901/0100** **Mixtard-EMEA/H/C/000428/WS1901/0087** **NovoMix-EMEA/H/C/000308/WS1901/0106** **NovoRapid-EMEA/H/C/000258/WS1901/0136** **Protaphane-EMEA/H/C/000442/WS1901/0083** **Ryzodeg-EMEA/H/C/002499/WS1901/0041** **Tresiba-EMEA/H/C/002498/WS1901/0048** **Xultophy-EMEA/H/C/002647/WS1901/0038** Novo Nordisk A/S, Lead Rapporteur: Kirstine Moll Harboe, "To implement the risk of cutaneous amyloidosis in sections 4.4 and 4.8 of the SmPC following a PRAC signal assessment (EPITT no. 19499). Sections 2 and 4 of the PL are updated accordingly and also changed to highlight to patients that the precautions they need to take (such as rotating the injection site) are important not only to help preventing cutaneous amyloidosis but also to help preventing lipodystrophy. Additionally, the annexes have been brought in line with the current QRD template (version 10.1)." Opinion adopted on 24.09.2020.	Positive Opinion adopted by consensus on 24.09.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1907/G **Galvus-EMEA/H/C/000771/WS1907/0065/G** **Jalra-EMEA/H/C/001048/WS1907/0067/G** **Xiliarx-EMEA/H/C/001051/WS1907/0065/G** Novartis Europharm Limited, Lead Rapporteur: Kristina Dunder Opinion adopted on 01.10.2020.	Positive Opinion adopted by consensus on 01.10.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1910/G **Filgrastim Hexal-EMEA/H/C/000918/**	Positive Opinion adopted by consensus on 15.10.2020. The Icelandic and Norwegian CHMP

WS1910/0058/G **Zarzio-EMEA/H/C/000917/WS1910/ 0059/G** Sandoz GmbH, Lead Rapporteur: Johann Lodewijk Hillege Opinion adopted on 15.10.2020.	Members were in agreement with the CHMP recommendation.
WS1916 **Cegfila-EMEA/H/C/005312/WS1916/0005** **Pelmeg-EMEA/H/C/004700/WS1916/0009** Mundipharma Corporation (Ireland) Limited, Lead Rapporteur: Karin Janssen van Doorn Opinion adopted on 08.10.2020.	Positive Opinion adopted by consensus on 08.10.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1925 **Filgrastim Hexal-EMEA/H/C/000918/ WS1925/0057** **Zarzio-EMEA/H/C/000917/WS1925/0058** Sandoz GmbH, Lead Rapporteur: Johann Lodewijk Hillege Opinion adopted on 15.10.2020.	Positive Opinion adopted by consensus on 15.10.2020. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

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

WS1894/G **Incresync-EMEA/H/C/002178/WS1894/ 0032/G** **Vipdomet-EMEA/H/C/002654/WS1894/ 0028/G** **Vipidia-EMEA/H/C/002182/WS1894/ 0023/G** Takeda Pharma A/S, Lead Rapporteur: Johann Lodewijk Hillege Request for Supplementary Information adopted on 03.09.2020. Withdrawal request submitted on 05.10.2020.	The MAH withdrew the procedure on 05.10.2020.
WS1918 **Kinzalkomb-EMEA/H/C/000415/WS1918/ 0115** **MicardisPlus-EMEA/H/C/000413/WS1918/ 0118** **PritorPlus-EMEA/H/C/000414/WS1918/ 0125** Boehringer Ingelheim International GmbH, Lead Rapporteur: Armando Genazzani, "Update of sections 4.2, 4.3, 4.5, 4.7, 4.8, 5.1 and 5.2 of the SmPC in order to amend posology recommendations in elderly population, add new contraindications, delete the mono components hydrochlorothiazide and telmisartan list of adverse drug reactions (ADRs), and update the	The MAH withdrew the procedure on 30.09.2020.

description of pharmacodynamic effects of individual components, following the update of the originator reference product information (PI). In addition, the MAH took the opportunity to introduce changes as per the Excipients Guideline and editorial changes to bring the PI in line with the latest QRD template version 10.1. The Package Leaflet and the Labelling are updated accordingly." Withdrawal request submitted on 30.09.2020.	

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

Mysimba - naltrexone hydrochloride / bupropion hydrochloride - EMEA/H/C/003687/II/0042 Orexigen Therapeutics Ireland Limited, Rapporteur: Kirstine Moll Harboe Request for Supplementary Information adopted on 23.07.2020.	Request by the applicant for an extension of the clock-stop to respond to the RSI adopted on 23.07.2020. The CHMP agreed to the request by the applicant.

B.6. START OF THE PROCEDURES TIMETABLES FOR INFORMATION

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

abrocitinib - EMEA/H/C/005452 Indicated for the treatment of moderate-to-severe atopic dermatitis in adults and adolescents 12 years and older who are candidates for systemic therapy.
artesunate - EMEA/H/C/005550, Orphan Amivas Ireland Ltd, treatment of malaria
avalglucosidase alfa - EMEA/H/C/005501, Orphan Genzyme Europe BV, for long-term enzyme replacement therapy for the treatment of patients with Pompe disease.
ranibizumab - EMEA/H/C/005545 treatment of neovascular age-related macular degeneration (AMD)
lonapegsomatropin - EMEA/H/C/005367, Orphan Ascendis Pharma Endocrinology Division A/S, Treatment of growth hormone deficiency
adalimumab - EMEA/H/C/005548 treatment of rheumatoid arthritis, juvenile idiopathic arthritis, axial spondyloarthritis,

psoriatic arthritis, psoriasis, paediatric plaque psoriasis, hidradenitis suppurativa, Crohn's disease, paediatric Crohn's disease, ulcerative colitis, uveitis, paediatric uveitis	
teriparatide - EMEA/H/C/004932 Treatment of osteoporosis in postmenopausal women and in men at increased risk of fracture	
pegcetacoplan - EMEA/H/C/005553, Orphan Apellis Ireland Limited, paroxysmal nocturnal haemoglobinuria (PNH)	
ripretinib - EMEA/H/C/005614, Orphan Deciphera Pharmaceuticals (Netherlands) B.V., Treatment of patients with advanced gastrointestinal stromal tumour (GIST)	
rivaroxaban - EMEA/H/C/005600 Prevention of venous thromboembolism in adult patients undergoing elective hip or knee replacement surgery and treatment of deep vein thrombosis and pulmonary embolism as well as prevention of recurrent DVT and PE in adults. Prevention of stroke and systemic embolism in adult patients with non-valvular atrial fibrillation with one or more risk factors.	
autologous glioma tumor cells, inactivated / autologous glioma tumor cell lysates, inactivated / allogeneic glioma tumor cells, inactivated / allogeneic glioma tumor cell lysates, inactivated - EMEA/H/C/003693, Orphan, ATMP Epitopoietic Research Corporation-Belgium (E.R.C.), treatment of glioma	
elivaldogene autotemcel - EMEA/H/C/003690, Orphan, ATMP bluebird bio (Netherlands) B.V, treatment of ABCD1 genetic mutation and cerebral adrenoleukodystrophy	**Accelerated review**

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

Akynzeo - fosnetupitant / netupitant / palonosetron - EMEA/H/C/003728/X/0031 Helsinn Birex Pharmaceuticals Limited, Rapporteur: Peter Kiely, PRAC Rapporteur: Ilaria Baldelli, "Extension application to introduce a new pharmaceutical form	

(concentrate for solution for infusion)."	
Noxafil - posaconazole - EMEA/H/C/000610/X/0063/G Merck Sharp & Dohme B.V., Rapporteur: Alexandre Moreau, PRAC Rapporteur: Adrien Inoubli, "Extension application to introduce a new pharmaceutical form (gastro-resistant powder and solvent for oral suspension), grouped with a type II variation (C.I.6.a) to extend the approved indications for Noxafil to the paediatric population. As a consequence, sections 4.1, 4.2, 4.8, 5.1 and 5.2 of the SmPC, as well as the package leaflet, are updated. The RMP (version 17.1) is updated in accordance."	

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

bevacizumab - EMEA/H/C/005327 Treatment of metastatic carcinoma of the colon or rectum, metastatic breast cancer and recurrence of platinum-sensitive epithelial ovarian, fallopian tube or primary peritoneal cancer. First-line treatment of patients with unresectable advanced, metastatic or recurrent non-small cell lung cancer. First-line treatment of patients with advanced and/or metastatic renal cell cancer. List of Questions adopted on 23.07.2020.	
bevacizumab - EMEA/H/C/005611 Treatment of metastatic carcinoma of the colon or rectum, metastatic breast cancer and recurrence of platinum-sensitive epithelial ovarian, fallopian tube or primary peritoneal cancer. First-line treatment of patients with unresectable advanced, metastatic or recurrent non-small cell lung cancer. First-line treatment of patients with advanced and/or metastatic renal cell cancer. List of Questions adopted on 23.07.2020.	
cenobamate - EMEA/H/C/005377 for the adjunctive treatment of focal onset seizures with or without secondary generalisation in adult patients with epilepsy who have not been adequately controlled despite a history of treatment with at least 2	

anti-epileptic products.
List of Questions adopted on 23.07.2020.

dostarlimab - EMEA/H/C/005204
treatment of mismatch repair deficient (dMMR)/microsatellite instability-high (MSI-H) endometrial cancer (EC)
List of Questions adopted on 23.06.2020.

trastuzumab deruxtecan - EMEA/H/C/005124
treatment of unresectable or metastatic HER2-positive breast cancer
List of Questions adopted on 15.09.2020.

berotralstat - EMEA/H/C/005138, Orphan
BioCryst Ireland Limited, prevention of hereditary angioedema (HAE)
List of Questions adopted on 23.07.2020.

sildenafil - EMEA/H/C/005439
treatment of erectile dysfunction
List of Questions adopted on 25.06.2020.

thiotepa - EMEA/H/C/005434
conditioning treatment prior to haematopoietic progenitor cell transplantation (HPCT), treatment of solid tumours
List of Questions adopted on 17.09.2020.

Trimbow - beclometasone dipropionate / formoterol fumarate dihydrate / glycopyrronium - EMEA/H/C/004257/X/0012
Chiesi Farmaceutici S.p.A., Rapporteur: Janet Koenig, Co-Rapporteur: Peter Kiely, PRAC Rapporteur: Jan Neuhauser, "Extension application to add a new pharmaceutical form (inhalation powder) associated with new strength (88µg / 5µg / 9µg).
The RMP (version 6.2) is updated in accordance."
List of Questions adopted on 23.07.2020.

Xerava - eravacycline - EMEA/H/C/004237/X/0009
Tetraphase Pharmaceuticals Ireland Limited, Rapporteur: Filip Josephson, PRAC Rapporteur: Adam Przybylkowski, "Extension application to add a new strength of 100 mg for eravacycline powder for concentrate for solution for infusion.
The RMP (version 3.0) is updated in accordance.
Additionally, the marketing authorisation holder took the opportunity to align the PI with the

latest QRD template."
List of Questions adopted on 17.09.2020.

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

Bosulif - bosutinib -
EMEA/H/C/002373/R/0045
Pfizer Europe MA EEIG, Rapporteur: Janet Koenig, Co-Rapporteur: Blanca Garcia-Ochoa, PRAC Rapporteur: Martin Huber

Cometriq - cabozantinib -
EMEA/H/C/002640/R/0042, Orphan
Ipsen Pharma, Rapporteur: Paula Boudewina van Hennik, Co-Rapporteur: Bjorg Bolstad, PRAC Rapporteur: Menno van der Elst

Deltyba - delamanid -
EMEA/H/C/002552/R/0047, Orphan
Otsuka Novel Products GmbH, Rapporteur: Christophe Focke, PRAC Rapporteur: Laurence de Fays

Ondexxya - andexanet alfa -
EMEA/H/C/004108/R/0015
Portola Netherlands B.V., Rapporteur: Jan Mueller-Berghaus, Co-Rapporteur: Maria Concepcion Prieto Yerro, PRAC Rapporteur: Menno van der Elst

WAYLIVRA - volanesorsen -
EMEA/H/C/004538/R/0009, Orphan
Akcea Therapeutics Ireland Limited, Rapporteur: Johann Lodewijk Hillege, Co-Rapporteur: Karin Janssen van Doorn, PRAC Rapporteur: Martin Huber

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

AMGEVITA - adalimumab -
EMEA/H/C/004212/II/0023
Amgen Europe B.V., Rapporteur: Kristina

Dunder

Beovu - brolucizumab -
EMEA/H/C/004913/II/0005/G
Novartis Europharm Limited, Rapporteur: Alexandre Moreau

Beromun - tasonermin -
EMEA/H/C/000206/II/0050
Belpharma s.a., Rapporteur: Sinan B. Sarac

Beromun - tasonermin -
EMEA/H/C/000206/II/0051
Belpharma s.a., Rapporteur: Sinan B. Sarac

Cerezyme - imiglucerase -
EMEA/H/C/000157/II/0118
Genzyme Europe BV, Rapporteur: Johann Lodewijk Hillege

Cinqaero - reslizumab -
EMEA/H/C/003912/II/0037/G
Teva B.V., Rapporteur: Johann Lodewijk Hillege

Cinryze - C1 esterase inhibitor (human) -
EMEA/H/C/001207/II/0082/G
Shire Services BVBA, Rapporteur: Jan Mueller-Berghaus

Hulio - adalimumab -
EMEA/H/C/004429/II/0021
Mylan S.A.S, Rapporteur: Christophe Focke

Kentera - oxybutynin -
EMEA/H/C/000532/II/0059
Teva B.V., Rapporteur: Karin Janssen van Doorn

Kyprolis - carfilzomib -
EMEA/H/C/003790/II/0050/G, Orphan
Amgen Europe B.V., Rapporteur: Blanca Garcia-Ochoa

Lonquex - lipegfilgrastim -
EMEA/H/C/002556/II/0060
Teva B.V., Rapporteur: Outi Mäki-Ikola

Lyumjev - insulin lispro -
EMEA/H/C/005037/II/0006/G
Eli Lilly Nederland B.V., Rapporteur: Outi Mäki-Ikola

M-M-RVAXPRO - measles, mumps and rubella vaccine (live) -
EMEA/H/C/000604/II/0103
MSD Vaccins, Rapporteur: Jan Mueller-Berghaus

Palynziq - pegvaliase -

EMEA/H/C/004744/II/0014, Orphan
BioMarin International Limited, Rapporteur: Johann Lodewijk Hillege

Pemetrexed Sandoz - pemetrexed - EMEA/H/C/004011/II/0009
Sandoz GmbH, Generic, Generic of Alimta, Rapporteur: Bjorg Bolstad

Privigen - human normal immunoglobulin - EMEA/H/C/000831/II/0169
CSL Behring GmbH, Rapporteur: Jan Mueller-Berghaus

ProQuad - measles, mumps, rubella and varicella vaccine (live) - EMEA/H/C/000622/II/0143
MSD Vaccins, Rapporteur: Jan Mueller-Berghaus

Puregon - follitropin beta - EMEA/H/C/000086/II/0111/G
Merck Sharp & Dohme B.V., Rapporteur: Peter Kiely

Puregon - follitropin beta - EMEA/H/C/000086/II/0112/G
Merck Sharp & Dohme B.V., Rapporteur: Peter Kiely

Reblozyl - luspatercept - EMEA/H/C/004444/II/0002/G, Orphan
Celgene Europe BV, Rapporteur: Milena Stain

Thyrogen - thyrotropin alfa - EMEA/H/C/000220/II/0106
Genzyme Europe BV, Rapporteur: Peter Kiely

Voncento - human coagulation factor viii / human von willebrand factor - EMEA/H/C/002493/II/0045
CSL Behring GmbH, Rapporteur: Paula Boudewina van Hennik

Xolair - omalizumab - EMEA/H/C/000606/II/0105/G
Novartis Europharm Limited, Rapporteur: Kristina Dunder

Yervoy - ipilimumab - EMEA/H/C/002213/II/0086
Bristol-Myers Squibb Pharma EEIG, Rapporteur: Paula Boudewina van Hennik

WS1906/G
Hexacima-EMEA/H/C/002702/WS1906/0108/G

Hexaxim-EMEA/H/W/002495/WS1906/0113/G
Hexyon-EMEA/H/C/002796/WS1906/0112/G
Sanofi Pasteur, Lead Rapporteur: Jan Mueller-Berghaus

WS1914/G
Mircera-EMEA/H/C/000739/WS1914/0080/G
NeoRecormon-EMEA/H/C/000116/WS1914/0109/G
Roche Registration GmbH, Lead Rapporteur: Martina Weise

WS1924
HyQvia-EMEA/H/C/002491/WS1924/0064
Kiovig-EMEA/H/C/000628/WS1924/0105
Takeda Manufacturing Austria AG, Lead Rapporteur: Jan Mueller-Berghaus

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

AUBAGIO - teriflunomide - EMEA/H/C/002514/II/0033
sanofi-aventis groupe, Rapporteur: Martina Weise, "To update sections 4.4 and 4.8 of the SmPC regarding skin reactions in particular to drug reaction with eosinophilia and systemic symptoms (DRESS) and to update the frequency of severe skin reactions from "Not known" to "Uncommon", following a review of the Sanofi global PV database. The Package Leaflet section 4 is updated as to add fever."

Darzalex - daratumumab - EMEA/H/C/004077/II/0041, Orphan
Janssen-Cilag International NV, Rapporteur: Sinan B. Sarac, "C.I.4
Update of section 4.8 of the SmPC in order to include CMV infections as a new adverse drug reaction (ADR) with frequency common following a comprehensive, cross-program evaluation of all potential cases of treatment-emergent cytomegalovirus (CMV) infections with use of daratumumab. The Package Leaflet is updated accordingly. Several minor linguistic improvements are also proposed."

Dengvaxia - dengue tetravalent vaccine (live, attenuated) - EMEA/H/C/004171/II/0013

Sanofi Pasteur, Rapporteur: Christophe Focke, "To update sections 4.2, 4.8 and 5.1 of the SmPC in order to introduce a new posology regimen from 3 doses to 2 doses for individuals from 9 years of age based on interim results from study CYD65 listed as a category 3 study in the RMP; this is a Phase II, observer-blind, placebo-controlled trial in order to assess Immunogenicity and Safety of Tetravalent Dengue Vaccine Given in 1-, 2-, or 3-Dose Schedules Followed by a Single Booster; section 3 of the Package Leaflet is updated accordingly"

Foclivia - influenza virus surface antigens (inactivated) of strain A/Vietnam/1194/2004 (H5N1) - EMEA/H/C/001208/II/0058

Seqirus S.r.l, Rapporteur: Armando Genazzani, "Update of sections 2, 4.2-4.8, 5.1, 6.4 and 6.5 of the SmPC based on data obtained from two clinical trials (V87_25 and V87_26) already assessed and approved for Aflunov, the corresponding H5N1 Zoonotic Influenza Vaccine (procedure EMEA/H/C/002094/II/0044/G approved in June 2019) in order to align both products. In addition, the MAH took the opportunity to also add in section 5.1 data from study V87P2 and study V87P11 (A/Turkey/turkey/1/2005) already evaluated for the RMP of both products and included in the label of Aflunov for further alignment; the Package Leaflet and Labelling are updated accordingly. Finally, the Marketing authorisation holder (MAH) makes additional changes based on the most recent
EU Guidelines and some additional minor editorial corrections.
The requested variation proposed amendments to the Summary of Product Characteristics, Labelling and Package Leaflet."

Genvoya - elvitegravir / cobicistat / emtricitabine / tenofovir alafenamide - EMEA/H/C/004042/II/0070/G

Gilead Sciences Ireland UC, Rapporteur: Bruno Sepodes, "- C.I.4 (Type II): Update of section 4.5 of the SmPC to include data on the drug-drug interaction between Genvoya and the thienopyridine anti-platelet drugs clopidogrel and prasugrel, based on a MAH cumulative safety review. The Package Leaflet is updated accordingly.

- C.I.4 (Type II): Update of section 4.5 of the SmPC to include data on the drug-drug interaction between Genvoya and medicinal products or oral supplements containing polyvalent cations, based on a MAH cumulative safety review. The Package Leaflet is updated accordingly.
In addition, the Marketing Authorisation Holder (MAH) took the opportunity to correct the amount of lactose stated in section 2 of the SmPC and make minor editorial changes throughout the PI."

Imfinzi - durvalumab - EMEA/H/C/004771/II/0024

AstraZeneca AB, Rapporteur: Sinan B. Sarac, "Update of sections 4.4 and 4.8 of the SmPC in order to add immune thrombocytopenia to the list of adverse drug reactions (ADRs) with frequency (rare) following the MAH internal review; the Package Leaflet (PL) is updated accordingly. The MAH took the opportunity to correct information in the PL and to make editorial changes to the names of the manufacturer in Annex II."

Invanz - ertapenem - EMEA/H/C/000389/II/0062

Merck Sharp & Dohme B.V., Rapporteur: Fátima Ventura, "Update of section 4.8 of the SmPC in order to add 'hypersensitivity vasculitis' to the list of adverse drug reactions (ADRs) with frequency 'Not known', based on post-marketing reports; the Package Leaflet is updated accordingly. In addition, the MAH took the opportunity to update the list of local representatives in the Package Leaflet, update the PI in line with the Annex to the European Commission guideline on `Excipients in the labelling and package leaflet of medicinal products for human use' (EMA/CHMP/302620/2017 Rev.1) regarding sodium content, and bring the PI in line with the latest QRD template version 10.1. The MAH also updated the package leaflet to add the missing adverse event "injection site induration" with frequency 'Rare'."

Jinarc - tolvaptan - EMEA/H/C/002788/II/0031

Otsuka Pharmaceutical Netherlands B.V., Rapporteur: Armando Genazzani, "Update of

sections 4.2 and 4.4 of the SmPC in order to include information on patients with CKD late stage 4 based on final results from study 156-12-211 listed as a category 3 study in the RMP; this is a Phase 3b, Multicenter, Open-label Trial to Evaluate the Long Term Safety of Immediate-release Tolvaptan (OPC-41061, 30 mg to 120 mg/day, Split dose) in Subjects with Autosomal Dominant Polycystic Kidney Disease."	
Keytruda - pembrolizumab - EMEA/H/C/003820/II/0094 Merck Sharp & Dohme B.V., Rapporteur: Armando Genazzani, "Update of sections 4.4 and 5.1 of the SmPC in order to update efficacy information based on final results from study KEYNOTE-361 listed as a PAES in the Annex II; this is a Phase III Randomised, Controlled Clinical Trial of Pembrolizumab with or without Platinum-based Combination Chemotherapy versus Chemotherapy in Subjects with Advanced or Metastatic Urothelial Carcinoma; Annex IID is updated accordingly."	
Lorviqua - lorlatinib - EMEA/H/C/004646/II/0009/G Pfizer Europe MA EEIG, Rapporteur: Sinan B. Sarac, "Update of sections 4.2 and 5.2 of the SmPC in order to change the posology recommendations in patients with severe renal impairment based on the results from Study B7461010 (a phase 1, single dose open-label study to evaluate the pharmacokinetics of lorlatinib in subjects with impaired renal function). The package leaflet has been updated accordingly. Update of sections 4.4 and 4.5 of the SmPC in order to include information regarding drug-drug interaction with moderate CYP3A4/5 inducers based on study B7461026 (Phase 1, open-label, fixed sequence, 2-period study to investigate the effect of multiple doses of modafinil on the pharmacokinetics of single dose lorlatinib in healthy participants). The MAH took the opportunity to update the list of local representatives in the Package Leaflet."	
Maviret - glecaprevir / pibrentasvir - EMEA/H/C/004430/II/0037 AbbVie Deutschland GmbH & Co. KG, Rapporteur: Jean-Michel Race, "Submission of the final clinical study report from study B16-	

439 (Phase 3b, a Multi-Center, Randomized, Open-Label, Pragmatic Study of Glecaprevir/Pibrentasvir (G/P) +/- Ribavirin for GT1 Subjects with Chronic Hepatitis C Previously Treated with an NS5A Inhibitor + Sofosbuvir Therapy)."

Mekinist - trametinib - EMEA/H/C/002643/II/0041

Novartis Europharm Limited, Rapporteur: Paula Boudewina van Hennik, "Update of sections 4.5, 4.6 and 5.2 of the SmPC in order to add drug-drug interaction information with hormonal contraceptives and to update relevant part of the SmPC regarding this interaction; the Package Leaflet is updated accordingly. Furthermore, the MAH took the occasion to include the information regarding the sodium content in the products in line with relevant guidelines and to bring the PI in line with the latest QRD template version 10.1. In addition, the MAH took the opportunity to introduce some editorial changes in the PI and to update the list of local representatives for the Netherlands in the Package Leaflet."

NINLARO - ixazomib - EMEA/H/C/003844/II/0025, Orphan

Takeda Pharma A/S, Rapporteur: Armando Genazzani, "Update of the SmPC section 4.9 with additional information on Ninlaro overdose. The Package leaflet is updated accordingly."

Palynziq - pegvaliase - EMEA/H/C/004744/II/0015, Orphan

BioMarin International Limited, Rapporteur: Johann Lodewijk Hillege, "Submission of the final report from the non-clinical study BMN-165-18-080 listed as a category 3 study in the RMP."

Parsabiv - etelcalcetide - EMEA/H/C/003995/II/0015

Amgen Europe B.V., Rapporteur: Johann Lodewijk Hillege, Co-Rapporteur: Andrea Laslop (AT) (MNAT with AT for Coordination, AT for Clinical Safety, AT for Non-Clinical, AT for Clinical Pharmacology, AT for Clinical Efficacy, DE-BfArM for Quality), "Update of the SmPC section 4.4 to remove anti-etelcalcetide antibodies testing, and update of the Product information in line with QRD template v10.1."

Perjeta - pertuzumab -
EMEA/H/C/002547/II/0053
Roche Registration GmbH, Rapporteur: Sinan B. Sarac, "Submission of the final report from study MO27775 (PERTAIN). This is a randomized, two-arm, open-label, multicenter Phase II trial assessing the efficacy and safety of pertuzumab given in combination with trastuzumab plus an aromatase inhibitor in first line patients with HER2-positive and hormone receptor-positive advanced (metastatic or locally advanced) breast cancer."

Qutenza - capsaicin -
EMEA/H/C/000909/II/0051/G
Grunenthal GmbH, Rapporteur: Bruno Sepodes, "Update of sections 4.2 and 4.4 to delete the explicit reference to pre-treatments used in clinical trials and to opioids. Update of section 4.4 of the SmPC to include more detail on unintended exposure to capsaicin."

Remsima - infliximab -
EMEA/H/C/002576/II/0095
Celltrion Healthcare Hungary Kft., Rapporteur: Outi Mäki-Ikola, PRAC Rapporteur: Kimmo Jaakkola, "Addition of a new posology for the rheumatoid arthritis indication that does not include IV induction doses."

Rubraca - rucaparib -
EMEA/H/C/004272/II/0024/G
Clovis Oncology Ireland Limited, Rapporteur: Blanca Garcia-Ochoa, "Submission of the final reports from four non-clinical studies (Report 181000, OPT-2018-074, 8388100 and CLO-P8799)."

Stelara - ustekinumab -
EMEA/H/C/000958/II/0083
Janssen-Cilag International NV, Rapporteur: Jayne Crowe, "Update of section 4.8 of the SmPC in order to add hypersensitivity vasculitis to the list of adverse drug reactions (ADRs) with frequency rare based on cumulative review from the literature and post-marketing reporting; the Package Leaflet is updated accordingly."

Stivarga - regorafenib -
EMEA/H/C/002573/II/0031
Bayer AG, Rapporteur: Paula Boudewina van Hennik, "Submission of final study report for study 15982, a randomized, double blind,

placebo-controlled, multicenter Phase 3 study that investigated regorafenib in subjects with hepatocellular carcinoma (HCC) after progression on sorafenib treatment."

Tecentriq - atezolizumab - EMEA/H/C/004143/II/0050

Roche Registration GmbH, Rapporteur: Sinan B. Sarac, "Update of section 5.1 of the SmPC in order to reflect efficacy results based on the final OS analysis from study WO29522 (IMpassion130) comparing atezolizumab in combination with nab-paclitaxel with placebo with nab-paclitaxel for patients with previously untreated metastatic triple-negative breast cancer."

Veklury - remdesivir - EMEA/H/C/005622/II/0012

Gilead Sciences Ireland UC, Rapporteur: Janet Koenig, "Submission of the final D28 mortality data by ordinal scale categories of study COUS-540-5776 (NIAID-ACTT1), listed as a Specific Obligation in the Annex II of the Product Information, in order to confirm the efficacy and safety of remdesivir in patients on Invasive Mechanical Ventilation and Extracorporeal Membrane Oxygenation (IMV/ECMO). In addition, the MAH discusses the potential imbalance in the use of corticosteroids and effect modification in study CO-US-540-5776. As a consequence, Annex II is updated accordingly."

Vyxeos liposomal - daunorubicin / cytarabine - EMEA/H/C/004282/II/0014, Orphan

Jazz Pharmaceuticals Ireland Limited, Rapporteur: Outi Mäki-Ikola, "Update of section 5.1 of the SmPC to reflect the 5-years overall survival data from the Follow-Up Phase of the Phase 3 Study CPX310-301. Additionally, the MAH has introduced minor editorial changes in the PI."

Xtandi - enzalutamide - EMEA/H/C/002639/II/0050

Astellas Pharma Europe B.V., Rapporteur: Maria Concepcion Prieto Yerro, "Update of section 4.8 of the SmPC in order to add severe skin reactions to the list of adverse drug reactions (ADRs) with frequency not known based on a safety review; the Package Leaflet is updated

accordingly. In addition, the MAH took the opportunity to make minor corrections in the SmPC."

Zerbaxa - ceftolozane / tazobactam - EMEA/H/C/003772/II/0032

Merck Sharp & Dohme B.V., Rapporteur: Bjorg Bolstad, "Update of section 5.1 of the SmPC for Zerbaxa to implement the EUCAST MIC breakpoints of ceftolozanetazobactam for Enterobacterales according to the EUCAST Clinical breakpoints table v. 10.0, valid from January 2020.
In addition, the MAH took the opportunity to update the List of Local representatives in the Package leaflet."

WS1877
Invega-EMEA/H/C/000746/WS1877/0068
Paliperidone Janssen-Cilag International-EMEA/H/C/005486/WS1877/0001
Trevicta-EMEA/H/C/004066/WS1877/0026
Xeplion-EMEA/H/C/002105/WS1877/0051

Janssen-Cilag International NV, Lead Rapporteur: Kristina Dunder, "to add a new post-marketing adverse drug reaction (ADR) " Stevens-Johnson syndrome/toxic epidermal necrolysis" to section 4.8 (Undesirable effects) of the Summary of Product Characteristics (SmPC) for 4 centrally authorised medicinal products (Invega/Xeplion/Trevicta/Paliperidone Janssen-Cilag International) and 2 medicinal products authorised via MRP (Risperdal Oral and Risperdal Consta). Section 4 of the Package Leaflet (PL) for each medicinal product is also amended accordingly."

WS1917
Kivexa-EMEA/H/C/000581/WS1917/0087
Triumeq-EMEA/H/C/002754/WS1917/0085
Trizivir-EMEA/H/C/000338/WS1917/0119
Ziagen-EMEA/H/C/000252/WS1917/0114

ViiV Healthcare B.V., Lead Rapporteur: Jean-Michel Race, "Update of sections 4.4 of the SmPC (for Ziagen, Kivexa, Trizivir and Triumeq) and 5.2 (for Triumeq only) to add new information about the drug-drug interactions between abacavir and riociguat. The Package Leaflet is updated accordingly. Furthermore, the

MAH took the opportunity to introduce an excipient update for Ziagen, Kivexa and Trizivir in line with the SmPC guideline, a syringe instruction update in the Package Leaflet of Ziagen and a revised statement in section 6.6 of the SmPC for Triumeq in line with the QRD template. Moreover, minor editorial updates have been introduced throughout the Product Information of all four products."

WS1922
Lixiana-EMEA/H/C/002629/WS1922/0028
Roteas-EMEA/H/C/004339/WS1922/0016
Daiichi Sankyo Europe GmbH, Lead Rapporteur: Maria Concepcion Prieto Yerro, "Update of sections 4.2 and 5.2 of the SmPC, based on data from the bioavailability study DU176b-A-U166, in order to add information about the bioavailability of edoxaban crushed tablet administered by nasogastric tube or in apple puree and ingested versus the current tablet formulation in healthy subjects. The Package Leaflet has been updated accordingly. In addition, the MAH took the opportunity to correct typos in Lithuanian, Slovakian and Portuguese versions of the SmPC, Labelling and Package Leaflet."

WS1939
Vfend-EMEA/H/C/000387/WS1939/0139
Pfizer Europe MA EEIG, Lead Rapporteur: Johann Lodewijk Hillege, "Update of sections 4.3 and 4.5 of the SmPC in order to add a new contraindication with ivabradine to the contraindications, and add drug-drug interaction information between voriconazole and ivabradine and venetoclax to the Interactions section. The Package Leaflet is updated accordingly.
In addition, the WSA took the opportunity to align with the current Annex to the European Commission guideline on `Excipients in the labelling and package leaflet of medicinal products for human use' (SANTE-2017-11668), 22 November 2019, EMA/CHMP/302620/2017 Rev. 1, for lactose, and to update the list of local representatives in the Package Leaflet."

B.6.10. CHMP-PRAC assessed procedures

Lynparza - olaparib -

EMEA/H/C/003726/II/0042

AstraZeneca AB, Rapporteur: Alexandre Moreau, PRAC Rapporteur: Amelia Cupelli, "Update of sections 4.4, 4.8 and 5.1 of the SmPC in order to add Myelodysplastic syndrome (MDS)/Acute myeloid leukaemia (AML) to the list of adverse drug reactions with the frequency uncommon, modify the existing warning on MDS/AML and update efficacy information based on final results from study SOLO-2 listed as a PAES in the Annex II; this is a phase III randomised, double blind, placebo controlled, multicentre study of olaparib maintenance monotherapy in platinum sensitive relapsed BRCA mutated ovarian cancer patients who are in complete or partial response following platinum based chemotherapy; the Package Leaflet and Annex II are updated accordingly. The RMP version 21 has also been submitted."

Lyxumia - lixisenatide -
EMEA/H/C/002445/II/0030

sanofi-aventis groupe, Rapporteur: Kristina Dunder, PRAC Rapporteur: Annika Folin, "Submission of the final report from study TDR14311 listed as a category 3 study in the RMP, and submitted in accordance with article 46. This is a randomized, double-blind, placebo-controlled, dose escalation study on safety, pharmacokinetics and pharmacodynamics of lixisenatide in paediatric patients with Type 2 diabetes mellitus not adequately controlled with metformin and/or basal insulin. The RMP version 6.0 has also been submitted."

Natpar - parathyroid hormone -
EMEA/H/C/003861/II/0026, Orphan

Shire Pharmaceuticals Ireland Limited, Rapporteur: Karin Janssen van Doorn, PRAC Rapporteur: Rhea Fitzgerald, "Submission of the final results of study PAR-C10-008; a long-term open-label study investigating the safety and tolerability of a rhPTH[1-84] for the treatment of adults with hypoparathyroidism – a clinical extension study (RACE). Update of SmPC section 5.1 to reflect 72 month data from the study. Update of the RMP (version 3.0) with the completed study results, to remove this study as an additional pharmacovigilance activity and to align with the GVP module V Rev 2."

Rubraca - rucaparib -

EMEA/H/C/004272/II/0023
Clovis Oncology Ireland Limited, Rapporteur: Blanca Garcia-Ochoa, PRAC Rapporteur: Annika Folin, "Update of sections 4.5, 4.6 and 5.2 of the SmPC to add drug-drug interaction (DDI) information with rosuvastatin and oral contraceptives based on the results of study CO-338-095 listed as a category 3 study in the RMP; Study CO-338-095 is a phase 1, open-label, DDI study to determine the effect of rucaparib on the pharmacokinetics of oral rosuvastatin (Arm A) and oral contraceptives (ethinylestradiol and levonorgestrel - Arm B) in patients with advanced solid tumors. The RMP version 4.1 has also been submitted."

Shingrix - herpes zoster vaccine (recombinant, adjuvanted) - EMEA/H/C/004336/II/0037
GlaxoSmithkline Biologicals SA, Rapporteur: Christophe Focke, PRAC Rapporteur: Sonja Hrabcik, "To update sections 4.4 and 5.1 of the SmPC following the final results from study ZOSTER-064, listed as a category 3 study in the RMP; this is an observational study to assess frailty and other prognostic factors for development of herpes zoster in adult subjects who participated in studies ZOSTER-006 and ZOSTER-022 and the HZ efficacy, immunogenicity and safety of Shingrix by frailty status, in order to fulfil the post-authorisation measure MEA/FSR 012. The updated RMP version 4.1 has also been submitted.
The MAH takes the opportunity to implement some editorial changes in sections 4.4 and 5.1. and correction of the abbreviation CHO cells from Chinese Hamster Ovarian cells to Chinese Hamster Ovary cells in the Annex A."

Xeljanz - tofacitinib - EMEA/H/C/004214/II/0027
Pfizer Europe MA EEIG, Rapporteur: Armando Genazzani, PRAC Rapporteur: Liana Gross-Martirosyan, "Update of sections 4.1, 4.2, 4.4 , 4.8, 5.1 and 5.2 of Xeljanz 11 mg prolonged-release tablets SmPC in order to include the treatment of active psoriatic arthritis (PsA) in adult patients who have had an inadequate response or who have been intolerant to a prior disease modifying antirheumatic drug therapy; as an alternative to the immediate release film-coated tablets; Section 4.2 of Xeljanz film-

coated tablets is also updated to include switching with the prolonged-release tablet in the treatment of PsA. The Package Leaflet is updated accordingly. The RMP version 13.1 has also been submitted."

Zykadia - ceritinib - EMEA/H/C/003819/II/0034

Novartis Europharm Limited, Rapporteur: Blanca Garcia-Ochoa, PRAC Rapporteur: Annika Folin, "Update of sections 4.2, 4.4, 4.8 and 5.1 of the SmPC in order to reflect the results of study CLDK378A2112 as recommended by the CHMP. The study assesses the steady-state PK of 450 mg or 600 mg ceritinib taken daily with a low-fat meal as compared with that of 750 mg ceritinib taken daily in the fasted state in patients with metastatic ALK-positive NSCLC. The Package Leaflet is updated accordingly. The RMP version 16 has also been submitted. In addition, the MAH took the opportunity to bring the PI in line with the latest QRD template version 10.1. Other editorial changes include the addition of the Sodium content in the SmPCs and PLs and the removal of the black triangle."

B.6.11. PRAC assessed procedures

PRAC Led

Brineura - cerliponase alfa - EMEA/H/C/004065/II/0027, Orphan

BioMarin International Limited, Rapporteur: Martina Weise, PRAC Rapporteur: Ulla Wändel Liminga, PRAC-CHMP liaison: Kristina Dunder, "An update to the RMP, to change the final date for the completion of the Post-authorisation efficacy study 190-203."

PRAC Led

Neuraceq - florbetaben (18F) - EMEA/H/C/002553/II/0033

Life Radiopharma Berlin GmbH, Rapporteur: Maria Concepcion Prieto Yerro, PRAC Rapporteur: Martin Huber, PRAC-CHMP liaison: Martina Weise, "C.I.13: Submission of the final report from study PASS2 listed as a category 3 study in the RMP. This is a cross-sectional safety study. The RMP version 5.9 has also been submitted."

PRAC Led

Ontruzant - trastuzumab -
EMEA/H/C/004323/II/0026
Samsung Bioepis NL B.V., Rapporteur: Karin Janssen van Doorn, PRAC Rapporteur: Brigitte Keller-Stanislawski, PRAC-CHMP liaison: Jan Mueller-Berghaus, "Submission of the RMP version 4.0 in order to propose the early termination of long-term observational follow up study for cardiac safety (SB3-G31-BC-E)."

PRAC Led
Pioglitazone Accord - pioglitazone -
EMEA/H/C/002277/II/0020
Accord Healthcare S.L.U., Generic, Generic of Actos, Rapporteur: Peter Kiely, PRAC Rapporteur: Rhea Fitzgerald, PRAC-CHMP liaison: Peter Kiely, "Update of the Risk Management Plan (RMP) for the removal of safety concerns and additional risk minimisation measures (ARRM) as per summary of RMP of Glidipion (pioglitazone; published on 20-Jul-2020), and content adapted to the new GVP Module V (Rev.2)."

PRAC Led
Piqray - alpelisib -
EMEA/H/C/004804/II/0001
Novartis Europharm Limited, Rapporteur: Blanca Garcia-Ochoa, PRAC Rapporteur: Menno van der Elst, PRAC-CHMP liaison: Johann Lodewijk Hillege, "Submission of an updated RMP version 2.0 in order to replace the category 3 studies CBYL719C2402 and CBYL719A0IC02 with a new non interventional safety study (CBYL719C2404). Additionally, a separated Health Care Professional Survey (CBYL719A0IC02) is proposed as part of the pharmacovigilance plan."

PRAC Led
Stelara - ustekinumab -
EMEA/H/C/000958/II/0082
Janssen-Cilag International NV, Rapporteur: Jayne Crowe, PRAC Rapporteur: Rhea Fitzgerald, PRAC-CHMP liaison: Jayne Crowe, "To submit the final safety registry report of CNTO1275PSO4005 "Nordic Database Initiative for Exposure to Ustekinumab: a Review and Analysis of Adverse Events from the Swedish and Danish National Registry Systems" listed as a category 3 in the RMP. An updated RMP version (18.2) has also been submitted."

PRAC Led

VPRIV - velaglucerase alfa - EMEA/H/C/001249/II/0049, Orphan

Shire Pharmaceuticals Ireland Limited, Rapporteur: Martina Weise, PRAC Rapporteur: Martin Huber, PRAC-CHMP liaison: Martina Weise, "Submission of final physician data study results for PASS study "Evaluation of the Effectiveness of Risk Minimisation Measures: A Survey among Health Care Professionals and Patient/Caregivers to Assess their Knowledge and Attitudes on Prescribing and Home Administration Conditions of Velaglucerase Alpha (VPRIV) in 6 European Countries" (EUPASS 14255)"

PRAC Led

Zinforo - ceftaroline fosamil - EMEA/H/C/002252/II/0055

Pfizer Ireland Pharmaceuticals, Rapporteur: Alar Irs, PRAC Rapporteur: Maia Uusküla, PRAC-CHMP liaison: Alar Irs, "Update of sections 4.4 and 5.2 of the SmPC in order to include information on the use of ceftaroline in patients with cystic fibrosis, based on a pooled population pharmacokinetic (Pop PK) analysis that included data from cystic fibrosis patients treated with ceftaroline fosamil. This submission fulfils the post-authorisation measure LEG 016.1. In addition, the Marketing Authorisation Holder (MAH) took the opportunity to make minor editorial changes."

PRAC Led

WS1944

Izba-EMEA/H/C/002738/WS1944/0014

Travatan-EMEA/H/C/000390/WS1944/0064

Novartis Europharm Limited, Lead Rapporteur: Maria Concepcion Prieto Yerro, Lead PRAC Rapporteur: Eva A. Segovia, "To submit an updated RMP for Travatan and Izba with the following proposed changes:
The following risks are proposed for removal in this RMP update:
important identified risks
- Macular oedema
- Hyperpigmentation
- Hypertrichoses
- Iris and uveal inflammations
- Cardiac and vascular disorders
- Respiratory disorders

- Hypersensitivity reactions important potential risks: - Melanoma - Corneal damage due to use of preserved eye drops - Use during pregnancy and lactation removal of the following missing information topics: - Long-term safety in the paediatric population - Potential interactions In addition, the format of the Risk management plan has been aligned with GVP Module V Revision 2 requirements (EMA 2017). The changes made to the RMP were performed following the PRAC recommendation dated 31-Oct-2019 for the recent PSUR with PSUR number PSUSA/003011/201902."	

B.6.12. CHMP-CAT assessed procedures

Zolgensma - onasemnogene abeparvovec - EMEA/H/C/004750/II/0008, Orphan, ATMP AveXis EU Limited, Rapporteur: Johannes Hendrikus Ovelgonne, CHMP Coordinator: Johann Lodewijk Hillege, "Update of SmPC for sections 4.4 (Special warnings and precautions for use), 4.8 (Undesirable Effects) and corresponding sections in the Package Leaflet to add a new safety signal of 'Thrombotic microangiopathy'."	
Zolgensma - onasemnogene abeparvovec - EMEA/H/C/004750/II/0009/G, Orphan, ATMP AveXis EU Limited, Rapporteur: Johannes Hendrikus Ovelgonne, CHMP Coordinator: Johann Lodewijk Hillege	
Zynteglo - betibeglogene autotemcel - EMEA/H/C/003691/II/0017, Orphan, ATMP bluebird bio (Netherlands) B.V, Rapporteur: Carla Herberts, CHMP Coordinator: Paula Boudewina van Hennik	

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

WS1854
Renvela-EMEA/H/C/000993/WS1854/0053
Sevelamer carbonate Winthrop-EMEA/H/C/003971/WS1854/0026
Genzyme Europe BV, Lead Rapporteur: Karin Janssen van Doorn, "To update section 2 of the SmPC, labelling and section 2 of the PL for the Powder for oral suspension presentations for Renvela and Sevelamer carbonate Winthrop with regards to clarify the exact quantity and threshold of propylene glycol per product following the European Commission guideline on 'Excipients in the labelling and package leaflet of medicinal products for human use' (SANTE-2017-11668).
The MAH took the opportunity to include an update about the local representatives in the Pl for Italy, Malta, the Netherlands and Slovakia."

WS1878
Ambirix-EMEA/H/C/000426/WS1878/0110
Fendrix-EMEA/H/C/000550/WS1878/0073
Infanrix hexa-EMEA/H/C/000296/WS1878/0284
Twinrix Adult-EMEA/H/C/000112/WS1878/0145
Twinrix Paediatric-EMEA/H/C/000129/WS1878/0146
GlaxoSmithkline Biologicals SA, Lead Rapporteur: Christophe Focke

WS1890
Infanrix hexa-EMEA/H/C/000296/WS1890/0286
GlaxoSmithkline Biologicals SA, Lead Rapporteur: Christophe Focke

WS1895/G
Lixiana-EMEA/H/C/002629/WS1895/0029/G

Roteas-EMEA/H/C/004339/WS1895/0017/G
Daiichi Sankyo Europe GmbH, Lead Rapporteur: Maria Concepcion Prieto Yerro

WS1905
Infanrix hexa-EMEA/H/C/000296/WS1905/0283
GlaxoSmithkline Biologicals SA, Lead Rapporteur: Christophe Focke

WS1912
Ambirix-EMEA/H/C/000426/WS1912/0111
Infanrix hexa-EMEA/H/C/000296/WS1912/0285
Rotarix-EMEA/H/C/000639/WS1912/0117
Twinrix Adult-EMEA/H/C/000112/WS1912/0146
Twinrix Paediatric-EMEA/H/C/000129/WS1912/0147
GlaxoSmithkline Biologicals SA, Lead Rapporteur: Christophe Focke

WS1930
Hexacima-EMEA/H/C/002702/WS1930/0107
Hexaxim-EMEA/H/W/002495/WS1930/0112
Hexyon-EMEA/H/C/002796/WS1930/0111
Sanofi Pasteur, Lead Rapporteur: Jan Mueller-Berghaus

WS1931
Hirobriz Breezhaler-EMEA/H/C/001211/WS1931/0060
Onbrez Breezhaler-EMEA/H/C/001114/WS1931/0058
Oslif Breezhaler-EMEA/H/C/001210/WS1931/0058
Novartis Europharm Limited, Lead Rapporteur: Kirstine Moll Harboe, "To update SmPC sections 4.4 and 5.1 (indacaterol maleate) on LABA components and to make changes related to QT interval prolongation.
In addition the MAH has taken this opportunity to bring the annexes in line with QRD version 10.1 and to update the instruction for use in SmPC section 6.6 and package leaflet."

WS1948
Viagra-EMEA/H/C/000202/WS1948/0107

Upjohn EESV, Lead Rapporteur: Johann Lodewijk Hillege, "To update section 4.4 of the SmPC and section 2 of the PL of the Product information for Viagra, Verventi and Sildenafil Pfizer SmPCs in line with the European Commission guideline on 'Excipients in the labelling and package leaflet of medicinal products for human use'. The MAH has proposed wording in the SmPC to correspond to the excipient guideline PIL wording relevant to patients with low sodium diets. The MAH has also taken the opportunity to correct an inaccuracy in the current Viagra 25 mg, 50 mg and 100 mg film coated tablet SmPC related to the quantity of lactose calculated to be present in each tablet. For Sildenafil Pfizer and Verventi, the SmPCs have been amended to specify the quantity of lactose rather than the quantity of lactose monohydrate thus aligning the text with that of the Viagra SmPC."	
WS1958/G **Aflunov-EMEA/H/C/002094/WS1958/0063/G** **Foclivia-EMEA/H/C/001208/WS1958/0059/G** Seqirus S.r.l, Lead Rapporteur: Armando Genazzani	

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 - EMA 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

F.1. Pursuant to Article 9 of Council Regulation (EC) No. 2743/98 of 14 December 1998, as amended

F.2. Request for scientific opinion on justification of exceptional circumstance and for imperative grounds of public health

G. ANNEX G

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

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 12-15 October 2020 CHMP plenary:

Neurology	
Treatment of Adrenoleukodystrophy (SME)	The CHMP denied eligibility to PRIME and adopted the critical summary report.
Treatment of idiopathic intracranial hypertension(SME)	The CHMP denied eligibility to PRIME and adopted the critical summary report.
Vaccines	
VLA1553 Prophylaxis against Chikungunya disease	The CHMP granted eligibility to PRIME and adopted the critical summary report.
Immunology-Rheumatology-Transplantation	
PF-06823859 Treatment of Dermatomyositis	The CHMP granted eligibility to PRIME and adopted the critical summary report.
Gastroenterology-Hepatology	
Efruxifermin Nonalcoholic steatohepatitis(SME)	The CHMP granted eligibility to PRIME and adopted the critical summary report.
Oncology	
Magrolimab Myelodysplastic Syndromes	The CHMP granted eligibility to PRIME and adopted the critical summary report.
Metabolic	
Treatment of Pearson Syndrome(SME)	The CHMP denied eligibility to PRIME and adopted the critical summary report.

G.3.2. List of procedures starting in October 2020 for November 2020 CHMP adoption of outcomes

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