30 April 2021
EMA/CHMP/176288/2021
Human Medicines Division

Committee for medicinal products for human use (CHMP)

Minutes for the meeting on 22-25 February 2021

Chair: Harald Enzmann – Vice-Chair: Bruno Sepodes

Disclaimers

Some of the information contained in this set of 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 Chairperson opened the meeting by welcoming all participants. Due to the current coronavirus (COVID-19) outbreak, and the associated EMA Business Continuity Plan (BCP), the meeting was held 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 February 2021 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 22 – 25 February 2021.

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

Discussions, deliberations and voting took place in full respect of the restricted involvement of Committee members and experts in line with the relevant provisions of the Rules of Procedure and as included in the list of participants. All decisions taken at this meeting were made in the presence of a quorum of members (i.e. 17 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 22-25 February 2021

The CHMP adopted the agenda.

1.3. Adoption of the minutes

CHMP minutes for 25-29 January 2021.

Minutes from PRocedural and Organisational Matters (PROM) meeting (previously called ORGAM meeting) held on 18 January 2021.

The CHMP adopted the CHMP minutes for 25-29 January 2021.

The CHMP adopted the Minutes from the PROM meeting held on 18 January 2021.

2. Oral Explanations

2.1. Pre-authorisation procedure oral explanations

2.1.1. Evrysdi - risdiplam - Orphan - EMEA/H/C/005145

Roche Registration GmbH; treatment of spinal muscular atrophy (SMA)

Scope: Oral explanation

Action: Oral explanation to be held on Wednesday 24 February 2021 at 15:30

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

List of Outstanding Issues adopted on 26.01.2021. List of Questions adopted on 10.11.2020.

The CHMP discussed the outstanding issues.

An oral explanation was held on Wednesday 24 February 2021. The presentation by the applicant focused on the clinical data in support of the application.

See 3.1

2.2. Re-examination procedure oral explanations

No items

2.3. Post-authorisation procedure oral explanations

No items

2.4. Referral procedure oral explanations

2.4.1. Regeneron (casirivimab and imdevimab) for the treatment of COVID-19 - EMEA/H/A-5(3)/1503

MAH: Regeneron Ireland DAC

Referral Rapporteur: Jan Mueller-Berghaus, Referral Co-Rapporteur: Jayne Crowe (MNAT: IE (clinical), PL (quality), NO (non-clinical))

Scope: Opinion

Rapporteurs were appointed via written procedures on 4 February 2021

Possible oral explanation to be held on Tuesday 23 February 2021 at 16:00

Action: For adoption

The Committee for Medicinal Products for Human Use (CHMP) has been asked by the EMA to give a scientific opinion under Article 5(3) of Regulation (EC) No 726/2004 on the currently available quality, preclinical and clinical data on the potential use of casirivimab and imdevimab for the treatment of confirmed COVID-19 in patients that do not require supplemental oxygen and who are at high risk of progressing to severe COVID-19.

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

See 10.2

3. Initial applications

3.1. Initial applications; Opinions

3.1.1. Abevmy - bevacizumab - EMEA/H/C/005327

Mylan IRE Healthcare Limited; 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: Opinion

Action: For adoption

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

List of Outstanding Issues adopted on 10.12.2020. 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 recommending the granting of a marketing authorisation by consensus together with the CHMP assessment report and translation timetable.

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

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

The CHMP noted the letter of recommendation dated 15 February 2021.

The summary of opinion was circulated for information.

The CHMP adopted the similarity assessment report.

3.1.2. Abiraterone Accord - abiraterone acetate - EMEA/H/C/005408

Accord Healthcare S.L.U.; treatment of metastatic prostate cancer

Scope: Opinion

Action: For adoption

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

List of Outstanding Issues adopted on 28.01.2021, 15.10.2020. 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 medical prescription.

The summary of opinion was circulated for information.

3.1.3. Evrysdi - risdiplam - Orphan - EMEA/H/C/005145

Roche Registration GmbH; treatment of spinal muscular atrophy (SMA)

Scope: Oral explanation, Opinion

Action: For adoption

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

List of Outstanding Issues adopted on 26.01.2021. List of Questions adopted on 10.11.2020.

The CHMP discussed the outstanding issues.

See 2.1

An oral explanation was held on Wednesday 24 February 2021. The presentation by the applicant focused on the clinical data 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 recommending the granting of a marketing authorisation by consensus together with the CHMP assessment report and translation timetable.

Furthermore, the CHMP considered that risdiplam 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 01.03.2021.

The summary of opinion was circulated for information.

The CHMP adopted the similarity assessment report.

3.1.4. Jemperli - dostarlimab - EMEA/H/C/005204

GlaxoSmithKline (Ireland) Limited; treatment of mismatch repair deficient (dMMR)/microsatellite instability-high (MSI-H) endometrial cancer (EC)

Scope: Opinion

Action: For adoption

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

List of Outstanding Issues adopted on 28.01.2021, 10.12.2020. List of Questions adopted on 23.06.2020.

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 conditional marketing authorisation by consensus together with the CHMP assessment report and translation timetable.

Furthermore, the CHMP considered that dostarlimab 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 23 February 2021.

The summary of opinion was circulated for information

3.1.5. Lextemy - bevacizumab - EMEA/H/C/005611

Mylan IRE Healthcare Limited; 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: Opinion

Action: For adoption

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

List of Outstanding Issues adopted on 10.12.2020. 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 recommending the granting of a marketing authorisation by consensus together with the CHMP assessment report and translation timetable.

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

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

The summary of opinion was circulated for information.

The CHMP adopted the similarity assessment report.

3.1.6. Orladeyo - berotralstat - Orphan - EMEA/H/C/005138

BioCryst Ireland Limited; prevention of hereditary angioedema (HAE)

Scope: Opinion

Action: For adoption

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

List of Outstanding Issues adopted on 10.12.2020. 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 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 25 February 2021.

The summary of opinion was circulated for information.

The CHMP adopted the similarity assessment report.

3.1.7. Pemazyre - pemigatinib - Orphan - EMEA/H/C/005266

Incyte Biosciences Distribution B.V.; treatment of locally advanced or metastatic cholangiocarcinoma

Scope: Revised opinion

Action: For adoption

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

Opinion adopted on 28.01.2021. List of Outstanding Issues adopted on 17.09.2020. List of Questions adopted on 30.04.2020.

The CHMP re-discussed the wording of the indication.

The Committee adopted a revised positive opinion recommending the granting of a marketing authorisation by majority (27 out of 29 votes) together with the CHMP assessment report and translation timetable.

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

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

The divergent position (Armando Genazzani and Martina Weise) was appended to the opinion.

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

The summary of opinion was circulated for information.

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

3.2.1. idecabtagene vicleucel - Orphan - ATMP - EMEA/H/C/004662

Celgene Europe BV; treatment of multiple myeloma

Scope: List of outstanding issues

Action: For information

List of Outstanding Issues adopted on 04.12.2020. List of Questions adopted on 11.09.2020.

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

The Committee endorsed the 2nd list of outstanding issues with a specific timetable as adopted by the CAT.

3.2.2. abiraterone acetate - EMEA/H/C/005649

treatment of prostate cancer in adult men

Scope: List of outstanding issues

Action: For adoption

List of Questions adopted on 12.11.2020.

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. abiraterone acetate - EMEA/H/C/005368

treatment of metastatic castration resistant prostate cancer

Scope: List of outstanding issues

Action: For adoption

List of Questions adopted on 23.07.2020.

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. tralokinumab - EMEA/H/C/005255

treatment of moderate-to-severe atopic dermatitis in adult patients who are candidates for systemic therapy

Scope: List of outstanding issues

Action: For adoption

List of Questions adopted on 17.09.2020.

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 Outstanding Issues adopted on 15.10.2020. 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 2nd list of outstanding issues with a specific timetable.

3.2.6. 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: List of outstanding issues

Action: For adoption

List of Outstanding Issues adopted on 10.12.2020, 17.09.2020. 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 3rd list of outstanding issues with a specific timetable.

3.2.7. dasatinib - EMEA/H/C/005446

treatment of leukaemia

Scope: List of outstanding issues

Action: For adoption

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

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

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

3.2.8. dasatinib - EMEA/H/C/005317

treatment of leukaemia

Scope: List of outstanding issues

Action: For adoption

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

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

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

3.2.9. satralizumab - Orphan - EMEA/H/C/004788

Roche Registration GmbH; treatment of adult and adolescent patients from 12 years of age with neuromyelitis optica spectrum disorders (NMOSD)

Scope: List of outstanding issues

Action: For adoption

List of Outstanding Issues adopted on 28.05.2020. List of Questions adopted on 10.12.2019.

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

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

3.2.10. evinacumab - EMEA/H/C/005449

Accelerated assessment

treatment of homozygous familial hypercholesterolemia (HoFH)

Scope: List of outstanding issues

Action: For adoption

List of Questions adopted on 08.12.2020.

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. roxadustat - EMEA/H/C/004871

treatment of anaemia

Scope: List of outstanding issues

Action: For adoption

List of Questions adopted on 17.09.2020.

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. istradefylline - EMEA/H/C/005308

indicated as an adjunctive treatment to levodopa-based regimens in patients with Parkinson's disease

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.

3.2.13. 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 Outstanding Issues adopted on 15.10.2020. 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 2nd list of outstanding issues with a specific timetable.

3.2.14. selumetinib - Orphan - EMEA/H/C/005244

AstraZeneca AB; treatment of neurofibromatosis type 1 (NF1)

Scope: List of outstanding issues

List of experts for the ad-hoc expert group meeting scheduled on 09 February 2021 adopted via written procedure on 09 February 2021

Ad-hoc expert group report

Action: For adoption

List of Questions adopted on 23.07.2020.

The CHMP noted the report from the ad-hoc expert group.

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

EigerBio Europe Limited; treatment of Hutchinson-Gilford Progeria Syndrome and Progeroid Laminopathies

Scope: List of outstanding issues

Action: For adoption

List of Questions adopted on 23.07.2020.

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.

3.2.16. azacitidine - EMEA/H/C/004761

treatement for acute myeloid leukaemia

Scope: List of outstanding issues

Action: For adoption

List of Questions adopted on 17.09.2020.

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. tirbanibulin mesilate - EMEA/H/C/005183

topical field treatment of non-hyperkeratotic, non-hypertrophic actinic keratosis

Scope: List of outstanding issues

Action: For adoption

List of Questions adopted on 25.06.2020.

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. arachis hypogaea extract - Article 28 - EMEA/H/C/004810

treatment of peanut allergy

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. aducanumab - EMEA/H/C/005558

Alzheimer’s disease

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. odevixibat - Orphan - EMEA/H/C/004691

Accelerated assessment

Albireo; treatment of progressive familial intrahepatic cholestasis (PFIC)

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. lenadogene nolparvovec - Orphan - ATMP - EMEA/H/C/005047

GenSight Biologics S.A.; treatment of vision loss due to Leber Hereditary Optic Neuropathy (LHON)

Scope: List of questions

Action: For information

The CHMP was updated on discussions at the CAT. The Committee discussed the issues identified in this application.

The Committee endorsed the list of questions as adopted by the CAT.

3.3.5. anifrolumab - EMEA/H/C/004975

indicated as an add-on therapy for the treatment of adult patients with moderate to severe systemic lupus erythematosus (SLE), despite standard therapy

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.

3.3.6. avacopan - Orphan - EMEA/H/C/005523

Vifor Fresenius Medical Care Renal Pharma France; Treatment of granulomatosis with polyangiitis (GPA) or microscopic polyangiitis (MPA)

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.

3.3.7. tecovirimat - EMEA/H/C/005248

treatment of orthopoxvirus disease

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. dabigatran etexilate - EMEA/H/C/005639

prevention of venous thromboembolic events

Scope: Letter from the applicant dated 04 February 2021 requesting an extension of clock-stop to respond to the list of questions adopted in November 2020.

Action: For adoption

List of Questions adopted on 12.11.2020.

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

3.4.2. risperidone- EMEA/H/C/005406

treatment of schizophrenia

Scope: Letter from the applicant dated 8 February 2021 requesting an extension of clock-stop to respond to the list of outstanding issues adopted in October 2020.

Action: For adoption

List of outstanding issues adopted on 15.10.2020. List of questions adopted on 28.05.2020.

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 October 2020.

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

No items

3.6. Initial applications in the decision-making phase

No items

3.7. Withdrawals of initial marketing authorisation application

No items

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

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

4.1.1. Elebrato Ellipta - fluticasone furoate / umeclidinium / vilanterol - EMEA/H/C/004781/X/0014/G

GlaxoSmithKline Trading Services Limited

Rapporteur: Peter Kiely, Co-Rapporteur: Janet Koenig, PRAC Rapporteur: Annika Folin

Scope: "Extension application to introduce a new strength (184 mcg/55 mcg/22 mcg) grouped with a type II variation (C.I.6.a) to add a new indication (asthma).
Whilst the new indication is applicable also to the existing presentations (EU/1/17/1237/001-3), the new strength is indicated only for the treatment of asthma.
The RMP (version 2.2) is updated in accordance."

Action: For adoption

Oral explanation was held in January 2021. List of Outstanding Issues adopted on 12.11.2020. List of Questions adopted on 25.06.2020.

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

The CHMP adopted a negative opinion by consensus recommending the refusal of the granting of an extension of the marketing authorisation concerning a new strength: 184 mcg/55 mcg/22 mcg.

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

4.1.2. Insulin aspart Sanofi - insulin aspart - EMEA/H/C/005033/X/0003

sanofi-aventis groupe

Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Annika Folin

Scope: "Extension application to introduce a new route of administration, intravenous use, for the 10 ml vial presentations only."

Action: For adoption

List of Questions adopted on 10.12.2020.

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 adopted the similarity assessment report.

4.1.3. Temybric Ellipta - fluticasone furoate / umeclidinium / vilanterol - EMEA/H/C/005254/X/0004/G

GlaxoSmithKline Trading Services Limited

Rapporteur: Peter Kiely, Co-Rapporteur: Janet Koenig, PRAC Rapporteur: Annika Folin

Scope: "Extension application to introduce a new strength (184 mcg/55 mcg/22 mcg) grouped with a type II variation (C.I.6.a) to add a new indication (asthma).
Whilst the new indication is applicable also to the existing presentations (EU/1/19/13781237/001-3), the new strength is indicated only for the treatment of asthma.
The RMP (version 2.2) is updated in accordance."

Action: For adoption

Oral explanation was held in January 2021. List of Outstanding Issues adopted on 12.11.2020. List of Questions adopted on 25.06.2020.

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

The CHMP adopted a negative opinion by consensus recommending the refusal of the granting of an extension of the marketing authorisation concerning a new strength: 184 mcg/55 mcg/22 mcg.

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

4.1.4. Trelegy Ellipta - fluticasone furoate / umeclidinium / vilanterol - EMEA/H/C/004363/X/0012/G

GlaxoSmithKline Trading Services Limited

Rapporteur: Peter Kiely, Co-Rapporteur: Janet Koenig, PRAC Rapporteur: Annika Folin

Scope: "Extension application to introduce a new strength (184 mcg/55 mcg/22 mcg) grouped with a type II variation (C.I.6.a) to add a new indication (asthma).
Whilst the new indication is applicable also to the existing presentations (EU/1/17/1236/001-3), the new strength is indicated only for the treatment of asthma.
The RMP (version 2.2) is updated in accordance."

Action: For adoption

Oral explanation was held in January 2021. List of Outstanding Issues adopted on 12.11.2020. List of Questions adopted on 25.06.2020.

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

The CHMP adopted a negative opinion by consensus recommending the refusal of the granting of an extension of the marketing authorisation concerning a new strength: 184 mcg/55 mcg/22 mcg.

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

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. Aubagio - teriflunomide - EMEA/H/C/002514/X/0031/G

Sanofi-aventis groupe

Rapporteur: Martina Weise, PRAC Rapporteur: Martin Huber

Scope: "1- Extension of a marketing authorisation for Aubagio to add the new strength, 7 mg film-coated tablet, for use in paediatric patients from 10 years of age and older with relapsing remitting multiple sclerosis (MS).
2- Type II (C.I.6) - Extension of indication to include treatment of paediatric patients aged 10 years and older with relapsing remitting multiple sclerosis (MS) for Aubagio 14 mg tablet. As a consequence, sections 4.1, 4.2, 4.4, 4.8, 5.1, 5.2 and 5.3 of the SmPC are updated. The Package Leaflet and Labelling are updated in accordance.
The MAH is requesting an extension of the market protection of one additional year in line with the guidance on elements required to support the significant clinical benefit in comparison with existing therapies of a new therapeutic indication in order to benefit from an extended (11-year) marketing protection period.
Version 6.0 of the RMP has also been submitted."

Action: For adoption

List of Questions adopted on 17.09.2020

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.

4.2.2. Bortezomib Accord - bortezomib - EMEA/H/C/003984/X/0023

Accord Healthcare S.L.U.

Rapporteur: Daniela Philadelphy, PRAC Rapporteur: Amelia Cupelli

Scope: "Extension application to introduce a new pharmaceutical form associated with new strength (2.5 mg/ml solution for injection)."

Action: For adoption

List of Questions adopted on 17.09.2020.

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

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

4.2.3. 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

List of Questions adopted on 15.10.2020.

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

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

4.2.4. Fabrazyme - agalsidase beta - EMEA/H/C/000370/X/0118/G

Genzyme Europe BV

Rapporteur: Johann Lodewijk Hillege

Scope: Quality changes

Action: For adoption

List of Questions adopted on 17.09.2020.

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

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

4.2.5. Ferriprox - deferiprone - EMEA/H/C/000236/X/0145

Chiesi Farmaceutici S.p.A.

Rapporteur: Alexandre Moreau, PRAC Rapporteur: Tiphaine Vaillant

Scope: "Extension application to introduce a new pharmaceutical form (gastro-resistant tablets). The RMP (version 14.0) is updated in accordance."

Action: For adoption

List of Questions adopted on 17.09.2020.

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.

4.2.6. Maviret - glecaprevir / pibrentasvir - EMEA/H/C/004430/X/0033/G

AbbVie Deutschland GmbH & Co. KG

Rapporteur: Jean-Michel Race, PRAC Rapporteur: Ana Sofia Diniz Martins

Scope: "Extension application to introduce a new pharmaceutical form (50/20 mg coated granules in sachet), grouped with a type II extension of indication variation (C.I.6.a) to include the treatment of children from 3 to 12 years of age for the approved Maviret 100 mg/40 mg film-coated tablets; as a consequence, sections 4.1, 4.2, 5.1 and 5.2 of the SmPC are updated. The Package Leaflet and Labelling are updated accordingly. Version 5.0 of the RMP has also been submitted. Furthermore, the PI is brought in line with the latest QRD template version 10.1."

Action: For adoption

List of Questions adopted on 17.09.2020.

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

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

4.2.7. Nitisinone MDK - nitisinone - EMEA/H/C/004281/X/0007

MendeliKABS Europe Limited

Rapporteur: Alar Irs, PRAC Rapporteur: Amelia Cupelli

Scope: "Extension application to add a new strength of 20 mg (hard capsule)."

Action: For adoption

List of Questions adopted on 17.09.2020.

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.

4.2.8. Skyrizi - risankizumab - EMEA/H/C/004759/X/0012

AbbVie Deutschland GmbH & Co. KG

Rapporteur: Peter Kiely, PRAC Rapporteur: Liana Gross-Martirosyan

Scope: "Extension application to add a new strength of 150 mg for solution for injection in a pre-filled syringe and pre-filled pen."

Action: For adoption

List of Questions adopted on 10.12.2020.

The Committee discussed the issues identified in this application, 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.

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

4.3.1. Cosentyx - secukinumab - EMEA/H/C/003729/X/0067

Novartis Europharm Limited

Rapporteur: Outi Mäki-Ikola, PRAC Rapporteur: Eva A. Segovia

Scope: "Extension application to introduce a new strength of 75 mg solution for injection."

Action: For adoption

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 questions.

4.3.2. Rinvoq - upadacitinib - EMEA/H/C/004760/X/0006/G

AbbVie Deutschland GmbH & Co. KG

Rapporteur: Kristina Dunder, PRAC Rapporteur: Nikica Mirošević Skvrce

Scope: "Extension application to introduce a new strength (30 mg prolonged-release tablet), grouped with a type II variation (C.I.6.a) to add a new indication (treatment of moderate to severe atopic dermatitis in adults and adolescents 12 years and older who are candidates for systemic therapy for Rinvoq). As a consequence, sections 4.1, 4.2, 4.5, 4.8, 5.1, 5.2 and 5.3 of the SmPC as well as the Package Leaflet are updated. The RMP (version 4.0) is updated in accordance. In addition, the marketing authorisation holder (MAH) took the opportunity to include a minor update in the Annex II."

Action: For adoption

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 questions.

4.3.3. Vosevi - sofosbuvir / velpatasvir / voxilaprevir - EMEA/H/C/004350/X/0045/G

Gilead Sciences Ireland UC

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

Scope: "Extension application to introduce a new strength (200 mg /50 mg /50 mg film-coated tablets). The new presentation is indicated for the treatment of chronic hepatitis C virus (HCV) infection in patients aged 12 years and older OR weighing at least 30 kg. In addition, the MAH took the opportunity to implement minor editorial updates.
The extension application is grouped with a type II variation (C.I.6.a) to include paediatric use in patients aged 12 years and older OR weighing at least 30 kg to the existing presentation. Sections 4.2, 4.8, 5.1 and 5.2 of the SmPC and the Package Leaflet are updated to support the extended indication. The RMP (version 3.2) is updated in accordance."

Action: For adoption

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 questions.

4.3.4. Xeljanz - tofacitinib - EMEA/H/C/004214/X/0030/G

Pfizer Europe MA EEIG

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

Scope: "Extension application to add a new strength (22 mg prolonged-release tablet) grouped with a type II variation C.I.4: 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 adult patients with moderately to severely active ulcerative colitis who have had an inadequate response, lost response, or were intolerant to either conventional therapy or a biologic agent; 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 UC. The Package Leaflet is updated accordingly. The RMP version 15.1 has also been submitted."

Action: For adoption

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 questions.

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

4.4.1. Lacosamide Accord - lacosamide - EMEA/H/C/004443/X/0007

Accord Healthcare S.L.U.

Rapporteur: John Joseph Borg, PRAC Rapporteur: Ulla Wändel Liminga

Scope: "Extension application to introduce a new pharmaceutical form (solution for infusion), a new strength (10 mg/ml) and a new route of administration (intravenous use)."

Scope: Letter from the applicant dated 17 February 2021 requesting an extension to the clock stop to respond to the list of outstanding issues adopted in November 2020.

Action: For adoption

List of Outstanding Issues adopted on 12.11.2020. List of Questions adopted on 26.03.2020.

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

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. Cabometyx - cabozantinib - EMEA/H/C/004163/II/0017

Ipsen Pharma

Rapporteur: Bjorg Bolstad, PRAC Rapporteur: Menno van der Elst

Scope: "Extension of indication to include in combination with nivolumab first line treatment of advanced renal cell carcinoma for Cabometyx; 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 5.0 of the RMP has also been submitted."

Action: For adoption

Request for Supplementary Information adopted on 10.12.2020.

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.

5.1.2. Darzalex - daratumumab - Orphan - EMEA/H/C/004077/II/0043

Janssen-Cilag International NV

Rapporteur: Sinan B. Sarac, Co-Rapporteur: Blanca Garcia-Ochoa, PRAC Rapporteur: Marcia Sofia Sanches de Castro Lopes Silva

Scope: "Extension of indication to include treatment of adult patients with systemic light chain (AL) amyloidosis for Darzalex; as a consequence, sections 4.1, 4.2, 4.4, 4.5, 4.8, 5.1 and 5.2 of the SmPC are updated. The Package Leaflet is updated in accordance. Version 8.2 of the RMP has also been submitted."

Action: For adoption

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

aspects.

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

5.1.3. Darzalex - daratumumab - Orphan - EMEA/H/C/004077/II/0044

Janssen-Cilag International NV

Rapporteur: Sinan B. Sarac, Co-Rapporteur: Blanca Garcia-Ochoa, PRAC Rapporteur: Marcia Sofia Sanches de Castro Lopes Silva

Scope: "Extension of indication for Darzalex subcutaneous formulation to include combination with pomalidomide and dexamethasone for the treatment of adult patients with relapsed and refractory multiple myeloma, whose prior therapy included a proteasome inhibitor and an immunomodulatory agent and who have demonstrated disease progression on the last therapy; as a consequence, sections 4.1, 4.2, 4.4, 4.5, 4.8, 5.1 and 5.2 of the SmPC are updated. In addition, section 4.8 of the SmPC for the intravenous formulation is also updated based on the pooled safety analysis. The Package Leaflet is updated in accordance. Version 8.2 of the RMP has also been submitted."

Action: For adoption

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

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

5.1.4. 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 2 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 2.0 is agreed.
The MAH 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. In addition, the list of local representatives in the PL has been revised to amend contact details for the representative(s) of Sweden.", 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 28.01.2021, 15.10.2020, 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 CHMP noted the letter of recommendation dated 22 February 2021.

The summary of opinion was circulated for information.

The CHMP adopted the similarity assessment report.

5.1.5. Keytruda - pembrolizumab - EMEA/H/C/003820/II/0097

Merck Sharp & Dohme B.V.

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

Scope: "Extension of indication to include in combination with chemotherapy, first-line treatment of locally advanced unresectable or metastatic carcinoma of the oesophagus or HER-2 negative gastroesophageal junction adenocarcinoma in adults for Keytruda, based on the results from the pivotal KEYNOTE-590 (KN590) trial. Phase 3, randomized, double-blind, placebo-controlled, multisite study to evaluate the efficacy and safety of pembrolizumab in combination with chemotherapy (cisplatin and 5-FU) versus chemotherapy (cisplatin with 5-FU) as first-line treatment in participants with locally advanced unresectable metastatic adenocarcinoma or squamous cell carcinoma of the oesophagus or advanced/metastatic Siewert type 1 adenocarcinoma of the gastroesophageal junction; as a consequence, sections 4.1, 4.8 and 5.1 of the SmPC are updated. The Package Leaflet is updated in accordance. Version of the RMP (Version 30.1) has also been submitted."

Action: For adoption

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

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

5.1.6. Opdivo - nivolumab - EMEA/H/C/003985/II/0092

Bristol-Myers Squibb Pharma EEIG

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

Scope: "Extension of indication to include in combination with cabozantinib for the first-line treatment of advanced renal cell carcinoma for Opdivo; as a consequence, sections 4.1, 4.2, 4.4, 4.8 and 5.1 of the SmPC are updated. The Package Leaflet is updated in accordance. Version 19.0 of the RMP has also been submitted."

Action: For adoption

Request for Supplementary Information adopted on 10.12.2020.

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.

5.1.7. Quofenix - delafloxacin - EMEA/H/C/004860/II/0003

A. Menarini Industrie Farmaceutiche Riunite s.r.l.

Rapporteur: Janet Koenig, Co-Rapporteur: Alar Irs, PRAC Rapporteur: Nikica Mirošević Skvrce

Scope: "Extension of indication to include treatment of Community Acquired Pneumonia (CAP) in adults when it is considered inappropriate to use other antibacterial agents that are commonly recommended for the initial treatment of these infections, for Quofenix 450 mg tablets and 300 mg powder for concentrate for solution for infusion; 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 2.1 of the RMP has also been submitted.
In addition, the list of local representatives in the PL has been revised to amend contact details for the representative to mention United Kingdom (Northern Ireland) in line with the current QRD template (10.2)."

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.

The summary of opinion was circulated for information.

5.1.8. Sarclisa - isatuximab - Orphan - EMEA/H/C/004977/II/0003

sanofi-aventis groupe

Rapporteur: Paula Boudewina van Hennik, PRAC Rapporteur: Eva A. Segovia

Scope: "An extension of indication for Sarclisa to add combination with carfilzomib and dexamethasone, for the treatment of patients with multiple myeloma who have received at least one prior therapy. As a consequence, the sections 4.1, 4.2, 4.4, 4.5, 4.8, 5.1 and 5.2 have been updated. The PL is updated accordingly. The MAH took the opportunity to introduce minor changes in the SmPC sections 4.9, 6.3 and 6.6 and update the details of local representatives. Revised RMP version 1 has been submitted."

Action: For adoption

Request for Supplementary Information adopted on 10.12.2020.

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.

The CHMP adopted the similarity assessment report.

5.1.9. Tecentriq - atezolizumab - EMEA/H/C/004143/II/0052

Roche Registration GmbH

Rapporteur: Sinan B. Sarac, Co-Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Marcia Sofia Sanches de Castro Lopes Silva

Scope: "Extension of indication to include Tecentriq in combination with nab-paclitaxel and anthracycline-based chemotherapy for the neoadjuvant treatment of adult patients with locally advanced or early Triple Negative Breast Cancer (TNBC) based on the results of the pivotal study WO39392 (IMpassion031); as a consequence, sections 4.1, 4.2, 4.8 and 5.1 of the Tecentriq 840 mg concentrate for solution for infusion SmPC and section 4.8 of the Tecentriq 1,200 mg concentrate for solution for infusion SmPC are updated. The Package Leaflet is updated in accordance. Version 18 of the RMP has also been submitted."

Action: For adoption

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

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

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

5.1.10. Verzenios - abemaciclib - EMEA/H/C/004302/II/0013

Eli Lilly Nederland B.V.

Rapporteur: Filip Josephson, PRAC Rapporteur: Marcia Sofia Sanches de Castro Lopes Silva

Scope: "Extension of indication to include Verzenios in combination with endocrine therapy for adjuvant treatment of patients with hormone receptor (HR)-positive, human epidermal growth factor receptor 2 (HER2)-negative, node-positive early breast cancer at high risk of recurrence; as a consequence, sections 4.1, 4.2, 4.4, 4.6, 4.8, 5.1 and 5.3 of the SmPC are updated. The Package Leaflet is updated in accordance. Version 1.1 of the RMP has also been submitted.", 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 clinical aspects and the request for 1 year of market protection.

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

5.1.11. WS1941
Edistride - dapagliflozin - EMEA/H/C/004161/WS1941/0043
Forxiga - dapagliflozin - EMEA/H/C/002322/WS1941/0062

AstraZeneca AB

Lead Rapporteur: Kristina Dunder, PRAC Rapporteur: Annika Folin

Scope: "Update of sections 4.1, 4.2, 4.4, 4.8, 5.1 and 5.2 of the SmPC for Forxiga and

Edistride based on the results from the renal outcomes study D169AC00001 (DAPA-CKD). The Annex II.B and Package Leaflet of these products are updated accordingly. The DAPA-CKD study is a category 3, Post-Authorisation Safety Study (PASS) listed in the dapagliflozin RMP to evaluate the potential risk of lower limb amputation; it is a multicentre, event-driven, randomized, double-blind, parallel group, placebo-controlled study, evaluating the effect of dapagliflozin versus placebo, given once daily in addition to standard of care, to prevent the progression of chronic kidney disease (CKD) or cardiovascular (CV)/renal death. In addition, the Risk Management Plan for dapagliflozin (version 22) has been updated."

Action: For adoption

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

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

5.1.12. WS1953 Segluromet - ertugliflozin / metformin hydrochloride - EMEA/H/C/004314/WS1953/0012 Steglatro - ertugliflozin - EMEA/H/C/004315/WS1953/0013

Merck Sharp & Dohme B.V.

Lead Rapporteur: Kristina Dunder, PRAC Rapporteur: Menno van der Elst

Scope: "Update of sections 4.1, 4.2, 4.4, 4.8 and 5.1 of the SmPC of Steglatro and Segluromet in order to modify the indication, update posology recommendations and include efficacy and safety information based on final results from the VERTIS CV study (protocol 8835-004/B1521021) listed as a category 3 study in the RMP. This is a multi-centre, multi-national, randomised, double-blind, placebo-controlled study to evaluate the effect of ertugliflozin on cardiovascular risk in adult patients with type 2 diabetes and established atherosclerotic cardiovascular disease. The Package Leaflet is updated accordingly. The RMP version 2.0 has also been submitted."

Action: For adoption

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

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

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

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. enfortumab vedotin - H0005392

Indicated in adults for the treatment of patients with locally advanced (LA) or metastatic urothelial cancer (mUC) who have received a programmed death receptor-1 (PD-1) or programmed death-ligand 1 (PD-L1) inhibitor and who have received a platinum-containing chemotherapy in the neoadjuvant/adjuvant, locally advanced or metastatic setting.

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

Action: For adoption

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

8.1.2. melphalan flufenamide - Orphan - H0005681

Oncopeptides AB, in combination with dexamethasone is indicated for the treatment of adult patients with multiple myeloma whose disease is refractory to at least one proteasome inhibitor, one immunomodulatory agent, and one anti-CD38 monoclonal antibody

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. tezepelumab – H0005588

add-on maintenance treatment in adults and adolescents 12 years and older with severe asthma

Scope: Adoption of the Briefing note and the Rapporteur's 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.

8.1.4. sacituzumab govitecan – H0005182

treatment of patients with metastatic triple-negative breast cancer (mTNBC) who previously received at least two prior therapies for metastatic disease

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

Action: For adoption

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

8.2. Priority Medicines (PRIME)

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 4 recommendations for eligibility to PRIME: all 4 were denied.

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

9. Post-authorisation issues

9.1. Post-authorisation issues

9.1.1. Ondexxya - andexanet alfa - EMEA/H/C/004108/II/0003

Alexion Europe SAS

Rapporteur: Jan Mueller-Berghaus

Scope: "Submission of the final report from Population PK and PK/PD Modelling Report (POR-PKPD-ADEX-321) listed as a category 2 study in the RMP/Specific Obligation 004 in Annex II."

Action: For discussion

Request for Supplementary Information adopted on 28.05.2020, 12.12.2019.

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

The Committee adopted a 3rd request for supplementary information with a specific timetable.

The CHMP agreed to consult the modelling and simulation working party and adopted a list of question to this group.

9.1.2. Strimvelis - autologous CD34+ enriched cell fraction that contains CD34+ cells transduced with retroviral vector that encodes for the human ADA cDNA sequence - EMEA/H/C/003854/R/0029

Orchard Therapeutics (Netherlands) BV

Rapporteur: Sol Ruiz, Co-Rapporteur: Egbert Flory, CHMP Coordinators: Maria Concepcion Prieto Yerro and Jan Mueller-Berghaus

Scope: Renewal of marketing authorisation requiring 2nd renewal

Action: For adoption

Request for Supplementary Information adopted on 04.12.2020.

The CHMP adopted a positive opinion 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.

9.1.3. Zynteglo - betibeglogene autotemcel - EMEA/H/C/003691/R/0018

Bluebird bio (Netherlands) B.V

Rapporteur: Carla Herberts, CHMP Coordinator: Paula Boudewina van Hennik

Scope: Renewal of conditional marketing authorisation

Action: For adoption

Request for Supplementary Information adopted on 22.01.2021.

The CHMP noted that all clinical trials are put on hold and that the marketing of the product was temporarily suspended due to possible safety issues. The CHMP was informed that the renewal procedure is put on hold, until the finalisation of an Article 20 referral procedure, which will be triggered to further assess the benefit/risk for Zynteglo.

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

10.2.1. Eli Lilly (bamlanivimab and etesevimab) for the treatment of COVID-19 - EMEA/H/A-5(3)/1502

Eli Lilly

Referral Rapporteur: Jan Mueller-Berghaus, Referral Co-Rapporteur: Tomas Radimersky

Scope: List of Outstanding Issues

Rapporteurs were appointed via written procedures on 4 February 2021

Action: For adoption

The Committee for Medicinal Products for Human Use (CHMP) has been asked by the EMA to give a scientific opinion under Article 5(3) of Regulation (EC) No 726/2004 on the currently available quality, preclinical and clinical data on the potential use of bamlanivimab and etesevimab for the treatment of confirmed COVID-19 in patients that do not require supplemental oxygen and who are at high risk of progressing to severe COVID-19.

The CHMP noted that an extraordinary CHMP will be scheduled on 4 March 2021.

10.2.2. Regeneron (casirivimab and imdevimab) for the treatment of COVID-19 - EMEA/H/A-5(3)/1503

Regeneron Ireland DAC

Referral Rapporteur: Jan Mueller-Berghaus, Referral Co-Rapporteur: Jayne Crowe (MNAT: IE (clinical), PL (quality), NO (non-clinical))

Scope: Opinion

Rapporteurs were appointed via written procedures on 4 February 2021

Possible oral explanation to be held on Tuesday 23 February 2021 at 16:00

See 2.4

Action: For adoption

The Committee for Medicinal Products for Human Use (CHMP) has been asked by the EMA to give a scientific opinion under Article 5(3) of Regulation (EC) No 726/2004 on the currently available quality, preclinical and clinical data on the potential use of casirivimab and imdevimab for the treatment of confirmed COVID-19 in patients that do not require supplemental oxygen and who are at high risk of progressing to severe COVID-19.

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

The CHMP adopted an opinion by consensus, recommending that Regeneron (casirivimab and imdevimab) can be used for the treatment of confirmed COVID-19 in patients who do not require supplemental oxygen and who are at high risk of progressing to severe COVID-19.

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

The EMA press release was circulated for information.

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

No items

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

No items

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

10.5.1. Varilrix - EMEA/H/A-30/1499

MAH: GlaxoSmithKline Biologicals

Referral Rapporteur: Jan Mueller-Berghaus, Referral Co-Rapporteur: Sol Ruiz

Scope: Opinion

Action: For adoption

Harmonisation exercise for Varilrix and associated names. Product information harmonisation was triggered by the MAH.

The CHMP adopted an opinion by consensus, recommending the revision and harmonisation of the product information for Varilrix and associated names.

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

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

No items

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

February 2021 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. Election of co-opted member

Election of CHMP co-opted member in light of the expiry of mandate of co-opted member Blanka Hirschlerova on 18 March 2021.

Agreed areas of expertise: quality (non-biologicals) and pharmacokinetics.

Nomination(s) received

Action: For election

The CHMP re-elected Blanka Hirschlerova as CHMP co-opted member for another 3 year mandate.

14.2. Coordination with EMA Scientific Committees

14.2.1. Pharmacovigilance Risk Assessment Committee (PRAC)

List of Union Reference Dates and frequency of submission of Periodic Safety Update Reports (EURD list) for February 2021

Action: For adoption

The CHMP adopted the EURD list.

14.2.2. Paediatric Committee (PDCO)

PIPs reaching D30 at February 2021 PDCO

Action: For information

The CHMP noted the information.

Report from the PDCO meeting held on 23-26 February 2021

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 February 2021 meeting to CHMP for adoption:

- 17 reports on products in scientific advice and protocol assistance
- 09 reports on products in pre-authorisation procedures
- 03 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 16-17 February 2021.

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 08-11 February 2021. Table of conclusions

Action: For information

The CHMP noted the report.

Scientific advice letters

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

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 COVID-19/ Regulatory consideration on the variants

Action: For information

The CHMP noted the update.

15.1.3. Reflection paper on the regulatory requirements for vaccines intended to provide protection against variant strain(s) of SARS-CoV-2

Action: For adoption

The CHMP adopted the reflection paper.

15.1.4. COVID-19 vaccine (Ad26.COV2-S [recombinant]) – EMEA/H/C/005737

prevention of coronavirus disease-2019 (COVID-19)

Scope: Conditional marketing authorisation timetable adopted via written procedure on 15 February 2021

Action: For information

The CHMP noted the CMA timetable which was adopted via written procedure.

15.1.5. COVID-19 vaccine (NVX-CoV2373) – EMEA/H/C/005808

prevention of COVID-19

Scope: Rolling review timetable adopted via written procedure on 11 February 2021

Action: For information

The CHMP noted the RR timetable which was adopted via written procedure.

15.1.6. COVID-19 mRNA vaccine – EMEA/H/C/005845

prevention of COVID-19

Scope: Rolling review timetable adopted via written procedure on 12 February 2021

Action: For information

The CHMP noted the RR timetable which was adopted via written procedure.

15.1.7. Regdanvimab – H0005854

mild to moderate coronavirus disease 2019 (COVID-19) in adults

Scope: Rolling Review timetable adopted via written procedure on 24 February 2021

Action: For information

The CHMP noted the RR timetable which was adopted via written procedure.

16. Lists of participants

List of participants including any restrictions with respect to involvement of members/alternates/experts following evaluation of declared interests for the 22-25 February 2021 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	
Daniela Philadelphy	Alternate	Austria	No interests declared	
Christophe Focke	Member	Belgium	No restrictions applicable to this meeting	
Karin Janssen van Doorn	Alternate	Belgium	No interests declared	
Ilko Getov	Member	Bulgaria	No interests declared	
Margareta Bego	Member	Croatia	No interests declared	
Selma Arapovic Dzakula	Alternate	Croatia	No interests declared	
Helena Panayiotopoulou	Member	Cyprus	No interests declared	
Ondřej Slanař	Member	Czechia	No restrictions applicable to this meeting	
Tomas Radimersky	Alternate	Czechia	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	
Johanna Lähteenvuo	Alternate	Finland	No interests declared	
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	
Konstantinos Markopoulos	Member	Greece	No interests declared	

Name	Role	Member State or affiliation	Outcome restriction following evaluation of e-DoI	Topics on agenda for which restrictions apply
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 participation in final deliberations and voting	COVID-19 vaccines
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 restrictions applicable to this meeting	
Nevenka Trsinar Brodt	Alternate	Slovenia	No interests declared	
Maria Concepcion Prieto Yerro	Member	Spain	No interests declared	
Blanca Garcia-Ochoa	Alternate	Spain	No interests declared	
Kristina Dunder	Member	Sweden	No interests declared	
Filip Josephson	Alternate	Sweden	No interests declared	

Name	Role	Member State or affiliation	Outcome restriction following evaluation of e-DoI	Topics on agenda for which restrictions apply
Christian Gartner	Co-opted member	Austria	No restrictions applicable to this meeting	
Carla Torre	Co-opted member	Portugal	No interests declared	
Jan Mueller-Berghaus	Co-opted member	Germany	No interests declared	
Blanka Hirschlerova	Co-opted member	Czechia	No interests declared	
Sol Ruiz	Co-opted member	Spain	No interests declared	
Rune Kjeken	Expert - via Adobe*	Norway	No restrictions applicable to this meeting	
Helga Haugom Olsen	Expert - via Adobe*	Norway	No interests declared	
Hilde Roshol	Expert - via Adobe*	Norway	No interests declared	
Sara Galluzzo	Expert - via Adobe*	Italy	No interests declared	
João Manuel Lopes de Oliveira	Expert - via Adobe*	Portugal	No interests declared	
Minne Casteels	Expert - via Adobe*	Belgium	No interests declared	
Linda Trauffler	Expert - via Adobe*	Austria	No interests declared	
Antero Kallio	Expert - via Adobe*	Finland	No restrictions applicable to this meeting	
Elina Asikanius	Expert - via Adobe*	Finland	No part in discussions, final deliberations and voting	Evrysdi - risdiplam - Orphan - EMEA/H/C/005145 Tecentriq - atezolizumab - EMEA/H/C/004143/II/0052
Susanne Mueller-Egert	Expert - via Adobe*	Germany	No interests declared	
Hilke Zander	Expert - via Adobe*	Germany	No interests declared	
Birgit Ahrens	Expert - via Adobe*	Germany	No interests declared	
Susanne Kaul	Expert - via Adobe*	Germany	No interests declared	
Ebru Karakoc Madsen	Expert - via Adobe*	Denmark	No restrictions applicable to this meeting	
Hester Peltenburg	Expert - via Adobe*	Netherlands	No interests declared	
Lieke Sandberg Smits	Expert - via Adobe*	Netherlands	No interests declared	
Taco Monster	Expert - via Adobe*	Netherlands	No interests declared	
Nienke Rodenhuis	Expert - via Adobe*	Netherlands	No interests declared	
Michel Kooijman	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
Steven Teerenstra	Expert - via Adobe*	Netherlands	No interests declared	
Johannes Petrus Theodorus Span	Expert - via Adobe*	Netherlands	No interests declared	
Frank Holtkamp	Expert - via Adobe*	Netherlands	No interests declared	
Hanja de Kooter	Expert - via Adobe*	Netherlands	No restrictions applicable to this meeting	
Hendrik Boersma	Expert - via Adobe*	Netherlands	No restrictions applicable to this meeting	
Ineke Havinga	Expert - via Adobe*	Netherlands	No interests declared	
Anna Cunney	Expert - via Adobe*	Ireland	No interests declared	
Alicia Pérez González	Expert - via Adobe*	Spain	No interests declared	
Annika Ekbom Schnell	Expert - via Adobe*	Sweden	No restrictions applicable to this meeting	
Annika Folin	Expert - via Adobe*	Sweden	No interests declared	
Mair Powell	Expert - via Adobe*	Ireland	No interests declared	
Sandra Bright	Expert - via Adobe*	Ireland	No interests declared	
Brian Aylward	Expert - via Adobe*	Ireland	No interests declared	
Catherine Byrne	Expert - via Adobe*	Ireland	No interests declared	
Finbarr Leacy	Expert - via Adobe*	Ireland	No interests declared	
Geraldine O'Dea	Expert - via Adobe*	Ireland	No interests declared	
Larissa Higgins	Expert - via Adobe*	Ireland	No interests declared	
Benita Cullen	Expert - via Adobe*	Ireland	No interests declared	
Laila Sortvik Nilssen	Expert - via Adobe*	Norway	No restrictions applicable to this meeting	
Frederike Lentz	Expert - via Adobe*	Germany	No interests declared	
Norbert Benda	Expert - via Adobe*	Germany	No interests declared	
Gabriele Schlosser-Weber	Expert - via Adobe*	Germany	No interests declared	
Bojana Divkovic	Expert - via Adobe*	Austria	No restrictions applicable to this meeting	
Ilona Reischl	Expert - via Adobe*	Austria	No interests declared	

Name	Role	Member State or affiliation	Outcome restriction following evaluation of e-DoI	Topics on agenda for which restrictions apply
Angelina Doriguzzi	Expert - via Adobe*	Austria	No restrictions applicable to this meeting	
Christoph Mueck	Expert - via Adobe*	Austria	No interests declared	
Christine Vaculik	Expert - via Adobe*	Austria	No interests declared	
Manfred Schuster	Expert - via Adobe*	Austria	No restrictions applicable to this meeting	
Martina Schussler-Lenz	Expert - via Adobe*	Germany	No interests declared	
Carla Herberts	Expert - via Adobe*	Netherlands	No interests declared	
Minne Casteels	Expert - via Adobe*	Belgium	No interests declared	
Serena Marchetti	Expert - via Adobe*	Netherlands	No restrictions applicable to this meeting	
Anna Vikerfors	Expert - via Adobe*	Sweden	No interests declared	
Linda Trauffler	Expert - via Adobe*	Austria	No interests declared	
Adriana Andrić	Expert - via Adobe*	Croatia	No interests declared	
Caomhim Concannon	Expert - via Adobe*	Ireland	No interests declared	
Mario Miguel da Silva Rosa	Expert - via Adobe*	Portugal	No interests declared	
Lukas Malte Aguirre Davila	Expert - via Adobe*	Germany	No interests declared	
Edward Bojtor	Expert - via Adobe*	Netherlands	No restrictions applicable to this meeting	
Erik Hergarden	Expert - via Adobe*	Netherlands	No interests declared	
Sinead Harrington	Expert - via Adobe*	Ireland	No interests declared	
Taina Mattila	Expert - via Adobe*	Netherlands	No interests declared	
Nathalie Morgensztejn	Expert - via Adobe*	France	No interests declared	
Ulrike Heissenberger	Expert - via Adobe*	Austria	No interests declared	
Sara Galluzzo	Expert - via Adobe*	Italy	No interests declared	
Alessandro Assisi	Expert - via Adobe*	Italy	No interests declared	
Paolo Foggi	Expert - via Adobe*	Italy	No interests declared	
Rosa Giuliani	Expert – via telephone*	Italy	No part in discussions, final deliberations and voting	Tecentriq - atezolizumab - EMEA/H/C/004143/II/0052 - satralizumab - EMEA/H/C/004788

Name	Role	Member State or affiliation	Outcome restriction following evaluation of e-DoI	Topics on agenda for which restrictions apply
				Evrysdi - risdiplam - Orphan - EMEA/H/C/005145 Keytruda - pembrolizumab - EMEA/H/C/003820/II/0097

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)*

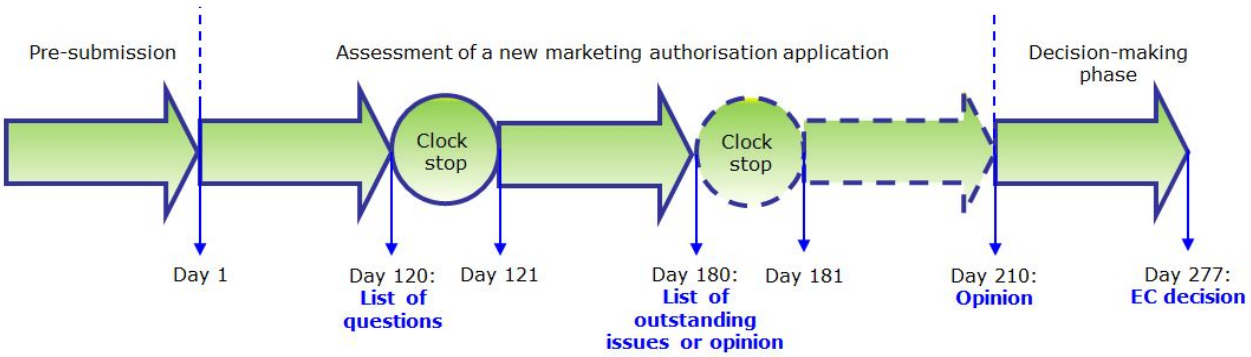
The items listed in this section are those for which marketing authorisation holders (MAHs) or applicants have been invited to the CHMP plenary meeting to address questions raised by the Committee. Oral explanations normally relate to on-going applications (section 3, 4 and 5) or referral procedures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

30 April 2021
EMA/CHMP/171230/2021

Annex to 22-25 February 2021 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 February 2021: **For adoption**	Adopted

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

Final Outcome of Rapporteurship allocation for February 2021: **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

Defitelio - defibrotide - EMEA/H/C/002393/S/0051, Orphan Gentium S.r.l., Rapporteur: Kristina Dunder, PRAC Rapporteur: Ulla Wändel Liminga	Positive Opinion adopted by consensus together with the CHMP assessment report. The Marketing Authorisation remains under exceptional circumstances. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP opinion.
Increlex - mecasermin - EMEA/H/C/000704/S/0064 Ipsen Pharma, Rapporteur: Outi Mäki-Ikola, PRAC Rapporteur: Kirsti Villikka Request for Supplementary Information adopted on 10.12.2020.	Positive Opinion adopted by consensus together with the CHMP assessment report and translation timetable. The Marketing Authorisation remains under exceptional circumstances. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP opinion.
Orphacol - cholic acid - EMEA/H/C/001250/S/0038, Orphan Laboratoires CTRS, Rapporteur: Konstantinos Markopoulos, PRAC Rapporteur: Sofia Trantza	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

Strimvelis - autologous CD34+ enriched cell fraction that contains CD34+ cells transduced with retroviral vector that encodes for the human ADA cDNA sequence - EMEA/H/C/003854/R/0029, Orphan, ATMP Orchard Therapeutics (Netherlands) BV, Rapporteur: Sol Ruiz, Co-Rapporteur: Egbert Flory, CHMP Coordinators: Maria Concepcion Prieto Yerro and Jan Mueller-Berghaus, PRAC Rapporteur: Menno van der Elst Request for Supplementary Information adopted on 04.12.2020.	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 an additional five-year renewal was required. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP Opinion. See 9.1

B.2.2. Renewals of Marketing Authorisations for unlimited validity

Atazanavir Mylan - atazanavir - EMEA/H/C/004048/R/0016 Mylan S.A.S, Generic, Generic of Reyataz, Rapporteur: Bjorg Bolstad, PRAC Rapporteur: Adrien Inoubli	Positive Opinion adopted by consensus together with the CHMP assessment report. Based on the review of the available information, the CHMP was of the opinion that the renewal of the marketing authorisation can be granted with unlimited validity. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP Opinion.
Bortezomib Hospira - bortezomib - EMEA/H/C/004207/R/0020 Pfizer Europe MA EEIG, Generic, Generic of VELCADE, Rapporteur: Daniela Philadelphy, PRAC Rapporteur: Amelia Cupelli	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.
Bortezomib SUN - bortezomib - EMEA/H/C/004076/R/0015 Sun Pharmaceutical Industries Europe B.V., Generic, Generic of VELCADE, Rapporteur: Margareta Bego, PRAC Rapporteur: Amelia Cupelli Request for Supplementary Information adopted on 25.02.2021.	Request for supplementary information adopted with a specific timetable.
CABOMETYX - cabozantinib - EMEA/H/C/004163/R/0018	Positive Opinion adopted by consensus together with the CHMP assessment report and

Ipsen Pharma, Rapporteur: Bjorg Bolstad, Co-Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Menno van der Elst Request for Supplementary Information adopted on 28.01.2021.	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.
Cinqaero - reslizumab - EMEA/H/C/003912/R/0038 Teva B.V., Rapporteur: Johann Lodewijk Hillege, Co-Rapporteur: Jayne Crowe, PRAC Rapporteur: Brigitte Keller-Stanislawski Request for Supplementary Information adopted on 25.02.2021.	Request for supplementary information adopted with a specific timetable.
Nordimet - methotrexate - EMEA/H/C/003983/R/0018 Nordic Group B.V., Rapporteur: Bruno Sepodes, PRAC Rapporteur: Martin Huber Request for Supplementary Information adopted on 25.02.2021.	Request for supplementary information adopted with a specific timetable.
Pemetrexed Fresenius Kabi - pemetrexed - EMEA/H/C/003895/R/0023 Fresenius Kabi Deutschland GmbH, Generic, Generic of Alimta, Rapporteur: Bjorg Bolstad, 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 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.
Zepatier - elbasvir / grazoprevir - EMEA/H/C/004126/R/0026 Merck Sharp & Dohme B.V., Rapporteur: Johann Lodewijk Hillege, Co-Rapporteur: Alexandre Moreau, PRAC Rapporteur: Ana Sofia Diniz Martins Request for Supplementary Information adopted on 28.01.2021.	Positive Opinion adopted by consensus together with the CHMP assessment report. Based on the review of the available information, the CHMP was of the opinion that the renewal of the marketing authorisation can be granted with unlimited validity. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP Opinion.
Zoely - nomegestrol acetate / estradiol - EMEA/H/C/001213/R/0055 Theramex Ireland Limited, Rapporteur: Jean-Michel Race, Co-Rapporteur: Agnes Gyurasics, PRAC Rapporteur: Adrien Inoubli Request for Supplementary Information adopted on 28.01.2021.	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

Deltyba - delamanid - EMEA/H/C/002552/R/0047, Orphan Otsuka Novel Products GmbH, Rapporteur: Christophe Focke, PRAC Rapporteur: Laurence de Fays Request for Supplementary Information adopted on 10.12.2020.	Positive Opinion adopted by consensus together with the CHMP assessment report. 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.
Natpar - parathyroid hormone - EMEA/H/C/003861/R/0027, Orphan Shire Pharmaceuticals Ireland Limited, Rapporteur: Karin Janssen van Doorn, PRAC Rapporteur: Rhea Fitzgerald Request for Supplementary Information adopted on 10.12.2020.	Positive Opinion adopted by consensus together with the CHMP assessment report. 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.
Zynteglo - betibeglogene autotemcel - EMEA/H/C/003691/R/0018, Orphan, ATMP bluebird bio (Netherlands) B.V, Rapporteur: Carla Herberts, CHMP Coordinator: Paula Boudewina van Hennik, PRAC Rapporteur: Brigitte Keller-Stanislawski Request for Supplementary Information adopted on 22.01.2021.	The CHMP noted that all clinical trials are put on hold and that the marketing of the product was temporarily suspended due to possible safety issues. The CHMP was informed that the renewal procedure is put on hold, until the finalisation of an Article 20 referral procedure, which will be triggered to further assess the benefit/risk for Zynteglo. See 9.1

B.3. POST-AUTHORISATION PHARMACOVIGILANCE OUTCOMES

PSUR procedures for which PRAC adopted a recommendation for variation of the terms of the MA at its February 2021 meeting:	
EMEA/H/C/PSUSA/00009255/202007 (perampanel) CAPS: **Fycompa** (EMEA/H/C/002434) (perampanel),	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,

Eisai GmbH, Rapporteur: Alexandre Moreau, PRAC Rapporteur: Tiphaine Vaillant, "23 July 2019 to 22 July 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 section 4.9 of the SmPC to update the adverse reactions reported in case of overdosage. 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/00010379/202007 (nivolumab) CAPS: **OPDIVO** (EMEA/H/C/003985) (nivolumab), Bristol-Myers Squibb Pharma EEIG, Rapporteur: Blanca Garcia-Ochoa, PRAC Rapporteur: Brigitte Keller-Stanislawski, "03/07/2019 To: 03/07/2020"	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 4.8 of the SmPC to add the adverse reaction Lichen Sclerosus and Other Lichen Disorders with a frequency not known (for nivolumab monotherapy) and not known (for nivolumab in combination with ipilimumab, including chemotherapy); update of section 4.4 of the SmPC to revise the warning on Immune-related adverse reactions including information on "simultaneous immune-mediated disorders". 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/00010405/202007 (evolocumab) CAPS: **Repatha** (EMEA/H/C/003766) (evolocumab), Amgen Europe B.V., Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Kimmo Jaakkola, "16/07/2019 To: 16/07/2020"	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 4.8 of the SmPC to add the adverse reaction headache with a frequency common. The Package leaflet is updated accordingly. The Icelandic and the Norwegian CHMP members agree with the above-mentioned recommendation of the CHMP.
EMEA/H/C/PSUSA/00010448/202007 (carfilzomib)	The CHMP, having considered in accordance with Article 28 of Regulation (EC) No 726/2004 the

CAPS: **Kyprolis** (EMEA/H/C/003790) (carfilzomib), Amgen Europe B.V., Rapporteur: Blanca Garcia-Ochoa, PRAC Rapporteur: Nikica Mirošević Skvrce, "18/07/2019 To: 18/07/2020"	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 in order: •to add a warning on bradycardia as symptom of infusion related reaction •to add a warning that reports of ventricular tachycardia have been reported in patients receiving carfilzomib •to reflect the occurrence of QT prolongation also in the post-marketing setting in addition to the existing wording about relevant reports only in clinical studies Update of section 4.8 of the SmPC in order: •to add ventricular tachycardia as an adverse event with frequency uncommon •to add the acute pancreatitis with a frequency uncommon 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/00010697/202007 (inotersen) CAPS: **Tegsedi** (EMEA/H/C/004782) (inotersen), Akcea Therapeutics Ireland Limited, Rapporteur: Martina Weise, PRAC Rapporteur: Rhea Fitzgerald, "04/01/2020 To: 04/07/2020"	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 4.8 of the SmPC to add hypersensitivity with a frequency uncommon. The Package leaflet is updated accordingly. The Icelandic and the Norwegian CHMP members agree with the above-mentioned recommendation of the CHMP.

B.4. EPARs / WPARs

Alymsys - bevacizumab - EMEA/H/C/005286 Mabxience Research SL, 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	For information only. Comments can be sent to the PL in case necessary.

unresectable advanced, metastatic or recurrent non-small cell lung cancer. First-line treatment of patients with advanced and/or metastatic renal cell cancer., Similar biological application (Article 10(4) of Directive No 2001/83/EC)	
BroPair Spiromax - salmeterol xinafoate / fluticasone propionate - EMEA/H/C/005591 Teva B.V., treatment of asthma, Duplicate, Duplicate of Seffalair Spiromax, 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.
Byfavo - remimazolam - EMEA/H/C/005246 PAION Netherlands B.V., indicated for procedural sedation, 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.
Kesimpta - ofatumumab - EMEA/H/C/005410 Novartis Ireland Ltd, treatment of relapsing forms of multiple sclerosis, 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.
Nexpovio - selinexor - EMEA/H/C/005127, Orphan Karyopharm Europe GmbH, treatment of patients with relapsed refractory multiple myeloma (RRMM), 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.
Ontozry - cenobamate - EMEA/H/C/005377 Arvelle Therapeutics Netherlands B.V., 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., 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.
Oyavas - bevacizumab - EMEA/H/C/005556 STADA Arzneimittel AG, 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.	For information only. Comments can be sent to the PL in case necessary.

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., Duplicate, Duplicate of Alymsys, Similar biological application (Article 10(4) of Directive No 2001/83/EC)	
Pemazyre - pemigatinib - EMEA/H/C/005266, Orphan Incyte Biosciences Distribution B.V., treatment of locally advanced or metastatic cholangiocarcinoma, 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.
Seffalair Spiromax - salmeterol xinafoate / fluticasone propionate - EMEA/H/C/004881 Teva B.V., treatment of asthma, 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.
Sogroya - somapacitan - EMEA/H/C/005030, Orphan Novo Nordisk A/S, indicated for the replacement of endogenous growth hormone (GH) in adults with growth hormone deficiency (AGHD), 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.
Thiotepa Riemser - thiotepa - EMEA/H/C/005434 Riemser Pharma GmbH, conditioning treatment prior to haematopoietic progenitor cell transplantation (HPCT), treatment of solid tumours, Generic, Generic of TEPADINA, Generic application (Article 10(1) of Directive No 2001/83/EC)	For information only. Comments can be sent to the PL in case necessary.
Tuzequiva (WD) - trastuzumab - EMEA/H/C/005883 Prestige Biopharma Belgium, treatment of metastatic and early breast cancer and metastatic gastric cancer (MGC), Duplicate, Duplicate of TUZNUE, Similar biological application (Article 10(4) of Directive No 2001/83/EC) **WPAR**	For information only. Comments can be sent to the PL in case necessary.
Vazkepa - icosapent ethyl - EMEA/H/C/005398 Amarin Pharmaceuticals Ireland Limited,	For information only. Comments can be sent to the PL in case necessary.

indicated to reduce cardiovascular risk as an adjunct to statin therapy, New active substance (Article 8(3) of Directive No 2001/83/EC)	

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

AYVAKYT - avapritinib - EMEA/H/C/005208/II/0003/G, Orphan Blueprint Medicines (Netherlands) B.V., Rapporteur: Blanca Garcia-Ochoa Request for Supplementary Information adopted on 04.02.2021.	Request for supplementary information adopted with a specific timetable.
Benlysta - belimumab - EMEA/H/C/002015/II/0090 GlaxoSmithKline (Ireland) Limited, Rapporteur: Kristina Dunder Opinion adopted on 04.02.2021.	Positive Opinion adopted by consensus on 04.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Cinryze - c1 esterase inhibitor (human) - EMEA/H/C/001207/II/0082/G Shire Services BVBA, Rapporteur: Jan Mueller-Berghaus Opinion adopted on 18.02.2021. Request for Supplementary Information adopted on 10.12.2020.	Positive Opinion adopted by consensus on 18.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
COMIRNATY - covid-19 mrna vaccine (nucleoside-modified) - EMEA/H/C/005735/II/0004 BioNTech Manufacturing GmbH, Rapporteur: Filip Josephson Opinion adopted on 04.02.2021. Request for Supplementary Information adopted on 27.01.2021.	Positive Opinion adopted by consensus on 04.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
COMIRNATY - covid-19 mrna vaccine (nucleoside-modified) - EMEA/H/C/005735/II/0005 BioNTech Manufacturing GmbH, Rapporteur: Filip Josephson Opinion adopted on 10.02.2021. Request for Supplementary Information adopted on 01.02.2021.	Positive Opinion adopted by consensus on 10.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
COMIRNATY - covid-19 mrna vaccine (nucleoside-modified) - EMEA/H/C/005735/II/0008/G	Positive Opinion adopted by consensus on 16.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP

BioNTech Manufacturing GmbH, Rapporteur: Filip Josephson Opinion adopted on 16.02.2021. Request for Supplementary Information adopted on 08.02.2021.	recommendation.
COMIRNATY - covid-19 mrna vaccine (nucleoside-modified) - EMEA/H/C/005735/II/0009 BioNTech Manufacturing GmbH, Rapporteur: Filip Josephson Request for Supplementary Information adopted on 09.02.2021.	See also B.6.8 Positive Opinion adopted by consensus on 19.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
COMIRNATY - covid-19 mrna vaccine (nucleoside-modified) - EMEA/H/C/005735/II/0010/G BioNTech Manufacturing GmbH, Rapporteur: Filip Josephson Opinion adopted on 22.02.2021.	See also B.6.8 Positive Opinion adopted by consensus on 22.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
COVID-19 Vaccine Moderna - covid-19 mrna vaccine (nucleoside-modified) - EMEA/H/C/005791/II/0001 Moderna Biotech Spain, S.L., Rapporteur: Jan Mueller-Berghaus Opinion adopted on 15.02.2021. Request for Supplementary Information adopted on 05.02.2021.	Positive Opinion adopted by consensus on 15.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
CRYSVITA - burosumab - EMEA/H/C/004275/II/0017, Orphan Kyowa Kirin Holdings B.V., Rapporteur: Kristina Dunder Opinion adopted on 18.02.2021. Request for Supplementary Information adopted on 12.11.2020, 10.09.2020.	Positive Opinion adopted by consensus on 18.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Cystagon - mercaptamine bitartrate - EMEA/H/C/000125/II/0062 Recordati Rare Diseases, Rapporteur: Maria Concepcion Prieto Yerro Request for Supplementary Information adopted on 04.02.2021.	Request for supplementary information adopted with a specific timetable.
Forsteo - teriparatide - EMEA/H/C/000425/II/0056 Eli Lilly Nederland B.V., Rapporteur: Alexandre Moreau Request for Supplementary Information adopted on 11.02.2021.	Request for supplementary information adopted with a specific timetable.
Forsteo - teriparatide - EMEA/H/C/000425/II/0057/G	Positive Opinion adopted by consensus on 11.02.2021. The Icelandic and Norwegian CHMP

Eli Lilly Nederland B.V., Rapporteur: Alexandre Moreau Opinion adopted on 11.02.2021.	Members were in agreement with the CHMP recommendation.
Hemoblast Bellows - thrombin - EMEA/H/D/002769/II/0008/G BSI Group, The Netherlands B.V, Rapporteur: Armando Genazzani Opinion adopted on 25.02.2021.	Positive Opinion adopted by consensus on 25.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Inflectra - infliximab - EMEA/H/C/002778/II/0094/G Pfizer Europe MA EEIG, Duplicate, Duplicate of Remsima, Rapporteur: Outi Mäki-Ikola Opinion adopted on 25.02.2021. Request for Supplementary Information adopted on 14.01.2021.	Positive Opinion adopted by consensus on 25.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Inhixa - enoxaparin sodium - EMEA/H/C/004264/II/0073/G Techdow Pharma Netherlands B.V., Duplicate, Duplicate of Thorinane (EXP), Rapporteur: Andrea Laslop Opinion adopted on 04.02.2021.	Positive Opinion adopted by consensus on 04.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Kadcyla - trastuzumab emtansine - EMEA/H/C/002389/II/0053 Roche Registration GmbH, Rapporteur: Sinan B. Sarac Opinion adopted on 18.02.2021. Request for Supplementary Information adopted on 14.01.2021.	Positive Opinion adopted by consensus on 18.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Kentera - oxybutynin - EMEA/H/C/000532/II/0059 Teva B.V., Rapporteur: Karin Janssen van Doorn Opinion adopted on 04.02.2021. Request for Supplementary Information adopted on 26.11.2020.	Positive Opinion adopted by consensus on 04.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Lyumjev - insulin lispro - EMEA/H/C/005037/II/0006/G Eli Lilly Nederland B.V., Rapporteur: Outi Mäki-Ikola Opinion adopted on 11.02.2021. Request for Supplementary Information adopted on 26.11.2020.	Positive Opinion adopted by consensus on 11.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Myalepta - metreleptin - EMEA/H/C/004218/II/0017/G, Orphan Amryt Pharmaceuticals DAC, Rapporteur: Karin Janssen van Doorn Request for Supplementary Information adopted	Request for supplementary information adopted with a specific timetable.

on 04.02.2021.	
Myozyme - alglucosidase alfa - EMEA/H/C/000636/II/0084 Genzyme Europe BV, Co-Rapporteur: Karin Janssen van Doorn Opinion adopted on 11.02.2021.	Positive Opinion adopted by consensus on 11.02.2021. 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 Opinion adopted on 11.02.2021. Request for Supplementary Information adopted on 10.12.2020, 15.10.2020.	Positive Opinion adopted by consensus on 11.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Palynziq - pegvaliase - EMEA/H/C/004744/II/0014, Orphan BioMarin International Limited, Rapporteur: Johann Lodewijk Hillege Opinion adopted on 04.02.2021. Request for Supplementary Information adopted on 03.12.2020.	Positive Opinion adopted by consensus on 04.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Privigen - human normal immunoglobulin - EMEA/H/C/000831/II/0169 CSL Behring GmbH, Rapporteur: Jan Mueller-Berghaus Opinion adopted on 11.02.2021. Request for Supplementary Information adopted on 10.12.2020.	Positive Opinion adopted by consensus on 11.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Privigen - human normal immunoglobulin - EMEA/H/C/000831/II/0170/G CSL Behring GmbH, Rapporteur: Jan Mueller-Berghaus Request for Supplementary Information adopted on 04.02.2021.	Request for supplementary information adopted with a specific timetable.
Remsima - infliximab - EMEA/H/C/002576/II/0096/G Celltrion Healthcare Hungary Kft., Rapporteur: Outi Mäki-Ikola Opinion adopted on 25.02.2021. Request for Supplementary Information adopted on 14.01.2021.	Positive Opinion adopted by consensus on 25.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Ruconest - conestat alfa - EMEA/H/C/001223/II/0058 Pharming Group N.V, Rapporteur: Andrea Laslop Opinion adopted on 04.02.2021.	Positive Opinion adopted by consensus on 04.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Vaxelis - diphtheria, tetanus, pertussis (acellular, component), hepatitis B (rDNA),	Positive Opinion adopted by consensus on 04.02.2021. The Icelandic and Norwegian CHMP

poliomyelitis (inact.) and Haemophilus type b conjugate vaccine (adsorbed) - EMEA/H/C/003982/II/0070 MCM Vaccine B.V., Rapporteur: Christophe Focke Opinion adopted on 04.02.2021.	Members were in agreement with the CHMP recommendation.
Veklury - remdesivir - EMEA/H/C/005622/II/0013/G Gilead Sciences Ireland UC, Rapporteur: Janet Koenig Request for Supplementary Information adopted on 25.02.2021.	Request for supplementary information adopted with a specific timetable.
VPRIV - velaglucerase alfa - EMEA/H/C/001249/II/0051, Orphan Shire Pharmaceuticals Ireland Limited, Rapporteur: Martina Weise Opinion adopted on 18.02.2021. Request for Supplementary Information adopted on 14.01.2021.	Positive Opinion adopted by consensus on 18.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Xadago - safinamide - EMEA/H/C/002396/II/0034 Zambon S.p.A., Rapporteur: Johann Lodewijk Hillege Opinion adopted on 11.02.2021. Request for Supplementary Information adopted on 26.11.2020, 02.04.2020, 16.01.2020.	Positive Opinion adopted by consensus on 11.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Xenical - orlistat - EMEA/H/C/000154/II/0083 CHEPLAPHARM Arzneimittel GmbH, Rapporteur: Jean-Michel Race Request for Supplementary Information adopted on 18.02.2021, 15.10.2020.	Request for supplementary information adopted with a specific timetable.
Zavicefta - ceftazidime / avibactam - EMEA/H/C/004027/II/0025/G Pfizer Ireland Pharmaceuticals, Rapporteur: Bjorg Bolstad Request for Supplementary Information adopted on 04.02.2021.	Request for supplementary information adopted with a specific timetable.
WS1926/G **Hexacima-EMEA/H/C/002702/WS1926/0106/G** **Hexaxim (SRD)-EMEA/H/W/002495/WS1926/0111/G** **Hexyon-EMEA/H/C/002796/WS1926/0110/G** Sanofi Pasteur, Lead Rapporteur: Jan Mueller-Berghaus	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 recommendation.

Request for Supplementary Information adopted on 12.11.2020.	
WS1960 **Infanrix hexa-EMEA/H/C/000296/ WS1960/0290** GlaxoSmithkline Biologicals SA, Lead Rapporteur: Christophe Focke Opinion adopted on 04.02.2021.	Positive Opinion adopted by consensus on 04.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1974 **Ratiograstim-EMEA/H/C/000825/ WS1974/0070** **Tevagrastim-EMEA/H/C/000827/ WS1974/0078** TEVA GmbH, Duplicate, Duplicate of Biograstim (SRD), Lead Rapporteur: Outi Mäki-Ikola Request for Supplementary Information adopted on 04.02.2021.	Request for supplementary information adopted with a specific timetable.
WS1991 **Hexacima-EMEA/H/C/002702/WS1991/ 0112** **Hexyon-EMEA/H/C/002796/WS1991/ 0116** Sanofi Pasteur, Lead Rapporteur: Jan Mueller-Berghaus Opinion adopted on 11.02.2021.	Positive Opinion adopted by consensus on 11.02.2021. 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

Adjupanrix - pandemic influenza vaccine (h5n1) (split virion, inactivated, adjuvanted) - EMEA/H/C/001206/II/0072/G GlaxoSmithkline Biologicals SA, Informed Consent of Pandemrix (EXP), Rapporteur: Johann Lodewijk Hillege, "A grouped application of 3 type II variations under category C.I.13: - Submission of a safety pharmacology study performed to assess the effect of AS03 alone and the adjuvanted influenza antigen on cardiovascular and respiratory of telemetered dogs (study MDS AA80120). - Submission of a biodistribution study (study GSK-CH-02-11) conducted in mice with the 3 components of the AS03 Adjuvant System radio-labelled ([14C]-α-tocopherol, [14C]-squalene, and [3H]-polysorbate) to support the understanding of mode of action of AS03.	Positive Opinion adopted by consensus on 04.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

- Submission of a GLP reproductive and developmental toxicity study (study HLS GVB/007/063710) conducted to evaluate the effect of AS03 on embryo-fetal and peri- and post-natal development in Crl:CD (SD) IGS BR rats following intramuscular administration." Opinion adopted on 04.02.2021.	
Beovu - brolucizumab - EMEA/H/C/004913/II/0006 Novartis Europharm Limited, Rapporteur: Alexandre Moreau, "Change(s) in the Summary of Product Characteristics, Labelling or Package Leaflet due to new quality, preclinical, clinical or pharmacovigilance data" Request for Supplementary Information adopted on 25.02.2021.	Request for supplementary information adopted with a specific timetable.
Budesonide/Formoterol Teva Pharma B.V. - budesonide / formoterol fumarate dihydrate - EMEA/H/C/004882/II/0001/G Teva Pharma B.V., Duplicate, Duplicate of DuoResp Spiromax, Rapporteur: John Joseph Borg, "C.I.2.b - Updates of section 4.2 to add information on the use as reliever for allergen- and exercise-induced bronchoconstriction, section 4.4 to revise the general warning on complete withdrawal of inhaled corticosteroids, and section 6.6 to update the statement on special precautions for disposal and other handling following assessment of the same changes for the reference product DuoResp Spiromax. C.I.3.z - update of the SmPC following a PSUR (PSUSA/00010585/201908) for the reference product DuoResp Spiromax to add 'dysphonia' as an adverse drug reaction with a frequency 'common' in section 4.8. The Package Leaflet (PL) and Labelling are updated accordingly. In addition, the Marketing authorisation holder (MAH) took the opportunity to update the list of local representatives in the PL." Opinion adopted on 11.02.2021.	Positive Opinion adopted by consensus on 11.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
CellCept - mycophenolate mofetil - EMEA/H/C/000082/II/0161 Roche Registration GmbH, Rapporteur: Kirstine Moll Harboe, "Update of section 4.4 of the SmPC to amend the existing warning on infections due to potential increase severity of COVID-19 in patients treated with Mycophenolic acid (MPA)	Positive Opinion adopted by consensus on 25.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

based on cumulative reviews from available data. Additionally, consideration of dose adjustment has been suggested in case of clinically significant COVID-19." Opinion adopted on 25.02.2021.	
Cosentyx - secukinumab - EMEA/H/C/003729/II/0069 Novartis Europharm Limited, Rapporteur: Outi Mäki-Ikola, "Update of section 5.1 of the SmPC based on final results from study CAIN457F3302; this is a randomised, double-blind, placebo-controlled study (MAXIMISE) which assessed the efficacy of secukinumab in PsA patients with axial manifestations who were naive to biologic treatment and responded inadequately to NSAIDs; the MAH took this opportunity to introduce minor editorial changes in section 5.1 of the SmPC." Opinion adopted on 18.02.2021. Request for Supplementary Information adopted on 14.01.2021.	Positive Opinion adopted by consensus on 18.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Cyramza - ramucirumab - EMEA/H/C/002829/II/0039 Eli Lilly Nederland B.V., Rapporteur: Paula Boudewina van Hennik, "Update of section 4.8 of the SmPC in order to add hypothyroidism to the list of adverse reactions with a frequency of common based on the updated reference safety information for ramucirumab. The Package Leaflet is also updated. In addition, minor updates are included to the list of local representatives in the product information." Opinion adopted on 04.02.2021.	Positive Opinion adopted by consensus on 04.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Dapivirine Vaginal Ring 25 mg - dapivirine - EMEA/H/W/002168/II/0003 International Partnership for Microbicides Belgium AISBL, Rapporteur: Paula Boudewina van Hennik, "Submission of the final bioanalytical (sub)reports or annexes for the long-term stability experiments for the plasma and vaginal fluid samples of studies IPM 027 and IPM 035." Opinion adopted on 04.02.2021.	Positive Opinion adopted by consensus on 04.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
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	Positive Opinion adopted by consensus on 04.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

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." Opinion adopted on 04.02.2021. Request for Supplementary Information adopted on 15.10.2020.	
Epidyolex - cannabidiol - EMEA/H/C/004675/II/0007, Orphan GW Pharma (International) B.V., Rapporteur: Kirstine Moll Harboe, "Update of section 4.2 of the SmPC in order to add information regarding enteral administration using nasogastric and gastrostomy tubes." Request for Supplementary Information adopted on 25.02.2021, 23.07.2020.	Request for supplementary information adopted with a specific timetable.
Epivir - lamivudine - EMEA/H/C/000107/II/0114 ViiV Healthcare B.V., Rapporteur: Jean-Michel Race, "Update of section 5.2 of the SmPC to add new information about the biotransformation of lamivudine. Furthermore, the MAH took the opportunity to introduce an excipient update in line with the SmPC guideline, a syringe and adapter instruction update in the Package Leaflet and a revision of Annex II in line with the QRD template. Moreover, minor editorial updates have been introduced throughout the Product Information." Opinion adopted on 25.02.2021. Request for Supplementary Information adopted on 14.01.2021.	Positive Opinion adopted by consensus on 25.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Eylea - aflibercept - EMEA/H/C/002392/II/0066 Bayer AG, Rapporteur: Alexandre Moreau, "Submission of final CSR for study 17514 (CENTERA). This was an international, multi-center, prospective, interventional, single-arm, open-label, phase 4 study on the efficacy, durability, posology, and safety of the T&E regimen in subjects with macular edema secondary to CRVO." Opinion adopted on 04.02.2021. Request for Supplementary Information adopted	Positive Opinion adopted by consensus on 04.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

on 29.10.2020.	
Fasenra - benralizumab - EMEA/H/C/004433/II/0031 AstraZeneca AB, Rapporteur: Fátima Ventura, "C.I.13: Submission of the final report from study D3250C00037 (MELTEMI), listed as a category 3 study in the RMP. This is an open-label safety extension study to evaluate the safety and tolerability of a fixed 30 mg dose of benralizumab s.c. in severe asthma patients" Request for Supplementary Information adopted on 11.02.2021.	Request for supplementary information adopted with a specific timetable.
Imfinzi - durvalumab - EMEA/H/C/004771/II/0026 AstraZeneca AB, Rapporteur: Sinan B. Sarac, "Update of section 5.1 of the SmPC in order to update efficacy information on Overall Survival based on the 4-years follow-up analysis of the PACIFIC study (D41991C00001) submitted as recommended by the CHMP; this is a phase III, randomised, double-blind, placebo-controlled, study of Durvalumab as sequential therapy in patients with locally advanced, unresectable non-small cell lung cancer (Stage III)." Opinion adopted on 11.02.2021.	Positive Opinion adopted by consensus on 11.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
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." Opinion adopted on 25.02.2021. Request for Supplementary Information adopted on 28.01.2021, 15.10.2020.	Positive Opinion adopted by consensus on 25.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Lyumjev - insulin lispro - EMEA/H/C/005037/II/0008/G	Positive Opinion adopted by consensus on 11.02.2021. The Icelandic and Norwegian CHMP

Eli Lilly Nederland B.V., Rapporteur: Outi Mäki-Ikola, "Update of sections 4.8 and 5.1 of the SmPC in order to update clinical information based on final results from study I8B-MC-ITRO (PRONTO-Pump-2); this is a Phase 3 prospective, randomized, double-blind trial, which compared Lyumjev to Humalog in adults with Type 1 Diabetes using continuous subcutaneous insulin infusion. The Package Leaflet is updated accordingly. The MAH also provides a phase 2 study evaluating Lyumjev in a Medtronic Pump (Study I8B-MC-ITSM) as a grouped variation." Opinion adopted on 11.02.2021.	Members were in agreement with the CHMP recommendation.
Nimenrix - Meningococcal group A, C, W135 and Y conjugate vaccine - EMEA/H/C/002226/II/0105 Pfizer Europe MA EEIG, Rapporteur: Bjorg Bolstad, "To update section 5.1 Pharmacodynamic properties of the SmPC with information regarding the effectiveness of Nimenrix, to include real-world data from the Netherlands describing the impact of a single dose of Nimenrix on the prevention of meningococcal disease. In addition, a cross-reference to section 4.2 Posology and method of administration of the SmPC was included, to direct the physicians attention to the robust persistence and booster data in section 5.1 and information in section 4.4 Special warnings and precautions for use. In addition, the MAH took the opportunity to include minor editorial changes to the SmPC and to bring the Product information in line with the latest QRD update." Opinion adopted on 11.02.2021.	Positive Opinion adopted by consensus on 11.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Odomzo - sonidegib - EMEA/H/C/002839/II/0035 Sun Pharmaceutical Industries Europe B.V., Rapporteur: Paula Boudewina van Hennik, "Submission of a pooled analysis of drug-related adverse reactions observed in 9 clinical studies with sonidegib, as reflected in the updated Core Data Sheet." Opinion adopted on 25.02.2021.	Positive Opinion adopted by consensus on 25.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Ondexxya - andexanet alfa - EMEA/H/C/004108/II/0003 Alexion Europe SAS, Rapporteur: Jan Mueller-Berghaus, "Submission of the final report from	Request for supplementary information adopted with a specific timetable. See 9.1

Population PK and PK/PD Modelling Report (POR-PKPD-ADEX-321) listed as a category 2 study in the RMP/Specific Obligation 004 in Annex II." Request for Supplementary Information adopted on 28.05.2020, 12.12.2019.	
Quofenix - delafloxacin - EMEA/H/C/004860/II/0009 A. Menarini Industrie Farmaceutiche Riunite s.r.l., Rapporteur: Janet Koenig, "Submission of the final report from study PAE-DELA-01 undertaken to evaluate the impact on the breakpoints of the postantibiotic effect and the delayed re-growth of bacteria following exposure to delafloxacin. The provision of the study report addresses the post-authorisation measure MEA 001." Opinion adopted on 11.02.2021.	Positive Opinion adopted by consensus on 11.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Slenyto - melatonin - EMEA/H/C/004425/II/0017 RAD Neurim Pharmaceuticals EEC SARL, Rapporteur: Kristina Dunder, "The update of the product information to reflect information from children treated with Circadin during the French RTU program and the known safety profile of Circadin authorised for adults." Opinion adopted on 18.02.2021. Request for Supplementary Information adopted on 14.01.2021, 10.09.2020.	Positive Opinion adopted by consensus on 18.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Tecentriq - atezolizumab - EMEA/H/C/004143/II/0054 Roche Registration GmbH, Rapporteur: Sinan B. Sarac, "Update of sections 4.2, 4.4 and 4.8 of the SmPC in order to add severe cutaneous adverse reactions (SCARs) to the list of adverse drug reactions (ADRs) with frequency uncommon, based on the review of safety data presented in a drug safety report (DSR 1105724); the Package Leaflet is updated accordingly. In addition, the MAH took the opportunity to add the term pemphigoid to the description of rash in section 4.8 of the SmPC. The MAH also took the opportunity to update minor typographical errors in the SmPC and PL." Opinion adopted on 25.02.2021.	Positive Opinion adopted by consensus on 25.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Tremfya - guselkumab - EMEA/H/C/004271/II/0026 Janssen-Cilag International N.V., Rapporteur: Agnes Gyurasics, "Update of sections 4.8 and	Positive Opinion adopted by consensus on 11.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP

5.1 of the SmPC in order to implement 1-year psoriatic arthritis clinical data from the pivotal Phase 3 studies CNTO1959PSA3001 and CNTO1959PSA3002. In addition, the MAH took the opportunity to make editorial changes to the product information. The requested variation proposed amendments to the Summary of Product Characteristics and Package Leaflet." Opinion adopted on 11.02.2021.	recommendation.
Vargatef - nintedanib - EMEA/H/C/002569/II/0038 Boehringer Ingelheim International GmbH, Rapporteur: Sinan B. Sarac, "Update of sections 4.4 and 4.8 of the SmPC in order to add nephrotic range proteinuria to the list of adverse drug reactions (ADRs) with frequency common, following the quarterly signal detection in EudraVigilance/EVDAS and based on MAH assessment of safety data retrieved from all completed ICTs conducted with nintedanib and the MAH Global Drug Safety System (GDSS); the Package Leaflet is updated accordingly." Opinion adopted on 11.02.2021.	Positive Opinion adopted by consensus on 11.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Venclyxto - venetoclax - EMEA/H/C/004106/II/0031 AbbVie Deutschland GmbH & Co. KG, Rapporteur: Filip Josephson, "to update venetoclax SmPC wording regarding Tumor lysis syndrome (TLS) prophylaxis and management following an update to the Company Core Data Sheet (CCDS) as result of a medical safety assessment conducted on TLS post-marketing reports. The proposed changes to the SmPC include sections 4.2 and 4.4: • Section 4.2: A more prescriptive table which replaces the text around the risk assessment, prophylaxis and monitoring measures based on the level of tumor burden. In addition, the text on the recommended dose modifications for toxicities is replaced by a table format for clarity. • Section 4.4: the text is revised to emphasise the fact that TLS occur in all patients and requires adequate risk assessment that considers comorbidities, particularly renal impairment, and other risk factors such as splenomegaly." Request for Supplementary Information adopted	Request for supplementary information adopted with a specific timetable.

on 25.02.2021, 12.11.2020.	
Venclyxto - venetoclax - EMEA/H/C/004106/II/0032 AbbVie Deutschland GmbH & Co. KG, Rapporteur: Filip Josephson, "Update of SPC, section 5.1 with new data of venetoclax in combination with rituximab patients with relapsed or refractory chronic lymphocytic leukemia (R/R CLL) from Study GO28667 (MURANO) interim CSR with a CCOD date of 8 May 2020. Study GO28667 is an ongoing open-label, international, multicenter, randomized, Phase III study to investigate the efficacy and safety of venetoclax in combination with rituximab (V+R) compared with bendamustine in combination with rituximab (BR) in patients with R/R CLL. The updated analysis included in this submission presents approximately 60 months of follow-up data. The applicant is also taking advantage of this opportunity to make the below correction and propose editorial changes in the SmPC: - Correcting the upper limit of the confidence interval of the venetoclax + obinituzumab 24-months PFS estimate (92.6 rather than 95.1) in table 5 of section 5.1 of the SmPC. Reference is made to the CSR for study BO25323. - Rounding the percentages across section 5.1 of the SmPC in line with the Rapporteur's comment during the initial MAA procedure." Opinion adopted on 25.02.2021.	Positive Opinion adopted by consensus on 25.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Xarelto - rivaroxaban - EMEA/H/C/000944/II/0079 Bayer AG, Rapporteur: Kristina Dunder, "Submission of the final report from the CASSINI study, an interventional phase III study comparing 10 mg rivaroxaban to placebo in the prevention of venous thromboembolism in ambulatory cancer patients." Request for Supplementary Information adopted on 18.02.2021, 03.09.2020.	Request for supplementary information adopted with a specific timetable.
Zejula - niraparib - EMEA/H/C/004249/II/0024, Orphan GlaxoSmithKline (Ireland) Limited, Rapporteur: Bjorg Bolstad, "Update of sections 4.2 and 5.2 of the SmPC in order to include information based on final results from hepatic study 3000-01-003 (HEPATIC). The Package Leaflet is	Request for supplementary information adopted with a specific timetable.

updated accordingly. In addition, the MAH took the opportunity to update the list of local representatives in the Package Leaflet and to introduce editorial changes in SmPC section 4.4." Request for Supplementary Information adopted on 25.02.2021.	
WS1874/G **Advagraf-EMEA/H/C/000712/WS1874/0058/G** **Modigraf-EMEA/H/C/000954/WS1874/0036/G** Astellas Pharma Europe B.V., Lead Rapporteur: Jayne Crowe, "C.I.4 Update of sections 4.4, and 4.5 of the SmPC on the drug-drug interaction with CYP3A4 based on a comprehensive review of available data. Section 4.5 of the SmPC is also updated to include impact of direct acting antiviral therapy. C.I.z Update of section 4.8 of the SmPC to add posterior reversible encephalopathy syndrome as an adverse reaction. The MAH took also the opportunity to change the SOC for febrile neutropenia from General disorders and administration site conditions to Blood and lymphatic system disorders in section 4.8 of the SmPC and to update the instruction for handling of the product in section 6.6 of the SmPC. The Package Leaflet is updated accordingly." Request for Supplementary Information adopted on 25.02.2021, 17.09.2020.	Request for supplementary information adopted with a specific timetable.
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	Positive Opinion adopted by consensus on 25.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

local representatives in the Package Leaflet." Opinion adopted on 25.02.2021. Request for Supplementary Information adopted on 10.12.2020.	

B.5.3. CHMP-PRAC assessed procedures

Aimovig - erenumab - EMEA/H/C/004447/II/0013/G Novartis Europharm Limited, Rapporteur: Kristina Dunder, PRAC Rapporteur: Kirsti Villikka, "Update of section 4.8 of the SmPC in line with revised clinical safety data. Submission of the study report from 5-year open-label study 20120178 with consequential changes to the sections 4.8 and 5.1 of the SmPC as well as an update of the EU RMP. Type IA variation to the include ATC code for erenumab. The Package Leaflet is updated accordingly." Request for Supplementary Information adopted on 11.02.2021.	Request for supplementary information adopted with a specific timetable.
CRYSVITA - burosumab - EMEA/H/C/004275/II/0021, Orphan Kyowa Kirin Holdings B.V., Rapporteur: Kristina Dunder, PRAC Rapporteur: Brigitte Keller-Stanislawski, "Update of section 4.2 of the SmPC in order to modify administration instructions to include the option of self/carer-administration based on results from two Phase 3 interventional clinical safety and efficacy studies; Study KRN23-003 in paediatric patients (final study report) and Study KRN23-004 in adult patients (interim report). The Package Leaflet has been updated accordingly and a new section with instructions for use has been added at the end. In addition, the MAH took the opportunity to implement editorial changes in the SmPC and Package Leaflet. The updated RMP version 3.0 has also been submitted." Request for Supplementary Information adopted on 11.02.2021.	Request for supplementary information adopted with a specific timetable.
Dengvaxia - dengue tetravalent vaccine (live, attenuated) - EMEA/H/C/004171/II/0016/G Sanofi Pasteur, Rapporteur: Christophe Focke, PRAC Rapporteur: Sonja Hrabcik, "Update of section 4.5. of the SmPC to include	Request for supplementary information adopted with a specific timetable.

coadministration data on Gardasil/Cervarix/Adacel from CYD67, CYD71 and CYD66 final study reports respectively (final reports from 3 PAM MEA studies listed as category 3 studies in the RMP); these studies are dedicated to immunogenicity and safety of the concomitant administration; the Package Leaflet is updated accordingly. The RMP version 6.1. 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 11.02.2021.	
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 is approved. In addition, the MAH took the opportunity to update the list of local representatives in the Package leaflet." Opinion adopted on 18.02.2021. Request for Supplementary Information adopted on 14.01.2021, 01.10.2020.	Positive Opinion adopted by consensus on 18.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Erleada - apalutamide - EMEA/H/C/004452/II/0009 Janssen-Cilag International N.V., Rapporteur: Blanca Garcia-Ochoa, PRAC Rapporteur: Tiphaine Vaillant, "Update of section 5.3 of the SmPC in order to include non-clinical information based on final results from a 26-week carcinogenicity study (TOX13540) listed as a category 3 study in the RMP. The RMP version 3.2 is approved." Opinion adopted on 18.02.2021. Request for Supplementary Information adopted on 14.01.2021.	Positive Opinion adopted by consensus on 18.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Eylea - aflibercept - EMEA/H/C/002392/II/0069 Bayer AG, Rapporteur: Alexandre Moreau, PRAC	Request for supplementary information adopted with a specific timetable.

Rapporteur: Tiphaine Vaillant, "This type II variation under category C.1.4 is to update the Posology section 4.2 of the Product Information for the indication DME based on results from the PAES VIOLET (Study 17613; (EMEA/H/C/002392/ANX/011) and to include study data to EU-PI section 5.1. The submission package also contains the AQUA CSR, a phase 4 study which served as run-in study for VIOLET." Request for Supplementary Information adopted on 25.02.2021.	
Herceptin - trastuzumab - EMEA/H/C/000278/II/0168 Roche Registration GmbH, Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Brigitte Keller-Stanislawski, "Update of sections 4.2 and 4.4 of the SmPC in order to modify the administration instructions by removing the observation time currently stipulated after administration and to amend the existing warning respectively based on final results from study SafeHER (MO28048) listed as a category 3 study in the RMP; this is a Phase III prospective, two Cohort nonrandomized, multicentre, multinational, open label study to assess the safety of assisted- and self-administered subcutaneous Herceptin as adjuvant therapy in patients with operable HER2- positive early breast cancer. The Package Leaflet is updated accordingly. The RMP version 21 has also been submitted." Request for Supplementary Information adopted on 11.02.2021.	Request for supplementary information adopted with a specific timetable.
Kaftrio - ivacaftor / tezacaftor / elexacaftor - EMEA/H/C/005269/II/0003, Orphan Vertex Pharmaceuticals (Ireland) Limited, Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Martin Huber, "C.I.13: Submission of the final clinical study report for study VX18-445-007 (study 007), listed as a category 3 study in the RMP with the aim to evaluate the pharmacokinetics of Kaftrio (elexacaftor/tezacaftor/ivacaftor) in subjects with moderate hepatic impairment. The RMP version 1.2 has also been submitted." Opinion adopted on 11.02.2021.	Positive Opinion adopted by consensus on 11.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Kisplyx - lenvatinib - EMEA/H/C/004224/II/0041	Request for supplementary information adopted with a specific timetable.

Eisai GmbH, Rapporteur: Karin Janssen van Doorn, PRAC Rapporteur: David Olsen, "Update of sections 4.5 and 5.1 of the SmPC in order to update the drug-drug interaction with everolimus and to update the efficacy information based on the results from the study E7080-M001-221. Study 221 is a single-arm, multicenter, Phase 2 trial to evaluate the safety and efficacy of lenvatinib in combination with everolimus in subjects with unresectable advanced or metastatic non clear cell renal cell carcinoma (nccRCC) who have not received any chemotherapy for advanced disease. (MEA 008.1). The RMP version 12.1 has also been submitted." Request for Supplementary Information adopted on 11.02.2021.	
Kisplyx - lenvatinib - EMEA/H/C/004224/II/0042 Eisai GmbH, Rapporteur: Karin Janssen van Doorn, PRAC Rapporteur: David Olsen, "Submission of the final Clinical Study Report for study E7080-G000-218. Study 218 is a randomized, open-label (formerly double-blind), Phase 2 trial to assess safety and efficacy of Lenvatinib at two different starting doses (18 mg vs 14 mg QD) in combination with Everolimus (5 mg QD) in Renal Cell Carcinoma following one prior VEGF-Targeted treatment. (MEA 007.3). The RMP 12.2 has also been submitted." Opinion adopted on 11.02.2021.	Positive Opinion adopted by consensus on 11.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Lorviqua - lorlatinib - EMEA/H/C/004646/II/0013 Pfizer Europe MA EEIG, Rapporteur: Sinan B. Sarac, PRAC Rapporteur: Nikica Mirošević Skvrce, "Update of sections 4.2, 4.4 and 4.8 of the SmPC in order to include hypertension and hyperglycaemia as new adverse drug reactions (ADRs) with frequency common and very common respectively together with recommended dose modifications and warnings, based on data from Phase 3 study B7461006, comparing lorlatinib versus crizotinib for the first line treatment of advanced ALK-positive NSCLC. In addition, the pooled safety dataset has been updated to include data from studies B7461001, a Phase 1/2 open-label, multiple-dose, dose-escalation, safety, pharmacokinetic,	Request for supplementary information adopted with a specific timetable.

pharmacodynamic, and anti-tumour efficacy exploration study and B7461006. As a consequence, the frequencies of ADRs have been updated in section 4.8 of the SmPC and existing warnings on Hyperlipidaemia and Lipase and amylase increase have been amended. The Package Leaflet is updated accordingly. The RMP version 2.0 has also been submitted." Request for Supplementary Information adopted on 25.02.2021.	
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 is approved." Opinion adopted on 11.02.2021. Request for Supplementary Information adopted on 26.11.2020.	Positive Opinion adopted by consensus on 11.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Maviret - glecaprevir / pibrentasvir - EMEA/H/C/004430/II/0039 AbbVie Deutschland GmbH & Co. KG, Rapporteur: Jean-Michel Race, PRAC Rapporteur: Ana Sofia Diniz Martins, "Update of section 5.1 of the SmPC based on results from study M13-576; a non-drug interventional follow-up study to assess resistance and durability of response to AbbVie direct-acting antiviral agent (DAA) therapy (ABT-493 and/or ABT-530) in subjects who participated in Phase 2 or 3 clinical studies for the treatment of chronic hepatitis C Virus (HCV) infection. The study is included as a category 3 study in the RMP, and an updated RMP version 6.0 has also been submitted." Request for Supplementary Information adopted	Request for supplementary information adopted with a specific timetable.

on 11.02.2021.	
Ocrevus - ocrelizumab - EMEA/H/C/004043/II/0021 Roche Registration GmbH, Rapporteur: Kirstine Moll Harboe, PRAC Rapporteur: Brigitte Keller-Stanislawski, "Update of sections 4.4 and 4.8 in order to include the term 'anaphylaxis' among the possible symptoms of infusion-related reactions (IRRs), following an analysis of cases retrieved by anaphylactic reaction MedDRA narrow SMQ. The MAH took the opportunity to update Annex II.C and D in line with the QRD template. The RMP version 7.0 has been submitted." Opinion adopted on 11.02.2021. Request for Supplementary Information adopted on 29.10.2020.	Positive Opinion adopted by consensus on 11.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
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 (1) the final 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 (2) interim results of 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 updated to remove study 12-266 A(12). The RMP (version 4.1) is updated accordingly. An editorial change has been added to the SmPC." Opinion adopted on 11.02.2021. Request for Supplementary Information adopted on 01.10.2020.	Positive Opinion adopted by consensus on 11.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Steglujan - ertugliflozin / sitagliptin - EMEA/H/C/004313/II/0015 Merck Sharp & Dohme B.V., Rapporteur: Kristina Dunder, PRAC Rapporteur: Menno van der Elst, "Update of sections 4.2, 4.4, 4.8 and 5.1 of the SmPC for Steglujan in order to update clinical information following final results from the VERTIS CV study (protocol 8835-004/B1521021) listed as a category 3 study in	Request for supplementary information adopted with a specific timetable.

the RMP. This is a multi-centre, multi-national, randomised, double-blind, placebo-controlled study to evaluate the effect of ertugliflozin on cardiovascular risk in adult patients with type 2 diabetes and established atherosclerotic cardiovascular disease. The Package Leaflet is updated accordingly. The RMP version 2.0 has also been submitted. In addition, the MAH took the opportunity to include an editorial change in section 4.1 of the SmPC." Request for Supplementary Information adopted on 25.02.2021.	
Tecentriq - atezolizumab - EMEA/H/C/004143/II/0053 Roche Registration GmbH, Rapporteur: Sinan B. Sarac, PRAC Rapporteur: Marcia Sofia Sanches de Castro Lopes Silva, "Submission of the final report from study GO29322 listed as a category 3 study in the RMP. This is a Phase Ib study investigating the safety and pharmacology of atezolizumab administered with ipilimumab, interferon-alpha, or other immunomodulating therapies in patients with locally advanced or metastatic solid tumours. The RMP version 19.1 has also been submitted to remove this category 3 study along with the related safety concern of concomitant use with other immuno-modulatory drugs." Opinion adopted on 11.02.2021.	Positive Opinion adopted by consensus on 11.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Zercepac - trastuzumab - EMEA/H/C/005209/II/0003 Accord Healthcare S.L.U., Rapporteur: Sol Ruiz, PRAC Rapporteur: Brigitte Keller-Stanislawski Opinion adopted on 11.02.2021.	Positive Opinion adopted by consensus on 11.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1965/G **Hexacima-EMEA/H/C/002702/WS1965/0110/G** **Hexaxim (SRD)-EMEA/H/W/002495/WS1965/0115/G** **Hexyon-EMEA/H/C/002796/WS1965/0114/G** Sanofi Pasteur, Lead Rapporteur: Jan Mueller-Berghaus, Lead PRAC Rapporteur: Brigitte Keller-Stanislawski, "C.I.4 (type II): Update of section 5.1 of the SmPC in order to describe the persistence of anti-HBs antibodies in subjects 6 years of age having received a hexavalent vaccine based on the final results from study A3L00052; this is a phase IV, open-label, multi-	Positive Opinion adopted by consensus on 18.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

centre study in children previously vaccinated in study A3L38a with 3 doses of either Hexacima/Hexyon/Hexaxim (Group 1) or Infanrix Hexa (Group 2). C.I.4 (type II): Update of sections 4.4 and 5.1 of the SmPC in order to reword safety and immunogenicity information regarding individuals with immunodeficiency based on the final results from study A3L44; this is a Phase III, single centre, open-label, two-arm study including HIV-exposed infected and uninfected infants vaccinated with a 3-dose infant primary series (at 6, 10, and 14 weeks of age) and a booster dose (at 15 to 18 months of age) with Hexacima/Hexyon/Hexaxim in Republic of South Africa. The updates to the SmPC were requested following assessment of these data by Article 46, EMEA/H/C/002702/P46/036 (Hexacima), EMEA/H/C/002796/P46/034 (Hexyon) and EMEA/H/W/002495/P46/036 (Hexaxim). C.I.z (type IB): Update of section 4.4 of the SmPC in order to include syncope within the precautions for use. The package leaflet is updated accordingly. In addition, the WSA took the opportunity to update the list of local representatives in the Package Leaflet. The RMP version 13.0 has also been submitted." Opinion adopted on 18.02.2021. Request for Supplementary Information adopted on 14.01.2021.	

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	Request for supplementary information adopted with a specific timetable.

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 11.02.2021, 01.10.2020.	
PRAC Led **Alecensa - alectinib - EMEA/H/C/004164/II/0030** Roche Registration GmbH, Rapporteur: Filip Josephson, PRAC Rapporteur: Jana Lukacisinova, PRAC-CHMP liaison: Ondřej Slanař, "Submission of the final report from study (BO40643) listed as an additional pharmacovigilance activity in the RMP. This is a non-interventional post authorisation safety study (PASS) aimed at evaluating the effectiveness of the risk minimisation measures (RMMs) for the important identified risks of ILD/pneumonitis, hepatotoxicity, bradycardia, photosensitivity, severe myalgia, and CPK elevations for Alecensa." Opinion adopted on 11.02.2021.	Positive Opinion adopted by consensus on 11.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
PRAC Led **Bexsero - meningococcal group B vaccine (recombinant, component, adsorbed) - EMEA/H/C/002333/II/0098** GSK Vaccines S.r.l, Rapporteur: Kristina Dunder, PRAC Rapporteur: Ulla Wändel Liminga, PRAC-CHMP liaison: Kristina Dunder, "Submission of the final report from study V72_36OB, an observational PASS conducted by the Public Health England to further characterise the important potential risks of seizures (including febrile seizures), vasculitis/Kawasaki syndrome (KD), anaphylaxis (including anaphylactic shock), Acute Disseminated Encephalomyelitis (ADEM), and Glillain-Barré Syndrome (GBS) in routine UK care. The study is listed as a category 3 study in the RMP. The revised RMP version 9.0 has also been submitted."	Positive Opinion adopted by consensus on 11.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

Opinion adopted on 11.02.2021.	
PRAC Led **Circadin - melatonin - EMEA/H/C/000695/II/0061** RAD Neurim Pharmaceuticals EEC SARL, 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"." Opinion adopted on 11.02.2021. Request for Supplementary Information adopted on 14.01.2021, 01.10.2020.	Positive Opinion adopted by consensus on 11.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
PRAC Led **Kineret - anakinra - EMEA/H/C/000363/II/0078** Swedish Orphan Biovitrum AB (publ), Rapporteur: Kirstine Moll Harboe, PRAC Rapporteur: Anette Kirstine Stark, PRAC-CHMP liaison: Kirstine Moll Harboe, "Submission of the final report from study (Sobi-ANAKIN-201) listed as a category 3 study in the RMP. This is a non-interventional post-authorisation safety study to evaluate the safety of Kineret in the treatment of Cryopyrin Associated Periodic Syndromes (CAPS) in routine clinical care with regard to serious infections, malignancies, injection site reactions, allergic reactions and medication errors, including reuse of syringe. The RMP version 5.4 has been submitted to reflect completion of this study. In addition, the RMP is updated to include information about a completed paediatric study (Sobi.ANAKIN-301) assessed as per Article 46 of Reg No 1901/2006 (EMEA/H/C/000363/P46/031). This was a randomised, double-blind, placebo-controlled, multicenter, phase 3 study which evaluated the efficacy, the safety, pharmacokinetics and immunogenicity of anakinra as compared to placebo in newly diagnosed Still's disease patients (including systemic juvenile idiopathic arthritis [SJIA] and adult-onset Still's disease [AOSD])." Request for Supplementary Information adopted on 11.02.2021.	Request for supplementary information adopted with a specific timetable.
PRAC Led	Request for supplementary information adopted

Levemir - insulin detemir - EMEA/H/C/000528/II/0101 Novo Nordisk A/S, Rapporteur: Kirstine Moll Harboe, PRAC Rapporteur: Anette Kirstine Stark, PRAC-CHMP liaison: Kirstine Moll Harboe, "Update of sections 4.6 and 5.1 of the SmPC in order to update information on pregnancy, based on final results from the non-interventional Post-Authorisation Safety Study, NN304-4016, listed as a category 3 study in the RMP. This is a diabetes pregnancy registry study conducted to assess the long-term safety of insulin use in pregnant women. The RMP version 21.0 has also been submitted." Request for Supplementary Information adopted on 11.02.2021.	with a specific timetable.
PRAC Led **Nerlynx - neratinib - EMEA/H/C/004030/II/0020** Pierre Fabre Medicament, Rapporteur: Bruno Sepodes, PRAC Rapporteur: Menno van der Elst, PRAC-CHMP liaison: Johann Lodewijk Hillege, "Submission of an updated RMP version 1.1 in order to add a new important identified risk, update data concerning the post authorisation safety studies and change of submission due date of the final Study Report of the PASS n°6201 (MEA 001) ." Request for Supplementary Information adopted on 11.02.2021.	Request for supplementary information adopted with a specific timetable.
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, "Submission of the final report from study FBB-01_03_13 (PASS-2) (listed as a category 3 study in the RMP): a non-interventional, cross-sectional, retrospective, multicentre, multi-country registry to observe usage pattern, safety and tolerability of the diagnostic agent NeuraCeq (florbetaben (18F)) in European clinical practice. The RMP (version 5.9) is updated accordingly" Opinion adopted on 11.02.2021. Request for Supplementary Information adopted on 26.11.2020.	Positive Opinion adopted by consensus on 11.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
PRAC Led	Positive Opinion adopted by consensus on

Orfadin - nitisinone - EMEA/H/C/000555/II/0074 Swedish Orphan Biovitrum International AB, PRAC Rapporteur: Amelia Cupelli, PRAC-CHMP liaison: Armando Genazzani, "Submission of the final report from study Sobi.NTBC-005 listed as a category 3 study in the RMP. This is a non-interventional Post Authorisation Safety Study (PASS) to evaluate long-term safety of Orfadin treatment in hypertyrosinemia type 1 (HT-1) patients in standard clinical care. The RMP version 5.5 has also been submitted." Opinion adopted on 11.02.2021. Request for Supplementary Information adopted on 26.11.2020, 03.09.2020.	11.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
PRAC Led **Repatha - evolocumab - EMEA/H/C/003766/II/0047** Amgen Europe B.V., Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Kimmo Jaakkola, PRAC-CHMP liaison: Outi Mäki-Ikola, "C.I.3, Update of section 4.8 of the SmPC in order to add myalgia to the list of adverse drug reactions (ADRs) with frequency (frequency category) common following the review of nonclinical, clinical, post-marketing safety, and external spontaneous reporting databases as requested in the PSUR. The Package Leaflet are updated accordingly. In addition, the MAH took the opportunity to add a traceability statement in line with a statement previously added to the SmPC and to propose minor updates to instructions for use of evolocumab SureClick pre-filled pen for enhanced usability." Opinion adopted on 11.02.2021.	Positive Opinion adopted by consensus on 11.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
PRAC Led **RotaTeq - rotavirus vaccine (live, oral) - EMEA/H/C/000669/II/0085** MSD Vaccins, Rapporteur: Kristina Dunder, PRAC Rapporteur: Ulla Wändel Liminga, PRAC-CHMP liaison: Kristina Dunder, "To update the RMP for RotaTeq to version 7.2 to meet the requirements and updated definitions in the Guideline on good pharmacovigilance practices (GVP) module V (EMA/838713/2011; Rev 2); consequently, the list of safety concerns is updated and a reclassification of important risks is proposed. In addition, the proposed RMP version 7.2 implements the removal of	Request for supplementary information adopted with a specific timetable.

hypersensitivity and severe combined immunodeficiency (SCID) from the list of safety concerns as requested by the PRAC in PSUR procedure (PSUSA/00002666/201911)." Request for Supplementary Information adopted on 11.02.2021.	
PRAC Led **SIRTURO - bedaquiline - EMEA/H/C/002614/II/0042, Orphan** Janssen-Cilag International NV, Rapporteur: Filip Josephson, PRAC Rapporteur: Ulla Wändel Liminga, PRAC-CHMP liaison: Filip Josephson, "Submission of the final report of the PASS TMC207TBC4002, a non-interventional multi-country multidrug-resistant tuberculosis patient registry in South Africa and South Korea to monitor bedaquiline safety, utilisation, and emergence of resistance. The study is listed as a category 3 study in the RMP, and with this submission the MAH fulfils the Post Authorisation Measure MEA 010.6. The updated RMP version 8.1 has also been submitted." Opinion adopted on 11.02.2021.	Positive Opinion adopted by consensus on 11.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
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." Opinion adopted on 11.02.2021. Request for Supplementary Information adopted on 26.11.2020.	Positive Opinion adopted by consensus on 11.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
PRAC Led **Tremfya - guselkumab - EMEA/H/C/004271/II/0025** Janssen-Cilag International N.V., Rapporteur: Agnes Gyurasics, PRAC Rapporteur: Brigitte Keller-Stanislawski, PRAC-CHMP liaison: Jan Mueller-Berghaus, "Submission of an updated RMP version 7.1 in order to amend the study population for Psoriasis registry C0168Z03 (PSOLAR) defined as Additional	Positive Opinion adopted by consensus on 11.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

Pharmacovigilance Activities in the RMP. The amended protocol of the registry is included for assessment." Opinion adopted on 11.02.2021.	
PRAC Led **Trulicity - dulaglutide - EMEA/H/C/002825/II/0051** Eli Lilly Nederland B.V., Rapporteur: Martina Weise, PRAC Rapporteur: Ilaria Baldelli, PRAC-CHMP liaison: Armando Genazzani, "Update of section 4.8 of the SmPC to add 'delayed gastric emptying' as a new ADR with a frequency of rare, based on the final study report for the PASS category 3 dulaglutide drug utilisation study H9X-MC-B009: a multi-database collaborative research program of observational studies to monitor the utilisation and safety of dulaglutide in the EU (ref. PAM MEA 002-002.5), and taking into account the data from the pooled clinical trials, REWIND trial, post-marketing surveillance and Eudravigilance. The Package Leaflet has been updated accordingly. An updated RMP version 6.2 was agreed during the procedure." Opinion adopted on 11.02.2021. Request for Supplementary Information adopted on 26.11.2020, 09.07.2020.	Positive Opinion adopted by consensus on 11.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
PRAC Led **Truvada - emtricitabine / tenofovir disoproxil - EMEA/H/C/000594/II/0169** Gilead Sciences Ireland UC, Rapporteur: Bruno Sepodes, PRAC Rapporteur: Ana Sofia Diniz Martins, PRAC-CHMP liaison: Bruno Sepodes, "Submission of an updated RMP (version 16.1) to remove from the Pharmacovigilance Plan - two completed category 3 studies (study GS-US-276-0103 and study GS-EU-276-4027) and - the category 3 additional pharmacovigilance activity for the registry study GS- EU- 276-4487 (a category 3 study in the RMP): a prospective, longitudinal, observational registry of emtricitabine/tenofovir disoproxil fumarate for human immunodeficiency virus 1 (HIV-1) pre-exposure prophylaxis (PrEP) in the European Union." Request for Supplementary Information adopted on 11.02.2021.	Request for supplementary information adopted with a specific timetable.
PRAC Led **VPRIV - velaglucerase alfa -**	Request for supplementary information adopted with a specific timetable.

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)" Request for Supplementary Information adopted on 11.02.2021, 26.11.2020.	
PRAC Led **Yondelis - trabectedin - EMEA/H/C/000773/II/0061** Pharma Mar, S.A., Rapporteur: Sinan B. Sarac, PRAC Rapporteur: Anette Kirstine Stark, PRAC-CHMP liaison: Sinan B. Sarac, "Submission of an updated RMP version 9.0 in order to reflect new available data from completed studies, removal of safety concerns, removal of a target follow-up questionnaire and update of the format in line with the guidance "EMA/164014/2018 Rev.2.0.1 accompanying GVP Module V Rev.2"." Opinion adopted on 11.02.2021. Request for Supplementary Information adopted on 29.10.2020.	Positive Opinion adopted by consensus on 11.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
PRAC Led **WS1805** **Advagraf-EMEA/H/C/000712/WS1805/0057** **Modigraf-EMEA/H/C/000954/WS1805/0035** Astellas Pharma Europe B.V., Lead Rapporteur: Jayne Crowe, Lead PRAC Rapporteur: Ronan Grimes, PRAC-CHMP liaison: Jayne Crowe, "Submission of an updated RMP version 3.2 in order to add a non-interventional post-authorisation safety study related to the safety concerns of use during pregnancy and use during lactation. Removal of DHPC on medication errors sent to HCPs in 2008. The RMP is being brought to EU RMP template revision 2." Opinion adopted on 11.02.2021. Request for Supplementary Information adopted	Positive Opinion adopted by consensus on 11.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

on 29.10.2020, 09.07.2020.	
PRAC Led **WS2000** **Leganto-EMEA/H/C/002380/WS2000/0035** **Neupro-EMEA/H/C/000626/WS2000/0089** UCB Pharma S.A., Lead Rapporteur: Bruno Sepodes, Lead PRAC Rapporteur: Ana Sofia Diniz Martins, PRAC-CHMP liaison: Bruno Sepodes, "Type II WS: C.I.11.b for RMP: Submission of an updated RMP version 5.0 in order to update RMP according to Good Pharmacovigilance Practices (GVP) Module V template (Rev 2)." Opinion adopted on 11.02.2021.	Positive Opinion adopted by consensus on 11.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

B.5.5. CHMP-CAT assessed procedures

Imlygic - talimogene laherparepvec - EMEA/H/C/002771/II/0041/G, ATMP Amgen Europe B.V., Rapporteur: Olli Tenhunen, CHMP Coordinator: Johanna Lähteenvuo Opinion adopted on 25.02.2021, 19.02.2021.	Positive Opinion adopted by consensus on 25.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Kymriah - tisagenlecleucel - EMEA/H/C/004090/II/0030, Orphan, ATMP Novartis Europharm Limited, Rapporteur: Rune Kjeken, CHMP Coordinator: Ingrid Wang, "Update of section 5.1 of the SmPC to include the complete data set (updated overall survival analysis) of study CCTL019B2205J, a Phase II, single arm, multicenter study to determine the efficacy and safety of Kymriah in paediatric subjects with relapsed or refractory B-cell ALL. The clinical results have already been assessed in procedure EMA/H/C/004090/P46/011. In addition, the final ATC code for tisagenlecleucel (L01XX71) has been added as an editorial change." Opinion adopted on 25.02.2021, 19.02.2021. Request for Supplementary Information adopted on 22.01.2021.	Positive Opinion adopted by consensus on 25.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
Spherox - spheroids of human autologous matrix-associated chondrocytes - EMEA/H/C/002736/II/0021/G, ATMP CO.DON AG, Rapporteur: Lisbeth Barkholt, CHMP Coordinator: Kristina Dunder	Request for supplementary information adopted with a specific timetable.

Request for Supplementary Information adopted on 19.02.2021.	
Zolgensma - onasemnogene abeparvovec - EMEA/H/C/004750/II/0008, Orphan, ATMP Novartis Gene Therapies EU Limited, Rapporteur: Carla Herberts, CHMP Coordinator: Johann Lodewijk Hillege, "Updates to SmPC sections 4.2, 4.4 and 4.8 to reflect the risk of Thrombotic Microangiopathy, recommend additional tests to help early identification of this risk and advice for prompt clinical management. The Package Leaflet is updated accordingly." Opinion adopted on 25.02.2021, 19.02.2021. Request for Supplementary Information adopted on 04.12.2020.	Positive Opinion adopted by consensus on 25.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

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 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 19.02.2021, 09.10.2020.	Request for supplementary information adopted with a specific timetable.

B.5.7. PRAC assessed ATMP procedures

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

WS1930 **Hexacima-EMEA/H/C/002702/WS1930/0107** **Hexaxim (SRD)-EMEA/H/W/002495/WS1930/0112** **Hexyon-EMEA/H/C/002796/WS1930/0111** Sanofi Pasteur, Lead Rapporteur: Jan Mueller-Berghaus Request for Supplementary Information adopted on 03.12.2020.	Request for supplementary information adopted with a specific timetable.
WS1973 **Infanrix hexa-EMEA/H/C/000296/WS1973/0291** GlaxoSmithkline Biologicals SA, Lead Rapporteur: Christophe Focke Opinion adopted on 04.02.2021.	Positive Opinion adopted by consensus on 04.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1977 **Enurev Breezhaler-EMEA/H/C/002691/WS1977/0033** **Seebri Breezhaler-EMEA/H/C/002430/WS1977/0033** **Tovanor Breezhaler-EMEA/H/C/002690/WS1977/0037** Novartis Europharm Limited, Lead Rapporteur: Kirstine Moll Harboe, Opinion adopted on 04.02.2021.	Positive Opinion adopted by consensus on 04.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1979/G **Actos-EMEA/H/C/000285/WS1979/0084/G** **Competact-EMEA/H/C/000655/WS1979/0076/G** **Glubrava-EMEA/H/C/000893/WS1979/0062/G** **Glustin-EMEA/H/C/000286/WS1979/0083/G** **Tandemact-EMEA/H/C/000680/WS1979/0065/G** Takeda Pharma A/S, Lead Rapporteur: Peter Kiely, "Type IB, Category C.I.z. - Update of the Product Information (PI) with Sodium content wording in line with the Annex to the European Commission guideline on `Excipients in the labelling and package leaflet of medicinal products for human use' in section 4.4 of the	Positive Opinion adopted by consensus on 04.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.

Summary of Product Characteristics (SmPC) and section 2 of the Package Leaflet (PL) for Competact, Glubrava and Tandemact. Type IAIN, Category A.1 - Change in the address of the marketing authorisation holder (MAH) Takeda Pharma A/S, Denmark for Actos, Glustin, Competact, Glubrava and Tandemact. Updates to the Product Information in line with the latest QRD template version 10.1 for Actos, Glustin, Competact, Glubrava and Tandemact. Minor editorial/typographical updates to local Product Information (PI), including updates to comply with EN PI and local QRD, for the following languages, for each product: - Actos: BG, CS, DA, FI, DE, HU, IS, PT, RO, ES, SV. - Competact: FI, FR, DE, PT, RO, SK, SV. - Glubrava: FI, DE, PT, ES. - Glustin: DA, FI, DE, HU, PT, SL. - Tandemact: FI, DE, PT, SL, ES. Updates to local representatives contact information in section 6 of the PL for the following countries for each of the following products: - Actos: DE, FR, PL - Competact: DE, FR, PL - Glubrava: DE, ES, FR, LT, NL, PL - Glustin: DE, ES, FR, LT, NL, PL - Tandemact: DE, ES, FR, LT, NL For the Danish (DA) PI only, for all products, the letters highlighted in the street name, Vallensbaek, is spelt in Danish in which the English letters "a" and "e" are replaced with the dipthong character "æ". In addition, for the German (DE) PI, for all products, due to restricted space on the carton and in order to implement the FMD printing features, the Expiry date ' Verwendbar bis' is shortened to 'Verw. bis'. This change is in alignment with local legislation for Germany and Austria." Opinion adopted on 04.02.2021.	
WS1980 **Axura-EMEA/H/C/000378/WS1980/0082** **Memantine Merz-EMEA/H/C/002711/WS1980/0018** Merz Pharmaceuticals GmbH, Lead Rapporteur: Maria Concepcion Prieto Yerro, "To align the wording regarding sorbitol, potassium and sodium in accordance to the standard wording	Request for supplementary information adopted with a specific timetable.

listed in the Annex to the European Commission guideline on `Excipients in the labelling and package leaflet of medicinal products for human use' in the national product information."" Request for Supplementary Information adopted on 04.02.2021.	
WS1984 **HyQvia-EMEA/H/C/002491/WS1984/0066** **Kiovig-EMEA/H/C/000628/WS1984/0107** Takeda Manufacturing Austria AG, Lead Rapporteur: Jan Mueller-Berghaus Opinion adopted on 11.02.2021.	Positive Opinion adopted by consensus on 11.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1985 **Aflunov-EMEA/H/C/002094/WS1985/0067** **Foclivia-EMEA/H/C/001208/WS1985/0063** Seqirus S.r.l, Lead Rapporteur: Armando Genazzani Request for Supplementary Information adopted on 25.02.2021, 14.01.2021.	Request for supplementary information adopted with a specific timetable.
WS1987 **Cervarix-EMEA/H/C/000721/WS1987/0111** **Infanrix hexa-EMEA/H/C/000296/WS1987/0292** **Mosquirix-EMEA/H/W/002300/WS1987/0053** **Rotarix-EMEA/H/C/000639/WS1987/0119** **Shingrix-EMEA/H/C/004336/WS1987/0041** **Synflorix-EMEA/H/C/000973/WS1987/0155** GlaxoSmithkline Biologicals SA, Lead Rapporteur: Christophe Focke Opinion adopted on 11.02.2021.	Positive Opinion adopted by consensus on 11.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS1996 **Rixathon-EMEA/H/C/003903/WS1996/0046** **Riximyo-EMEA/H/C/004729/WS1996/0046** Sandoz GmbH, Lead Rapporteur: Jan Mueller-Berghaus Opinion adopted on 04.02.2021.	Positive Opinion adopted by consensus on 04.02.2021. The Icelandic and Norwegian CHMP Members were in agreement with the CHMP recommendation.
WS2003 **Silodyx-EMEA/H/C/001209/WS2003/0043** **Urorec-EMEA/H/C/001092/WS2003/0047** Recordati Ireland Ltd, Lead Rapporteur:	Request for supplementary information adopted with a specific timetable.

Armando Genazzani Request for Supplementary Information adopted on 11.02.2021.	
WS1961 **Mosquirix-EMEA/H/W/002300/WS1961/0052** **Shingrix-EMEA/H/C/004336/WS1961/0040** GlaxoSmithkline Biologicals SA, Lead Rapporteur: Jan Mueller-Berghaus Request for Supplementary Information adopted on 11.02.2021.	Request for supplementary information adopted with a specific timetable.

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

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

B.6. START OF THE PROCEDURES TIMETABLES FOR INFORMATION

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

COVID-19 vaccine (Ad26.COV2-S [recombinant]) - EMEA/H/C/005737 prevention of coronavirus disease-2019 (COVID-19)	See 15.1.5
ertapenem - EMEA/H/C/005815 treatment of bacterial infections and prophylaxis of surgical site infection following elective colorectal surgery	
gefapixant - EMEA/H/C/005476 treatment of refractory or unexplained chronic cough	
trastuzumab - EMEA/H/C/005880 treatment of metastatic and early breast cancer and metastatic gastric cancer (MGC)	
somatrogon - EMEA/H/C/005633, Orphan Pfizer Europe MA EEIG, indicated for the long-term treatment of paediatric patients with growth disturbance due to insufficient secretion of growth hormone.	
pneumococcal polysaccharide conjugate vaccine (20-valent, adsorbed) - EMEA/H/C/005451 prevention of invasive disease and pneumonia	

caused by Streptococcus pneumoniae serotypes 1, 3, 4, 5, 6A, 6B, 7F, 8, 9V, 10A, 11A, 12F, 14, 15B, 18C, 19A, 19F, 22F, 23F and 33F

eptacog beta (activated) - EMEA/H/C/005655
treatment and for the prevention of bleeding

rimegepant - EMEA/H/C/005725
management of migraine

retifanlimab - EMEA/H/C/005632, Orphan
Incyte Biosciences Distribution B.V., Treatment of locally advanced or metastatic squamous carcinoma of the anal canal (SCAC) who have progressed on or who are intolerant of platinum-based chemotherapy

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

AYVAKYT - avapritinib - EMEA/H/C/005208/X/0004/G, Orphan
Blueprint Medicines (Netherlands) B.V., Rapporteur: Blanca Garcia-Ochoa, PRAC Rapporteur: Menno van der Elst"Extension application to add 2 new strengths of film-coated tablets (25 mg and 50 mg), grouped with a type II variation (C.I.6.a) to introduce a new therapeutic indication for AYVAKYT. Extension of indication to include treatment of adult patients with advanced systemic mastocytosis (AdvSM), including aggressive systemic mastocytosis (ASM), systemic mastocytosis with an associated haematological neoplasm (SM-AHN), and mast cell leukaemia (MCL), after at least one systemic therapy for AYVAKYT based on the results of the BLU-285-2101 and BLU-285-2202 studies.As a consequence, sections 1, 2, 3, 4.1, 4.2, 4.4, 4.5, 4.6, 4.8, 5.1, 5.2, 5.3, 6.1 and 8 of the SmPC are updated. The Labelling and Package Leaflet are updated in accordance. Version 1.1 of the RMP has also been submitted."
Request for 1 year of market protection for a new indication (Article 14(11) of Regulation (EC) 726/2004)

Nuwiq - simoctocog alfa - EMEA/H/C/002813/X/0042
Octapharma AB, Rapporteur: Jan Mueller-Berghaus, "Extension application to add a new

strength of 1500 IU for simoctocog alfa powder and solvent for solution for injection, for Intravenous use.
The marketing authorisation holder took the opportunity to align the PI to the latest QRD template (version 10.2)."

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

bimekizumab - EMEA/H/C/005316
treatment of plaque psoriasis
List of Questions adopted on 10.12.2020.

lisocabtagene maraleucel / lisocabtagene maraleucel - EMEA/H/C/004731, Orphan, ATMP
Celgene Europe BV, treatment of large B-cell lymphoma, diffuse large B-cell lymphoma (DLBCL), primary mediastinal large B-cell lymphoma (PMBCL) and follicular lymphoma grade 3B (FL3B)
List of Questions adopted on 06.11.2020.

zanubrutinib - EMEA/H/C/004978, Orphan
BeiGene Ireland Ltd, treatment of Waldenström's macroglobulinaemia (WM)
List of Questions adopted on 15.10.2020.

Pheburane - sodium phenylbutyrate - EMEA/H/C/002500/X/0026
Eurocept International B.V., Rapporteur: Jayne Crowe, PRAC Rapporteur: Rhea Fitzgerald, "Extension application to introduce a new pharmaceutical form associated with new strength (350 mg/ml oral solution). The RMP (version 0.1) is updated in accordance."
List of Questions adopted on 12.11.2020.

Pheburane - sodium phenylbutyrate - EMEA/H/C/002500/X/0028
Eurocept International B.V., Rapporteur: Jayne Crowe, PRAC Rapporteur: Rhea Fitzgerald, "Extension application to introduce a new pharmaceutical form associated with new strength (500 mg film-coated tablets). The RMP (version 0.1) is updated in accordance."
List of Questions adopted on 12.11.2020.

eladocagene exuparvovec - EMEA/H/C/005352, Orphan, ATMP
PTC Therapeutics International Limited,

treatment of aromatic L-amino aciddecarboxylase (AADC) deficiency
List of Questions adopted on 20.05.2020.

vosoritide - EMEA/H/C/005475, Orphan
BioMarin International Limited, Indicated for the treatment of achondroplasia.
List of Questions adopted on 10.12.2020.

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

Ceplene - histamine dihydrochloride - EMEA/H/C/000796/S/0042
Noventia Pharma S.r.l., Rapporteur: Jayne Crowe, PRAC Rapporteur: Rhea Fitzgerald

SCENESSE - afamelanotide - EMEA/H/C/002548/S/0035, Orphan
Clinuvel Europe Limited, Rapporteur: Janet Koenig, PRAC Rapporteur: Martin Huber

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

Blenrep - belantamab mafodotin - EMEA/H/C/004935/R/0003, Orphan
GlaxoSmithKline (Ireland) Limited, Rapporteur: Johanna Lähteenvuo, PRAC Rapporteur: Annika Folin

Dovprela - pretomanid - EMEA/H/C/005167/R/0005, Orphan
Mylan IRE Healthcare Limited, Rapporteur: Filip Josephson, Co-Rapporteur: Ingrid Wang, PRAC Rapporteur: Liana Gross-Martirosyan

Emtricitabine/Tenofovir disoproxil Zentiva - emtricitabine / tenofovir disoproxil - EMEA/H/C/004137/R/0019
Zentiva k.s., Generic, Generic of Truvada, Rapporteur: Alar Irs, PRAC Rapporteur: Ana Sofia Diniz Martins

Glyxambi - empagliflozin / linagliptin - EMEA/H/C/003833/R/0039
Boehringer Ingelheim International GmbH, Rapporteur: Johann Lodewijk Hillege, Co-Rapporteur: Karin Janssen van Doorn, PRAC Rapporteur: Eva A. Segovia

Hepcludex - bulevirtide -

EMEA/H/C/004854/R/0003, Orphan
MYR GmbH, Rapporteur: Filip Josephson, PRAC Rapporteur: Adam Przybylkowski

IBRANCE - palbociclib - EMEA/H/C/003853/R/0034
Pfizer Europe MA EEIG, Rapporteur: Filip Josephson, Co-Rapporteur: Alexandre Moreau, PRAC Rapporteur: Anette Kirstine Stark

Idefirix - imlifidase - EMEA/H/C/004849/R/0003, Orphan
Hansa Biopharma AB, Rapporteur: Martina Weise, PRAC Rapporteur: Menno van der Elst

Ivabradine Zentiva - ivabradine - EMEA/H/C/004117/R/0008
Zentiva k.s., Generic, Generic of Procoralan, Rapporteur: Tomas Radimersky, PRAC Rapporteur: Menno van der Elst

Onivyde pegylated liposomal - irinotecan hydrochloride trihydrate - EMEA/H/C/004125/R/0025, Orphan
Les Laboratoires Servier, Rapporteur: Filip Josephson, Co-Rapporteur: Armando Genazzani, PRAC Rapporteur: David Olsen

Rekovelle - follitropin delta - EMEA/H/C/003994/R/0028
Ferring Pharmaceuticals A/S, Rapporteur: Jean-Michel Race, PRAC Rapporteur: Menno van der Elst

Translarna - ataluren - EMEA/H/C/002720/R/0061, Orphan
PTC Therapeutics International Limited, Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Liana Gross-Martirosyan

XALKORI - crizotinib - EMEA/H/C/002489/R/0071
Pfizer Europe MA EEIG, Rapporteur: Alexandre Moreau, PRAC Rapporteur: Tiphaine Vaillant

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

Briviact - brivaracetam - EMEA/H/C/003898/II/0032/G

UCB Pharma S.A., Rapporteur: Filip Josephson, Co-Rapporteur: Armando Genazzani, PRAC Rapporteur: Adam Przybylkowski, "- Extension of indication to include patients from 1 month to 4 years of age for the Briviact treatment. As a consequence, sections 4.1, 4.2, 4.8, 5.1 and 5.2 of the SmPC are updated. The RMP version 8.0 has also been submitted. Furthermore, the PI is brought in line with the latest QRD template version 10.2 and the MAH took the opportunity to implement minor editorial updates. - B.II.f.1.b.2 - - B.IV.1.a.1 The Package Leaflet and Labelling are updated in accordance."	
Lorviqua - lorlatinib - EMEA/H/C/004646/II/0015 Pfizer Europe MA EEIG, Rapporteur: Sinan B. Sarac, Co-Rapporteur: Armando Genazzani, PRAC Rapporteur: Nikica Mirošević Skvrce, "Extension of indication to include the treatment of adult patients with anaplastic lymphoma kinase (ALK)-positive advanced non-small cell lung cancer (NSCLC) previously not treated with an ALK inhibitor based on results from the phase III randomised CROWN (1006) study listed as a specific obligation (SOB) in the Annex II; as a consequence, sections 4.1, 4.2, and 5.1 of the SmPC are updated. The Package leaflet is updated accordingly. Version 3.0 of the RMP has also been submitted. In addition, the applicant proposes to downgrade the specific obligation to conduct a single arm study in patients who progressed after alectinib or ceritinib to a recommendation and convert the conditional MA to a full MA."	
TOOKAD - padeliporfin - EMEA/H/C/004182/II/0013 STEBA Biotech S.A, Rapporteur: Bruno Sepodes, PRAC Rapporteur: Maia Uusküla, "Modification of the wording of the existing indication. The new wording will be the treatment of adult patients with previously untreated, unilateral, low-risk, adenocarcinoma of the prostate with a life expectancy ≥ 10 years and Clinical stage T1c or T2a, ISUP Grade Group ≤ 2, based on high-resolution biopsy strategies, PSA ≤ 10 ng/mL, Low core positivity for TOOKAD; as a consequence, section 4.1 of the SmPC is updated. Version 6.0 of the RMP has also been	

submitted."

Veklury - remdesivir - EMEA/H/C/005622/II/0016

Gilead Sciences Ireland UC, Rapporteur: Janet Koenig, PRAC Rapporteur: Eva Jirsová, "Extension of indication to include treatment of adults with pneumonia not requiring supplemental oxygen (moderate COVID-19), based on Part A of Study GS-US-540-5774, a Phase 3, randomized, open-label, multicenter study comparing 2 RDV regimens (5 days and 10 days) versus standard of care in 584 participants with moderate COVID 19, and Study CO US 540 5776 [Adaptive COVID-19 Treatment Trial (ACTT) 1, a National Institute of Allergy and Infectious Diseases (NIAID)-sponsored Phase 3, randomized, double blind, placebo controlled, multicenter study]. As a consequence, sections 4.1 and 5.1 of the SmPC are being updated, and the Package Leaflet is updated in accordance. A revised version 1.2 of the RMP has also been submitted."

Vyxeos liposomal - daunorubicin / cytarabine - EMEA/H/C/004282/II/0018/G, Orphan

Jazz Pharmaceuticals Ireland Limited, Rapporteur: Johanna Lähteenvuo, Co-Rapporteur: Janet Koenig, PRAC Rapporteur: Marcia Sofia Sanches de Castro Lopes Silva, "Extension of indication to add treatment of relapsed/refractory AML in paediatric patients with subsequent updates to sections 4.1, 4.2, 4.8, 5.1, 5.2 of the SmPC based on the new safety and efficacy data from the paediatric clinical study AAML1421. The package leaflet is updated accordingly. The RMP version 1.1 has also been submitted. In addition, the PI is updated in line with the latest QRD template 10.2.
Submission of the final data from paediatric clinical study CPX-MA-1201 in support of the extension of indication."
Request for 1 year of market protection for a new indication (Article 14(11) of Regulation (EC) 726/2004)

Xeljanz - tofacitinib - EMEA/H/C/004214/II/0035

Pfizer Europe MA EEIG, Rapporteur: Armando Genazzani, Co-Rapporteur: Johann Lodewijk

Hillege, PRAC Rapporteur: Liana Gross-Martirosyan, "Extension of indication to include treatment of adult patients with active ankylosing spondylitis (AS) who have responded inadequately to conventional therapy for XELJANZ film-coated tablets; 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 17.1 of the RMP has also been submitted."	

B.6.8. CHMP assessed procedures scope: Pharmaceutical aspects

ADYNOVI - rurioctocog alfa pegol - EMEA/H/C/004195/II/0020 Baxalta Innovations GmbH, Rapporteur: Andrea Laslop	
COMIRNATY - covid-19 mrna vaccine (nucleoside-modified) - EMEA/H/C/005735/II/0009 BioNTech Manufacturing GmbH, Rapporteur: Filip Josephson Opinion adopted on 19.02.2021. Request for Supplementary Information adopted on 09.02.2021.	See also B.5.1
COMIRNATY - covid-19 mrna vaccine (nucleoside-modified) - EMEA/H/C/005735/II/0010/G BioNTech Manufacturing GmbH, Rapporteur: Filip Josephson Opinion adopted on 22.02.2021.	See also B.5.1
COMIRNATY - covid-19 mrna vaccine (nucleoside-modified) - EMEA/H/C/005735/II/0011/G BioNTech Manufacturing GmbH, Rapporteur: Filip Josephson	
Cufence - trientine dihydrochloride - EMEA/H/C/004111/II/0007/G Univar Solutions BV, Rapporteur: Daniela Philadelphy	
Elonva - corifollitropin alfa - EMEA/H/C/001106/II/0058/G Merck Sharp & Dohme B.V., Rapporteur: Paula Boudewina van Hennik	
Empliciti - elotuzumab - EMEA/H/C/003967/II/0026	

Bristol-Myers Squibb Pharma EEIG, Rapporteur: Paula Boudewina van Hennik

GIVLAARI - givosiran - EMEA/H/C/004775/II/0004/G, Orphan
Alnylam Netherlands B.V., Rapporteur: Paula Boudewina van Hennik

MenQuadfi - Meningococcal group A, C, W135 and Y conjugate vaccine - EMEA/H/C/005084/II/0001/G
Sanofi Pasteur, Rapporteur: Andrea Laslop

Omnitrope - somatropin - EMEA/H/C/000607/II/0070
Sandoz GmbH, Rapporteur: Johann Lodewijk Hillege

Palynziq - pegvaliase - EMEA/H/C/004744/II/0017, Orphan
BioMarin International Limited, Rapporteur: Johann Lodewijk Hillege

Repatha - evolocumab - EMEA/H/C/003766/II/0051
Amgen Europe B.V., Rapporteur: Johann Lodewijk Hillege

Revestive - teduglutide - EMEA/H/C/002345/II/0052/G, Orphan
Shire Pharmaceuticals Ireland Limited, Rapporteur: Kirstine Moll Harboe

Rybelsus - semaglutide - EMEA/H/C/004953/II/0012
Novo Nordisk A/S, Rapporteur: Johann Lodewijk Hillege

Tecentriq - atezolizumab - EMEA/H/C/004143/II/0057/G
Roche Registration GmbH, Rapporteur: Sinan B. Sarac

Vaxelis - diphtheria, tetanus, pertussis (acellular, component), hepatitis B (rDNA), poliomyelitis (inact.) and Haemophilus type b conjugate vaccine (adsorbed) - EMEA/H/C/003982/II/0075
MCM Vaccine B.V., Rapporteur: Christophe Focke

Voncento - human coagulation factor viii / human von willebrand factor - EMEA/H/C/002493/II/0047/G
CSL Behring GmbH, Rapporteur: Paula

Boudewina van Hennik

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

Bridion - sugammadex - EMEA/H/C/000885/II/0039

Merck Sharp & Dohme B.V., Rapporteur: Outi Mäki-Ikola, "C.I.4 type II Update of sections 4.8 and 5.1 of the SmPC in order to update information on safety profile in American Society of Anesthesiologists (ASA) Class 3 or 4 patients (patients with severe systemic disease or patients with severe systemic disease that is a constant threat to life) based on final results from study 8616-P145, an interventional safety study of sugammadex for the reversal of neuromuscular blockage induced by rocuronium or vecuronium in adult ASA 3-4 participants."

Cholib - fenofibrate / simvastatin - EMEA/H/C/002559/II/0029/G

Mylan IRE Healthcare Limited, Rapporteur: Alar Irs, "Update of section 4.4 of the SmPC in order to amend the existing warning on immune-mediated necrotizing myopathy (IMNM) and section 4.5 of the SmPC to add drug-drug interaction information with cobicistat, following the update of the company's core data sheet (CCDS) due to new data; the Package Leaflet is updated accordingly. In addition, the MAH took the opportunity to update the list the UK (Northern Ireland) local representative in the Package Leaflet."

Darzalex - daratumumab - EMEA/H/C/004077/II/0047, Orphan

Janssen-Cilag International NV, Rapporteur: Sinan B. Sarac, "C.I.4
Update of section 4.4 of the SmPC in order to include a fatal outcome for IRRs following a systematic cross-programmatic review of fatal cases of Infusion Related Reaction (IRR) with use of daratumumab. In addition, the MAH has taken the opportunity to correct in section 4.8 the reported incidence rate of Grade 3 or 4 treatment-emergent infections from study MMY3003 for DRd from 27% to 28%."

Dovprela - pretomanid - EMEA/H/C/005167/II/0004/G, Orphan

Mylan IRE Healthcare Limited, Rapporteur: Filip Josephson, "Grouped application including three type II variations under category C.I.4.
Update of sections 4.5 and 5.3 of the SmPC based on data from three non-clinical studies:
- Assessment of pretomanid as an inhibitor of human OCT2-, OATP1B3-, BCRP-, and P-gp mediated transport;
- In vitro evaluation of induction/inhibition of CYP 2C8, 2C9, and 2C19 in human hepatocytes;
- A 24-Month Oral (Gavage) Carcinogenicity Study in Rats."

Eliquis - apixaban - EMEA/H/C/002148/II/0080

Bristol-Myers Squibb / Pfizer EEIG, Rapporteur: Johann Lodewijk Hillege, "C.I.4
Update of section 4.4 of the SmPC in order to update the existing warning regarding patients with active cancer in line with the final results of the study CARAVAGGIO (NCT03045406), which is a randomized open-label non-inferiority clinical trial assessing apixaban for the treatment of acute proximal DVT and/or PE in ambulatory patients with active cancer or history of cancer. In addition, the MAH took the opportunity to correct a typo in section 5.1 of the SmPC."

Esbriet - pirfenidone - EMEA/H/C/002154/II/0070, Orphan

Roche Registration GmbH, Rapporteur: Peter Kiely, "Update of section 4.8 of the SmPC to revise the MeDRA frequency categories for some adverse drug reactions (ADR) and to combine the ADR 'anorexia' with the preferred term 'decreased appetite' based on a safety update report previously submitted in variation EMEA/H/C/2154/II/0021. The package leaflet is updated accordingly."

Keytruda - pembrolizumab - EMEA/H/C/003820/II/0102

Merck Sharp & Dohme B.V., Rapporteur: Armando Genazzani, "Update of sections 4.2 and 5.1 of the SmPC in order to introduce an alternative dosing regimen of 400 mg every 6 weeks (Q6W) for all approved indications based on interim results from study KEYNOTE-555; this is an interventional, PK study in patients with advanced melanoma. Additional data/analysis from studies KEYNOTE-021, -048,

-189, -407 and -426 were provided."

Maviret - glecaprevir / pibrentasvir - EMEA/H/C/004430/II/0040

AbbVie Deutschland GmbH & Co. KG, Rapporteur: Jean-Michel Race, "Update of sections 4.8 and 5.1 of the SmPC to include information related to the safety and efficacy of Maviret for people who inject drugs (PWID) and those who are on medication-assisted treatment (MAT) for opioid use disorder based on data from Phase 2 and 3 clinical trials.
In addition, the MAH took the opportunity to include an editorial change and corrected the number of subjects stated in Footnote B, Table 8 of the SmPC section 5.1."

Talzenna - talazoparib - EMEA/H/C/004674/II/0009

Pfizer Europe MA EEIG, Rapporteur: Filip Josephson, "Update of sections 4.2 and 5.2 based on the results from PK study MDV3800-02 (C3441002), a phase 1 open-label pharmacokinetics and safety study of talazoparib (MDV3800) in patients with advanced solid tumors and normal or varying degrees of hepatic impairment."

Zostavax - varicella vaccine (live) - EMEA/H/C/000674/II/0132

MSD Vaccins, Rapporteur: Jan Mueller-Berghaus, "C.I.13: Submission of the final study report from the post-licensure observational study of the long-term effectiveness of Zostavax (Protocol 024) listed as category 3 study in the RMP. With this application, the post-authorisation measure REC 23 is fulfilled."

WS2027
OFEV-EMEA/H/C/003821/WS2027/0042
Vargatef-EMEA/H/C/002569/WS2027/0039

Boehringer Ingelheim International GmbH, Lead Rapporteur: Peter Kiely, "Update of sections 4.4 and 6.6. of the SmPC in order to include improved label instructions for the handling of the capsules: Inclusion of a general recommendation not to open the capsules and addition of a statement to wash hands in the hypothetical event of getting in contact with the content of the capsule, respectively. This update is based on post-marketing experience and in line with regulatory guidance. The Package

Leaflet is updated accordingly. In addition, the MAH took the opportunity to correct the name of the local representative in Portugal.”	
WS2035 **Prezista-EMEA/H/C/000707/WS2035/0110** **Rezolsta-EMEA/H/C/002819/WS2035/0041** **Symtuza-EMEA/H/C/004391/WS2035/0032** Janssen-Cilag International NV, Lead Rapporteur: Johann Lodewijk Hillege, “To update section 4.5 of the SmPC to provide guidance on drug-drug interaction between cutaneously-administered corticosteroids and boosted darunavir, darunavir/cobicistat and darunavir/cobicistat/emtricitabine/tenofovir alafenamide, based on recent scientific literature publication. In addition, the MAH took the opportunity to include minor editorial updates in the Product Information consisting of formatting, spelling and typo corrections.”	

B.6.10. CHMP-PRAC assessed procedures

Accofil - filgrastim - EMEA/H/C/003956/II/0046/G Accord Healthcare S.L.U., Rapporteur: Outi Mäki-Ikola, PRAC Rapporteur: Kirsti Villikka, “- Type II B.II.e.5.c- To introduce a new presentation, to cater to low-weight patients as the clinical administration of Filgrastim is based on body weight. RMP and PI are updated to include this new strength. - Type II B.II.e.5.c- To introduce a new presentation, based on the dosing regimen to avoid multiple administrations. RMP and PI are updated to include this new strength.”	
Jyseleca - filgotinib - EMEA/H/C/005113/II/0003 Gilead Sciences Ireland UC, Rapporteur: Kristina Dunder, PRAC Rapporteur: Nikica Mirošević Skvrce, “Update to sections 4.5 and 5.2 of the SmPC to update the wording on the inhibition of P-gp and BCRP by the primary metabolite of filgotinib (GS-829845) based upon results from an in vitro study (AD-417-2028) which assessed	

in vitro inhibition of human P-gp and BCRP by GS-829845. The Package Leaflet has been updated accordingly. A consequential update of the RMP has been submitted (version 1.2)."

Lojuxta - lomitapide - EMEA/H/C/002578/II/0046

Amryt Pharmaceuticals DAC, Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Menno van der Elst, "To propose an alternative study (LILITH) to the currently agreed protocol for the CAPTURE study. The new study proposes an evaluation of the effect of lomitapide treatment on major adverse cardiovascular events (MACE) in patients with homozygous familial hypercholesterolemia. Consequential submission of an updated RMP (version 6.4) and Annex IID of the Product Information. In addition, the MAH took the opportunity to bring the PI in line with the latest QRD template version 10.2"

Piqray - alpelisib - EMEA/H/C/004804/II/0005/G

Novartis Europharm Limited, Rapporteur: Blanca Garcia-Ochoa, PRAC Rapporteur: Menno van der Elst, "C.1.4 Update of sections 4.4 and 4.8 of the SmPC in order to add hyperglycaemic hyperosmolar non-ketotic syndrome to the list of adverse drug reactions (ADRs) with frequency "unknown" and to update the warning on hyperglycaemia and ketoacidosis based on a review of the safety database. The package leaflet and Annex II are updated accordingly. The RMP version 3.0 has also been submitted. C.1.4 Update of sections 4.2 and 4.8 of the SmPC to modify the management of hyperglycaemia, rash and diarrhoea and add information about osteonecrosis of the jaw based on the pivotal trial SOLAR-1. The MAH also took the opportunity to make minor editorial changes to the SmPC."

Vemlidy - tenofovir alafenamide - EMEA/H/C/004169/II/0030

Gilead Sciences Ireland UC, Rapporteur: Janet Koenig, PRAC Rapporteur: Ilaria Baldelli, "Submission of the final report from study GS-US-320-4018, listed as a category 3 study in the RMP. This is a phase 3, randomized, double blind study to evaluate the efficacy and safety of switching from tenofovir disoproxil fumarate

300 mg once daily to tenofovir alafenamide 25 mg once daily in subjects with chronic hepatitis B who are virologically suppressed. The RMP (v 6.1) has also been submitted."

B.6.11. PRAC assessed procedures

PRAC Led

Constella - linaclotide - EMEA/H/C/002490/II/0053

Allergan Pharmaceuticals International Limited, Rapporteur: Martina Weise, PRAC Rapporteur: Martin Huber, PRAC-CHMP liaison: Martina Weise, "C.I.3b: Update of sections 4.4 and 4.8 of the SmPC in order to add a new warning on intestinal perforation and to add gastrointestinal perforation to the list of adverse drug reactions (ADRs) with frequency rare based on the Truven Market Scan study and as requested by the PRAC in procedure EMEA/H/C/002490/LEG/015; the Package Leaflet is updated accordingly. In addition, the MAH took the opportunity to update the list of local representatives in the Package Leaflet"

PRAC Led

Gardasil - human papillomavirus vaccine [types 6, 11, 16, 18] (recombinant, adsorbed) - EMEA/H/C/000703/II/0091

MSD Vaccins, Rapporteur: Kristina Dunder, PRAC Rapporteur: Ulla Wändel Liminga, PRAC-CHMP liaison: Kristina Dunder, "Submission of the final report from study 070 listed as a category 3 study in the RMP in order to address MEA 86.2. This is a post-licensure observational study of the safety of Gardasil in males. The RMP version 14.1 has been updated. The MAH took the opportunity to update the RMP with the protocol synopsis of the 2-dose effectiveness in Sweden (MEA 82.6 assessed by CHMP)."

PRAC Led

Ocrevus - ocrelizumab - EMEA/H/C/004043/II/0024

Roche Registration GmbH, Rapporteur: Kirstine Moll Harboe, PRAC Rapporteur: Brigitte Keller-Stanislawski, PRAC-CHMP liaison: Jan Mueller-Berghaus, "Type II variation application, category C.I.4, to amend the wording on Progressive Multifocal Leukoencephalopathy

(PML) in the SmPC, section 4.4 Special warnings and precautions, for compliance with PRAC Recommendations.”
PRAC Led **WS2009/G Edistride-EMEA/H/C/004161/WS2009/0045/G Forxiga-EMEA/H/C/002322/WS2009/0064/G** AstraZeneca AB, Lead Rapporteur: Kristina Dunder, Lead PRAC Rapporteur: Annika Folin, PRAC-CHMP liaison: Kristina Dunder, “Submission of the final report from studies MB102103, MB102104 and MB102110 listed as a category 3 study in the RMP. These are observational studies comparing the risk of severe complications of UTI, acute liver injury and acute kidney injury respectively, between patients with Type 2 Diabetes exposed to dapagliflozin and those exposed to other antidiabetic treatments. The RMP version 23.1 for Forxiga/Edistride has also been submitted.”

B.6.12. CHMP-CAT assessed procedures

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 - EMEA/H/C/005102/II/0001, Orphan, ATMP Kite Pharma EU B.V., Rapporteur: Jan Mueller-Berghaus, CHMP Coordinator: Jan Mueller-Berghaus
Yescarta - axicabtagene ciloleucel - EMEA/H/C/004480/II/0035, Orphan, ATMP Kite Pharma EU B.V., Rapporteur: Jan Mueller-Berghaus, CHMP Coordinator: Jan Mueller-Berghaus
Zynteglo - betibeglogene autotemcel - EMEA/H/C/003691/II/0022, 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

WS2025
Hefiya-EMEA/H/C/004865/WS2025/0028
Hyrimoz-EMEA/H/C/004320/WS2025/0028
Sandoz GmbH, Lead Rapporteur: Daniela Philadelphy

WS2034
Hexacima-EMEA/H/C/002702/WS2034/0115
Hexyon-EMEA/H/C/002796/WS2034/0119
MenQuadfi-EMEA/H/C/005084/WS2034/0002
Sanofi Pasteur, Lead Rapporteur: Jan Mueller-Berghaus

WS2041/G
Copalia-EMEA/H/C/000774/WS2041/0118/G
Dafiro-EMEA/H/C/000776/WS2041/0122/G
Exforge-EMEA/H/C/000716/WS2041/0117/G
Novartis Europharm Limited, Lead Rapporteur: Kirstine Moll Harboe

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

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

B.7.2. Monthly Line listing for Type I variations

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

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

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

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

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

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

E. Annex E - EMEA CERTIFICATION OF PLASMA MASTER FILES

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

E.1. PMF Certification Dossiers:

E.1.1. Annual Update

E.1.2. Variations:

E.1.3. Initial PMF Certification:

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

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

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

G. ANNEX G

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

G.2. PRIME

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

G.2.1. List of procedures concluding at 22-25 February 2021 CHMP plenary:

Oncology	
Treatment of adult patients with ROS1-positive, advanced non-small cell lung cancer (NSCLC) not previously treated with ROS1 inhibitors	The CHMP denied eligibility to PRIME and adopted the critical summary report
Adjuvant treatment of stage IIB/IIC melanoma after surgical resection	The CHMP denied eligibility to PRIME and adopted the critical summary report
Musculoskeletal disorders	
Treatment of Fibrodysplasia Ossificans Progressiva	The CHMP denied eligibility to PRIME and adopted the critical summary report
Endocrinology-Gynaecology-Fertility-Metabolism	
Stem cell therapy for ovarian insufficiency includes Diminished Ovarian Reserve (DOR), Premature Ovarian Failure (POF), Primary Ovarian Insufficiency (POI) and Poor Ovarian Response (POR). (SME)	The CHMP denied eligibility to PRIME and adopted the critical summary report

G.2.2. List of procedures starting in February 2021 for March 2021 CHMP adoption of outcomes

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