



EUROPEAN MEDICINES AGENCY  
SCIENCE MEDICINES HEALTH

18 April 2024  
EMA/CHMP/106558/2024  
Human Medicines Division

## Committee for medicinal products for human use (CHMP)

Minutes for the meeting on 19-22 February 2024

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, the 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. 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 meeting, the Committee Secretariat announced the restricted involvement of some Committee members, alternates and experts for concerned agenda topics.

Participants 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. Restrictions applicable to this meeting are captured in the list of participants included in the minutes.

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. All decisions, recommendations and advice were agreed by consensus, unless otherwise specified.

### 1.2. Adoption of agenda

CHMP agenda for 19-22 February 2024.

The CHMP adopted the agenda.

### 1.3. Adoption of the minutes

CHMP minutes for 22-25 January 2024.

The CHMP adopted the minutes for the 22-25 January 2024 plenary.

Minutes from PReparatory and Organisational Matters (PROM) meeting held on 12 February 2024.

The CHMP adopted the minutes from the PROM meeting held on 12 February 2024.

## 2. Oral Explanations

### 2.1. Pre-authorisation procedure oral explanations

#### 2.1.1. Bevacizumab - EMEA/H/C/005723

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Treatment of neovascular (wet) age-related macular degeneration (nAMD).

Scope: Oral explanation

**Action:** Possible Oral explanation to be held on 20 February 2024 at 14:00

List of Outstanding Issues adopted on 09.11.2023. List of Questions adopted on 26.04.2023.

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

See 3.2

## 2.2. Re-examination procedure oral explanations

No items

## 2.3. Post-authorisation procedure oral explanations

### 2.3.1. Keytruda - Pembrolizumab - EMEA/H/C/003820/II/0134

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Merck Sharp & Dohme B.V.

Rapporteur: Paolo Gasparini, PRAC Rapporteur: Bianca Mulder

Scope: "Extension of indication to include in combination with platinum-containing chemotherapy as neoadjuvant treatment, and then continued as monotherapy as adjuvant, treatment of resectable Stage II, IIIA, or IIIB (T3 4N2) non-small cell lung carcinoma in adults for Keytruda based on study KEYNOTE-671, a phase III, randomized, double-blind trial of platinum doublet chemotherapy +/- pembrolizumab as neoadjuvant/adjuvant therapy for participants with resectable stage II, IIIA, and resectable IIIB (T3-4N2) non-small cell lung cancer. As a consequence, sections 4.1, 4.2 and 5.1 of the SmPC are updated. The Package Leaflet is updated in accordance. Version 41.1 of the RMP has also been submitted."

Scope: Oral explanation

**Action:** Oral explanation to be held on 21 February 2024 at 14:00

Request for Supplementary Information adopted on 14.12.2023, 22.06.2023.

An oral explanation was held on 21 February 2024. The presentation by the MAH focused on clinical aspects.

See 5.1

### 2.3.2. Orencia - Abatacept - EMEA/H/C/000701/II/0152

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Bristol-Myers Squibb Pharma EEIG

Rapporteur: Outi Mäki-Ikola, Co-Rapporteur: Beata Maria Jakline Ullrich, PRAC Rapporteur: Kimmo Jaakkola

Scope: "Extension of indication to include the prophylaxis of acute Graft versus Host Disease (aGvHD) in the adult and paediatric population for Orencia, based on final results from studies IM101311 - Abatacept Combined With a Calcineurin Inhibitor and Methotrexate for Graft Versus Host Disease Prophylaxis and IM101841 - Overall Survival In 7/8 HLA-

Matched Hematopoietic Stem Cell Transplantation Patients Treated With Abatacept Combined With A Calcineurin Inhibitor And Methotrexate - An Analysis Of The Center For International Blood And Marrow Transplant Research (Cibmtr) Database. As a consequence, sections 4.1, 4.2, 4.4, 4.8, 5.1 and 5.2 of the SmPC are updated. The Package Leaflet and Labelling are updated in accordance. Version 28.0 of the RMP has also been submitted. In addition, the MAH took the opportunity to introduce minor editorial changes to the PI.”

Scope: Oral explanation

**Action:** Oral explanation to be held on 20 February 2024 at 16:00

Request for Supplementary Information adopted on 12.10.2023, 30.03.2023.

The CHMP noted the withdrawal of the extension on indication application.

See 9.1

## 2.4. Referral procedure oral explanations

No items

## 3. Initial applications

### 3.1. Initial applications; Opinions

#### 3.1.1. Apremilast Accord - Apremilast - EMEA/H/C/006208

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Accord Healthcare S.L.U.; treatment of psoriatic arthritis, psoriasis, Behçet’s disease

Scope: Opinion

**Action:** For adoption

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

List of Outstanding Issues adopted on 14.12.2023. List of Questions adopted on 20.07.2023.

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

The summary of opinion was circulated for information.

#### 3.1.2. Celldemic - Zoonotic influenza vaccine (H5N1) (surface antigen, inactivated, adjuvanted, prepared in cell cultures) - EMEA/H/C/006052

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Seqirus Netherlands B.V.; active immunisation for the prevention of disease caused by the influenza A virus H5N1 subtype contained in the vaccine

Scope: Opinion

**Action:** For adoption

List of Outstanding Issues adopted on 09.11.2023. List of Questions adopted on 26.04.2023.

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

The CHMP noted the letter of recommendations dated 21 February 2024.

The summary of opinion was circulated for information.

### 3.1.3. [FILSPARI - Sparsentan - Orphan - EMEA/H/C/005783](#)

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Vifor France; for the treatment of primary immunoglobulin A nephropathy (IgAN).

Scope: Opinion

**Action:** For adoption

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

List of Outstanding Issues adopted on 14.12.2023, 12.10.2023, 25.05.2023. List of Questions adopted on 15.12.2022.

The Committee confirmed that all issues previously identified in this application have 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 sparsentan is a new active substance, as claimed by the applicant.

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.

### 3.1.4. [Incellipan - Pandemic influenza vaccine \(H5N1\) \(surface antigen, inactivated, adjuvanted, prepared in cell cultures\) - EMEA/H/C/006051](#)

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Seqirus Netherlands B.V.; prophylaxis of influenza

Scope: Opinion

**Action:** For adoption

List of Outstanding Issues adopted on 09.11.2023. List of Questions adopted on 26.04.2023.

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

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

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

The CHMP noted the letter of recommendations dated 21 February 2024.

The summary of opinion was circulated for information.

### 3.1.5. Nintedanib Accord - Nintedanib - EMEA/H/C/006179

Accord Healthcare S.L.U.; treatment of idiopathic pulmonary fibrosis (IPF), chronic fibrosing interstitial lung diseases (ILDs) and lung diseases (ILDs) systemic sclerosis associated interstitial lung disease (SSc-ILD)

Scope: Opinion

**Action:** For adoption

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

List of Outstanding Issues adopted on 14.12.2023. List of Questions adopted on 26.04.2023.

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

The summary of opinion was circulated for information.

### 3.1.6. Pyzchiva - Ustekinumab - EMEA/H/C/006183

Samsung Bioepis NL B.V.; treatment of Crohn's disease, Ulcerative colitis, Plaque psoriasis, Paediatric plaque psoriasis and Psoriatic arthritis (PsA)

Scope: Opinion

**Action:** For adoption

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

List of Outstanding Issues adopted on 14.12.2023. List of Questions adopted on 14.09.2023.

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 legal status was agreed as medicinal product subject to restricted medical prescription.  
The summary of opinion was circulated for information.

### 3.1.7. QALSODY - Tofersen - Orphan - EMEA/H/C/005493

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Biogen Netherlands B.V.; treatment of adults with amyotrophic lateral sclerosis (ALS), associated with a mutation in the superoxide dismutase 1 (SOD1) gene.

Scope: Opinion

**Action:** For adoption

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

List of Outstanding Issues adopted on 09.11.2023, 14.09.2023. List of Questions adopted on 30.03.2023.

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 under exceptional circumstances by majority (25 positive out of 31 votes) together with the CHMP assessment report and translation timetable.

The divergent positions (Peter Mol, Kristina Dunder, Sol Ruiz, Outi Maki-Ikola, Maria Concepcion Prieto Yerro, Maria Grazia Evandri) were appended to the opinion.

Furthermore, the CHMP considered that tofersen is a new active substance, as claimed by the Applicant.

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

The summary of opinion was circulated for information.

The CHMP noted the EMA news announcement.

### 3.1.8. Tizveni - Tislelizumab - EMEA/H/C/005542

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Beigene Ireland Limited; treatment of locally advanced or metastatic non-squamous non-small cell lung cancer in adults, treatment of locally advanced or metastatic squamous non-small cell lung cancer in adults, locally advanced or metastatic non-small cell lung cancer after prior chemotherapy in adults

Scope: Opinion

**Action:** For adoption

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

List of Outstanding Issues adopted on 30.03.2023. List of Questions adopted on 21.07.2022.

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 tislelizumab is not a new active substance, as claimed by the applicant.

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

The summary of opinion was circulated for information.

The CHMP adopted the new active substance assessment report for Tizveni.

### 3.1.9. [Voydeya - Danicopan - PRIME - Orphan - EMEA/H/C/005517](#)

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Alexion Europe; Treatment of extravascular haemolysis (EVH) in patients with paroxysmal nocturnal haemoglobinuria

Scope: Opinion

**Action:** For adoption

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

List of Outstanding Issues adopted on 25.01.2024, 09.11.2023. List of Questions adopted on 20.07.2023.

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

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

The summary of opinion was circulated for information.

The CHMP adopted the similarity assessment report.

### 3.1.10. [ZYNYZ - Retifanlimab - Orphan - EMEA/H/C/006194](#)

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Incyte Biosciences Distribution B.V.; Treatment of Merkel cell carcinoma (MCC).

Scope: Opinion

**Action:** For adoption

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

List of Outstanding Issues adopted on 14.12.2023. List of Questions adopted on 20.07.2023.

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 retifanlimab is a new active substance, as claimed

by the applicant.

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. Dantrolene sodium, hemiheptahydrate - Orphan - EMEA/H/C/006009

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Norgine B.V.; treatment of malignant hyperthermia (including suspected cases)

Scope: List of outstanding issues

**Action:** For adoption

List of Outstanding Issues adopted on 22.06.2023. List of Questions adopted on 10.11.2022.

The Committee was reminded of the status of this application.

The Committee adopted a 2<sup>nd</sup> list of outstanding issues with a specific timetable.

### 3.2.2. Concizumab - EMEA/H/C/005938

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routine prophylaxis to prevent or reduce the frequency of bleeding in patients with: haemophilia A (congenital factor VIII deficiency) with FVIII inhibitors  $\geq$  12 years of age; haemophilia B (congenital factor IX deficiency) with FIX inhibitors of any age

Scope: List of outstanding issues; the applicant requested an extension to the clock stop to respond to the list of outstanding issues

**Action:** For adoption

List of Outstanding Issues adopted on 14.12.2023, 12.10.2023. List of Questions adopted on 25.05.2023.

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

The Committee adopted a 3<sup>rd</sup> list of outstanding issues.

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

### 3.2.3. Insulin icodex - EMEA/H/C/005978

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treatment of diabetes mellitus in adults

Scope: List of outstanding issues

**Action:** For adoption

List of Questions adopted on 14.09.2023.

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. Capiwasertib - EMEA/H/C/006017

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is indicated in combination with fulvestrant for the treatment of adult patients with hormone receptor (HR) positive, human epidermal growth factor receptor 2 (HER2) negative (defined as IHC 0 or 1+, or IHC 2+/ISH-) locally advanced or metastatic breast cancer following recurrence or progression on or after an endocrine based regimen

Scope: List of outstanding issues

**Action:** For adoption

List of Questions adopted on 14.09.2023.

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. Aztreonam / Avibactam - EMEA/H/C/006113

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##### **Accelerated assessment**

treatment of complicated Intra-Abdominal Infection (cIAI), complicated Urinary Tract Infection (cUTI), including pyelonephritis, Hospital-acquired pneumonia (HAP), including ventilator associated pneumonia (VAP), and aerobic Gram-negative infections with limited treatment options

Scope: List of outstanding issues

**Action:** For adoption

List of Questions adopted on 12.12.2023.

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

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

#### 3.2.6. Eribulin - EMEA/H/C/006191

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treatment of breast cancer and liposarcoma

Scope: List of outstanding issues

**Action:** For adoption

List of Questions adopted on 14.09.2023.

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

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

### 3.2.7. Iptacopan - PRIME - Orphan - EMEA/H/C/005764

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Novartis Europharm Limited; treatment of paroxysmal nocturnal haemoglobinuria

Scope: List of outstanding issues

**Action:** For adoption

List of Questions adopted on 14.09.2023.

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

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

### 3.2.8. Fruquintinib - EMEA/H/C/005979

---

treatment of metastatic colorectal cancer

Scope: List of outstanding issues

**Action:** For adoption

List of Questions adopted on 12.10.2023.

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

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

### 3.2.9. Bevacizumab - EMEA/H/C/005723

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Treatment of neovascular (wet) age-related macular degeneration (nAMD).

Scope: List of outstanding issues

**Action:** For adoption

List of Outstanding Issues adopted on 09.11.2023. List of Questions adopted on 26.04.2023.

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

See 2.1

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

The Committee adopted a 2<sup>nd</sup> list of outstanding issues with a specific timetable.

### 3.2.10. Omalizumab - EMEA/H/C/005958

---

treatment of asthma

Scope: List of outstanding issues

**Action:** For adoption

List of Questions adopted on 14.09.2023.

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. Flortaucipir (18F) - EMEA/H/C/006064

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indicated for Positron Emission Tomography (PET) imaging of the brain

Scope: List of outstanding issues

**Action:** For adoption

List of Outstanding Issues adopted on 14.12.2023. List of Questions adopted on 20.07.2023.

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

The Committee adopted a 2<sup>nd</sup> list of outstanding issues.

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

### 3.2.12. Ustekinumab - EMEA/H/C/006415

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treatment of moderate to severe plaque psoriasis in adults, children and adolescents, active psoriatic arthritis in adults and Crohn's Disease, treatment of Crohn's Disease

Scope: List of outstanding issues

**Action:** For adoption

List of Questions adopted on 14.09.2023.

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. Vibegron - EMEA/H/C/005957

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treatment of micturition frequency and/or urgency incontinence as may occur in adult patients with Over Active Bladder (OAB) syndrome.

Scope: List of outstanding issues

**Action:** For adoption

List of Questions adopted on 14.09.2023.

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

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

### 3.2.14. Ustekinumab - EMEA/H/C/006132

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treatment of moderate to severe plaque psoriasis in adults, children and adolescents, active psoriatic arthritis in adults, Crohn's Disease and ulcerative colitis

Scope: List of outstanding issues

**Action:** For adoption

List of Questions adopted on 14.09.2023.

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. Guanfacine - EMEA/H/C/006312

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treatment of ADHD

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. Apremilast - EMEA/H/C/006193

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treatment of psoriatic arthritis, psoriasis, Behçet'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. Troriluzole - Orphan - EMEA/H/C/006068

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Biohaven Bioscience Ireland Limited; is indicated for the treatment of adult patients with spinocerebellar ataxia genotype 3 (SCA3)

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. Mirvetuximab soravtansine - Orphan - EMEA/H/C/005036

Immunogen Biopharma (Ireland) Limited; treatment of ovarian, fallopian tube, or primary peritoneal cancer

Scope: List of questions

**Action:** For adoption

The Committee discussed the issues identified in this application.

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

#### 3.3.5. Tiratricol - Orphan - EMEA/H/C/005220

Rare Thyroid Therapeutics International AB; treatment of monocarboxylate transporter 8 (MCT8) deficiency

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. Eplontersen - Orphan - EMEA/H/C/006295

AstraZeneca AB; indicated for the treatment of adult patients with polyneuropathy associated with hereditary transthyretin-mediated amyloidosis (ATTRv).

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. Marstacimab - Orphan - EMEA/H/C/006240

Pfizer Europe Ma EEIG; is indicated for routine prophylaxis of bleeding episodes in patients with haemophilia A or haemophilia B

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.

### 3.3.8. Elafibranor - Orphan - EMEA/H/C/006231

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Ipsen Pharma; treatment of primary biliary cholangitis (PBC)

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.

### 3.3.9. Chikungunya virus, strain CHIKV LR2006-OPY1, live attenuated - PRIME – OPEN - EMEA/H/C/005797

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#### **Accelerated assessment**

prevention of disease caused by chikungunya (CHIKV) virus

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.

### 3.3.10. Clascoterone - EMEA/H/C/006138

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indicated for the topical treatment of acne vulgaris in adults and adolescents

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. givinostat - Orphan - EMEA/H/C/006079

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Italfarmaco S.p.A.; treatment of Duchenne muscular dystrophy (DMD)

Scope: Letter by the applicant dated 16.02.2024 requesting an extension to the clock stop to respond to the list of questions adopted in December 2023.

**Action:** For adoption

List of Questions adopted on 14.12.2023.

The CHMP agreed to the request by the applicant for an extension to the clock stop to respond to the list of questions adopted in December 2023.

### 3.4.2. vilobelimab - EMEA/H/C/006123

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treatment of adult patients with SARS-CoV-2 induced septic acute respiratory distress syndrome (ARDS) receiving invasive mechanical ventilation (IMV) or extracorporeal membrane oxygenation (ECMO).

Scope: Letter by the applicant dated 13.02.2024 requesting an extension to the clock stop to respond to the list of questions adopted in December 2023.

**Action:** For adoption

List of Questions adopted on 14.12.2023.

The CHMP agreed to the request by the applicant for an extension to the clock stop to respond to the list of questions adopted in December 2023.

### 3.4.3. serplulimab - Orphan - EMEA/H/C/006170

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Henlius Europe GmbH, first-line treatment of adult patients with extensive-stage small cell lung cancer (ES-SCLC)

Scope: Update on procedure

**Action:** For discussion

List of Outstanding Issues adopted on 14.12.2023. List of questions adopted on 20.03.2023.

The CHMP noted the update on the procedure.

### 3.4.4. lecanemab - EMEA/H/C/005966

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a disease modifying treatment in adult patients with Mild Cognitive Impairment due to Alzheimer's disease and Mild Alzheimer's disease (Early Alzheimer's disease)

Scope: Draft list of experts for SAG

**Action:** For adoption

List of Outstanding Issues adopted on 09.11.2023. List of Questions adopted on 25.05.2023.

The CHMP adopted the list of experts for the SAG.

### 3.4.5. amino acids - Orphan - EMEA/H/C/005557

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Recordati Rare Diseases; treatment of decompensation episodes in MSUD patients

Scope: Letter by the applicant dated 15.02.2024 requesting an extension to the clock stop to respond to the list of questions adopted in January 2024.

**Action:** For adoption

List of Questions adopted on 25.01.2024.

The CHMP agreed to the request by the applicant for an extension to the clock stop to respond to the list of questions adopted in January 2024.

### 3.4.6. [epinephrine - EMEA/H/C/006139](#)

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Treatment of allergic reactions (anaphylaxis) and idiopathic or exercise induced anaphylaxis

Scope: Letter by the applicant dated 23.01.2024 requesting an extension to the clock stop to respond to the list of outstanding issues adopted in October 2023.

**Action:** For adoption

List of Outstanding Issues adopted on 12.10.2023. List of Questions adopted on 23.02.2023.

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 October 2023.

### 3.4.7. [teriparatide - EMEA/H/C/005687](#)

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treatment of osteoporosis

Scope: Letter by the applicant dated 08.02.2024 requesting an clock stop to respond to the list of questions adopted in November 2023.

**Action:** For adoption

List of Questions adopted on 09.11.2023.

The CHMP agreed to the request by the applicant for an clock stop to respond to the list of questions adopted in November 2023.

### 3.4.8. [tocilizumab - EMEA/H/C/005984](#)

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treatment of rheumatoid arthritis, active systemic juvenile idiopathic arthritis (sJIA), juvenile idiopathic polyarthritis (pJIA) and COVID-19

Scope: Letter by the applicant dated 13.02.2024 requesting an extension to the clock stop to respond to the list of outstanding issues adopted in May 2023.

**Action:** For adoption

List of outstanding issues adopted on 25.05.2023. List of Questions adopted on 26.01.2023.

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 May 2023.

### 3.4.9. [Bevacizumab - EMEA/H/C/005574](#)

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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: Letter by the applicant dated 20.02.2024 requesting an extension to the clock stop to respond to the list of outstanding issues adopted in February 2022.

**Action:** For adoption

List of Outstanding Issues adopted on 24.02.2022. List of Questions adopted on 22.04.2021.

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 February 2022.

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

#### **3.5.1. Nezglial - Ierigitazone - Orphan - EMEA/H/C/005757**

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Minoryx Therapeutics S.L.; the treatment of cerebral progression and myelopathy in male patients with adrenoleukodystrophy (ALD)

Re-examination Rapporteur: TBC, Re-examination Co-Rapporteur: TBC

Scope: Re-examination request, appointment of re-examination rapporteurs

**Action:** For adoption

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

Opinion adopted on 25.01.2023. List of Outstanding Issues adopted on 14.09.2023. List of Questions adopted on 15.12.2022.

The CHMP noted the call for re-examination rapporteurs.

#### **3.5.2. Syfovre - Pegcetacoplan - EMEA/H/C/005954**

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Apellis Netherlands B.V.; Treatment of geographic atrophy (GA) secondary to age-related macular degeneration (AMD)

Scope: Re-examination request, appointment of re-examination rapporteurs

**Action:** For adoption

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

Opinion adopted on 25.01.2023. List of Outstanding Issues adopted on 12.10.2023. List of Questions adopted on 25.05.2023.

The CHMP adopted a Re-examination Rapporteur and a Re-examination Co-Rapporteur.

### **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. Kalydeco - Ivacaftor - EMEA/H/C/002494/X/0115/G

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Vertex Pharmaceuticals (Ireland) Limited

Rapporteur: Maria Concepcion Prieto Yerro, Co-Rapporteur: Beata Maria Jakline Ullrich, PRAC Rapporteur: Monica Martinez Redondo

Scope: "Extension application to introduce a new strength (13.4 mg of ivacaftor granules in sachet), grouped with a type II variation (C.I.6.a) in order to extend the indication of the granule presentations to include children with cystic fibrosis aged 1 to less than 4 months of age and weighing 3 kg or more who have an R117H CFTR mutation or one of the approved 9 gating (class III) mutations based on interim results from study VX15-770-124 (study 124); this is a phase 3, 2-part, open-label study to evaluate the safety, pharmacokinetics, and pharmacodynamics of ivacaftor (IVA) in subjects with CF who are less than 24 months of age at treatment initiation and have a CFTR gating mutation. As a consequence, sections 1, 2, 4.1, 4.2, 4.4, 4.5, 4.8, 5.1, 5.2, 6.3 and 8 of the SmPC of the granules presentations and sections 4.2, 4.8, 5.1 and 5.2 of the SmPC of the tablets presentations are updated. The Labelling for the 13.4 mg granule presentation and the Package Leaflet of the granules and tablets presentations are updated in accordance. Version 15.2 of the RMP has also been submitted. In addition, the MAH took the opportunity to introduce minor editorial changes to the PI.

Type IA A.5.b

Type IA B.II.b.2.a"

**Action:** For adoption

List of Outstanding Issues adopted on 14.12.2023, 12.10.2023. List of Questions adopted on 25.05.2023.

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.

#### 4.1.2. Mounjaro - Tirzepatide - EMEA/H/C/005620/X/0015

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Eli Lilly Nederland B.V.

Rapporteur: Martina Weise

Scope: "Extension application to add 6 new strengths of 2.5 mg (4.17 mg/ml), 5 mg (8.33 mg/ml), 7.5 mg (12.5 mg/ml), 10 mg (16.67 mg/ml), 12.5 mg (20.83 mg/ml) and 15 mg (25 mg/ml) for Mounjaro solution for injection in pre-filled pen (KwikPen), multidose. The Package Leaflet and Labelling are updated in accordance."

**Action:** For adoption

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

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

#### 4.1.3. [Teriflunomide Accord - Teriflunomide - EMEA/H/C/005960/X/0002](#)

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Accord Healthcare S.L.U.

Rapporteur: Kristina Nadrah, PRAC Rapporteur: Martin Huber

Scope: "Extension application to add a new strength of 7 mg film-coated tablets. The bioequivalence study data were submitted."

**Action:** For adoption

List of Outstanding Issues adopted on 14.12.2023. List of Questions adopted on 20.07.2023.

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.

## 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. [Betmiga - Mirabegron - EMEA/H/C/002388/X/0039/G](#)

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Astellas Pharma Europe B.V.

Rapporteur: Maria Concepcion Prieto Yerro, PRAC Rapporteur: Maria del Pilar Rayon

Scope: "Extension application to introduce a new pharmaceutical form associated with a new strength (8 mg/ml prolonged-release granules for oral suspension), grouped with a type II variation (C.I.6.a) to include treatment of neurogenic detrusor overactivity (NDO) in pediatric patients aged 3 to less than 18 years. The RMP (version 9.0) is updated in accordance."

**Action:** For adoption

List of Outstanding Issues adopted on 15.09.2022. List of Questions adopted on 22.04.2022.

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

The Committee adopted a list of outstanding issues.

The CHMP agreed to the request by the MAH for an extension to the clock stop to respond to the list of outstanding issues.

Roche Registration GmbH

Rapporteur: Paolo Gasparini, PRAC Rapporteur: Bianca Mulder

Scope: "Extension application to:

1) Introduce a new pharmaceutical form (coated granules) associated with a new strength (50 mg).

2) Introduce a new route of administration (gastroenteral use) for the already authorised 100 mg and 200 mg hard capsules presentations.

The above two line extensions are grouped with 3 type II variations:

- C.I.6.a - To extend the currently approved indication in solid tumours with NTRK gene fusion to patients from birth to 12 years of age (both for the coated granules and already approved hard capsules presentations).

- C.I.6.a - To add a new paediatric indication from birth to 18 years of age for patients with solid tumours with a ROS1 gene fusion (both for the coated granules and already approved hard capsules presentations).

Based on final results from studies CO40778 (STARTRK-NG), GO40782 (STARTRK-2) and BO41932 (TAPISTRY). Study CO40778 is a Phase I/II open-label, dose-escalation and expansion study of entrectinib in pediatrics with locally advanced or metastatic solid or primary CNS tumors and/or who have no satisfactory treatment options; study GO40782 is an open-label, multicenter, global Phase II basket study of entrectinib for the treatment of patients with solid tumors that harbor an NTRK1/2/3, ROS1, or ALK gene rearrangement (fusion), and study BO41932 is a Phase II, global, multicenter, open-label, multi-cohort study designed to evaluate the safety and efficacy of targeted therapies or immunotherapy as single agents or in rational, specified combinations in participants with unresectable, locally advanced or metastatic solid tumors determined to harbor specific oncogenic genomic alterations or who are tumor mutational burden (TMB)-high as identified by a validated next-generation sequencing (NGS) assay.

As a consequence, sections 4.1, 4.2, 4.4, 4.5, 4.8, 5.1, 5.2, 6.3, 6.4 and 6.6 of the SmPC are updated accordingly. The Package Leaflet and Labelling are updated in accordance.

- C.I.4 - To add a wording regarding the option of suspension in water of the content of the capsules to be used orally or via the e.g., gastric or nasogastric tube (in sections 4.2 and 5.2 of the SmPC).

The RMP (version 5) is updated in accordance.

The MAH took the opportunity to introduce minor editorial changes to the PI and to update Annex II of the SmPC."

**Action:** For adoption

List of Questions adopted on 14.09.2023.

The Committee was reminded of the status of this application and its remaining outstanding issues relating to clinical aspects.

The Committee adopted a list of outstanding issues with 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. Ofev - Nintedanib - EMEA/H/C/003821/X/0057/G

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Boehringer Ingelheim International GmbH

Rapporteur: Finbarr Leacy, Co-Rapporteur: Ewa Balkowiec Iskra, PRAC Rapporteur: Barbara Kovacic Bytyqi

Scope: "Extension application to add a new strength of 25 mg hard capsules, grouped with an extension of indication (C.I.6.a) to include treatment of fibrosing Interstitial Lung Diseases (ILDs) in children and adolescents from 6 to 17 years of age for Ofev, following the assessment of procedure X/0052/G, based on final results from study 1199-0337 (A Double Blind, Randomised, Placebo-controlled Trial to Evaluate the Dose-exposure and Safety of Nintedanib Per os on Top of Standard of Care for 24 Weeks, Followed by Open Label Treatment With Nintedanib of Variable Duration, in Children and Adolescents (6 to 17 Year-old) With Clinically Significant Fibrosing Interstitial Lung Disease), which is supplemented by the currently ongoing prospective Phase III extension trial 1199-0378 (An Open-label Trial of the Long-term Safety and Tolerability of Nintedanib Per os, on Top of Standard of Care, Over at Least 2 Years, in Children and Adolescents With Clinically Significant Fibrosing Interstitial Lung Disease). The main objective of the study 1199-0337 was to evaluate dose-exposure and safety of nintedanib in children and adolescents with fibrosing Interstitial Lung Disease (ILD). As a consequence, sections 4.1, 4.2, 4.4, 4.5, 4.8, 5.1 and 5.2 of the SmPC are updated. The Package Leaflet and Labelling are updated in accordance. Version 12.0 of the RMP has also been submitted."

**Action:** For adoption

The Committee discussed the issues identified in this application relating to quality and clinical aspects.

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

The CHMP agreed to consult a SAG and adopted the list of questions to these experts.

#### 4.3.2. Opsumit - Macitentan - EMEA/H/C/002697/X/0051/G

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Janssen-Cilag International N.V.

Rapporteur: Maria Concepcion Prieto Yerro, Co-Rapporteur: Patrick Vrijlandt, PRAC Rapporteur: Maria del Pilar Rayon

Scope: "Extension application to introduce a new pharmaceutical form associated with new strengths (1 and 2.5 mg dispersible tablet) grouped with an extension of indication (C.I.6.a) to include, as monotherapy or in combination, the long-term treatment of pulmonary arterial hypertension (PAH) in paediatric patients aged 1 month to less than 18 years of age of WHO Functional Class (FC) I to III for OPSUMIT, based on interim results from AC-055-312 study (TOMORROW). This is a multicenter, open-label, randomized study with single-arm extension period to assess the pharmacokinetics, safety, and efficacy of macitentan versus standard of care in children with pulmonary arterial hypertension. As a consequence,

sections 4.1, 4.2, 4.4, 4.8, 4.9, 5.1 and 5.2 of the SmPC for film-coated tablets are updated. The Package Leaflet and Labelling are updated in accordance. Version 14.1 of the RMP has also been submitted.”

**Action:** For adoption

The Committee discussed the issues identified in this application relating to clinical aspects.

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

#### 4.3.3. Ozempic - Semaglutide - EMEA/H/C/004174/X/0043

Novo Nordisk A/S

Rapporteur: Patrick Vrijlandt

Scope: quality

**Action:** For adoption

The Committee discussed the issues identified in this application relating to quality aspects.

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

#### 4.3.4. Rybelsus - Semaglutide - EMEA/H/C/004953/X/0038

Novo Nordisk A/S

Rapporteur: Patrick Vrijlandt

Scope: “Extension application to introduce three new strengths of tablets (1.5 mg, 4 mg and 9 mg) for semaglutide.”

**Action:** For adoption

The Committee discussed the issues identified in this application relating to clinical and quality aspects.

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

#### 4.3.5. Rybelsus - Semaglutide - EMEA/H/C/004953/X/0039

Novo Nordisk A/S

Rapporteur: Patrick Vrijlandt

Scope: “Extension application to add two new strengths (25 mg and 50 mg) tablets.”

**Action:** For adoption

The Committee discussed the issues identified in this application relating to quality aspects.

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

#### 4.3.6. Wegovy - Semaglutide - EMEA/H/C/005422/X/0016

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Novo Nordisk A/S

Rapporteur: Patrick Vrijlandt

Scope: quality

**Action:** For adoption

The Committee discussed the issues identified in this application relating to quality 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**

No items

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

No items

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

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

##### 5.1.1. CARVYKTI - Ciltacabtagene autoleucel - Orphan - ATMP - EMEA/H/C/005095/II/0021

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Janssen-Cilag International NV

Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Jo Robays

Scope: "Extension of indication to include treatment of adult patients with relapsed and refractory multiple myeloma, who have received at least 1 prior therapy, including an IMiD and a PI, have demonstrated disease progression on or after the last therapy and are refractory to lenalidomide for CARVYKTI, based on interim results from study MMY3002 listed as a specific obligation (SOB/006) in the Annex II. This is an ongoing, Phase 3, randomized, open-label, multicentre study to determine whether treatment with cilta-cel provides an efficacy benefit compared to standard therapy in participants with relapsed and lenalidomide-refractory multiple myeloma. As a consequence, sections 4.1, 4.4, 4.5, 4.8, 5.1 and 5.2 of the SmPC are updated. The Package Leaflet is updated in accordance. Version 4.1 of the RMP has also been submitted. In addition, the MAH took the opportunity

to update Annex II of the PI. As part of the application the MAH is requesting a 1-year extension of the market protection.”, 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 08.12.2023, 08.09.2023.

The CHMP was updated on discussions at CAT.

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

Based on the draft opinion from CAT the Committee adopted a positive opinion by consensus together with the CHMP Assessment Report and translation timetable.

The CHMP also recommended converting of the conditional marketing authorisation to a standard marketing authorisation, as the obligations of the conditional approval have now been met.

The CHMP noted the letter of recommendation dated 19 February.

The summary of opinion was circulated for information.

#### 5.1.2. [Cibinqo - Abrocitinib - EMEA/H/C/005452/II/0010](#)

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Pfizer Europe MA EEIG

Rapporteur: Kristina Dunder, PRAC Rapporteur: Petar Mas

Scope: “Extension of indication to include treatment of adolescents 12 to < 18 years of age with moderate to severe atopic dermatitis for CIBINQO based on final results from non-clinical study 00655292 [21GR211] and interim results from clinical study B7451015; this is a Phase III multi-center, long-term extension study investigating the efficacy and safety of abrocitinib, with or without topical medications, administered to subjects aged 12 years and older with moderate to severe atopic dermatitis. As a consequence, sections 4.1, 4.2, 4.8 and 5.1 of the SmPC are updated. The Package Leaflet is updated in accordance. Version 4.1 of the RMP has also been submitted.”

**Action:** For adoption

Request for Supplementary Information adopted on 14.12.2023, 14.09.2023.

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 summary of opinion was circulated for information.

#### 5.1.3. [Hepcludex - Bulevirtide - Orphan - EMEA/H/C/004854/II/0031](#)

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Gilead Sciences Ireland Unlimited Company

Rapporteur: Filip Josephson, PRAC Rapporteur: Adam Przybylkowski

Scope: “Extension of indication to include treatment of chronic hepatitis delta virus (HDV)

infection in paediatric patients 3 years of age and older weighing at least 10 kg with compensated liver disease for Hepcludex, based on a modelling and simulation study and an extrapolation study to evaluate the use of Bulevirtide for the treatment of chronic hepatitis D infection in children from 3 to less than 18 years of age. As a consequence, sections 4.1, 4.2, 5.1 and 5.2 of the SmPC are updated. The Package Leaflet has been updated accordingly. Version 4.1 of the RMP has also been submitted. In addition, the marketing authorisation holder (MAH) took the opportunity to introduce minor editorial changes to the PI.”

**Action:** For adoption

The Committee discussed the issues identified in this application relating to clinical aspects.

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

#### 5.1.4. Keytruda - Pembrolizumab - EMEA/H/C/003820/II/0134

Merck Sharp & Dohme B.V.

Rapporteur: Paolo Gasparini, PRAC Rapporteur: Bianca Mulder

Scope: “Extension of indication to include in combination with platinum-containing chemotherapy as neoadjuvant treatment, and then continued as monotherapy as adjuvant, treatment of resectable Stage II, IIIA, or IIIB (T3 4N2) non-small cell lung carcinoma in adults for Keytruda based on study KEYNOTE-671, a phase III, randomized, double-blind trial of platinum doublet chemotherapy +/- pembrolizumab as neoadjuvant/adjuvant therapy for participants with resectable stage II, IIIA, and resectable IIIB (T3-4N2) non-small cell lung cancer. As a consequence, sections 4.1, 4.2 and 5.1 of the SmPC are updated. The Package Leaflet is updated in accordance. Version 41.1 of the RMP has also been submitted.”

**Action:** For adoption

Request for Supplementary Information adopted on 14.12.2023, 22.06.2023.

See 2.3

An oral explanation was held on 21 February 2024. The presentation by the MAH focused on clinical aspects.

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 summary of opinion was circulated for information.

#### 5.1.5. Nilemdo - Bempedoic acid - EMEA/H/C/004958/II/0031

Daiichi Sankyo Europe GmbH

Rapporteur: Patrick Vrijlandt, Co-Rapporteur: Alar Irs, PRAC Rapporteur: Kimmo Jaakkola

Scope: “Extension of indication to include treatment of adults with established or at high risk for atherosclerotic cardiovascular disease to reduce cardiovascular risk, based on results from study 1002-043 (CLEAR). CLEAR Outcomes Study is a phase 3 multi-centre

randomised, double-blind, placebo-controlled study to evaluate whether long-term treatment with bempedoic acid reduces the risk of major adverse cardiovascular events (MACE) in patients with, or at high risk for, cardiovascular disease who are statin intolerant. As a consequence, sections 4.1, 4.8 and 5.1 of the SmPC are updated. The Package Leaflet is updated accordingly. Version 4.0 of the RMP has also been submitted. In addition, the MAH took the opportunity to introduce minor formatting changes to the PI. As part of the application, the MAH is requesting a 1-year extension of the market protection.", 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 09.11.2023.

The Committee discussed the issues identified in this application relating to clinical aspects.

The Committee adopted a 2<sup>nd</sup> request for supplementary information with a specific timetable.

#### 5.1.6. Nustendi - Bempedoic acid / Ezetimibe - EMEA/H/C/004959/II/0035

Daiichi Sankyo Europe GmbH

Rapporteur: Patrick Vrijlandt, Co-Rapporteur: Alar Irs, PRAC Rapporteur: Kimmo Jaakkola

Scope: "Extension of indication to include treatment of adults with established or at high risk for atherosclerotic cardiovascular disease to reduce cardiovascular risk for NUSTENDI, based on results from study 1002-043, known as the CLEAR [Cholesterol Lowering via Bempedoic Acid, an ATP citrate lyase (ACL) Inhibiting Regimen] Outcomes Trial; this is a Phase 3, randomized, double-blind, placebo-controlled study to assess the effects of bempedoic acid (ETC-1002) on the occurrence of major cardiovascular events in patients with, or at high risk for, cardiovascular disease who are statin intolerant; As a consequence, sections 4.1, 4.8 and 5.1 of the SmPC are updated. The Package Leaflet is updated in accordance. Version 4.0 of the RMP has also been submitted. As part of the application the MAH is requesting a 1-year extension of the market protection.", 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 09.11.2023.

The Committee discussed the issues identified in this application relating to clinical aspects.

The Committee adopted a 2<sup>nd</sup> request for supplementary information with a specific timetable.

#### 5.1.7. Pegasys - Peginterferon alfa-2a - EMEA/H/C/000395/II/0119/G

Pharmaand GmbH

Rapporteur: Filip Josephson, PRAC Rapporteur: Ulla Wändel Liminga

Scope: "Grouped application consisting of:

Extension of indication to include treatment of Polycythaemia Vera (PV) and Essential thrombocytopenia (ET) for PEGASYS, based on published data of clinical studies conducted in support of the efficacy and safety of Pegasys for the treatment of ET and PV. As a

consequence, sections 4.1, 4.2, 4.8 and 5.2 of the SmPC are updated. The Package Leaflet is updated in accordance. Version 10.1 of the RMP has also been submitted. Furthermore, the PI is brought in line with the latest QRD template version 10.3.”

**Action:** For adoption

The Committee discussed the issues identified in this application relating to clinical and the RMP.

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

#### 5.1.8. Reblozyl - Luspatercept - Orphan - EMEA/H/C/004444/II/0021

Bristol-Myers Squibb Pharma EEIG

Rapporteur: Daniela Philadelphia, Co-Rapporteur: Ewa Balkowiec Iskra, PRAC Rapporteur: Jo Robays

Scope: “Extension of indication to include treatment of adult patients with anaemia due to very low, low and intermediate-risk myelodysplastic syndromes (MDS), who may require RBC transfusions for Reblozyl, based on results from study ACE-536-MDS-002 (COMMANDS), an active-controlled, open-label, randomized Phase 3 study comparing the efficacy and safety of luspatercept vs epoetin alfa in adult subjects with anemia due to IPSS-R very low, low or intermediate risk MDS, who are ESA naïve and require RBC transfusions, and studies ACE-536-MDS-001(MEDALIST), ACE-536-MDS-004, A536-03, A536-05 and ACE-536-LTFU-001. 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 2.0 of the RMP has also been submitted.”

**Action:** For adoption

Request for Supplementary Information adopted on 12.10.2023, 22.06.2023.

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 summary of opinion was circulated for information.

#### 5.1.9. SIRTURO - Bedaquiline - Orphan - EMEA/H/C/002614/II/0056

Janssen-Cilag International N.V.

Rapporteur: Filip Josephson, PRAC Rapporteur: Ulla Wändel Liminga

Scope: “Extension of indications by removal of the restriction for use of SIRTURO (bedaquiline [BDQ]), based on final results from study STREAM Stage 2; this is a multicenter, open-label, parallel-group, randomized, active-controlled study in participants aged 15 years or older with RR/MDR-TB to evaluate an investigational BDQ-containing, all-oral, 40-week regimen of anti-TB drugs (Regimen C) compared to an injectable-containing 40-week control regimen (Regimen B). As a consequence of the data emerging from the submitted study, sections 2, 4.1, 4.2, 4.4, 4.5, 4.8, 4.9, 5.1, 5.2 and 5.3 of the SmPC are updated. In addition, section E of Annex II has also been updated. The Labelling and Package Leaflet are updated in accordance. Version 10.1 of the RMP has also been

submitted. Furthermore, the PI is brought in line with the latest QRD template version 10.3. As part of the application, the MAH is requesting the switch from a conditional MA to standard MA.”

**Action:** For adoption

The Committee discussed the issues identified in this application relating to clinical aspects and the RMP.

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

#### 5.1.10. Tepkinly - Epcoritamab - Orphan - EMEA/H/C/005985/II/0001

AbbVie Deutschland GmbH & Co. KG

Rapporteur: Peter Mol, Co-Rapporteur: Ingrid Wang, PRAC Rapporteur: Monica Martinez Redondo

Scope: “Extension of indication to include treatment of adult patients with relapsed or refractory (R/R) follicular lymphoma (FL) after two or more lines of systemic therapy for TEPKINLY, based on results from the indolent Non-Hodgkins Lymphoma (iNHL) expansion cohort of Study GCT3013-01, the First In Human (FIH) Phase 1/2 study in R/R B-NHL, with key supportive data from the Phase 1b/2 Study GCT3013-04 in Japanese subjects. Study GCT3013-01 is an ongoing global, single-arm, Phase 1/2 study designed to evaluate epcoritamab as monotherapy in R/R B-NHL. As a consequence, sections 1, 3, 4.1, 4.2, 4.4, 4.8, 5.1, 5.2, 6.3, 6.4, 6.5 and 6.6 of the SmPC are updated. The Package Leaflet and Labelling are updated in accordance. Version 2.0 of the RMP has also been submitted. In addition, the MAH took the opportunity to introduce minor changes to the PI.”, Request for 1 year of market protection for a new indication (Article 14(11) of Regulation (EC) 726/2004)

**Action:** For adoption

The Committee discussed the issues identified in this application relating to clinical aspects.

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

#### 5.1.11. Xromi - Hydroxycarbamide - EMEA/H/C/004837/II/0019

Nova Laboratories Ireland Limited

Rapporteur: Anastasia Mountaki, Co-Rapporteur: Karin Janssen van Doorn, PRAC Rapporteur: Jo Robays

Scope: “Extension of indication to include the prevention of vaso-occlusive complications of sickle cell disease in children from 6 months to 2 years of age for Xromi, based on final results from the paediatric study INV543, listed as a category 3 study in the RMP; this is a single-arm, open-label, multi-center study in children with sickle cell anaemia over 6 months of age.

As a consequence, sections 4.1, 4.2, 4.8, 5.1 and 5.2 of the SmPC are updated. The Package Leaflet is updated in accordance. Version 4.1 of the RMP has also been submitted.”

**Action:** For adoption

Request for Supplementary Information adopted on 09.11.2023, 26.04.2023.

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 summary of opinion was circulated for information.

5.1.12. **WS2551**  
**Kaftrio - Ivacaftor / Tezacaftor / Elexacaftor - EMEA/H/C/005269/WS2551/0043**  
**Kalydeco - Ivacaftor - EMEA/H/C/002494/WS2551/0121**

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Vertex Pharmaceuticals (Ireland) Limited

Lead Rapporteur: Peter Mol, PRAC Rapporteur: Martin Huber

Scope: "Extension of the indication for Kaftrio (ivacaftor/tezacaftor/elexacaftor) and Kalydeco (ivacaftor) in a combination regimen to include the treatment of patients with cystic fibrosis (CF) aged 2 years and older who do not carry any F508del mutations and have at least one ivacaftor/tezacaftor/elexacaftor-responsive mutation in the cystic fibrosis transmembrane conductance regulator (CFTR) gene based on study VX21-445-124, study VX21-445-125 and study VX22-CFD-016. As a consequence, sections 4.1, 4.2, 4.8 and 5.1 of the Kaftrio SmPC are updated; sections 4.1 and 5.1 of the Kalydeco SmPC are updated. The Package Leaflet is updated in accordance. In addition, the MAH took this opportunity to introduce editorial changes to the PI."

**Action:** For adoption

The Committee discussed the issues identified in this application relating to non-clinical and 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. Medical devices**

### **6.1. Ancillary medicinal substances - initial consultation**

No items

## 6.2. Ancillary medicinal substances – post-consultation update

No items

## 6.3. Companion diagnostics - initial consultation

No items

## 6.4. Companion diagnostics – follow-up consultation

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. Sepiapterin - H0006331

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For the treatment of hyperphenylalaninaemia (HPA) in adults and paediatric patients of all ages with phenylketonuria (PKU).

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

**Action:** For adoption

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

#### 8.1.2. Chikungunya Virus Virus-Like Particle Vaccine – H0005470

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Active immunisation to prevent disease caused by chikungunya virus infection in individuals age 12 years and older

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

The CHMP adopted the recommendations for PRIME eligibility.

The individual outcomes are listed in the PRIME Monthly Report on the EMA website, in the PRIME homepage, under Outcome of eligibility section.

## 9. Post-authorisation issues

### 9.1. Post-authorisation issues

#### 9.1.1. Truvelog Mix 30 - insulin aspart - EMEA/H/C/005635

Sanofi Winthrop Industrie; treatment of diabetes mellitus

Rapporteur: Martina Weise, Co-Rapporteur: Thalia Marie Estrup Blicher, PRAC Rapporteur: Mari Thorn

Scope: Withdrawal of marketing authorisation

**Action:** For information

The CHMP noted the withdrawal of marketing authorisation.

#### 9.1.2. Zoledronic acid Actavis – zoledronic acid – EMEA/H/C/002488

Actavis Group PTC ehf.; prevention of skeletal related events and treatment of tumour-induced hypercalcaemia (TIH)

Rapporteur: Christian Gartner

Scope: Withdrawal of marketing authorisation

**Action:** For information

The CHMP noted the withdrawal of marketing authorisation.

#### 9.1.3. Zoledronic acid Hospira – zoledronic acid – EMEA/H/C/002365

Pfizer Europe MA EEIG; prevention of skeletal related events and treatment of tumour-induced hypercalcaemia (TIH)

Rapporteur: Kristina Dunder

Scope: Withdrawal of marketing authorisation

**Action:** For information

The CHMP noted the withdrawal of marketing authorisation.

#### 9.1.4. Orenzia - Abatacept - EMEA/H/C/000701/II/0152

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Bristol-Myers Squibb Pharma EEIG

Rapporteur: Outi Mäki-Ikola, Co-Rapporteur: Beata Maria Jakline Ullrich, PRAC Rapporteur: Kimmo Jaakkola

Scope: "Extension of indication to include the prophylaxis of acute Graft versus Host Disease (aGvHD) in the adult and paediatric population for Orenzia, based on final results from studies IM101311 - Abatacept Combined With a Calcineurin Inhibitor and Methotrexate for Graft Versus Host Disease Prophylaxis and IM101841 - Overall Survival In 7/8 HLA-Matched Hematopoietic Stem Cell Transplantation Patients Treated With Abatacept Combined With A Calcineurin Inhibitor And Methotrexate - An Analysis Of The Center For International Blood And Marrow Transplant Research (Cibmtr) Database. As a consequence, sections 4.1, 4.2, 4.4, 4.8, 5.1 and 5.2 of the SmPC are updated. The Package Leaflet and Labelling are updated in accordance. Version 28.0 of the RMP has also been submitted. In addition, the MAH took the opportunity to introduce minor editorial changes to the PI."

Withdrawal of extension of indication application.

**Action:** For information

Request for Supplementary Information adopted on 12.10.2023, 30.03.2023.

See 2.3

The CHMP noted the withdrawal of extension of indication application.

## 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. Colistimethate sodium (CMS) – EMEA/H/A-5(3)/1524

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Various MAHs

Referral Rapporteur: Martina Weise, Referral Co-Rapporteur: Ewa Balkowiec Iskra

Scope: List of outstanding issues

**Action:** For adoption

Review of the ratio of polymyxins E1 and E2 in colistin starting material and of the (sulfomethylation) composition profile of CMS finished product.

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

CHMP list of outstanding issues: February 2024 CHMP

Submission of responses: 29 February 2024

Re-start of the procedure: 04 March 2024

Rapporteurs' joint assessment report circulated to CHMP: 07 March 2024

Comments: 12 March 2024

Updated Rapporteurs' joint assessment report circulated to CHMP: 14 March 2024

CHMP list of outstanding issues/CHMP opinion: March 2024 CHMP

### **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**

#### **10.4.1. Ibuprofen NVT – ibuprofen - EMEA/H/A-29(4)/1533**

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Laboratorios Liconsa, S.A.

Referral Rapporteur: Vilma Petrikaite, Referral Co-Rapporteur: Maria Concepcion Prieto Yerro

Scope: Opinion

**Action:** For adoption

Mutual Recognition Procedure number: LT/H/0162/002/E/001, notification sent by the Agency of Lithuania dated 17 November 2023 notifying of the start of a referral under Article 29(4) of Directive 2001/83/EC.

The CHMP adopted an opinion by consensus, concluding that the benefits of Ibuprofen NVT 400 mg do not outweigh its risks and the marketing authorisation granted in Lithuania cannot be recognised in Spain, where the company had applied for a marketing authorisation.

In addition, the marketing authorisations in Lithuania and other Member States where the medicine is authorised (Estonia, France, Latvia, Poland and Romania) should be suspended.

The EMA question and answer document was circulated for information.

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

No items

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

No items

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

### **10.7.1. Synapse Labs Pvt. Ltd. – various – EMEA/H/A-31/1529**

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Various MAHs

Scope: Appointment of re-examination rapporteurs at the PROM meeting on 12 February 2024

**Action:** For information

Article 31 procedure triggered by the Agency of Medicines and Medical Devices (AEMPS) in Spain, concerning the contract research organisation (CRO) Synapse Labs Pvt. Ltd., located in Kharadi, Pune, India.

At the February PROM meeting, the CHMP appointed a re-examination referral rapporteur and a re-examination referral co-rapporteur.

## **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**

### **10.11.1. Lorazepam Macure – lorazepam - EMEA/H/A-13/1536**

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Macure Pharma ApS

Referral Rapporteur: Peter Mol, Referral Co-Rapporteur: Kristina Dunder

Scope: Appointment of Rapporteurs, list of questions (TBC), timetable

**Action:** For adoption

Variation number in decentralised procedure: NL/H/4353/001/II/004, notification sent by the Agency of The Netherlands dated 01 February 2024 notifying of the start of a referral under Article 13(1) of Regulation No 1234/2008.

The CHMP appointed Peter Mol as referral rapporteur and Kristina Dunder as referral co-rapporteur.

The CHMP agreed on a rapporteur-led procedure (i.e. no list of questions to the MAHs adopted in February 2024) together with a timetable.

Notification: 01.02.2024

Start of the procedure (CHMP): February 2024 CHMP

Rapporteur/co-rapporteur assessment reports circulated to CHMP: 07 March 2024

Comments: 12.03.2024

Updated Rapporteur/co-rapporteur assessment reports circulated to CHMP: 14 March 2024

CHMP list of questions or CHMP opinion: March 2024 CHMP

## 11. Pharmacovigilance issue

### 11.1. Early Notification System

February 2024 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. CHMP co-opted membership**

---

The 3-year co-opted member mandate for Carla Torre comes to an end on 21.02.2024.

The 3-year co-opted member mandate for Blanka Hirschlerova comes to an end on 18.03.2024.

The CHMP agreed that co-opted members should be appointed in the following areas of expertise:

- Position 1: Quality (non-biologicals),
- Position 2: Pharmacoepidemiology; especially for methodological analysis and interpretation of data in particular study designs\*.

\*The experience in pharmacoepidemiology should be applied to regulatory decision-making processes. The interpretation of data in particular study designs should include strengths and weaknesses (observational studies, RWD from different sources).

A call for nomination of co-opted members was launched following the January 2024 plenary.

Nomination(s) received

**Action:** For election

The CHMP elected Blanka Hirschlerova as co-opted member with expertise in quality (non-biologicals), and Carla Torre with expertise in pharmacoepidemiology; especially for methodological analysis and interpretation of data in particular study designs.

## 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 2024

**Action:** For adoption

The CHMP adopted the EURD list.

### 14.2.2. Paediatric Committee (PDCO)

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Agenda of the February 2024 PDCO plenary meeting.

**Action:** For information

The CHMP noted the PDCO agenda.

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

### 14.3.1. Biologics Working Party (BWP)

---

Chair: Sean Barry, Vice-Chair: Francesca Luciani

Reports from BWP February 2024 meeting to CHMP for adoption:

- 16 reports on products in scientific advice and protocol assistance
- 14 reports on products in pre-authorisation procedures
- 4 reports on products in post-authorisation procedures
- 3 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 13-14 February 2024.

**Action:** For adoption

The CHMP adopted the table of decisions.

### 14.3.3. Scientific Advice Working Party (SAWP)

---

Chair: Paolo Foggi

Report from the SAWP meeting held on 05-08 February 2024. Table of conclusions

**Action:** For information

Scientific advice letters:

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

The CHMP noted the update.

### 14.3.4. Guideline on allergen products development in moderate to low-sized study populations - Rheumatology and Immunology Working Party (RIWP)

---

Follow-up from the February PROM. The CHMP had discussed the guideline at the February PROM meeting and agreed to the content. Following the adoption of the guideline by the PDCO, the CHMP is asked to finally adopt the guideline.

**Action:** For adoption

The CHMP adopted the guideline on allergen products development in moderate to low-sized study populations.

## 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**

### **14.9.1. CHMP Learnings**

---

CHMP: Outi Mäki-Ikola

Collection, discussion and recording of CHMP learnings.

**Action:** For information

The CHMP noted the information.

## **15. Any other business**

### **15.1. AOB topic**

No items

## List of participants

List of participants including any restrictions with respect to involvement of members/alternates/experts following evaluation of declared interests for the 19-22 February 2024 CHMP meeting, which was held remotely.

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	
Daniela Philadelphia	Member	Austria	No interests declared	
Christian Gartner	Alternate	Austria	No interests declared	
Christophe Focke	Member	Belgium	No restrictions applicable to this meeting	
Karin Janssen van Doorn	Alternate	Belgium	No interests declared	
Lyubina Racheva Todorova	Member	Bulgaria	No interests declared	
Gergana Lazarova	Alternate	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	
Tomas Radimersky	Member	Czechia	No interests declared	
Petr Vrbata	Alternate	Czechia	No interests declared	
Thalia Marie Estrup Blicher	Member	Denmark	No interests declared	
Aaron Sosa Mejia	Alternate	Denmark	No participation in final deliberations and voting on:	Concizumab - EMEA/H/C/005938 Insulin icodec - EMEA/H/C/005978 Ozempic - Semaglutide - EMEA/H/C/004174/X/0043 Rybelsus - Semaglutide - EMEA/H/C/004953/X/0038, EMEA/H/C/004953/X/0039 Wegovy - Semaglutide - EMEA/H/C/005422/X/0016
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	
Anastasia Mountaki	Alternate	Greece	No interests declared	
Robert Porszasz	Member	Hungary	No interests declared	
Beata Maria Jakline Ullrich	Alternate	Hungary	No interests declared	
Hrefna Gudmundsdottir	Member	Iceland	No interests declared	
HjalTI Kristinsson	Alternate	Iceland	No interests declared	
Jayne Crowe	Member	Ireland	No interests declared	
Finbarr Leacy	Alternate	Ireland	No interests declared	
Paolo Gasparini	Member	Italy	No interests declared	
Maria Grazia Evandri	Alternate	Italy	No interests declared	
Elita Poplavska	Member	Latvia	No interests declared	
Vilma Petrikaite	Member	Lithuania	No interests declared	
Martine Trauffler	Member	Luxembourg	No interests declared	
Alexandra Branchu	Alternate	Luxembourg	No participation in discussion, final deliberations and voting on:	Truvelog Mix 30 - insulin aspart - EMEA/H/C/005635
John Joseph Borg	Member	Malta	No interests declared	
Peter Mol	Member	Netherlands	No interests declared	
Patrick Vrijlandt	Alternate	Netherlands	No interests declared	
Ingrid Wang	Member	Norway	No interests declared	
Eva Skovlund	Alternate	Norway	No interests declared	
Ewa Balkowiec Iskra	Member	Poland	No interests declared	
Grzegorz Cessak	Alternate	Poland	No interests declared	
Bruno Sepodes	Member (Vice-Chair)	Portugal	No interests declared	
Fatima Ventura	Alternate	Portugal	No restrictions applicable to this meeting	
Simona Badoi	Member	Romania	No interests declared	
Dana Gabriela Marin	Alternate	Romania	No interests declared	
Frantisek Drafi	Member	Slovakia	No interests declared	
Jana Klimasová	Alternate	Slovakia	No restrictions applicable to this meeting	
Andreja Kranjc	Alternate	Slovenia	No interests declared	
Maria Concepcion Prieto Yerro	Member	Spain	No interests declared	
Carolina Prieto Fernandez	Alternate	Spain	No interests declared	
Kristina Dunder	Member	Sweden	No interests declared	
Filip Josephson	Alternate	Sweden	No interests declared	
Bruno Delafont	Co-opted member	France	No interests declared	
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	
Jana Schweigertova	Expert	Slovakia	No interests declared	
Jenny-Maria Jönsson	Expert	Sweden	No participation in discussion, final deliberations and voting on:	Keytruda - Pembrolizumab - EMEA/H/C/003820/II/0 134
Daiana Vasilcanu	Expert	Sweden	No interests declared	
Helena Back	Expert	Sweden	No interests declared	
Per Albertsson	Expert	Sweden	No restrictions applicable to this meeting	
Maria Winqvist	Expert	Sweden	No interests declared	
Agnieszka Przybyszewska	Expert	Ireland	No interests declared	
Antero Kallio	Expert	Finland	No restrictions applicable to this meeting	
Riitta Niittyvuopio	Expert	Finland	No restrictions applicable to this meeting	
Tommi Nurminen	Expert	Finland	No restrictions applicable to this meeting	
Maija Tarkkanen	Expert	Finland	No interests declared	
Paolo Foggi	Expert	Italy	No interests declared	
Karri Penttila	Expert	Finland	No interests declared	
Rosalía Ruano Camps	Expert	Spain	No interests declared	
Viktorii Starokozhko	Expert	Netherlands	No restrictions applicable to this meeting	
Mikael Andersson	Expert	Sweden	No interests declared	
Andrea Laslop	Expert	Austria	No interests declared	
Hilke Zander	Expert	Germany	No interests declared	
Gabriele Maurer	Expert	Germany	No restrictions applicable to this meeting	
Igor Guljašević	Expert	Croatia	No interests declared	
Susanne Müller-Egert	Expert	Germany	No interests declared	
Kay-Martin Hanschmann	Expert	Germany	No restrictions applicable to this meeting	
Edwige Haelterman	Expert	Belgium	No interests declared	
Ingrid Bourges	Expert	Belgium	No interests declared	
Marta Romano	Expert	Belgium	No interests declared	
Kaatje Smits	Expert	Belgium	No interests declared	
Gaelle De Meyer	Expert	Belgium	No restrictions applicable to this meeting	
Violette Dirix	Expert	Belgium	No interests declared	
Anna Zadrozna	Expert	Poland	No interests declared	
Paweł Pawłowski	Expert	Poland	No interests declared	
Anna Mari Lone	Expert	Norway	No restrictions applicable to this meeting	

Aina Jannicke Øvrebust	Expert	Norway	No interests declared
Sara Tognarelli	Expert	Germany	No restrictions applicable to this meeting
Antonio Gomez Outes	Expert	Spain	No interests declared
Vincent Gazin	Expert	France	No interests declared
Attila Megyeri	Expert	Hungary	No interests declared
Luca Fancsalszky	Expert	Hungary	No interests declared
Anna Katalin Barane Gilicze	Expert	Hungary	No restrictions applicable to this meeting
Harald Bernsteiner	Expert	Austria	No interests declared
Tobias Glüxam	Expert	Austria	No interests declared
Melanie Ramberger	Expert	Austria	No interests declared
Helga Sgardelli	Expert	Austria	No interests declared
Sabrina Jenull	Expert	Austria	No restrictions applicable to this meeting
Bernhard Majer	Expert	Austria	No interests declared
Richard Cejka-Scheidl	Expert	Austria	No interests declared
Alexander Knieps	Expert	Austria	No interests declared
Johanna de Groot	Expert	Netherlands	No interests declared
Johannes Petrus Theodorus Span	Expert	Netherlands	No interests declared
Jacobus Johannes Christianus Maria Romme	Expert	Netherlands	No interests declared
Hinke Johanna Van der Woude	Expert	Netherlands	No interests declared
Frank Holtkamp	Expert	Netherlands	No interests declared
Cristina Migali	Expert	Italy	No interests declared
Sara Galluzzo	Expert	Italy	No interests declared
Cinzia Ciceroni	Expert	Italy	No interests declared
Michela Piezzo	Expert	Italy	No interests declared
Hannah Münch	Expert	Austria	No interests declared
Sabine Mayrhofer	Expert	Germany	No interests declared
Friederike Marei Feldmann	Expert	Germany	No interests declared
Franziska Wolter	Expert	Germany	No interests declared
Irene Bachmann	Expert	Germany	No interests declared
Georgios Aislaitner	Expert	Germany	No interests declared
Marion Haberkamp	Expert	Germany	No interests declared
Christine Greiner	Expert	Germany	No interests declared
Magdalena Kremer	Expert	Germany	No interests declared
Sofia Kapanadze	Expert	Germany	No restrictions applicable to this meeting
Uwe Lipke	Expert	Germany	No interests declared
Ulrike Hermes	Expert	Germany	No interests declared
Lothar Bergmann	Expert	EMA	No restrictions against giving the SAG report for Carvykti
Susanne Brendler-Schwaab	Expert	Germany	No interests declared
Sabine Straus	Expert	Netherlands	No interests declared
Maja Sommerfelt Grønvold	Expert	Norway	No interests declared

Juta Kraav	Expert	Estonia	No interests declared
Katia Pauwels	Expert	Belgium	No interests declared
Miriam Fürst-Wilmes	Expert	Germany	No interests declared

A representative from the European Commission attended the meeting.

Observers from ANVISA (Brazil), MHLW/PMDA (Japan), and FDA (USA) attended the meeting.

Meeting run with the help of EMA staff.

Experts were evaluated against the agenda topics or activities they participated in.

Experts from international organisations or regulatory authorities in third countries cannot participate in the adoption of any procedural decision, scientific opinion or recommendation by the Committee at any step of the procedure.

## 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** (section 5.3)

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 Pharmacovigilance 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/](http://www.ema.europa.eu/)



18 April 2024  
EMA/CHMP/77965/2024

## Annex to 19-22 February 2024 CHMP Minutes

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## A. PRE-SUBMISSION ISSUES

### A.1. ELIGIBILITY REQUESTS

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Report on Eligibility to Centralised Procedure for February 2024: **For adoption** Adopted

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

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Final Outcome of Rapporteurship allocation for February 2024: **For adoption** Adopted

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### 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

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<b>IMVANEX - Smallpox vaccine (live modified vaccinia virus Ankara) - EMEA/H/C/002596/S/0095</b> Bavarian Nordic A/S, Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Gabriele Maurer Request for Supplementary Information adopted on 14.12.2023.	Positive Opinion adopted by consensus together with the CHMP assessment report.  The Marketing Authorisation remains under exceptional circumstances.
<b>Lojuxta - Lomitapide - EMEA/H/C/002578/S/0057</b> Amryt Pharmaceuticals DAC, Rapporteur: Patrick Vrijlandt, Co-Rapporteur: Paolo Gasparini, PRAC Rapporteur: Bianca Mulder Request for Supplementary Information adopted on 14.12.2023.	Positive Opinion adopted by consensus together with the CHMP assessment report.  The Marketing Authorisation remains under exceptional circumstances.
<b>NULIBRY - Fosdenopterin - EMEA/H/C/005378/S/0006, Orphan</b> TMC Pharma (EU) Limited, Rapporteur: Patrick Vrijlandt, PRAC Rapporteur: Martin Huber	Positive Opinion adopted by consensus together with the CHMP assessment report.  The Marketing Authorisation remains under exceptional circumstances.
<b>NYXTHRACIS - Obiltoxaximab - EMEA/H/C/005169/S/0013, Orphan</b> SFL Pharmaceuticals Deutschland GmbH, Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Liana Martirosyan	Positive Opinion adopted by consensus together with the CHMP assessment report.  The Marketing Authorisation remains under exceptional circumstances.

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<p><b>Orphacol - Cholic acid - EMA/H/C/001250/S/0053</b></p> <p>Theravia, Rapporteur: Anastasia Mountaki, PRAC Rapporteur: Sofia Trantza</p>	<p>Positive Opinion adopted by consensus together with the CHMP assessment report.</p> <p>The Marketing Authorisation remains under exceptional circumstances.</p>
<p><b>Raxone - Idebenone - EMA/H/C/003834/S/0035, Orphan</b></p> <p>Chiesi Farmaceutici S.p.A., Rapporteur: John Joseph Borg, PRAC Rapporteur: Amelia Cupelli</p>	<p>Positive Opinion adopted by consensus together with the CHMP assessment report.</p> <p>The Marketing Authorisation remains under exceptional circumstances.</p>
<p><b>Upstaza - Eladocagene exuparvovec - EMA/H/C/005352/S/0017, Orphan, ATMP</b></p> <p>PTC Therapeutics International Limited, Rapporteur: Joseph DeCoursey, PRAC Rapporteur: Gabriele Maurer Request for Supplementary Information adopted on 08.12.2023.</p>	<p>Positive Opinion adopted by consensus together with the CHMP assessment report.</p> <p>The Marketing Authorisation remains under exceptional circumstances.</p>
<p><b>Vedrop - Tocofersolan - EMA/H/C/000920/S/0049</b></p> <p>Recordati Rare Diseases, Rapporteur: Beata Maria Jakline Ullrich, PRAC Rapporteur: Melinda Palfi</p>	<p>Positive Opinion adopted by consensus together with the CHMP assessment report.</p> <p>The Marketing Authorisation remains under exceptional circumstances.</p>

## B.2. RENEWALS OF MARKETING AUTHORISATIONS OUTCOMES

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

### B.2.2. Renewals of Marketing Authorisations for unlimited validity

<p><b>Grasustek - Pegfilgrastim - EMA/H/C/004556/R/0014</b></p> <p>Juta Pharma GmbH, Rapporteur: Karin Janssen van Doorn, Co-Rapporteur: Martina Weise, PRAC Rapporteur: Bianca Mulder Request for Supplementary Information adopted on 14.12.2023.</p>	<p>Positive Opinion adopted by consensus together with the CHMP assessment report and translation timetable.</p> <p>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.</p>
<p><b>Lacosamide UCB - Lacosamide - EMA/H/C/005243/R/0020</b></p> <p>UCB Pharma S.A., Informed Consent of Vimpat, Rapporteur: Filip Josephson, Co-Rapporteur: Paolo Gasparini, PRAC Rapporteur: Ulla Wändel Liminga</p>	<p>Positive Opinion adopted by consensus together with the CHMP assessment report.</p> <p>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.</p>
<p><b>LysaKare - L-lysine hydrochloride / L-arginine hydrochloride - EMA/H/C/004541/R/0016</b></p>	<p>Positive Opinion adopted by consensus together with the CHMP assessment report and translation timetable.</p>

<p>Advanced Accelerator Applications, Rapporteur: Janet Koenig, Co-Rapporteur: Aaron Sosa Mejia, PRAC Rapporteur: Adam Przybylkowski Request for Supplementary Information adopted on 25.01.2024.</p>	<p>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.</p>
<p><b>Palynziq - Pegvaliase - EMEA/H/C/004744/R/0038, Orphan</b> BioMarin International Limited, Rapporteur: Patrick Vrijlandt, Co-Rapporteur: Alexandre Moreau, PRAC Rapporteur: Rhea Fitzgerald Request for Supplementary Information adopted on 09.11.2023.</p>	<p>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.</p>
<p><b>Posaconazole Accord - Posaconazole - EMEA/H/C/005005/R/0014</b> Accord Healthcare S.L.U., Generic, Generic of Noxafil, Rapporteur: Hrefna Gudmundsdottir, PRAC Rapporteur: Nathalie Gault</p>	<p>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.</p>
<p><b>Posaconazole AHCL - Posaconazole - EMEA/H/C/005028/R/0011</b> Accord Healthcare S.L.U., Generic, Generic of Noxafil, Rapporteur: Hrefna Gudmundsdottir, PRAC Rapporteur: Nathalie Gault</p>	<p>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.</p>
<p><b>Talzenna - Talazoparib - EMEA/H/C/004674/R/0017</b> Pfizer Europe MA EEIG, Rapporteur: Filip Josephson, Co-Rapporteur: Hrefna Gudmundsdottir, PRAC Rapporteur: Carla Torre Request for Supplementary Information adopted on 25.01.2024.</p>	<p>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.</p>
<p><b>Ultomiris - Ravulizumab - EMEA/H/C/004954/R/0040</b> Alexion Europe SAS, Rapporteur: Carolina Prieto Fernandez, Co-Rapporteur: Robert Porszasz, PRAC Rapporteur: Kimmo Jaakkola Request for Supplementary Information adopted on 25.01.2024.</p>	<p>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.</p>
<p><b>Zydelig - Idelalisib - EMEA/H/C/003843/R/0059</b> Gilead Sciences Ireland UC, Rapporteur: Filip Josephson, Co-Rapporteur: Peter Mol, PRAC Rapporteur: Martin Huber Request for Supplementary Information adopted on 22.02.2024.</p>	<p>Request for supplementary information adopted with a specific timetable.</p>

### B.2.3. Renewals of Conditional Marketing Authorisations

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**Koselugo - Selumetinib -****EMA/H/C/005244/R/0015, Orphan**AstraZeneca AB, Rapporteur: Alexandre Moreau,  
PRAC Rapporteur: Ulla Wändel Liminga

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.

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**Lunsumio - Mosunetuzumab -****EMA/H/C/005680/R/0008, Orphan**

Roche Registration GmbH, Rapporteur: Aaron Sosa Mejia, PRAC Rapporteur: Ulla Wändel Liminga

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.

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**Pandemic influenza vaccine H5N1****AstraZeneca - Pandemic influenza vaccine (H5N1) (live attenuated, nasal) -****EMA/H/C/003963/R/0071**

AstraZeneca AB, Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Sonja Hrabcik

Positive Opinion adopted by consensus together with the CHMP assessment report and translation timetable.

The CHMP was of the opinion that the renewal for this conditional Marketing Authorisation can be granted.

The Marketing Authorisation remains conditional.

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### 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 2024 meeting:

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**EMA/H/C/PSUSA/00010066/202306**

(avanafil)

CAPS:

**Spedra** (EMA/H/C/002581) (Avanafil), Menarini International Operations Luxembourg S.A., Rapporteur: Maria Concepcion Prieto Yerro, PRAC Rapporteur: Maria del Pilar Rayon, "22/06/2020 To: 21/06/2023"The CHMP, having considered in accordance with Article 28 of Regulation (EC) No 726/2004 the PSUR on the basis of the PRAC recommendation and the PRAC assessment report as appended, recommends by consensus, the variation to the terms of the marketing authorisation(s) for the above mentioned medicinal product(s), concerning the following change(s):  
Update of section 4.4 of the SmPC to add a warning/precaution regarding central serous corioretinopathy. The package leaflet is updated accordingly.

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**EMA/H/C/PSUSA/00010369/202306**

(tedizolid phosphate)

CAPS:

**Sivextro** (EMA/H/C/002846) (Tedizolid)

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,

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phosphate), Merck Sharp & Dohme B.V.,  
Rapporteur: Bruno Sepodes, PRAC Rapporteur:  
Maria del Pilar Rayon, "20/06/2020 To:  
20/06/2023"

recommends by consensus, the variation to the terms of the marketing authorisation for the above mentioned medicinal product, concerning the following change(s):  
Update of sections 4.4 and 4.5 of the SmPC to amend a warning/precaution regarding serotonin syndrome. The package leaflet is updated accordingly.

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**EMA/H/C/PSUSA/00010742/202307**  
(voretigene neparvovec)  
CAPS:  
**Luxturna** (EMA/H/C/004451) (Voretigene neparvovec), Novartis Europharm Limited, Rapporteur: Sol Ruiz, CHMP Coordinator: Maria Concepcion Prieto Yerro, PRAC Rapporteur: Gabriele Maurer, "24/07/2022 To: 23/07/2023"

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 "chorioretinal atrophy" to the list of those related to voretigene neparvovec with a frequency "not known", and additional details on chorioretinal atrophy in the Description of selected adverse reactions. The package leaflet is updated accordingly.

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**EMA/H/C/PSUSA/00010903/202307**  
(brexucabtagene autoleucl)  
CAPS:  
**Tecartus** (EMA/H/C/005102) (Brexucabtagene autoleucl), Kite Pharma EU B.V., Rapporteur: Jan Mueller-Berghaus, CHMP Coordinator: Jan Mueller-Berghaus, PRAC Rapporteur: Bianca Mulder, "24/01/2023 To: 23/07/2023"

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 ICANS with a frequency very common, and to amend the warning/precaution regarding ICANS. In addition, update of Annex II to include ICANS in the key elements of the educational materials.

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#### **B.4. EPARs / WPARs**

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**EXBLIFEP - Cefepime / Enmetazobactam - EMA/H/C/005431**  
Advanz Pharma Limited, treatment of complicated urinary tract infections (cUTI), including pyelonephritis; hospital-acquired pneumonia (HAP), including ventilator associated pneumonia (VAP); patients with bacteraemia that occurs in association with, or

For information only. Comments can be sent to the PL in case necessary.

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is suspected to be associated with, any of the infections listed above, New active substance (Article 8(3) of Directive No 2001/83/EC)

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**Niapelf - Paliperidone - EMEA/H/C/006185** For information only. Comments can be sent to the PL in case necessary.  
Neuraxpharm Pharmaceuticals S.L., Treatment of schizophrenia, Generic, Generic of Xeplion, Generic application (Article 10(1) of Directive No 2001/83/EC)

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**Ryzneuta - Efbemalenograstim alfa - EMEA/H/C/005828** For information only. Comments can be sent to the PL in case necessary.  
Evide Biotechnology Ireland Limited, Reduction in the duration of neutropenia and the incidence of febrile neutropenia., New active substance (Article 8(3) of Directive No 2001/83/EC)

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## **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**

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**Abiraterone Krka - Abiraterone acetate - EMEA/H/C/005649/II/0004** Positive Opinion adopted by consensus on 15.02.2024.  
KRKA, d.d., Novo mesto, Generic, Generic of Zytiga, Rapporteur: Andreja Kranjc  
Opinion adopted on 15.02.2024.  
Request for Supplementary Information adopted on 14.12.2023.

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**Abrysvo - Respiratory syncytial virus vaccine (bivalent, recombinant) - EMEA/H/C/006027/II/0001** Positive Opinion adopted by consensus on 22.02.2024.  
Pfizer Europe Ma EEIG, Rapporteur: Jayne Crowe  
Opinion adopted on 22.02.2024.  
Request for Supplementary Information adopted on 14.12.2023.

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**Accofil - Filgrastim - EMEA/H/C/003956/II/0060/G** Positive Opinion adopted by consensus on 15.02.2024.  
Accord Healthcare S.L.U., Rapporteur: Outi Mäki-Ikola  
Opinion adopted on 15.02.2024.

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**Adtralza - Tralokinumab - EMEA/H/C/005255/II/0015** Request for supplementary information adopted with a specific timetable.  
LEO Pharma A/S, Rapporteur: Jayne Crowe  
Request for Supplementary Information adopted on 01.02.2024.

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<p><b>Apretude - Cabotegravir -</b>  <b>EMA/H/C/005756/II/0002/G</b>  ViiV Healthcare B.V., Duplicate, Duplicate of  Vocabria, Rapporteur: Bruno Sepodes  Opinion adopted on 08.02.2024.</p>	<p>Positive Opinion adopted by consensus on  08.02.2024.</p>
<p><b>Aybintio - Bevacizumab -</b>  <b>EMA/H/C/005106/II/0019/G</b>  Samsung Bioepis NL B.V., Rapporteur: Christian  Gartner  Opinion adopted on 08.02.2024.</p>	<p>Positive Opinion adopted by consensus on  08.02.2024.</p>
<p><b>Bevespi Aerosphere - Glycopyrronium /</b>  <b>Formoterol fumarate dihydrate -</b>  <b>EMA/H/C/004245/II/0019/G</b>  AstraZeneca AB, Rapporteur: Kristina Dunder  Opinion adopted on 22.02.2024.</p>	<p>Positive Opinion adopted by consensus on  22.02.2024.</p>
<p><b>Bortezomib SUN - Bortezomib -</b>  <b>EMA/H/C/004076/II/0022</b>  Sun Pharmaceutical Industries Europe B.V.,  Generic, Generic of VELCADE, Rapporteur:  Margareta Bego  Opinion adopted on 01.02.2024.</p>	<p>Positive Opinion adopted by consensus on  01.02.2024.</p>
<p><b>Cablivi - Caplacizumab -</b>  <b>EMA/H/C/004426/II/0047/G, Orphan</b>  Ablynx NV, Rapporteur: Filip Josephson  Opinion adopted on 08.02.2024.</p>	<p>Positive Opinion adopted by consensus on  08.02.2024.</p>
<p><b>Cerezyme - Imiglucerase -</b>  <b>EMA/H/C/000157/II/0131</b>  Sanofi B.V., Rapporteur: Patrick Vrijlandt  Opinion adopted on 15.02.2024.  Request for Supplementary Information adopted  on 14.12.2023.</p>	<p>Positive Opinion adopted by consensus on  15.02.2024.</p>
<p><b>Cetrotide - Cetrorelix -</b>  <b>EMA/H/C/000233/II/0090</b>  Merck Europe B.V., Rapporteur: Martina Weise  Request for Supplementary Information adopted  on 01.02.2024, 29.06.2023.</p>	<p>Request for supplementary information adopted  with a specific timetable.</p>
<p><b>Circadin - Melatonin -</b>  <b>EMA/H/C/000695/II/0071/G</b>  RAD Neurim Pharmaceuticals EEC SARL,  Rapporteur: Bruno Sepodes  Request for Supplementary Information adopted  on 08.02.2024.</p>	<p>Request for supplementary information adopted  with a specific timetable.</p>
<p><b>Diacomit - Stiripentol -</b>  <b>EMA/H/C/000664/II/0045/G</b>  BIOCODEX, Rapporteur: Alar Irs  Opinion adopted on 15.02.2024.</p>	<p>Positive Opinion adopted by consensus on  15.02.2024.</p>

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Request for Supplementary Information adopted on 23.11.2023.

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**Elfabrio - Pegunigalsidase alfa -  
EMA/H/C/005618/II/0002**

Chiesi Farmaceutici S.p.A., Rapporteur:  
Alexandre Moreau

Request for Supplementary Information adopted on 08.02.2024.

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Request for supplementary information adopted with a specific timetable.

**Elonva - Corifollitropin alfa -  
EMA/H/C/001106/II/0067**

Organon N.V., Rapporteur: Patrick Vrijlandt  
Opinion adopted on 15.02.2024.

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Positive Opinion adopted by consensus on 15.02.2024.

**Empliciti - Elotuzumab -  
EMA/H/C/003967/II/0037/G**

Bristol-Myers Squibb Pharma EEIG, Rapporteur:  
Peter Mol

Request for Supplementary Information adopted on 15.02.2024.

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Request for supplementary information adopted with a specific timetable.

**Empliciti - Elotuzumab -  
EMA/H/C/003967/II/0038/G**

Bristol-Myers Squibb Pharma EEIG, Rapporteur:  
Peter Mol  
Opinion adopted on 15.02.2024.

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Positive Opinion adopted by consensus on 15.02.2024.

**Enhertu - Trastuzumab -  
EMA/H/C/005124/II/0043/G**

Daiichi Sankyo Europe GmbH, Rapporteur:  
Aaron Sosa Mejia

Request for Supplementary Information adopted on 15.02.2024.

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Request for supplementary information adopted with a specific timetable.

**Eptifibatide Accord - Eptifibatide -  
EMA/H/C/004104/II/0015/G**

Accord Healthcare S.L.U., Generic, Generic of  
Integrilin, Rapporteur: Jayne Crowe

Request for Supplementary Information adopted on 08.02.2024, 26.10.2023, 12.05.2023.

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Request for supplementary information adopted with a specific timetable.

**Fasturtec - Rasburicase -  
EMA/H/C/000331/II/0069**

Sanofi Winthrop Industrie, Rapporteur: Peter  
Mol

Opinion adopted on 22.02.2024.

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Positive Opinion adopted by consensus on 22.02.2024.

**Flucelvax Tetra - Influenza vaccine  
(surface antigen, inactivated, prepared in  
cell cultures) -  
EMA/H/C/004814/II/0044**

Seqirus Netherlands B.V., Rapporteur: Sol Ruiz  
Opinion adopted on 22.02.2024.

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Positive Opinion adopted by consensus on 22.02.2024.

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Request for Supplementary Information adopted on 18.01.2024.

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**Fulphila - Pegfilgrastim -  
EMA/H/C/004915/II/0046**

Biosimilar Collaborations Ireland Limited,  
Rapporteur: Martina Weise  
Opinion adopted on 08.02.2024.

Positive Opinion adopted by consensus on 08.02.2024.

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**Hetlioz - Tasimelteon -  
EMA/H/C/003870/II/0037, Orphan**

Vanda Pharmaceuticals Netherlands B.V.,  
Rapporteur: Jayne Crowe  
Request for Supplementary Information adopted on 08.02.2024.

Request for supplementary information adopted with a specific timetable.

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**Iasibon - Ibandronic acid -  
EMA/H/C/002025/II/0025**

Pharmathen S.A., Generic, Generic of Bondronat, Rapporteur: Thalia Marie Estrup Blicher  
Opinion adopted on 01.02.2024.  
Request for Supplementary Information adopted on 21.09.2023.

Positive Opinion adopted by consensus on 01.02.2024.

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**Insuman - Insulin human -  
EMA/H/C/000201/II/0146**

Sanofi-Aventis Deutschland GmbH, Rapporteur: Karin Janssen van Doorn  
Request for Supplementary Information adopted on 15.02.2024.

Request for supplementary information adopted with a specific timetable.

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**Ixiaro - Japanese encephalitis vaccine (inactivated, adsorbed) -  
EMA/H/C/000963/II/0116**

Valneva Austria GmbH, Rapporteur: Jan Mueller-Berghaus  
Opinion adopted on 08.02.2024.

Positive Opinion adopted by consensus on 08.02.2024.

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**Keytruda - Pembrolizumab -  
EMA/H/C/003820/II/0144**

Merck Sharp & Dohme B.V., Rapporteur: Paolo Gasparini  
Opinion adopted on 15.02.2024.  
Request for Supplementary Information adopted on 14.12.2023.

Positive Opinion adopted by consensus on 15.02.2024.

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**Latuda - Lurasidone -  
EMA/H/C/002713/II/0041**

Aziende Chimiche Riunite Angelini Francesco A.C.R.A.F. S.p.A., Rapporteur: Filip Josephson  
Request for Supplementary Information adopted on 08.02.2024.

Request for supplementary information adopted with a specific timetable.

<p><b>Nordimet - Methotrexate -</b>  <b>EMA/H/C/003983/II/0033/G</b>  Nordic Group B.V., Rapporteur: Bruno Sepodes  Opinion adopted on 08.02.2024.</p>	<p>Positive Opinion adopted by consensus on  08.02.2024.</p>
<p><b>Nplate - Romiplostim -</b>  <b>EMA/H/C/000942/II/0090</b>  Amgen Europe B.V., Rapporteur: Maria  Concepcion Prieto Yerro  Opinion adopted on 08.02.2024.</p>	<p>Positive Opinion adopted by consensus on  08.02.2024.</p>
<p><b>Ontruzant - Trastuzumab -</b>  <b>EMA/H/C/004323/II/0049</b>  Samsung Bioepis NL B.V., Rapporteur: Karin  Janssen van Doorn  Opinion adopted on 08.02.2024.</p>	<p>Positive Opinion adopted by consensus on  08.02.2024.</p>
<p><b>Palynziq - Pegvaliase -</b>  <b>EMA/H/C/004744/II/0039/G, Orphan</b>  BioMarin International Limited, Rapporteur:  Patrick Vrijlandt  Opinion adopted on 08.02.2024.  Request for Supplementary Information adopted  on 09.11.2023.</p>	<p>Positive Opinion adopted by consensus on  08.02.2024.</p>
<p><b>Pedmarqsi - Sodium thiosulfate -</b>  <b>EMA/H/C/005130/II/0002/G</b>  Fennec Pharmaceuticals (EU) Limited,  Rapporteur: Elita Poplavska  Request for Supplementary Information adopted  on 01.02.2024.</p>	<p>Request for supplementary information adopted  with a specific timetable.</p>
<p><b>Pemetrexed Accord - Pemetrexed -</b>  <b>EMA/H/C/004072/II/0028</b>  Accord Healthcare S.L.U., Generic, Generic of  Alimta, Rapporteur: John Joseph Borg  Request for Supplementary Information adopted  on 15.02.2024.</p>	<p>Request for supplementary information adopted  with a specific timetable.</p>
<p><b>Qarziba - Dinutuximab beta -</b>  <b>EMA/H/C/003918/II/0056/G, Orphan</b>  Recordati Netherlands B.V., Rapporteur: Peter  Mol  Request for Supplementary Information adopted  on 15.02.2024.</p>	<p>Request for supplementary information adopted  with a specific timetable.</p>
<p><b>Reblozyl - Luspatercept -</b>  <b>EMA/H/C/004444/II/0027, Orphan</b>  Bristol-Myers Squibb Pharma EEIG, Rapporteur:  Daniela Philadelphia  Request for Supplementary Information adopted  on 15.02.2024.</p>	<p>Request for supplementary information adopted  with a specific timetable.</p>
<p><b>Ryzodeg - Insulin aspart / Insulin degludec</b></p>	<p>Positive Opinion adopted by consensus on</p>

<p><b>- EMEA/H/C/002499/II/0054</b> Novo Nordisk A/S, Rapporteur: Kristina Dunder Opinion adopted on 15.02.2024. Request for Supplementary Information adopted on 14.12.2023.</p>	15.02.2024.
<p><b>Skyrizi - Risankizumab - EMEA/H/C/004759/II/0046/G</b> AbbVie Deutschland GmbH &amp; Co. KG, Rapporteur: Finbarr Leacy Request for Supplementary Information adopted on 08.02.2024.</p>	Request for supplementary information adopted with a specific timetable.
<p><b>Skytrofa - Lonapegsomatropin - EMEA/H/C/005367/II/0024, Orphan</b> Ascendis Pharma Endocrinology Division A/S, Rapporteur: Patrick Vrijlandt Request for Supplementary Information adopted on 01.02.2024.</p>	Request for supplementary information adopted with a specific timetable.
<p><b>Skytrofa - Lonapegsomatropin - EMEA/H/C/005367/II/0025/G, Orphan</b> Ascendis Pharma Endocrinology Division A/S, Rapporteur: Patrick Vrijlandt Request for Supplementary Information adopted on 08.02.2024.</p>	Request for supplementary information adopted with a specific timetable.
<p><b>Spectrila - Asparaginase - EMEA/H/C/002661/II/0036</b> medac Gesellschaft für klinische Spezialpräparate mbH, Rapporteur: Christian Gartner Request for Supplementary Information adopted on 22.02.2024.</p>	Request for supplementary information adopted with a specific timetable.
<p><b>Sugammadex Mylan - Sugammadex - EMEA/H/C/005403/II/0010/G</b> Mylan Ireland Limited, Generic, Generic of Bridion, Rapporteur: Hrefna Gudmundsdottir Request for Supplementary Information adopted on 22.02.2024.</p>	Request for supplementary information adopted with a specific timetable.
<p><b>Suliqua - Insulin glargine / Lixisenatide - EMEA/H/C/004243/II/0037/G</b> Sanofi Winthrop Industrie, Rapporteur: Kristina Dunder Opinion adopted on 15.02.2024. Request for Supplementary Information adopted on 14.12.2023.</p>	Positive Opinion adopted by consensus on 15.02.2024.
<p><b>Supemtek - Influenza quadrivalent vaccine (rDNA) - EMEA/H/C/005159/II/0015/G</b> Sanofi Pasteur, Rapporteur: Jan Mueller-Berghaus</p>	Request for supplementary information adopted with a specific timetable.

Request for Supplementary Information adopted on 08.02.2024.	
<b>TEPMETKO - Tepotinib - EMA/H/C/005524/II/0012</b> Merck Europe B.V., Rapporteur: Filip Josephson Opinion adopted on 15.02.2024. Request for Supplementary Information adopted on 18.01.2024.	Positive Opinion adopted by consensus on 15.02.2024.
<b>Tresiba - Insulin degludec - EMA/H/C/002498/II/0060</b> Novo Nordisk A/S, Rapporteur: Kristina Dunder Opinion adopted on 15.02.2024. Request for Supplementary Information adopted on 14.12.2023.	Positive Opinion adopted by consensus on 15.02.2024.
<b>Trumenba - Meningococcal group B vaccine (recombinant, adsorbed) - EMA/H/C/004051/II/0050/G</b> Pfizer Europe MA EEIG, Rapporteur: Patrick Vrijlandt Request for Supplementary Information adopted on 22.02.2024, 14.12.2023.	Request for supplementary information adopted with a specific timetable.
<b>Zebinix - Eslicarbazepine acetate - EMA/H/C/000988/II/0089/G</b> Bial - Portela & C <sup>a</sup> , S.A., Rapporteur: Martina Weise Request for Supplementary Information adopted on 15.02.2024.	Request for supplementary information adopted with a specific timetable.
<b>Zirabev - Bevacizumab - EMA/H/C/004697/II/0032</b> Pfizer Europe MA EEIG, Rapporteur: Eva Skovlund Request for Supplementary Information adopted on 01.02.2024.	Request for supplementary information adopted with a specific timetable.
<b>Zoonotic Influenza Vaccine Seqirus - Zoonotic influenza vaccine (H5N1) (surface antigen, inactivated, adjuvanted) - EMA/H/C/006375/II/0001</b> Seqirus S.r.l., Informed Consent of Aflunov, Rapporteur: Maria Grazia Evandri Request for Supplementary Information adopted on 22.02.2024, 25.01.2024.	Request for supplementary information adopted with a specific timetable.
<b>WS2598/G Ambirix- EMA/H/C/000426/WS2598/0131/G Fendrix- EMA/H/C/000550/WS2598/0084/G Infanrix hexa-</b>	Positive Opinion adopted by consensus on 15.02.2024.

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**EMA/H/C/000296/WS2598/0338/G**

**Twinrix Adult-**

**EMA/H/C/000112/WS2598/0166/G**

**Twinrix Paediatric-**

**EMA/H/C/000129/WS2598/0167/G**

GlaxoSmithkline Biologicals SA, Lead

Rapporteur: Christophe Focke

Opinion adopted on 15.02.2024.

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**WS2600**

**Infanrix hexa-**

**EMA/H/C/000296/WS2600/0339**

GlaxoSmithkline Biologicals SA, Lead

Rapporteur: Christophe Focke

Opinion adopted on 15.02.2024.

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Positive Opinion adopted by consensus on 15.02.2024.

**WS2608/G**

**Apretude-**

**EMA/H/C/005756/WS2608/0001/G**

**Vocabria-**

**EMA/H/C/004976/WS2608/0020/G**

ViiV Healthcare B.V., Duplicate, Duplicate of

Vocabria, Lead Rapporteur: Bruno Sepodes

Request for Supplementary Information adopted on 08.02.2024.

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Request for supplementary information adopted with a specific timetable.

**WS2616/G**

**Hexacima-**

**EMA/H/C/002702/WS2616/0153/G**

**Hexyon-**

**EMA/H/C/002796/WS2616/0157/G**

Sanofi Pasteur, Lead Rapporteur: Jan Mueller-

Berghaus

Opinion adopted on 15.02.2024.

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Positive Opinion adopted by consensus on 15.02.2024.

**WS2625**

**Hukyndra-**

**EMA/H/C/005548/WS2625/0020**

**Libmyris-**

**EMA/H/C/005947/WS2625/0009**

STADA Arzneimittel AG, Lead Rapporteur: Outi

Mäki-Ikola

Opinion adopted on 08.02.2024.

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Positive Opinion adopted by consensus on 08.02.2024.

**Mosquirix-**

**EMA/H/W/002300/WS2585/0078**

**Shingrix-**

**EMA/H/C/004336/WS2585/0071**

GlaxoSmithkline Biologicals SA, Lead

Rapporteur: Jan Mueller-Berghaus

Opinion adopted on 01.02.2024.

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Positive Opinion adopted by consensus on 01.02.2024.

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

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**BIMERVAX - SARS-CoV-2, variant XBB.1.16, spike protein, receptor binding domain fusion homodimer / Selvacovatein - EMEA/H/C/006058/II/0004** Positive Opinion adopted by consensus on 22.02.2024.

Hipra Human Health S.L., Rapporteur: Beata Maria Jakline Ullrich, "Update of section 4.2 of the SmPC to introduce a homologous booster of Bimervax (PHH-1V) following a previous booster dose of PHH-1V based on interim results from clinical study HIPRA HH-2. Update of sections 4.4, 4.8 and 5.1 of the SmPC to add safety and immunogenicity information after a fourth dose based on interim results from study HIPRA-HH-2. Study HIPRA-HH2 is a Phase IIB, double-blind, randomised, active -controlled, multicentre, non-inferiority trial followed by a Phase III, single-arm, open-label trial to assess immunogenicity and safety of a booster vaccination with a PHH-1V against SARS-COV-2 in adults fully vaccinated against Covid-19 followed by an extension period to study a fourth dose administration of PHH-1V. The study is listed as a category 3 study in the RMP. The Package Leaflet is updated accordingly. In addition, the MAH submitted the full user consultation with target patient groups." Opinion adopted on 22.02.2024. Request for Supplementary Information adopted on 14.12.2023, 14.09.2023.

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**BLINCYTO - Blinatumomab - EMEA/H/C/003731/II/0053/G, Orphan** Positive Opinion adopted by consensus on 22.02.2024.

Amgen Europe B.V., Rapporteur: Alexandre Moreau, "A grouped application consisting of: Type II (C.I.4): Update of section 6.6 of the SmPC in order to add a statement that the administration of Blincyto for BSA of less than 0.4 m<sup>2</sup> has not been established. In addition, the MAH took the opportunity to update the name of ATC pharmacological subgroup according to WHO ATC Index and to delete "intravenous catheter" from the important note statement regarding flushing and to introduce minor editorial changes to the PI. The Package Leaflet is updated accordingly. Type IB (C.I.11.z): Update of the due dates for post-authorisation safety studies 20150136 and 20180130 in the Annex II D in order to align with the RMP version 16.0, following

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commitment agreed on during procedure  
EMA/H/C/003731/IB/0050.”  
Opinion adopted on 22.02.2024.  
Request for Supplementary Information adopted  
on 14.09.2023.

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**Brilique - Ticagrelor -  
EMA/H/C/001241/II/0061**

Positive Opinion adopted by consensus on  
22.02.2024.

AstraZeneca AB, Rapporteur: Patrick Vrijlandt,  
“Update of sections 4.2 and 4.4 of the SmPC in  
order to include a warning related to Single  
Antiplatelet Therapy (SAPT) in Patients with  
Acute Coronary Syndrome (ACS) who have  
undergone a Percutaneous Coronary  
Intervention (PCI) procedure and who have an  
increased risk of bleeding based on literature.”  
Opinion adopted on 22.02.2024.  
Request for Supplementary Information adopted  
on 14.09.2023.

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**BYANLI - Paliperidone -  
EMA/H/C/005486/II/0005**

Positive Opinion adopted by consensus on  
01.02.2024.

Janssen-Cilag International N.V., Informed  
Consent of Xeplion, Rapporteur: Kristina  
Dunder, “Submission of the Environmental Risk  
Assessment Report and environmental risk  
studies (OECD 232, OECD 307 and OECD 308).”  
Opinion adopted on 01.02.2024.

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**Darzalex - Daratumumab -  
EMA/H/C/004077/II/0070, Orphan**

Positive Opinion adopted by consensus on  
15.02.2024.

Janssen-Cilag International N.V., Rapporteur:  
Aaron Sosa Mejia, “Update of section 5.1 of the  
SmPC in order to update efficacy information  
based on the final overall survival analysis  
results from study MMY3007. In addition, the  
marketing authorisation holder (MAH) took the  
opportunity to introduce minor editorial  
changes.”  
Opinion adopted on 15.02.2024.

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**Evrysdi - Risdiplam -  
EMA/H/C/005145/II/0021**

Positive Opinion adopted by consensus on  
01.02.2024.

Roche Registration GmbH, Rapporteur: Bruno  
Sepodes, “Update of sections 4.8, 5.1 and 5.2 of  
the SmPC in order to update efficacy and safety  
information based on primary analysis results  
from study BN40703 (RAINBOWFISH); this is an  
open-label, single-arm, multicenter clinical  
study to investigate the efficacy, safety,  
pharmacokinetics, and pharmacodynamics of  
risdiplam in patients aged from birth to 6 weeks

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(at first dose) who are genetically diagnosed with SMA (SMN1 deletion and any SMN2 copies) but not yet presenting with symptoms. In addition, the MAH took the opportunity to introduce minor editorial changes to the PI and to update the Instructions for Use.”  
Opinion adopted on 01.02.2024.

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**Jentaduetto - Linagliptin / Metformin hydrochloride - EMEA/H/C/002279/II/0070**

Boehringer Ingelheim International GmbH, Rapporteur: Patrick Vrijlandt, “Update of sections 4.2, 4.8, 5.1 and 5.2 of the SmPC in order to update information on paediatric population based on final results from study DINAMO 1218-0091; this is a double-blind, randomised, placebo-controlled, parallel group trial to evaluate the efficacy and safety of empagliflozin and linagliptin over 26 weeks, with a double-blind active treatment safety extension period up to 52 weeks, in children and adolescents with type 2 diabetes mellitus. The Package Leaflet and Labelling are updated accordingly. In addition, the MAH took the opportunity to update the list of local representatives in the Package Leaflet.”

Positive Opinion adopted by consensus on 22.02.2024.

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**Keytruda - Pembrolizumab - EMEA/H/C/003820/II/0147**

Merck Sharp & Dohme B.V., Rapporteur: Paolo Gasparini, “Update of section 5.1 of the SmPC in order to update efficacy information based on final results from study KEYNOTE-B61; this is a Phase 2, Single-arm, Open-label Clinical Trial of Pembrolizumab Plus Lenvatinib in Participants with First-line Advanced/Metastatic Non-clear Cell Renal Cell Carcinoma (nccRCC).”  
Request for Supplementary Information adopted on 01.02.2024.

Request for supplementary information adopted with a specific timetable.

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**Kineret - Anakinra - EMEA/H/C/000363/II/0092**

Swedish Orphan Biovitrum AB (publ), Rapporteur: Thalia Marie Estrup Blicher, “Update of section 4.8 of the SmPC in order to add ‘Injection site amyloid deposits’ to the list of adverse drug reactions (ADRs) with frequency not known, based on a review of the clinical study and post-marketing data to evaluate a possible causal association between anakinra (Kineret) and amyloidosis. The Package Leaflet

Positive Opinion adopted by consensus on 08.02.2024.

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is updated accordingly.”  
Opinion adopted on 08.02.2024.

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**Kisplyx - Lenvatinib -  
EMA/H/C/004224/II/0058**

Eisai GmbH, Rapporteur: Karin Janssen van Doorn, “Update of section 5.1 of the SmPC in order to update efficacy information based on final results from study KEYNOTE-B61; this is a phase 2, single-arm, open-label clinical trial of pembrolizumab plus lenvatinib in participants with first-line advanced/metastatic non-clear cell Renal Cell Carcinoma (nccRCC). In addition, the MAH took the opportunity to introduce minor editorial changes to the PI.”

Request for Supplementary Information adopted on 01.02.2024.

Request for supplementary information adopted with a specific timetable.

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**Kisqali - Ribociclib -  
EMA/H/C/004213/II/0041/G**

Novartis Europharm Limited, Rapporteur: Filip Josephson, “Grouped application comprising two type II variations as follows:

- Update of section 5.2 of the SmPC in order to update absorption information based on final results from study CLEE011A2117, a Phase I, single center, two-period, two-treatment, open label, randomized crossover study to investigate the absolute bioavailability of a single oral dose of 600 mg of ribociclib relative to an intravenous (i.v.) infusion of 150 mg ribociclib in healthy subjects.

- Update of section 4.5 of the SmPC in order to update the drug-drug interaction information on substances that may increase ribociclib plasma concentrations based on the updated PBPK modelling.

In addition, the MAH took this opportunity to introduce minor editorial changes to the Package Leaflet.”

Opinion adopted on 22.02.2024.

Request for Supplementary Information adopted on 25.01.2024, 12.10.2023, 22.06.2023.

Positive Opinion adopted by consensus on 22.02.2024.

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**Mavenclad - Cladribine -  
EMA/H/C/004230/II/0032**

Merck Europe B.V., Rapporteur: Thalia Marie Estrup Blicher, “Update of section 4.4 of the SmPC in order to update an existing warning on infections. The Package Leaflet is updated accordingly. In addition, the MAH took the opportunity to introduce an editorial update to

Request for supplementary information adopted with a specific timetable.

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the PI.”

Request for Supplementary Information adopted on 08.02.2024.

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**Nexviadyme - Avalglucosidase alfa - EMEA/H/C/005501/II/0015**

Sanofi B.V., Rapporteur: Christian Gartner, “Update of sections 4.8, 5.1 and 5.2 of the SmPC in order to update safety and efficacy information based on final results from study EFC14028 - COMparative Enzyme replacement Trial with neoGAA versus rhGAA (COMET), listed as a category 3 study in the RMP. This is a phase 3 randomized, multicenter, multinational, double-blinded study comparing the efficacy and safety of repeated biweekly infusions of avalglucosidase alfa (neoGAA, GZ402666) and alglucosidase alfa in treatment naive patients with late onset Pompe disease. In addition, the MAH took this opportunity to update the list of local representatives in the Package Leaflet.” Request for Supplementary Information adopted on 08.02.2024.

Request for supplementary information adopted with a specific timetable.

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**Olumiant - Baricitinib - EMEA/H/C/004085/II/0046**

Eli Lilly Nederland B.V., Rapporteur: Peter Mol, “Update of section 5.1 of the SmPC in order to add information on JIA-associated uveitis or chronic anterior antibody positive uveitis based on interim results from study I4VMC-JAHW; this is an open-label, active-controlled, safety, and efficacy study of oral baricitinib in patients from 2 years to less than 18 years old with active juvenile idiopathic arthritis-associated uveitis or chronic anterior antinuclear antibody-positive uveitis.” Request for Supplementary Information adopted on 08.02.2024.

Request for supplementary information adopted with a specific timetable.

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**Oxlumo - Lumasiran - EMEA/H/C/005040/II/0017, Orphan**

Alnylam Netherlands B.V., Rapporteur: Martina Weise, “Submission of the final report from study ALN-GO1-002 (Study 002), listed as a category 3 study in the RMP. This is a phase 2, multicenter, open-label, extension study to evaluate the long-term administration of ALN-GO1 in patients with primary hyperoxaluria type 1.” Opinion adopted on 08.02.2024.

Positive Opinion adopted by consensus on 08.02.2024.

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**OZAWADE - Pitolisant -  
EMA/H/C/005117/II/0007**

Bioprojet Pharma, Rapporteur: Peter Mol,  
"Submission of the final report from study P21-03. This is an open label, single center, drug-drug interaction study to evaluate the effect of a combination of itraconazole and paroxetine treatment on the pitolisant pharmacokinetics at steady-state in eighteen healthy male Caucasian subjects."

Request for Supplementary Information adopted on 01.02.2024.

Request for supplementary information adopted with a specific timetable.

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**PONVORY - Ponesimod -  
EMA/H/C/005163/II/0014**

Janssen-Cilag International N.V., Rapporteur: Peter Mol, "Update of section 4.5 of the SmPC to amend an existing interaction wording for carbamazepine under the sub-heading "Effect of other medicinal products on ponesimod" based on study 67896153MSC1001. This is a Phase 1, Open-label, Parallel-group Study to Assess the Effect of Steady-state Carbamazepine on the Pharmacokinetics of Ponesimod in Healthy Adult Participants. In addition, the MAH took the opportunity to update the contact details of local representatives in the Package Leaflet."

Opinion adopted on 08.02.2024.

Positive Opinion adopted by consensus on 08.02.2024.

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**QUVIVIQ - Daridorexant -  
EMA/H/C/005634/II/0013/G**

Idorsia Pharmaceuticals Deutschland GmbH, Rapporteur: Alexandre Moreau, "Update of sections 4.4, 4.5 and 5.1 of the SmPC in order to reflect the conclusions of studies ID-075-121, ID-078-122 and ID-078-118, respectively. The Package Leaflet was updated accordingly. Study ID-078-121 is a randomized, double-blind, placebo-controlled, 2-way crossover study to investigate the effects of daridorexant on nighttime respiratory function and sleep in subjects with severe obstructive sleep apnea; Study ID-078-122 is a prospective, open-label, single-dose Phase 1 study to measure daridorexant in breast milk of healthy lactating women; and Study ID-078-118 is a single-center, randomized, double-blind, single-dose, 3-way crossover study to compare the effects of daridorexant and placebo on postural stability, the auditory awakening threshold, and cognitive function in the middle of the night following

Request for supplementary information adopted with a specific timetable.

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evening administration to healthy adult and elderly subjects.”  
Request for Supplementary Information adopted on 01.02.2024.

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**Reagila - Cariprazine -  
EMA/H/C/002770/II/0034**

Gedeon Richter Plc., Rapporteur: Kristina Dunder, PRAC Rapporteur: Ana Sofia Diniz Martins, “Update of sections 4.3 and 4.5 of the SmPC in order to update an existing contraindication and update drug-drug interaction information with CYP3A4 inhibitors, based on final results from study RGH-188-301 (CYPRESS) listed as a category 3 study in the RMP; this is an open-label, single-arm, fixed-sequence study to investigate the effect of erythromycin, a moderate CYP3A4 inhibitor on the pharmacokinetics of cariprazine in male patients with schizophrenia. The Package Leaflet is updated accordingly. The RMP version 4.0 has also been submitted. In addition, the MAH took the opportunity to introduce minor editorial changes to the PI.”

Request for Supplementary Information adopted on 08.02.2024, 26.10.2023.

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Request for supplementary information adopted with a specific timetable.

**Spinraza - Nusinersen -  
EMA/H/C/004312/II/0032, Orphan**

Biogen Netherlands B.V., Rapporteur: Bruno Sepodes, “Update of section 4.8 of the SmPC in order to add ‘Arachnoiditis’ to the list of adverse drug reactions (ADRs) with frequency not known, based on post-marketing review. The Package Leaflet is updated accordingly.”

Request for Supplementary Information adopted on 08.02.2024.

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Request for supplementary information adopted with a specific timetable.

**TAGRISSO - Osimertinib -  
EMA/H/C/004124/II/0054**

AstraZeneca AB, Rapporteur: Carolina Prieto Fernandez, “Update of section 4.8 of the SmPC to add ‘Skin Hyperpigmentation’ to the list of adverse drug reactions (ADRs) with frequency ‘uncommon’ based on literature. The Package Leaflet has been updated accordingly. In addition, the MAH took the opportunity to update the list of local representatives in the Package Leaflet.”

Opinion adopted on 08.02.2024.

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Positive Opinion adopted by consensus on 08.02.2024.

**TEPMETKO - Tepotinib -**

Request for supplementary information adopted

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**EMA/H/C/005524/II/0011**

Merck Europe B.V., Rapporteur: Filip Josephson, "Update of section 4.2 of the SmPC in order to add alternative methods of administration dispersed in water, as oral drinking suspension or via feeding tubes based on the available physicochemical and clinical pharmacology data. The Package Leaflet is updated accordingly." Request for Supplementary Information adopted on 22.02.2024.

with a specific timetable.

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**Translarna - Ataluren -****EMA/H/C/002720/II/0074, Orphan**

PTC Therapeutics International Limited, Rapporteur: Peter Mol, "The CHMP is of the opinion that the requested update of sections 4.2, 4.8, 5.1 and 5.2 of the SmPC in order to change posology information in paediatric population, to update the summary of safety profile and to update pharmacokinetic information on paediatric population based on the final results from study PTC124-GD-048-DMD "A Phase 2, multiple-dose, open-label study evaluating the safety and PK of ataluren in patients with nmDMD aged  $\geq 6$  months to  $< 2$  years old" (as well as the subsequent updates to the Package Leaflet and the additional editorial changes to the PI) are supported by the data submitted. However, in view of the CHMP opinion for the annual renewal EMA/H/C/002720/R/071 adopted on 24 January 2024 to not recommend the renewal of the marketing authorisation of Translarna, as a favourable benefit-risk balance was not confirmed in the treatment of ambulant patients with nmDMD aged 2 years or older, no changes to the marketing authorisation can be recommended at this stage." Opinion adopted on 22.02.2024. Request for Supplementary Information adopted on 14.12.2023.

Positive Opinion adopted by consensus on 22.02.2024.

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**Venclyxto - Venetoclax -****EMA/H/C/004106/II/0047**

AbbVie Deutschland GmbH & Co. KG, Rapporteur: Filip Josephson, "Submission of the final report from study GO28667 (MURANO) listed as a category 3 study in the RMP. This is a Multicenter, Phase III, Open-Label, Randomized Study in Relapsed/Refractory Patients with Chronic Lymphocytic Leukemia to Evaluate the

Request for supplementary information adopted with a specific timetable.

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Benefit of GDC-0199 (ABT-199) Plus Rituximab Compared with Bendamustine Plus Rituximab.” Request for Supplementary Information adopted on 08.02.2024.

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**VidPrevtyn Beta - SARS-CoV-2, B.1.351 variant, prefusion Spike delta TM protein, recombinant - EMEA/H/C/005754/II/0007/G**

Positive Opinion adopted by consensus on 15.02.2024.

Sanofi Pasteur, Rapporteur: Jan Mueller-Berghaus, “A grouped application consisting of: Type II (C.I.4): Update of section 4.8 of the SmPC in order to include additional safety data based on safety update reports from studies VAT00008 booster extension and VAT00002 Cohort 2, in order to fulfill REC 20. Type IA (A.6): To change the ATC Code of the COVID-19 protein subunit vaccine from J07BX03 to J07BN04.” Opinion adopted on 15.02.2024. Request for Supplementary Information adopted on 14.12.2023.

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**Vocabria - Cabotegravir - EMEA/H/C/004976/II/0019**

Positive Opinion adopted by consensus on 08.02.2024.

ViiV Healthcare B.V., Rapporteur: Jean-Michel Race, “Update of section 4.2 of the SmPC in order to update administration instructions to mitigate product leakage related to the correct use of the vial adapter, based on Human Factor studies. The Package Leaflet (Instructions for Use) is updated accordingly.” Opinion adopted on 08.02.2024.

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**Xevudy - Sotrovimab - EMEA/H/C/005676/II/0024**

Positive Opinion adopted by consensus on 15.02.2024.

Glaxosmithkline Trading Services Limited, Rapporteur: Thalia Marie Estrup Blicher, “Update of section 5.1 of the SmPC in order to include virology information based on data from various pharmacology studies on the in vitro activity of sotrovimab in a pseudotyped virus assay against the SARS-CoV-2 Omicron variants named XBB.1.16 and XBB.2.3, XBB.1.16.1, XBB.1.5.10 as well as data on the in vitro activity of sotrovimab in an authentic virus assay against the SARS-CoV-2 XBB.1.16, BA.2.75, BA.4.6 and BQ.1.1 variants. Based on the data reported in PC-23-0139, under this variation application, in addition to the proposed SmPC updates, the MAH also proposed a change to the current methodology

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for assessment of the in vitro neutralization potency of sotrovimab against SARS-CoV-2 variants (change in target cells used for the authentic virus neutralization assay, from the currently used Vero-TMPRSS2 cells, back to the previously used VeroE6 cells)."

Opinion adopted on 15.02.2024.

Request for Supplementary Information adopted on 11.01.2024.

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**Zeffix - Lamivudine -  
EMA/H/C/000242/II/0087**

Positive Opinion adopted by consensus on 15.02.2024.

GlaxoSmithKline (Ireland) Limited, Duplicate, Duplicate of Epivir, Rapporteur: Jean-Michel Race, "Update of section 4.4 of the SmPC in order to amend an existing warning on HIV co-infection. The Package Leaflet is updated accordingly. In addition, the MAH took the opportunity to introduce editorial changes to the PI.

The requested variation proposed amendments to the Summary of Product Characteristics and Package Leaflet."

Opinion adopted on 15.02.2024.

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**WS2544  
Ebymect-  
EMA/H/C/004162/WS2544/0064  
Komboglyze-  
EMA/H/C/002059/WS2544/0057  
Xigduo-EMA/H/C/002672/WS2544/0074**

Positive Opinion adopted by consensus on 08.02.2024.

AstraZeneca AB, Lead Rapporteur: Kristina Dunder, "Update of sections 4.4 and 4.8 of the SmPC in order to add a new warning on 'Vitamin B12 decrease/deficiency' and to change the frequency of 'Vitamin B12 decrease/deficiency' in the list of adverse drug reactions (ADRs) from frequency 'very rare' to 'common'. The Package Leaflet is updated accordingly. In addition, the MAH took the opportunity to introduce minor editorial changes to the PI and to update the contact details of the local representative in the Netherlands in the Package Leaflet."

Opinion adopted on 08.02.2024.

Request for Supplementary Information adopted on 21.09.2023.

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**WS2583  
Stayveer-  
EMA/H/C/002644/WS2583/0040  
Tracleer-  
EMA/H/C/000401/WS2583/0105**

Positive Opinion adopted by consensus on 08.02.2024.

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Janssen-Cilag International N.V., Lead Rapporteur: Alexandre Moreau, "Update of section 4.6 of the SmPC to update the wording concerning breast feeding based on literature and post-marketing data. The Package Leaflet is updated accordingly. In addition, the MAH took the opportunity to update the list of local representatives in the Package Leaflet." Opinion adopted on 08.02.2024.

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**WS2597**  
**OPDIVO-**  
**EMA/H/C/003985/WS2597/0138**  
**Yervoy-EMA/H/C/002213/WS2597/0107**  
Bristol-Myers Squibb Pharma EEIG, Lead Rapporteur: Carolina Prieto Fernandez, "Update of sections 4.4 and 4.8 of the SmPC in order to add 'myelitis' as a warning under the subsection "Other immune-mediated adverse reactions" and to the list of adverse drug reactions (ADRs) with their calculated frequencies for monotherapy (not known) and in combination (rare), based on post-marketing data and literature; the Package Leaflet is updated accordingly. In addition, the MAH took the opportunity to implement editorial changes to the SmPC in line with the QRD." Opinion adopted on 15.02.2024.  
Request for Supplementary Information adopted on 11.01.2024.

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Positive Opinion adopted by consensus on 15.02.2024.

### **B.5.3. CHMP-PRAC assessed procedures**

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**Beyfortus - Nirsevimab -**  
**EMA/H/C/005304/II/0018/G**  
Sanofi Winthrop Industrie, Rapporteur: Thalia Marie Estrup Blicher, PRAC Rapporteur: Kimmo Jaakkola, "Grouped application comprising two type II variations as follows:  
C.I.13: Submission of the final report from study D5290C00004 (MELODY) listed as a category 3 study in the RMP. This is a phase III study, randomized, double-blind, placebo-controlled study to evaluate the safety and efficacy of MEDI8897, a monoclonal antibody with an extended half-life against respiratory syncytial virus, in healthy late preterm and term infants.  
C.I.13: Submission of the final report from study D5290C00005 (MEDLEY) listed as a category 3 study in the RMP. This is a phase

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Request for supplementary information adopted with a specific timetable.

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II/III study, randomized, double-blind, placebo-controlled study to evaluate the safety of Beyfortus (nirsevimab) in high-risk children. The RMP version 2.3 has also been submitted.” Request for Supplementary Information adopted on 08.02.2024.

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**GAVRETO - Pralsetinib -  
EMA/H/C/005413/II/0012**

Positive Opinion adopted by consensus on 22.02.2024.

Roche Registration GmbH, Rapporteur: Aaron Sosa Mejia, PRAC Rapporteur: Ulla Wändel Liminga, “Update of sections 4.2, 4.4 and 4.5 of the SmPC in order to amend posology recommendations, warnings and drug-drug interaction information regarding the co-administration with CYP3A4 inhibitors, P-gp inhibitors and CYP3A4 inducers based on final results from the DDI study GP43162, listed as a category 3 study in the RMP, as well as results from the physiologically based pharmacokinetic (PBPK) analyses summarised in the PBPK Report 1120689. Study GP43162 is a phase 1, open-label, fixed-sequence study to evaluate the effect of a single dose of cyclosporine on the single dose pharmacokinetics of pralsetinib in healthy subjects. The package leaflet has been updated accordingly. The RMP version 1.8 is agreed.”

Opinion adopted on 22.02.2024.

Request for Supplementary Information adopted on 09.11.2023, 22.06.2023, 30.03.2023.

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**GAVRETO - Pralsetinib -  
EMA/H/C/005413/II/0017**

Positive Opinion adopted by consensus on 22.02.2024.

Roche Registration GmbH, Rapporteur: Aaron Sosa Mejia, PRAC Rapporteur: Ulla Wändel Liminga, “Update of sections 4.2 and 5.2 of the SmPC in order to include information regarding moderate and severe hepatic impairment based on final results from study GP43163 listed as a category 3 study in the RMP; this is a Phase I, open-label, single-dose study to evaluate the pharmacokinetics and safety of pralsetinib in subjects with moderate or severe hepatic impairment compared to healthy subjects. The RMP version 1.8 is agreed. In addition, the MAH took the opportunity to update the list of local representatives in the Package Leaflet and to update the marketing authorisation renewal date in Annex I.”

Opinion adopted on 22.02.2024.

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Request for Supplementary Information adopted on 14.12.2023.

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**Inrebic - Fedratinib -  
EMA/H/C/005026/II/0019, Orphan**

Bristol-Myers Squibb Pharma EEIG, Rapporteur: Peter Mol, PRAC Rapporteur: Sonja Hrabcik, "Update of sections 4.2 and 5.2 of the SmPC in order to update posology recommendations in patients with severe hepatic impairment and to update pharmacokinetic information based on final results from study FEDR-CP-001 listed as a category 3 study in the RMP; this is a phase 1, open-label, single-dose study to assess the pharmacokinetics, safety, and tolerability of fedratinib in subjects with moderate and severe hepatic impairment compared with healthy subjects. The RMP version 2.0 has also been submitted."

Request for Supplementary Information adopted on 22.02.2024.

Request for supplementary information adopted with a specific timetable.

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**Onpattro - Patisiran -  
EMA/H/C/004699/II/0034, Orphan**

Alnylam Netherlands B.V., Rapporteur: Kristina Dunder, PRAC Rapporteur: Rhea Fitzgerald, "Submission of the final report from study ALN-TTR02-006 (study 006), listed a category 3 study in the RMP. This is a multicenter, open-label, extension study to evaluate the long-term safety and efficacy of patisiran in patients with familial amyloidotic polyneuropathy who have completed a prior clinical study with patisiran. The RMP version 2.2 has also been submitted."

Opinion adopted on 08.02.2024.

Request for Supplementary Information adopted on 28.09.2023.

Positive Opinion adopted by consensus on 08.02.2024.

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**Piqray - Alpelisib -  
EMA/H/C/004804/II/0022/G**

Novartis Europharm Limited, Rapporteur: Carolina Prieto Fernandez, PRAC Rapporteur: Bianca Mulder, "Grouped application comprising two type II variations (C.I.4) as follows:  
- Update of sections 4.2, 4.4 and 4.8 of the SmPC in order to update information on prophylactic use of metformin for hyperglycaemia based on the results from study CBYL719CES01T (METALLICA). METALLICA is a Phase II study aimed to evaluate the effect of prophylactic use of metformin for hyperglycaemia in HR-positive, HER2-negative,

Request for supplementary information adopted with a specific timetable.

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PIK3CA-mutated advanced breast cancer patients treated with alpelisib plus endocrine therapy.

- Update of section 4.8 of the SmPC in order to add "uveitis" to the list of adverse drug reactions (ADRs) with frequency "Not known" based on a cumulative review of the MAH safety database and literature.

The Package Leaflet and Annex II are updated accordingly. The RMP version 7.0 has also been submitted."

Request for Supplementary Information adopted on 30.11.2023.

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**SARCLISA - Isatuximab -  
EMA/H/C/004977/II/0026**

Sanofi Winthrop Industrie, Rapporteur: Peter Mol, PRAC Rapporteur: Monica Martinez Redondo, "Update of sections 4.2, 4.4 and 5.2 of the SmPC based on final results from study TED16414, listed as a category 3 study in the RMP; this is a phase 1b/2 open label, non-randomized, multi center study to evaluate the safety, pharmacokinetics, and preliminary efficacy of isatuximab (SAR650984) in patients awaiting kidney transplantation. The Package Leaflet is updated accordingly. The RMP version 1.4 has also been submitted. In addition, the MAH took the opportunity to introduce minor editorial changes to the PI."  
Opinion adopted on 22.02.2024.

Positive Opinion adopted by consensus on 22.02.2024.

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**Tecentriq - Atezolizumab -  
EMA/H/C/004143/II/0083/G**

Roche Registration GmbH, Rapporteur: Aaron Sosa Mejia, PRAC Rapporteur: Carla Torre, "A grouped application comprising of 2 Type II variations, as follows:  
C.I.4: Update of section 5.1 of the SmPC in order to update efficacy information based on final results from study IMvigora210 (GO29293) listed as a PAES in the Annex II; this is a Phase II, multicenter, single-arm study of atezolizumab in patients with locally advanced or metastatic urothelial bladder cancer. The Annex II is updated accordingly. In addition, the MAH took the opportunity to introduce minor editorial changes to the PI and to bring it in line with the latest QRD template.  
C.I.13: Submission of the final report from study SAUL (MO29983) listed as a category 3

Positive Opinion adopted by consensus on 08.02.2024.

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study in the RMP. This is an open-label, single arm, multicenter, safety study of atezolizumab in locally advanced or metastatic urothelial or non-urothelial carcinoma of the urinary tract. The RMP version 30.0 has also been submitted.”  
Opinion adopted on 08.02.2024.

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**Vabysmo - Faricimab -  
EMA/H/C/005642/II/0009**

Positive Opinion adopted by consensus on 08.02.2024.

Roche Registration GmbH, Rapporteur: Jayne Crowe, PRAC Rapporteur: Carla Torre, “Update of section 4.8 of the SmPC in order to add ‘Retinal Vasculitis’ and ‘Retinal Occlusive Vasculitis’ to the list of adverse drug reactions (ADRs) with frequency not known, based on a drug safety report and post-marketing data; the Package Leaflet is updated accordingly. The RMP version 4.0 has also been submitted. In addition, the MAH took the opportunity to add study CR45271 as a category 3 study in the RMP, to introduce minor changes and corrections to the PI and to update the list of local representatives in the Package Leaflet.”  
Opinion adopted on 08.02.2024.

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**Vaxzevria - COVID 19 Vaccine (ChAdOx1 S [recombinant]) -  
EMA/H/C/005675/II/0096**

Request for supplementary information adopted with a specific timetable.

AstraZeneca AB, Rapporteur: Sol Ruiz, PRAC Rapporteur: Jean-Michel Dogné, “Update of sections 4.8 and 5.1 of the SmPC based on final results from study D7220C00001; this is a phase 2/3 partially double-blinded, randomised, multinational, active-controlled study in both previously vaccinated and unvaccinated adults to determine the safety and immunogenicity of AZD2816, a vaccine for the prevention of COVID-19 caused by variant strains of SARS-CoV-2. The RMP version 8 s1 has also been submitted.”  
Request for Supplementary Information adopted on 08.02.2024.

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**Vaxzevria - COVID 19 Vaccine (ChAdOx1 S [recombinant]) -  
EMA/H/C/005675/II/0097**

Request for supplementary information adopted with a specific timetable.

AstraZeneca AB, Rapporteur: Sol Ruiz, PRAC Rapporteur: Jean-Michel Dogné, “Submission of the final report from study D8110C00001 listed as a category 3 study in the RMP (SOB/020). This is a phase III, randomised, placebo-controlled study of AZD1222 (Vaxzevria)

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conducted in the US, Peru and Chile. The purpose of the final CSR addendum is to provide long-term safety data through to study completion and include the second year of follow-up post-first dose and final day 730 visit. The RMP version 8 s2 has also been submitted." Request for Supplementary Information adopted on 08.02.2024.

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**Xevudy - Sotrovimab -  
EMA/H/C/005676/II/0026**

Glaxosmithkline Trading Services Limited,  
Rapporteur: Thalia Marie Estrup Blicher, PRAC  
Rapporteur: Liana Martirosyan, "To update sections 4.2, 4.8 and 5.2 of the SmPC in order to update information on paediatric population based on final results from study COMET-PACE (215226), a category 3 study in the RMP; this is an open-label, non-comparator, multicentre study to describe the pharmacokinetics (PK), pharmacodynamics (PD; viral load) and safety following a single intravenous or intramuscular dose of sotrovimab in paediatric participants with mild to moderate COVID-19 at high risk of disease progression. The updated RMP version 1.1 has also been submitted." Request for Supplementary Information adopted on 08.02.2024.

Request for supplementary information adopted with a specific timetable.

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**WS2631  
Kispilyx-EMA/H/C/004224/WS2631/0059  
Lenvima-  
EMA/H/C/003727/WS2631/0054**

Eisai GmbH, Lead Rapporteur: Karin Janssen van Doorn, Lead PRAC Rapporteur: Ulla Wändel Liminga, "Update of sections 4.2, 4.8, 5.1 and 5.2 of the SmPC for Kispilyx and sections 4.8 and 5.1 of the SmPC for Lenvima, in order to reflect the results of two completed paediatric clinical studies E7080-G000-216 and E7080-G000-231. Study 231 is a Phase 2, open-label, multicenter basket study to evaluate the antitumor activity and safety of Lenvatinib in children, adolescents, and young adults with relapsed or refractory solid malignancies. Study 216 is a Phase 1/2, multicenter, open-label, single arm study of lenvatinib in combination with everolimus in pediatric subjects (and young adults aged ≤21 years) with relapsed or refractory malignant solid tumors. The Package Leaflet for Kispilyx is updated accordingly. The

Request for supplementary information adopted with a specific timetable.

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RMP version 15.3 has also been submitted.”  
Request for Supplementary Information adopted  
on 22.02.2024.

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#### **B.5.4. PRAC assessed procedures**

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PRAC Led  
**BLINCYTO - Blinatumomab -  
EMA/H/C/003731/II/0054, Orphan**  
Amgen Europe B.V., PRAC Rapporteur: Jana  
Lukacisinova, PRAC-CHMP liaison: Petr Vrbata,  
“To update sections 4.2, 4.4 and 4.8 of the  
SmPC to include Immune Effector Cell-  
Associated Neurotoxicity Syndrome (ICANS);  
and to update section D of Annex II to remove  
educational materials for physicians,  
pharmacists and nurses and to include ICANS  
within neurologic events in educational material  
for patient/caregivers and patient alert card  
following the outcome of PSUR procedure  
EMA/H/C/PSUSA/00010460/202212. The  
Package Leaflet is updated accordingly. The RMP  
version 17.0 has also been submitted.”  
Request for Supplementary Information adopted  
on 08.02.2024.

Request for supplementary information adopted  
with a specific timetable.

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PRAC Led  
**Entyvio - Vedolizumab -  
EMA/H/C/002782/II/0081**  
Takeda Pharma A/S, PRAC Rapporteur: Adam  
Przybylkowski, PRAC-CHMP liaison: Ewa  
Balkowiec Iskra, “Update of section 4.6 of the  
SmPC in order to update information on  
pregnancy based on final results from study  
Vedolizumab-5001 (OTIS Entyvio Pregnancy  
Exposure Registry); this is a non-interventional  
study to monitor planned and unplanned  
pregnancies in female patients with ulcerative  
colitis or Crohn’s disease. The Package Leaflet is  
updated accordingly. In addition, the MAH took  
the opportunity to introduce minor changes and  
corrections to the PI and bring it in line with the  
latest QRD template.”  
Request for Supplementary Information adopted  
on 08.02.2024.

Request for supplementary information adopted  
with a specific timetable.

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PRAC Led  
**HEPLISAV B - Hepatitis B surface antigen  
(rDNA) - EMA/H/C/005063/II/0031**  
Dynavax GmbH, PRAC Rapporteur: Gabriele  
Maurer, PRAC-CHMP liaison: Jan Mueller-

Positive Opinion adopted by consensus on  
08.02.2024.

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Berghaus, "Update of section 4.6 of the SmPC in order to update information on pregnancy based on final results from study DV2-HBV-28 - Post-marketing observational surveillance study to evaluate pregnancy outcomes among women who receive HEPLISAV-B or Engerix-B; HBV-28 was conducted using the same patient population as two observational post-marketing surveillance studies designed to evaluate the incidence of AMI (HBV-25) or new-onset immunemediated diseases, herpes zoster, and anaphylaxis (HBV-26) in recipients of HEPLISAV-B compared with recipients of Engerix-B. The primary objective of this study was to describe and compare pregnancy outcomes in recipients of HEPLISAV-B and recipients of Engerix-B. The Package Leaflet is updated accordingly. The RMP version 1.4 has also been submitted. In addition, the MAH took the opportunity to bring the PI in line with the latest QRD template version 10.3." Opinion adopted on 08.02.2024. Request for Supplementary Information adopted on 30.11.2023.

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PRAC Led

**MabThera - Rituximab -**

**EMA/H/C/000165/II/0201/G**

Roche Registration GmbH, PRAC Rapporteur: Karin Erneholm, PRAC-CHMP liaison: Aaron Sosa Mejia, "A grouped application comprising of: Type II (C.I.3.b): Update of sections 4.1, 4.2, 4.3, 4.8, 5.1, 6.2, 6.4 and 6.5 of the SmPC in order to introduce several structural and editorial changes to align with the current SmPC guideline and to remove the educational materials for HCPs and patients, following the request by the PRAC in the AR for the PSUSA procedure EMA/PRAC/257005/2023. The Annex II, Labelling and Package Leaflet are updated accordingly. The RMP version 25.0 has also been submitted. In addition, the MAH took the opportunity to introduce minor editorial changes to the PI and to update the list of local representatives in the Package Leaflet. Type I (A.6): To change the ATC Code of rituximab from L01XC02 to L01FA01." Request for Supplementary Information adopted on 08.02.2024.

Request for supplementary information adopted with a specific timetable.

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PRAC Led

Request for supplementary information adopted

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**Mysimba - Naltrexone hydrochloride /  
Bupropion hydrochloride -  
EMA/H/C/003687/II/0063**

Orexigen Therapeutics Ireland Limited,  
Rapporteur: Thalia Marie Estrup Blicher, PRAC  
Rapporteur: Martin Huber, PRAC-CHMP liaison:  
Janet Koenig, "To update sections 4.3, 4.4 and  
4.5 of the SmPC to update and streamline the  
relevant wording on opioids following the  
assessment of PSUSA/00010366/202209  
procedure. The Package Leaflet is updated  
accordingly. The RMP version 12.9 has also  
been submitted."

Request for Supplementary Information adopted  
on 09.02.2024, 31.08.2023.

with a specific timetable.

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PRAC Led

**RAYVOW - Lasmiditan -  
EMA/H/C/005332/II/0005**

Eli Lilly Nederland B.V., PRAC Rapporteur: Anna  
Mareková, PRAC-CHMP liaison: Frantisek Drafi,  
"Submission of an updated RMP version 1.1 in  
order to include a descriptive interim analysis in  
the study design of study H8H-MC-B006, listed  
as a category 3 study in the RMP. This is a non-  
interventional study titled 'Lasmiditan Use and  
Motor Vehicle Accidents in Real-World Settings  
in the US'."

Opinion adopted on 08.02.2024.

Positive Opinion adopted by consensus on  
08.02.2024.

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PRAC Led

**SCENESSE - Afamelanotide -  
EMA/H/C/002548/II/0049, Orphan**

Clinuvel Europe Limited, PRAC Rapporteur:  
Martin Huber, PRAC-CHMP liaison: Janet Koenig,  
"To remove study CUV-RCR-001 (Scenesse  
(Afamelanotide 16mg) Retrospective Chart  
Review) listed as an obligation in the Annex II  
of the Product Information. This is a  
retrospective study comparing long-term safety  
data and outcome endpoints in patients  
receiving and not receiving Scenesse or having  
discontinued Scenesse use. The Annex II and  
the RMP (version 9.6) are updated accordingly."

Request for Supplementary Information adopted  
on 08.02.2024.

Request for supplementary information adopted  
with a specific timetable.

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PRAC Led

**Spravato - Esketamine -  
EMA/H/C/004535/II/0021**

Janssen-Cilag International N.V., PRAC  
Rapporteur: Kirsti Villikka, PRAC-CHMP liaison:

Positive Opinion adopted by consensus on  
08.02.2024.

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Outi Mäki-Ikola, "Submission of an updated RMP version 5.2 in order to remove "use during pregnancy" as missing information from the list of safety concerns, with the consequential removal of the associated category 3 additional pharmacovigilance activity, the National Pregnancy Registry for Antidepressants ("Massachusetts General Hospital (MGH) pregnancy registry)."

Opinion adopted on 08.02.2024.

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PRAC Led

**Stelara - Ustekinumab -**

**EMA/H/C/000958/II/0101/G**

Janssen-Cilag International N.V., PRAC Rapporteur: Rhea Fitzgerald, PRAC-CHMP liaison: Jayne Crowe, "Update of section 4.4 of the SmPC in order to remove a warning on cardiovascular events based on final results from non-interventional PASS studies NDI-MACE (CNT01275PSO4005) and Quantify MACE (PCSIMM004697), listed as category 3 studies in the RMP (MEA/053 and MEA/054). NDI-MACE is a Nordic Database Initiative for Exposure to Ustekinumab: A Review and Analysis of Major Adverse Cardiovascular Events from the Swedish and Danish National Registry Systems; Quantify MACE is an Observational Longitudinal Post-authorisation Safety Study of STELARA in the Treatment of Psoriasis and Psoriatic Arthritis: Analysis of Major Adverse Cardiovascular Events (MACE) using Swedish National Health Registers. The Package Leaflet is updated accordingly. The RMP version 27.1 has also been submitted."

Opinion adopted on 08.02.2024.

Request for Supplementary Information adopted on 28.09.2023.

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Positive Opinion adopted by consensus on 08.02.2024.

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PRAC Led

**TachoSil - Human thrombin / Human fibrinogen - EMA/H/C/000505/II/0124**

Corza Medical GmbH, PRAC Rapporteur: Gabriele Maurer, PRAC-CHMP liaison: Jan Mueller-Berghaus, "Submission of an updated RMP version 9.1 in order to reflect the extension of indication to include the paediatric population and to update the details of the planned non-interventional post-authorisation safety study: PASS-TachoSil Evaluation (PasTel)."

Request for Supplementary Information adopted

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Request for supplementary information adopted with a specific timetable.

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on 08.02.2024.

PRAC Led

**Zessly - Infliximab -**

**EMA/H/C/004647/II/0033**

Sandoz GmbH, PRAC Rapporteur: Mari Thorn,

PRAC-CHMP liaison: Kristina Dunder,

“Submission of an updated RMP version 4.0 in order to remove the UKIBD (UK) registry from the additional pharmacovigilance activities.”

Request for Supplementary Information adopted on 08.02.2024.

Request for supplementary information adopted with a specific timetable.

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PRAC Led

**WS2587**

**TECFIDERA-**

**EMA/H/C/002601/WS2587/0085**

**Vumerity-**

**EMA/H/C/005437/WS2587/0015**

Biogen Netherlands B.V., Lead PRAC

Rapporteur: Martin Huber, PRAC-CHMP liaison:

Martina Weise, “Submission of the final report

from study 109MS401, a multicenter, global,

observational study to collect information on

safety and to document the drug utilization of

Tecfidera (Dimethyl Fumarate) when used in

routine medical practice in the treatment of

Multiple Sclerosis (ESTEEM), listed as a category

3 study in the RMP (MEA007.6). The RMPs

version 16.1 for Tecfidera and version 2.1 for

Vumerity, have also been submitted.”

Request for Supplementary Information adopted on 08.02.2024.

Request for supplementary information adopted with a specific timetable.

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PRAC Led

**WS2615**

**Abseamed-**

**EMA/H/C/000727/WS2615/0108**

**Binocrit-**

**EMA/H/C/000725/WS2615/0108**

**Epoetin alfa Hexal-**

**EMA/H/C/000726/WS2615/0108**

Sandoz GmbH, Lead PRAC Rapporteur: Tiphaine

Vaillant, PRAC-CHMP liaison: Alexandre Moreau,

“Submission of the final report from Non-

Interventional Post Authorisation Safety Study,

NI-PASS HX575-507 listed as a category 3

study in the RMP. The non-interventional (NIS

PASS) study HX575-507 was conducted to

address a post-approval requirement (MEA

13.5) to evaluate the safety profile of HX575

administered s.c. in patients with CKD-induced

Request for supplementary information adopted with a specific timetable.

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anemia under real-life conditions, in order to increase confidence on the safe use of s.c. HX575. The RMP version 19.0 has also been submitted.”

Request for Supplementary Information adopted on 08.02.2024.

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PRAC Led  
**WS2620**

**Dovato-EMEA/H/C/004909/WS2620/0047**

**Juluca-EMEA/H/C/004427/WS2620/0056**

**Tivicay-EMEA/H/C/002753/WS2620/0092**

**Triumeq-**

**EMEA/H/C/002754/WS2620/0118**

ViiV Healthcare B.V., Lead PRAC Rapporteur: Martin Huber, PRAC-CHMP liaison: Martina Weise, “Update of section 4.6 of the SmPC in order to update information about the use of DTG-containing regimens in pregnancy and at conception based on final results from non-interventional Tsepamo study and the Eswatini Birth Outcomes Surveillance study. In addition, data from other cohort studies and pregnancy registries, including the APR, DOLOMITE-EPPICC (Study 208613) and DOLOMITE-NEAT-ID Network study (Study 208759) both listed as category 3 studies in the RMP; and the US Chart Review (Study 212976) as well as data from literature are included. DOLOMITE-EPPICC (Study 208613) is a non-interventional study to Assess “real-world” maternal and foetal outcomes following DTG use during pregnancy and to describe patterns of DTG utilization; DOLOMITE NEAT ID Network Study (208759) is a non-interventional, multi-site observational study to define the safety and effectiveness of Dolutegravir use in HIV positive pregnant women. The Package Leaflet is updated accordingly. The RMP version 19 has also been submitted. In addition, the MAH took the opportunity to implement editorial changes to sections 4.4 and 4.5 of the SmPC.”

Request for Supplementary Information adopted on 08.02.2024.

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Request for supplementary information adopted with a specific timetable.

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#### **B.5.5. CHMP-CAT assessed procedures**

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**Breyanzi - Lisocabtagene maraleucel /  
Lisocabtagene maraleucel -**

**EMEA/H/C/004731/II/0032, ATMP**

Bristol-Myers Squibb Pharma EEIG, Rapporteur:

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Positive Opinion adopted by consensus on 22.02.2024.

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Concetta Quintarelli, CHMP Coordinator: Paolo Gasparini  
Opinion adopted on 22.02.2024, 16.02.2024.  
Request for Supplementary Information adopted on 08.12.2023.

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**Breyanzi - Lisocabtagene maraleucel / Lisocabtagene maraleucel - EMEA/H/C/004731/II/0036/G, ATMP**

Bristol-Myers Squibb Pharma EEIG, Rapporteur: Concetta Quintarelli, CHMP Coordinator: Paolo Gasparini, "Grouped application comprising two variations as follows:

C.I.4 – Update of sections 4.4 and 4.8 of the SmPC in order to add immune effector cell-associated neurotoxicity syndrome (ICANS) as an adverse drug reaction (ADR) based on the cumulative review of MAH safety database and literature. The Package Leaflet is updated accordingly. In addition, the MAH took this opportunity to introduce editorial changes.

A.6 – To include the ATC Code L01XL08 in section 5.1 of the SmPC."

Request for Supplementary Information adopted on 16.02.2024.

Request for supplementary information adopted with a specific timetable.

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**Kymriah - Tisagenlecleucel - EMEA/H/C/004090/II/0071, Orphan, ATMP**

Novartis Europharm Limited, Rapporteur: Rune Kjekken, CHMP Coordinator: Ingrid Wang, "Update of sections 5.1 and 5.2 of the SmPC to update efficacy and pharmacokinetic information based on final results from study CCTL019B2202 (a phase II, single arm, multicenter trial to determine the efficacy and safety of CTL019 in paediatric patients with relapsed and refractory B-cell acute lymphoblastic leukemia). Submission of cellular kinetic report for the B-cell acute lymphoblastic leukaemia (ALL) indication based on data from pivotal study CCTL019B2202 and the supportive study CCTL019B2205J involving paediatric ALL patients (partially fulfil REC). In addition, the MAH took this opportunity to introduce editorial changes."

Opinion adopted on 22.02.2024, 16.02.2024.

Request for Supplementary Information adopted on 31.10.2023.

Positive Opinion adopted by consensus on 22.02.2024.

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**Kymriah - Tisagenlecleucel - EMEA/H/C/004090/II/0079/G, Orphan,**

Positive Opinion adopted by consensus on

<p><b>ATMP</b> Novartis Europharm Limited, Rapporteur: Rune Kjekken, CHMP Coordinator: Ingrid Wang Opinion adopted on 22.02.2024, 16.02.2024.</p>	22.02.2024.
<p><b>Upstaza - Eladocagene exuparvovec - EMEA/H/C/005352/II/0013, Orphan, ATMP</b> PTC Therapeutics International Limited, Rapporteur: Joseph DeCoursey, CHMP Coordinator: Finbarr Leacy, "- To submit the final report with the results of the active substance and finished product concurrent process validation batches, including hold time data for the finished product batch, in order to further assess process consistency and maintain patient's safety (Annex II condition)." Opinion adopted on 22.02.2024, 16.02.2024. Request for Supplementary Information adopted on 19.01.2024, 08.09.2023.</p>	Positive Opinion adopted by consensus on 22.02.2024.
<p><b>WS2500 Tecartus- EMEA/H/C/005102/WS2500/0040 Yescarta- EMEA/H/C/004480/WS2500/0068</b> Kite Pharma EU B.V., Lead Rapporteur: Jan Mueller-Berghaus Request for Supplementary Information adopted on 16.02.2024.</p>	Request for supplementary information adopted with a specific timetable.
<p><b>B.5.6. CHMP-PRAC-CAT assessed procedures</b></p>	
<p><b>Kymriah - Tisagenlecleucel - EMEA/H/C/004090/II/0075, Orphan, ATMP</b> Novartis Europharm Limited, Rapporteur: Rune Kjekken, CHMP Coordinator: Ingrid Wang, PRAC Rapporteur: Gabriele Maurer, "Update of sections 5.1 and 5.2 of the SmPC to update efficacy and pharmacokinetic information based on final results from study Cl.CTL019C2201 PAES in the Annex II (ANX008); this is a Phase II, single arm, multicenter trial to determine the efficacy and safety of CTL019 in adult patients with relapsed or refractory diffuse large B-cell lymphoma (DLBCL). The RMP version 6 has also been submitted. In addition, Annex II.D of the product information is updated to reflect that the obligation to conduct the mention study is fulfilled."</p>	Positive Opinion adopted by consensus on 22.02.2024.

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Opinion adopted on 22.02.2024, 16.02.2024.  
Request for Supplementary Information adopted  
on 31.10.2023.

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#### **B.5.7. PRAC assessed ATMP procedures**

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

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**WS2533** Request for supplementary information adopted  
**Jentaduetto-** with a specific timetable.  
**EMA/H/C/002279/WS2533/0071**  
**Trajenta-**  
**EMA/H/C/002110/WS2533/0053**  
Boehringer Ingelheim International GmbH, Lead  
Rapporteur: Patrick Vrijlandt  
Request for Supplementary Information adopted  
on 01.02.2024.

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**WS2576** Positive Opinion adopted by consensus on  
**Elebrato Ellipta-** 15.02.2024.  
**EMA/H/C/004781/WS2576/0037**  
**Relvar Ellipta-**  
**EMA/H/C/002673/WS2576/0064**  
**Revinty Ellipta-**  
**EMA/H/C/002745/WS2576/0061**  
**Trelegy Ellipta-**  
**EMA/H/C/004363/WS2576/0034**  
GlaxoSmithKline (Ireland) Limited, Lead  
Rapporteur: Maria Concepcion Prieto Yerro  
Opinion adopted on 15.02.2024.  
Request for Supplementary Information adopted  
on 09.11.2023.

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**WS2588** Positive Opinion adopted by consensus on  
**Mircera-EMA/H/C/000739/WS2588/0097** 01.02.2024.  
**NeoRecormon-**  
**EMA/H/C/000116/WS2588/0122**  
Roche Registration GmbH, Lead Rapporteur:  
Martina Weise  
Opinion adopted on 01.02.2024.

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**WS2594/G** Positive Opinion adopted by consensus on  
**Ambirix-** 01.02.2024.  
**EMA/H/C/000426/WS2594/0132/G**  
**Twinrix Adult-**  
**EMA/H/C/000112/WS2594/0167/G**  
**Twinrix Paediatric-**  
**EMA/H/C/000129/WS2594/0168/G**  
GlaxoSmithkline Biologicals SA, Lead  
Rapporteur: Christophe Focke  
Opinion adopted on 01.02.2024.

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<p><b>WS2595/G</b>  <b>Riltrava Aerosphere-</b>  <b>EMA/H/C/005311/WS2595/0009/G</b>  <b>Trixeo Aerosphere-</b>  <b>EMA/H/C/004983/WS2595/0016/G</b>  AstraZeneca AB, Lead Rapporteur: Finbarr Leacy  Opinion adopted on 08.02.2024.</p>	Positive Opinion adopted by consensus on 08.02.2024.
<p><b>WS2605</b>  <b>HyQvia-EMA/H/C/002491/WS2605/0095</b>  <b>Kiovig-EMA/H/C/000628/WS2605/0126</b>  Takeda Manufacturing Austria AG, Lead Rapporteur: Jan Mueller-Berghaus  Opinion adopted on 29.02.2024.  Request for Supplementary Information adopted on 18.01.2024.</p>	Positive Opinion adopted by consensus on 29.02.2024
<p><b>WS2618</b>  <b>Dengue Tetravalent Vaccine (Live, Attenuated) Takeda-</b>  <b>EMA/H/W/005362/WS2618/0013</b>  <b>Qdenga-</b>  <b>EMA/H/C/005155/WS2618/0014</b>  Takeda GmbH, Lead Rapporteur: Sol Ruiz  Request for Supplementary Information adopted on 15.02.2024.</p>	Request for supplementary information adopted with a specific timetable.
<p><b>WS2621/G</b>  <b>Suboxone-</b>  <b>EMA/H/C/000697/WS2621/0059/G</b>  Indivior Europe Limited, Lead Rapporteur: Janet Koenig  Opinion adopted on 01.02.2024.</p>	Positive Opinion adopted by consensus on 01.02.2024.
<p><b>WS2629/G</b>  <b>Eviplera-</b>  <b>EMA/H/C/002312/WS2629/0115/G</b>  <b>Stribild-</b>  <b>EMA/H/C/002574/WS2629/0122/G</b>  <b>Truvada-</b>  <b>EMA/H/C/000594/WS2629/0180/G</b>  <b>Viread-</b>  <b>EMA/H/C/000419/WS2629/0211/G</b>  Gilead Sciences Ireland UC, Lead Rapporteur: Jean-Michel Race  Request for Supplementary Information adopted on 08.02.2024.</p>	Request for supplementary information adopted with a specific timetable.
<p><b>WS2643</b>  <b>Nuwiq-EMA/H/C/002813/WS2643/0059</b>  <b>Vihuma-</b>  <b>EMA/H/C/004459/WS2643/0041</b></p>	Positive Opinion adopted by consensus on 15.02.2024.

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Octapharma AB, Lead Rapporteur: Jan Mueller-Berghaus  
Opinion adopted on 15.02.2024.

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**WS2649** Positive Opinion adopted by consensus on  
**Luveris-EMA/H/C/000292/WS2649/0099** 22.02.2024.

**Pergoveris-EMA/H/C/000714/WS2649/0090**  
Merck Europe B.V., Lead Rapporteur: Thalia Marie Estrup Blicher  
Opinion adopted on 22.02.2024.

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#### **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**

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**Acoramidis - EMA/H/C/006333, Orphan**  
BridgeBio Europe B.V., for the treatment of wild-type or variant transthyretin amyloidosis in adult patients with cardiomyopathy (ATTR-CM).

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**Trastuzumab - EMA/H/C/006219**  
treatment of metastatic and early breast cancer

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**Diflunisal - EMA/H/C/006248, Orphan**  
AO Pharma AB, Treatment of ATTR amyloidosis

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**Ivermectin / Albendazole - EMA/H/W/005186, Article 58**  
prevention and treatment of lymphatic filariasis, and soil-transmitted helminths infections.

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**Lazertinib - EMA/H/C/006074**  
treatment of adult patients with advanced non-small cell lung cancer (NSCLC)

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**Linvoseltamab - EMA/H/C/006370**  
monotherapy for the treatment of adult patients with relapsed or refractory multiple myeloma

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**Nemolizumab - EMA/H/C/006149**  
for the treatment of moderate-to-severe atopic dermatitis and for the treatment of prurigo nodularis

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**Pegfilgrastim - EMA/H/C/006348, PUMA**  
treatment of neutropenia in paediatric patients

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**Tisotumab vedotin - EMEA/H/C/005363**

treatment of adult patients with recurrent or metastatic cervical  
treatment of adult patients with recurrent or metastatic cervical cancer with disease progression on or after systemic therapy

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**Trabectedin - EMEA/H/C/006433**

treatment of soft tissue sarcoma and combination with PLD treatment of relapsed platinum-sensitive ovarian cancer

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**Human albumin solution -****EMEA/H/D/006410**

vitrification of human MII-phase oocytes and embryos for assisted reproductive technology (ART).

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

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**Lyrica - Pregabalin -****EMEA/H/C/000546/X/0127**

Upjohn EESV, Rapporteur: Peter Mol, PRAC  
Rapporteur: Liana Martirosyan, "Extension application to introduce a new pharmaceutical form (orodispersible tablet)"

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**B.6.3. Restart of procedure - responses received to Day 120 List of Questions timetables: for information****B.6.4. Annual Re-assessments: timetables for adoption**

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**Ceplene - Histamine dihydrochloride -****EMEA/H/C/000796/S/0048**

Laboratoires Delbert, Rapporteur: Jayne Crowe,  
PRAC Rapporteur: Eamon O Murchu

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**LIVMARLI - Maralixibat -****EMEA/H/C/005857/S/0012, Orphan**

Mirum Pharmaceuticals International B.V.,  
Rapporteur: Martina Weise, PRAC Rapporteur:  
Adam Przybylkowski

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**SCENESSE - Afamelanotide -****EMEA/H/C/002548/S/0050, Orphan**

Clinuvel Europe Limited, Rapporteur: Janet  
Koenig, PRAC Rapporteur: Martin Huber

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

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#### **Epidyolex - Cannabidiol -**

##### **EMA/H/C/004675/R/0031, Orphan**

Jazz Pharmaceuticals Ireland Limited,  
Rapporteur: Thalia Marie Estrup Blicher, Co-  
Rapporteur: Tomas Radimersky, PRAC  
Rapporteur: Ana Sofia Diniz Martins

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#### **Inbrija - Levodopa -**

##### **EMA/H/C/004786/R/0022**

Acorda Therapeutics Ireland Limited,  
Rapporteur: Peter Mol, Co-Rapporteur: Jayne  
Crowe, PRAC Rapporteur: Barbara Kovacic  
Bytyqi

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#### **XOSPATA - Gilteritinib -**

##### **EMA/H/C/004752/R/0017, Orphan**

Astellas Pharma Europe B.V., Rapporteur: Ingrid  
Wang, Co-Rapporteur: Elita Poplavska, PRAC  
Rapporteur: Martin Huber

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### **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**

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#### **AREXVY - Respiratory syncytial virus, glycoprotein F, recombinant, stabilised in the pre-fusion conformation, adjuvanted with AS01E - EMA/H/C/006054/II/0008**

GlaxoSmithkline Biologicals S.A., Rapporteur:  
Patrick Vrijlandt, PRAC Rapporteur: Maria del  
Pilar Rayon, "Extension of indication to include  
treatment of adults 50-59 years of age who are  
at increased risk for RSV disease for AREXVY,  
based on results from study 219238 (RSV  
OA=ADJ-018); this is a phase 3, observer-blind,  
placebo-controlled, randomized, multi-country,  
multi-center, non-inferiority study with 2  
cohorts to evaluate immunogenicity,  
reactogenicity and safety of a single dose of  
RSVPreF3 OA in adults 50-59 years of age. As a  
consequence, sections 4.1, 4.6, 4.8 and 5.1 of  
the SmPC are updated. The Package Leaflet is  
updated in accordance. Version 1.1 of the RMP  
has also been submitted. In addition, the  
marketing authorisation holder (MAH) took the  
opportunity to introduce minor editorial changes  
to the PI, to bring it in line with the latest QRD

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template version 10.3, and to update the list of local representatives in the Package Leaflet. As part of the application, the MAH is requesting a 1-year extension of the market protection.”  
Request for 1 year of market protection for a new indication (Article 14(11) of Regulation (EC) 726/2004)

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**BUCCOLAM - Midazolam -  
EMA/H/C/002267/II/0061**

Neuraxpharm Pharmaceuticals S.L., Rapporteur: Peter Mol, Co-Rapporteur: Alexandre Moreau, PRAC Rapporteur: Liana Martirosyan, “Extension of indication to include treatment of adults to Buccolam 10 mg, based on the results from study 2023-504903-10-00; this is an Interventional Study, Relative Bioavailability to investigate the pharmacokinetics of a single dose of midazolam oromucosal solution (Buccolam) compared to midazolam solution for intramuscular injection (Hypnovel) in healthy volunteers under fasting conditions. As a consequence, sections 4.1, 4.2, 4.8, 5.1, 5.2, 6.5 and 6.6 of the SmPC are updated. The Package Leaflet and Labelling are updated in accordance. Version 8.1 of the RMP has also been submitted.”

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**Inaqovi - Decitabine / Cedazuridine -  
EMA/H/C/005823/II/0002**

Otsuka Pharmaceutical Netherlands B.V., Rapporteur: Filip Josephson, PRAC Rapporteur: Marie Louise Schougaard Christiansen, “Grouped application consisting of:  
C.I.6: Extension of indication to include treatment of adult patients with myelodysplastic syndromes (MDS) for INAQOVI.  
C.I.6: Extension of indication to include treatment of adult patients with chronic myelomonocytic leukaemia (CMML) for INAQOVI.  
Based on final results from studies ASTX727-01, ASTX727-02, ASTX727-04, E7727-01, and E7727-02. As a consequence, sections 4.1, 4.2, 4.8, 5.1 and 5.2 of the SmPC are updated. The Package Leaflet is updated in accordance. Version 1.1 of the RMP has also been submitted. Furthermore, the PI is brought in line with the latest QRD template version 10.3. As part of the application the MAH is requesting a 1-year extension of the market protection.”

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

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**Infanrix hexa - Diphtheria, tetanus, pertussis (acellular, component), hepatitis B (rDNA), poliomyelitis (inact.) and haemophilus type B conjugate vaccine (adsorbed) -**

**EMA/H/C/000296/II/0340/G**

GlaxoSmithkline Biologicals SA, Rapporteur: Christophe Focke, "A grouped application consisting of two type II variations, as follows: C.I.6.a: To modify the approved therapeutic indication to include treatment from the age of 6 weeks for the administration of the primary vaccination, section 4.1 of the SmPC is updated accordingly.

C.I.4: Update of section 4.2 of the SmPC for the use of mixed hexavalent/pentavalent primary vaccination schedule and vaccine interchangeability. The Package Leaflet is updated accordingly. In addition, the MAH took the opportunity to implement editorial changes to the SmPC and the Package Leaflet. "

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**Keytruda - Pembrolizumab -**

**EMA/H/C/003820/II/0150**

Merck Sharp & Dohme B.V., Rapporteur: Paolo Gasparini, PRAC Rapporteur: Bianca Mulder, "Extension of indication to include in combination with enfortumab vedotin, the first-line treatment of locally advanced or metastatic urothelial carcinoma in adults, based on the final results from KEYNOTE-A39/EV-302: "An open label, randomized, controlled phase 3 study of enfortumab vedotin in combination with pembrolizumab versus chemotherapy alone in previously untreated locally advanced (LA) or metastatic urothelial cancer (mUC)"; As a consequence, sections 4.1, 4.2, 4.8 and 5.1 of the SmPC are updated. The Package Leaflet is updated in accordance. Version 45.1 of the RMP has also been submitted."

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**OPDIVO - Nivolumab -**

**EMA/H/C/003985/II/0140**

Bristol-Myers Squibb Pharma EEIG, Rapporteur: Carolina Prieto Fernandez, PRAC Rapporteur: Martin Huber, "Extension of indication to include OPDIVO for the treatment of patients with resectable stage II-IIIB non-small cell lung

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cancer, based on results from study CA209977T; a phase 3, randomised, double-blind study of neoadjuvant chemotherapy plus nivolumab versus neoadjuvant chemotherapy plus placebo, followed by surgical resection and adjuvant treatment with nivolumab or placebo for participants with resectable stage II-IIIB non-small cell lung cancer. 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 36.0 of the RMP has also been submitted.”

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**Padcev - Enfortumab vedotin -  
EMA/H/C/005392/II/0013**

Astellas Pharma Europe B.V., Rapporteur: Aaron Sosa Mejia, Co-Rapporteur: Alexandre Moreau, PRAC Rapporteur: Eva Jirsová, “Extension of indication to include in combination with pembrolizumab, the first-line treatment of adult patients with locally advanced or metastatic urothelial cancer who are eligible for platinum-containing chemotherapy for PADCEV, based on the final results from study KEYNOTE-A39/EV-302: “An open label, randomized, controlled phase 3 study of enfortumab vedotin in combination with pembrolizumab versus chemotherapy alone in previously untreated locally advanced (LA) or metastatic urothelial cancer (mUC)””; As a consequence, sections 4.1, 4.2, 4.4, 4.6, 4.8, 5.1, 5.2, 5.3 and 6.6 of the SmPC are updated. The Package Leaflet is updated in accordance. Version 3.0 of the RMP has also been submitted. As part of the application the MAH is requesting a 1-year extension of the market protection.”

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

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**Pemazyre - Pemigatinib -  
EMA/H/C/005266/II/0015, Orphan**

Incyte Biosciences Distribution B.V., Rapporteur: Alexandre Moreau, Co-Rapporteur: Janet Koenig, PRAC Rapporteur: Bianca Mulder, “Extension of indication to include treatment of adults with myeloid/lymphoid neoplasms (MLNs) with Fibroblast Growth Factor Receptor1 (FGFR1) rearrangement for PEMAZYRE, based on final results from study INCB 54828-203 (FIGHT-203); this is a phase 2, open-label,

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monotherapy, multicenter study to evaluate the efficacy and safety of INCB054828 in subjects with myeloid/lymphoid neoplasms with FGFR1 rearrangement. As a consequence, sections 4.1, 4.2, 4.4, 4.5, 4.8 and 5.1 of the SmPC are updated. The Package Leaflet is updated in accordance. Version 3.0 of the RMP has also been submitted. In addition, the MAH took the opportunity to introduce minor changes to the PI. As part of the application, the MAH is requesting a 1-year extension of the market protection.”

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

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**Tevimbra - Tislelizumab -  
EMA/H/C/005919/II/0003**

Beigene Ireland Limited, Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Bianca Mulder, “Extension of indication to include in combination with platinum-based chemotherapy the first-line treatment of adult patients with unresectable, locally advanced or metastatic oesophageal squamous cell carcinoma (OSCC) for TEVIMBRA, based on results from study BGB-A317-306; this is a multi-regional, randomized, placebo-controlled, double-blind phase 3 study evaluating the efficacy and safety of tislelizumab in combination with chemotherapy compared to placebo in combination with chemotherapy as first-line treatment in patients with unresectable or locally advanced recurrent or metastatic OSCC. As a consequence, sections 4.1, 4.2, 4.4, 4.5, 4.6, 4.8 and 5.1 of the SmPC are updated. The Package Leaflet is updated in accordance. Version 1.1 of the RMP has also been submitted. In addition, the MAH took the opportunity to introduce minor editorial changes to the PI.”

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**Zavicefta - Ceftazidime / Avibactam -  
EMA/H/C/004027/II/0035**

Pfizer Ireland Pharmaceuticals, Rapporteur: Ingrid Wang, Co-Rapporteur: Larisa Gorobets, PRAC Rapporteur: Rugile Pilviniene, “Extension of indication to include treatment of paediatric patients from birth to less than 3-months of age in the following infections: complicated intra-abdominal infection (cIAI), complicated urinary tract infection (cUTI), including pyelonephritis,

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hospital-acquired pneumonia (HAP), including ventilator associated pneumonia (VAP) and in the treatment of infections due to aerobic Gram-negative organisms in patients with limited treatment options, for ZAVICEFTA, based on final results from study C3591024 and the population PK modelling/simulation analyses. Study C3591024 is a Phase 2a, 2-part, open-label, non-randomized, multicenter, single and multiple dose trial to evaluate pharmacokinetics, safety and tolerability of ceftazidime and avibactam in neonates and infants from birth to less than 3 months of age with suspected or confirmed infections due to gram-negative pathogens requiring intravenous antibiotic treatment. As a consequence, sections 4.1, 4.2, 4.4, 4.8, 5.1, 5.2, 6.3 and 6.6 of the SmPC are updated. The Package Leaflet is updated in accordance. Version 3.3 of the RMP has also been submitted. In addition, the MAH took the opportunity to introduce minor editorial changes to the PI.”

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#### **B.6.8. CHMP assessed procedures scope: Pharmaceutical aspects**

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##### **Adtralza - Tralokinumab -**

**EMA/H/C/005255/II/0018**

LEO Pharma A/S, Rapporteur: Jayne Crowe

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##### **Advate - Octocog alfa -**

**EMA/H/C/000520/II/0122/G**

Takeda Manufacturing Austria AG, Rapporteur:  
Jan Mueller-Berghaus

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##### **Aimovig - Erenumab -**

**EMA/H/C/004447/II/0030**

Novartis Europharm Limited, Rapporteur:  
Kristina Dunder

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##### **AQUIPTA - Atogepant -**

**EMA/H/C/005871/II/0001/G**

AbbVie Deutschland GmbH & Co. KG,  
Rapporteur: Janet Koenig

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##### **Elfabrio - Pegunigalsidase alfa -**

**EMA/H/C/005618/II/0004/G**

Chiesi Farmaceutici S.p.A., Rapporteur:  
Alexandre Moreau

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##### **Eylea - Aflibercept -**

**EMA/H/C/002392/II/0088**

Bayer AG, Rapporteur: Jean-Michel Race

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**GONAL-f - Follitropin alfa -  
EMA/H/C/000071/II/0168/G**

Merck Europe B.V., Rapporteur: Patrick Vrijlandt

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**Herzuma - Trastuzumab -  
EMA/H/C/002575/II/0061/G**

Celltrion Healthcare Hungary Kft., Rapporteur:  
Jan Mueller-Berghaus

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**Hizentra - Human normal immunoglobulin -  
EMA/H/C/002127/II/0151/G**

CSL Behring GmbH, Rapporteur: Jan Mueller-  
Berghaus

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**Jivi - Damoctocog alfa pegol -  
EMA/H/C/004054/II/0031/G**

Bayer AG, Rapporteur: Thalia Marie Estrup  
Blicher

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**Ogivri - Trastuzumab -  
EMA/H/C/004916/II/0060**

Biosimilar Collaborations Ireland Limited,  
Rapporteur: Karin Janssen van Doorn

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**Origio A.R.T. Media - Gentamicin sulfate /  
Sargramostim / Heparin sodium / Insulin  
human - EMA/H/D/006090/II/0002**

Coopersurgical Inc., Rapporteur: Jayne Crowe

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**Privigen - Human normal immunoglobulin -  
EMA/H/C/000831/II/0202/G**

CSL Behring GmbH, Rapporteur: Jan Mueller-  
Berghaus

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**Ruxience - Rituximab -  
EMA/H/C/004696/II/0015**

Pfizer Europe MA EEIG, Rapporteur: Peter Mol

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**Saxenda - Liraglutide -  
EMA/H/C/003780/II/0038**

Novo Nordisk A/S, Rapporteur: Patrick Vrijlandt

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**TEPADINA - Thiotepa -  
EMA/H/C/001046/II/0050/G**

ADIENNE S.r.l. S.U., Rapporteur: Alexandre  
Moreau

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**Vyvgart - Efgartigimod alfa -  
EMA/H/C/005849/II/0016, Orphan**

Argenx, Rapporteur: Thalia Marie Estrup Blicher

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**WS2634  
Hexacima-  
EMA/H/C/002702/WS2634/0154  
Hexyon-  
EMA/H/C/002796/WS2634/0158**

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Sanofi Pasteur Europe, Duplicate, Duplicate of  
Hexacima, Lead Rapporteur: Jan Mueller-  
Berghaus

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**WS2651**

**Nilemdo-**

**EMA/H/C/004958/WS2651/0036**

**Nustendi-**

**EMA/H/C/004959/WS2651/0041**

Daiichi Sankyo Europe GmbH, Lead Rapporteur:  
Patrick Vrijlandt

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**WS2655/G**

**GONAL-f-**

**EMA/H/C/000071/WS2655/0167/G**

**Pergoveris-**

**EMA/H/C/000714/WS2655/0091/G**

Merck Europe B.V., Lead Rapporteur: Patrick  
Vrijlandt

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

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**Cresemba - Isavuconazole -**

**EMA/H/C/002734/II/0045, Orphan**

Basilea Pharmaceutica Deutschland GmbH,  
Rapporteur: Patrick Vrijlandt, "Update of section  
4.5 of the SmPC in order to revise the  
interactions table to improve guidance for  
health care professionals in relation to the co-  
administration of cyclophosphamide with  
isavuconazole based on literature and post-  
marketing data. In addition, the MAH took the  
opportunity to correct a mistake in section 4.5  
of the SmPC."

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**Cuprior - Trientine -**

**EMA/H/C/004005/II/0028**

Orphalan, Rapporteur: Jayne Crowe,  
"Submission of the final report from study  
TRIUMPH-2: Trientine dihydrochloride (Syprine  
capsules) vs. tetrahydrochloride (tablets): a  
Phase 1, single centre, randomised,  
interventional, open-label, 4-way crossover  
study in adult healthy male and female subjects  
to evaluate the pharmacokinetics and the safety  
and tolerability of 2 different oral formulations."

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**Glyxambi - Empagliflozin / Linagliptin -**

**EMA/H/C/003833/II/0057**

Boehringer Ingelheim International GmbH,  
Rapporteur: Patrick Vrijlandt, "Update of  
sections 4.2, 5.1 and 5.2 of the SmPC in order

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to update the information on paediatric population based on final results from study DINAMO 1218-0091 - A double-blind, randomised, placebo-controlled, parallel group trial to evaluate the efficacy and safety of empagliflozin and linagliptin over 26 weeks, with a double-blind active treatment safety extension period up to 52 weeks, in children and adolescents with type 2 diabetes mellitus. The Package Leaflet is updated accordingly. In addition, the MAH took the opportunity to update the list of local representatives in the Package Leaflet.”

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**Lysodren - Mitotane -**

**EMA/H/C/000521/II/0029/G**

HRA Pharma Rare Diseases, Rapporteur: Carolina Prieto Fernandez, “A grouped application consisting of two Type II variations: Update of sections 4.4, 4.5, 4.6, 4.8 and 4.9 of the SmPC in order to update the special warnings information and to update the pregnancy information, as well as, to add “Corticosteroid binding globulin increased” and “Thyroxin binding globulin increased” to the list of adverse drug reactions (ADRs) with frequency ‘Not Known’; based on clinical practice guidance and post-marketing data.

The Package Leaflet is updated accordingly. In addition, the MAH took the opportunity to bring the PI in line with the latest QRD template version 10.3, and to implement editorial changes to the SmPC.”

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**MenQuadfi - Meningococcal Group A, C, W and Y conjugate vaccine -**

**EMA/H/C/005084/II/0030**

Sanofi Pasteur, Rapporteur: Daniela Philadelphia, “Update of sections 4.5 and 5.1 of the SmPC in order to update immunogenicity and safety information based on final results from study MEQ00071; this is a parallel, multi-center, multinational, randomized, active-controlled phase 3b immunogenicity and safety study of a quadrivalent meningococcal conjugate vaccine versus Nimenrix, and when administered alone or concomitantly with 9vHPV and Tdap-IPV vaccines in healthy adolescents aged 10 to 17 years. In addition, the MAH took the opportunity to introduce minor updates to the PI and to update the list of local

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representatives in the Package Leaflet.”

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**NUVAXOVID - Covid-19 Vaccine  
(recombinant, adjuvanted) -  
EMA/H/C/005808/II/0062**

Novavax CZ, a.s., Rapporteur: Patrick Vrijlandt,  
“Submission of the final report from clinical  
study 2019nCoV-505 listed as a category 3  
study in the RMP. This is a Phase 2,  
Randomized, Observer-Blinded Study to  
Evaluate the Safety and Immunogenicity of a  
SARS CoV-2 Recombinant Spike Protein  
Nanoparticle Vaccine (SARS-CoV-2 rS) with  
Matrix M Adjuvant in People Living with HIV.”

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**Pombiliti - Cipaglucosidase alfa -  
EMA/H/C/005703/II/0010**

Amicus Therapeutics Europe Limited,  
Rapporteur: Patrick Vrijlandt, “Update of  
sections 4.6 and 5.3 of the SmPC in order to  
provide information regarding pre-implantation  
loss based on the reassessment of non-clinical  
data. In addition, the MAH took the opportunity  
to bring the PI in line with the latest QRD  
template version 10.3 and to introduce editorial  
changes.”

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**REKAMBYS - Rilpivirine -  
EMA/H/C/005060/II/0020**

Janssen-Cilag International N.V., Rapporteur:  
Patrick Vrijlandt, “Update of section 4.2 of the  
SmPC in order to update administration  
instructions to mitigate product leakage related  
to the correct use of the vial adapter, based on  
Human Factor studies. The Package Leaflet  
(Instructions for Use) is updated accordingly.”

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**RINVOQ - Upadacitinib -  
EMA/H/C/004760/II/0050**

AbbVie Deutschland GmbH & Co. KG,  
Rapporteur: Kristina Dunder, “Update of section  
5.1 of the SmPC in order to include long-term  
efficacy and safety information (up to week 104  
data) from study M19-944 (Study 1); this is a  
phase 3 randomized, placebo-controlled,  
double-blind program to evaluate efficacy and  
safety of upadacitinib in adult subjects with  
axial spondyloarthritis followed by a remission-  
withdrawal period.”

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**SARCLISA - Isatuximab -  
EMA/H/C/004977/II/0028**

Sanofi Winthrop Industrie, Rapporteur: Peter

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Mol, "Update of sections 4.2, 5.1 and 5.2 of the SmPC in order to update information on paediatric population based on final results from study ACT15378 (ISAKIDS). This was a Phase 2, single-arm, multicenter, open-label study evaluating the antitumor activity, safety, and PK of isatuximab in combination with standard salvage chemotherapies in pediatric participants with R/R ALL (including both T-ALL and B-ALL) and AML conducted in 3 separate cohorts. Male and female children from 28 days to less than 18 years of age with R/R T-ALL, B-ALL, or AML in first or second relapse were eligible. Participants under 2 years of age could only be enrolled after the dose reassessment is completed on the first 20 participants who were 2 to less than 18 years of age. The Package Leaflet is updated accordingly."

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**Tecentriq - Atezolizumab -  
EMA/H/C/004143/II/0084**

Roche Registration GmbH, Rapporteur: Aaron Sosa Mejia, "Update of section 4.8 of the SmPC in order to add 'hypophysitis' to the list of adverse drug reactions (ADRs) with frequency 'uncommon' based on interim results from study WO39391 (IMpassion030). This is a Phase III, randomised, open label study comparing atezolizumab in combination with adjuvant anthracycline/taxane-based chemotherapy versus chemotherapy alone in patients with operable triple-negative breast cancer; the Package Leaflet is updated accordingly. In addition, the MAH took the opportunity to implement editorial changes to the SmPC, Labelling and Package Leaflet."

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**Wegovy - Semaglutide -  
EMA/H/C/005422/II/0019**

Novo Nordisk A/S, Rapporteur: Patrick Vrijlandt, "Update of sections 4.1, 4.4, 4.8 and 5.1 in order to include information in patients with obesity-related HFpEF, with and without type 2 diabetes based on the final reports from studies EX9536-4665 STEP-HFpEF, EX9536-4773 STEP HFpEF-DM and EX9536-4388 SELECT. In addition, the MAH took this opportunity to introduce editorial changes to the PI."

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**Xevudy - Sotrovimab -  
EMA/H/C/005676/II/0027**

Glaxosmithkline Trading Services Limited,

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Rapporteur: Thalia Marie Estrup Blicher,  
"Update of section 5.1 of the SmPC with data on the in vitro activity of sotrovimab in a pseudotyped virus assay against the Omicron HV.1 and BA.2.86 spike variants (PC-23-0165) and the Omicron HK.3 spike variant (PC-23-0179) as well as data on the in vitro activity of sotrovimab in an authentic virus assay against the SARS-COV-2 EG.5.1 variant (PC-23-0176) based on the relevant pharmacology study reports."

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**XGEVA - Denosumab -  
EMA/H/C/002173/II/0084**

Amgen Europe B.V., Rapporteur: Kristina Dunder, "Submission of the final report from study 20140114, listed as a category 3 study in the RMP. This is a long-term safety follow up study, that was conducted to continue to follow subjects with GCTB who were treated in Study 20062004 for an additional 5 or more years of long-term safety follow up and to further evaluate denosumab treatment in subjects with GCTB."

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**B.6.10. CHMP-PRAC assessed procedures**

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**Ilumetri - Tildrakizumab -  
EMA/H/C/004514/II/0055**

Almirall S.A, Rapporteur: Jan Mueller-Berghaus,  
PRAC Rapporteur: Adam Przybylkowski

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**JCOVDEN - COVID-19 Vaccine Janssen  
(Ad26.COV2.S) -  
EMA/H/C/005737/II/0076**

Janssen-Cilag International N.V., Rapporteur: Christophe Focke, PRAC Rapporteur: Ulla Wändel Liminga, "Update of sections 4.5, 4.8 and 5.1 of the SmPC in order to update information regarding the co-administration of JCOVDEN with influenza vaccine based on the final report from study VAC31518COV3005 listed as a category 3 study in the RMP; this is a randomized, double-blind, Phase 3 study to evaluate safety, reactogenicity, and immunogenicity of co-administration of Ad26.COV2.S and influenza vaccines in healthy adults 18 years of age and older. The Package Leaflet is updated accordingly. Version 8.1 of the RMP has also been submitted."

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**Leqvio - Inclisiran -**

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**EMA/H/C/005333/II/0021**

Novartis Europharm Limited, Rapporteur:  
Martina Weise, PRAC Rapporteur: Kimmo  
Jaakkola, "Submission of the final report from  
study ORION-8 - A long-term extension trial of  
the Phase III lipid-lowering trials to assess the  
effect of long-term dosing of inclisiran given as  
subcutaneous injections in subjects with high  
cardiovascular risk and elevated LDL-C, listed as  
a category 3 study in the RMP. The RMP version  
3.0 has also been submitted."

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**Lupkynis - Voclosporin -****EMA/H/C/005256/II/0013**

Otsuka Pharmaceutical Netherlands B.V.,  
Rapporteur: Kristina Dunder, PRAC Rapporteur:  
Adam Przybylkowski, "Update of section 4.6 of  
the SmPC in order to updated breast-feeding  
information based on final results from study  
AUR-VCS-2021-04. This study is a single-  
center, open-label, Phase 1, lactation study to  
investigate the amount of voclosporin excreted  
in breast milk following a single oral dose of  
23.7 mg voclosporin in healthy, lactating,  
female volunteers. The Package Leaflet is  
updated accordingly. The updated RMP version  
5.0 has also been submitted."

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**Prevenar 20 - Pneumococcal  
polysaccharide conjugate vaccine (20-  
valent, adsorbed) -****EMA/H/C/005451/II/0023**

Pfizer Europe MA EEIG, Rapporteur: Daniela  
Philadelphly, PRAC Rapporteur: Jean-Michel  
Dogné, "Submission of the final current  
B7471015 study protocol, the Statistical  
Analysis Plan (SAP) and the final country  
feasibility assessment report for Apexnar. The  
RMP (version 5.0) is updated accordingly."

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**Spikevax - COVID-19 mRNA vaccine  
(nucleoside-modified) -****EMA/H/C/005791/II/0120**

Moderna Biotech Spain S.L., Rapporteur: Jan  
Mueller-Berghaus, PRAC Rapporteur: Marie  
Louise Schougaard Christiansen, "Submission of  
the final report from study mRNA-1273-P301  
(Phase 3, Randomized, Stratified, Observer-  
Blind, Placebo-Controlled Study to Evaluate the  
Efficacy, Safety, and Immunogenicity of mRNA-  
1273 SARS-CoV-2 Vaccine in Adults Aged 18  
Years and Older) listed as a category 3 study in

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the RMP. The RMP version 8.2 has also been submitted.”

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**TAKHZYRO - Lanadelumab -  
EMA/H/C/004806/II/0040, Orphan**

Takeda Pharmaceuticals International AG  
Ireland Branch, Rapporteur: Kristina Dunder,  
PRAC Rapporteur: Kirsti Villikka, “Update of  
section 4.4 of the SmPC in order to remove the  
information related to non-availability of clinical  
data on the use of lanadelumab in HAE patients  
with normal C1-INH activity, based on results  
from studies CASPIAN (SHP643-303) and  
CASPIAN OLE (TAK-743-3001). CASPIAN  
(SHP643-303) is a Phase 3, multicenter,  
randomized, placebo-controlled, double-blind  
study to evaluate the efficacy and safety of  
lanadelumab for prevention against acute  
attacks of NONHISTAMINERGIC ANGIOEDEMA  
with Normal C1 Inhibitor (C1-INH); and  
CASPIAN OLE (TAK-743-3001) is an open-label  
study to evaluate the long term safety and  
efficacy of lanadelumab for prevention against  
acute attacks of Nonhistaminergic Angioedema  
with Normal C1-Inhibitor (C1-INH).  
The RMP version 4.0 has also been submitted.  
In addition, the MAH took the opportunity to  
implement editorial changes to the SmPC and  
the Package Leaflet.”

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**Tecvayli - Teclistamab -  
EMA/H/C/005865/II/0009**

Janssen-Cilag International N.V., Rapporteur:  
Johanna Lähteenvuo, PRAC Rapporteur: Jana  
Lukacisinova, “Update of section 4.4 of the  
SmPC in order to update the warning on  
Progressive Multifocal Leukoencephalopathy  
(PML) based on a cumulative safety review. The  
Package Leaflet is updated accordingly. The RMP  
version 4.1 has also been submitted. In  
addition, the MAH took the opportunity to  
introduce minor updates to the PI and to update  
the list of local representatives in the Package  
Leaflet.”

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**ZTALMY - Ganaxolone -  
EMA/H/C/005825/II/0004/G, Orphan**

Marinus Pharmaceuticals Emerald Limited,  
Rapporteur: Peter Mol, PRAC Rapporteur: Adam  
Przybylkowski, “A grouped application  
comprised of 8 Type II variations as follows:  
1 Type II (C.I.4): Update of section 5.2 of the

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SmPC in order to update ganaxolone metabolite pattern at steady state based on re-analysis of 1042-TQT-1001 listed as a category 3 study in the RMP to evaluate the ganaxolone steady-state metabolite.

7 Type II (C.I.13): Submission of the final non-clinical study reports for the in vitro DDI potential and in vivo PK of the metabolite M17 listed as category 3 studies in the RMP.

The RMP version 1.2 has also been submitted. In addition, the MAH took the opportunity to introduce updates to the PI that reflect clarifications and typographical corrections, including to sections 4.2 and 4.4 of the SmPC.”

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**ZTALMY - Ganaxolone -**

**EMA/H/C/005825/II/0006, Orphan**

Marinus Pharmaceuticals Emerald Limited,  
Rapporteur: Peter Mol, PRAC Rapporteur: Adam Przybylkowski, “Update of section 5.1 of the SmPC in order to update open-label data based on the final report from study 1042-CDD-3001 OLE listed as a category 3 study in the RMP. This was the open-label portion of the pivotal study 1042-CDD-3001; a double-blind, randomized, placebo-controlled trial of adjunctive ganaxolone treatment in children and young adults with cyclin-dependent kinase-like 5 (CDKL5) deficiency disorder (CDD) followed by long-term open-label treatment. The RMP version 1.4 has also been submitted.”

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**WS2619/G**

**Invokana-**

**EMA/H/C/002649/WS2619/0066/G**

**Vokanamet-**

**EMA/H/C/002656/WS2619/0073/G**

Janssen-Cilag International N.V., Lead  
Rapporteur: Martina Weise, Lead PRAC  
Rapporteur: Martin Huber, “A grouped application consisting of two Type II variations, as follows:

C.I.4: Update of section 4.4 of the SmPC in order to amend an existing warning on Diabetic Ketoacidosis based on literature. The Package Leaflet is updated accordingly.

C.I.4: Update of sections 4.6 and 5.3 of the SmPC in order to update information on pregnancy based on literature.

The RMP version 11.1 has also been submitted.”

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### **B.6.11. PRAC assessed procedures**

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PRAC Led

**AJOVY - Fremanezumab -  
EMA/H/C/004833/II/0047**

TEVA GmbH, PRAC Rapporteur: Kirsti Villikka,  
PRAC-CHMP liaison: Outi Mäki-Ikola,  
"Submission of the final report from the PASS  
study TV48125-MH-50039 listed as a category 3  
study in the RMP. This is a long-term,  
prospective, observational study to evaluate the  
safety, including cardiovascular safety, of  
fremanezumab in patients with migraine in  
routine clinical practice. The RMP version 6.0  
has also been submitted."

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PRAC Led

**Bavencio - Avelumab -  
EMA/H/C/004338/II/0044/G**

Merck Europe B.V., PRAC Rapporteur: Karin  
Erneholm, PRAC-CHMP liaison: Thalia Marie  
Estrup Blicher, "Grouped application comprising  
four variations as follows:

Type II (C.I.11.b): To update Annex II and the  
RMP version 7.1 for Bavencio to change the  
classification of "safety in patients with  
autoimmune disease" to the important identified  
risk "other immune mediated adverse reactions"  
along with removal of the patient information  
brochure from the educational material,  
following the PRAC assessment report  
PSUSA/00010635/202303.

Type IA (A.6): To change ATC level name from  
Other antineoplastic agents, monoclonal  
antibodies to Antineoplastic agents, monoclonal  
antibodies, PD-1/PDL-1 (Programmed cell death  
protein 1/death ligand 1) inhibitors in section  
5.1 of the Summary of Product Characteristics  
(SmPC). The ATC code remains unchanged.

Type IA (C.I.z): To update the statement for  
"infusion-related reactions" in section 4.4 of the  
SmPC and to align terminology with the RMP for  
the term "immune-related" versus "immune-  
mediated".

Type IAIN (C.I.12): To remove from the Product  
Information the black symbol and explanatory  
statements for medicinal products subject to  
additional monitoring.

In addition, the MAH took this opportunity to  
introduce editorial changes and to bring the PI  
in line with the latest QRD template version

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PRAC Led

**COMIRNATY - COVID-19 mRNA vaccine  
(nucleoside-modified) -  
EMA/H/C/005735/II/0206/G**

BioNTech Manufacturing GmbH, PRAC

Rapporteur: Liana Martirosyan, PRAC-CHMP

liaison: Patrick Vrijlandt, “A grouped application  
comprised of 3 Type II variations as follows:

C.I.13: Submission of the final report from  
study C4591012 listed as a category 3 study in  
the RMP. This is a non-interventional Post-  
Emergency Use Authorisation active safety  
surveillance study among individuals in the  
Veteran’s Affairs health system receiving Pfizer-  
BioNTech Coronavirus Disease 2019 (COVID-19)  
vaccine. The RMP version 11.2 has also been  
submitted.

C.I.11.b: Submission of an updated RMP version  
11.2 in order to implement changes to an  
agreed post-authorisation study (C4591052  
protocol amendments 1 & 2) in the RMP, where  
there is an impact on the description of the  
study.

C.I.11.b: Submission of an updated RMP version  
11.2 in order to implement changes to an  
agreed post-authorisation study (C4591021  
protocol amendment 4) in the RMP, where there  
is an impact on the description of the study.

In addition, the MAH took the opportunity to  
update the milestones for the two studies  
C4591022 and C4591051 in the RMP.”

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PRAC Led

**DaTSCAN - Ioflupane (123I) -  
EMA/H/C/000266/II/0067**

GE Healthcare B.V., Rapporteur: Alexandre  
Moreau, PRAC Rapporteur: Tiphaine Vaillant,  
PRAC-CHMP liaison: Alexandre Moreau, “To  
update sections 4.4 and 4.5 of the SmPC and  
section 2 of the Package Leaflet to implement  
the recommendation of the PRAC following the  
PSUSA procedure  
(EMA/H/C/PSUSA/00001767/202207). In  
addition, the MAH took the opportunity to  
update the list of local representatives in the  
Package Leaflet.”

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PRAC Led

**Flixabi - Infliximab -  
EMA/H/C/004020/II/0084/G**

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Samsung Bioepis NL B.V., PRAC Rapporteur:  
Mari Thorn, PRAC-CHMP liaison: Kristina  
Dunder, "A grouped application comprised of  
two Type II variations as follows:  
C.I.13: Submission of the final report from  
study CEDUR listed as a category 3 study in the  
RMP. This is a nationwide German IBD registry  
to describe the long-term effectiveness of  
treatment with IBD therapies such as drug  
survival, effectiveness, side effects of treatment  
combination, and disease activity achieved.  
C.I.13: Submission of the final report from  
study CREDIT listed as a category 3 study in the  
RMP. This is a Czech Register of IBD Patients on  
Biological Therapy to monitor effectiveness of  
total population of IBD patients on biological  
medication in the Czech Republic and regular  
analytical evaluation of the effectiveness.  
The RMP version 13.0 has also been submitted."

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PRAC Led

**Humira - Adalimumab -  
EMA/H/C/000481/II/0218**

AbbVie Deutschland GmbH & Co. KG, PRAC  
Rapporteur: Mari Thorn, PRAC-CHMP liaison:  
Kristina Dunder, "Submission of the final report  
for study P10-023 listed as a category 3 study  
in the RMP. This is a 10-year, post-marketing,  
observational registry to assess long term safety  
of Humira (adalimumab) in adult patients with  
chronic plaque psoriasis (Ps)."

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PRAC Led

**MenQuadfi - Meningococcal Group A, C, W  
and Y conjugate vaccine -  
EMA/H/C/005084/II/0031**

Sanofi Pasteur, PRAC Rapporteur: Jean-Michel  
Dogné, PRAC-CHMP liaison: Karin Janssen van  
Doorn, "Update of section 4.8 of the SmPC in  
order to add 'Hypersensitivity including  
anaphylaxis' to the list of adverse drug reactions  
(ADRs) with frequency not known, based on a  
cumulative review of cases of  
hypersensitivity/allergic reaction (including  
anaphylaxis) following the request by PRAC in  
the Assessment Report for  
PSUSA/00010044/202304. The Package Leaflet  
is updated accordingly."

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PRAC Led

**RAYVOW - Lasmiditan -  
EMA/H/C/005332/II/0007**

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Eli Lilly Nederland B.V., PRAC Rapporteur: Anna Mareková, PRAC-CHMP liaison: Frantisek Drafi, "Submission of the final report from study H8H-MC-B005, listed as a category 3 study in the RMP (MEA/003). This is a Real-World Observational Study to Assess Drug Utilisation Patterns in the US Among Migraine Patients Treated with Lasmiditan. The RMP version 2.1 is submitted alongside the final study report."

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PRAC Led

**VEYVONDI - Vonicog alfa -**

**EMA/H/C/004454/II/0033**

Baxalta Innovations GmbH, PRAC Rapporteur: Mari Thorn, PRAC-CHMP liaison: Kristina Dunder, "Submission of the final report from study TAK-577-4005 listed as a category 3 PASS in the RMP. This is a non-interventional retrospective cohort study that evaluated the safety of VEYVONDI in real-world clinical practice. The RMP version 5.0 has also been submitted."

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PRAC Led

**Vyndaqel - Tafamidis -**

**EMA/H/C/002294/II/0091/G, Orphan**

Pfizer Europe MA EEIG, PRAC Rapporteur: Tiphaine Vaillant, PRAC-CHMP liaison: Jean-Michel Race, "A grouped application comprised of two Type II Variations, as follows:  
C.I.4: Update of the Annex II based on final results from study B3461001 (THAOS) listed as a category 3 study in the RMP. This is a global, multi-center, longitudinal, observational survey of patients with documented transthyretin gene mutations or wild-type transthyretin amyloidosis.  
C.I.13: Submission of the final report from study B3461042 listed as a category 3 study in the RMP. This is a post-marketing safety surveillance study in Japanese patients with ATTR-PN.  
The RMP version 10.0 has also been submitted. In addition, the MAH took the opportunity to provide B3461028 Clinical Study Report (CSR) Errata."

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**B.6.12. CHMP-CAT assessed procedures**

**B.6.13. CHMP-PRAC-CAT assessed procedures**

**B.6.14. PRAC assessed ATMP procedures**

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

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**WS2635**

**Hexacima-**

**EMA/H/C/002702/WS2635/0155**

**Hexyon-**

**EMA/H/C/002796/WS2635/0159**

**MenQuadfi-**

**EMA/H/C/005084/WS2635/0032**

Sanofi Pasteur, Lead Rapporteur: Jan Mueller-Berghaus

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**WS2644/G**

**Entresto-**

**Neparvis-**

Novartis Europharm Limited, Lead Rapporteur:  
Patrick Vrijlandt

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**WS2661**

**Mirapexin-**

**EMA/H/C/000134/WS2661/0108**

**Sifrol-EMA/H/C/000133/WS2661/0099**

Boehringer Ingelheim International GmbH, Lead  
Rapporteur: Thalia Marie Estrup Blicher

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**WS2666/G**

**Ongentys-**

**EMA/H/C/002790/WS2666/0065/G**

**Ontilyv-**

**EMA/H/C/005782/WS2666/0020/G**

Bial - Portela & Ca, S.A., Lead Rapporteur:  
Martina Weise

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

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

### **B.7.2. Monthly Line listing for Type I variations**

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

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

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

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

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

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

## **E. Annex E - EMA CERTIFICATION OF PLASMA MASTER FILES**

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

### **E.1. PMF Certification Dossiers:**

#### **E.1.1. Annual Update**

#### **E.1.2. Variations:**

#### **E.1.3. Initial PMF Certification:**

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

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

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## **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):**

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

### **G.2. PRIME**

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

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