13 May 2024
EMA/PRAC/151850/2024
Human Medicines Division

Pharmacovigilance Risk Assessment Committee (PRAC)

Draft agenda for the meeting on 13 - 16 May 2024

Chair: Sabine Straus – Vice-Chair: Martin Huber

13 May 2024, 10:30 – 19:30, via teleconference

14 May 2024, 08:30 – 19:30, via teleconference

15 May 2024, 08:30 – 19:30, via teleconference

16 May 2024, 08:30 – 16:00, via teleconference

Organisational, regulatory and methodological matters (ORGAM)

30 May 2024, 09:00 – 12:00, via teleconference

Health and safety information

In accordance with the Agency's health and safety policy, delegates are to be briefed on health, safety and emergency information and procedures prior to the start of the meeting.

Disclaimers

Some of the information contained in this agenda is considered commercially confidential or sensitive and therefore not disclosed. With regard to intended therapeutic indications or procedure scopes listed against products, it must be noted that these may not reflect the full wording proposed by applicants and may also change during the course of the review. Additional details on some of these procedures will be published in the PRAC meeting highlights once the procedures are finalised.

Of note, this agenda is a working document primarily designed for PRAC members and the work the Committee undertakes.

Note on access to documents

Some documents mentioned in the agenda cannot be released at present following a request for access to documents within the framework of Regulation (EC) No 1049/2001 as they are subject to on-going procedures for which a final decision has not yet been adopted. They will become public when adopted or considered public according to the principles stated in the Agency policy on access to documents (EMA/127362/2006 Rev.1).

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1. Introduction

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

Pre-meeting list of participants and restrictions in relation to declarations of interests applicable to the items of the agenda for the PRAC plenary session to be held 13-16 May 2024. See May 2024 minutes (to be published post June 2024 PRAC meeting).

1.2. Agenda of the meeting on 13-16 May 2024

Action: For adoption

1.3. Minutes of the previous meeting on 08-11 April 2024

Action: For adoption

2. EU referral procedures for safety reasons: urgent EU procedures

2.1. Newly triggered procedures

None

2.2. Ongoing procedures

None

2.3. Procedures for finalisation

None

3. EU referral procedures for safety reasons: other EU referral procedures

3.1. Newly triggered procedures

None

3.2. Ongoing procedures

None

3.3. Procedures for finalisation

3.3.1. Hydroxyprogesterone (NAP) - EMEA/H/A-31/1528

Applicant(s): various

PRAC Rapporteur: Amelia Cupelli; PRAC Co-rapporteur: Nathalie Gault

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

Action: For adoption of a recommendation to CMDh

3.4. Re-examination procedures[1]

None

3.5. Others

None

4. Signals assessment and prioritisation[2]

4.1. New signals detected from EU spontaneous reporting systems and/or other sources

4.1.1. Adagrasib – KRAZATI (CAP)

Applicant: Bristol-Myers Squibb Pharma EEIG

PRAC Rapporteur: Kimmo Jaakkola

Scope: Signal of febrile neutropenia

Action: For adoption of PRAC recommendation

EPITT 20080– New Signal

Lead Member State(s): FI

4.1.2. Eptinezumab – VYEPTI (CAP); erenumab – AIMOVIG (CAP); fremanezumab – AJOVY (CAP); galcanezumab – EMGALITY (CAP)

Applicant(s): H. Lundbeck A/S (Vyepti), Novartis Europharm Limited (Aimovig), TEVA GmbH (Ajovy), Eli Lilly Nederland B.V. (Emgality)

PRAC Rapporteur: to be appointed

[1] Re-examination of PRAC recommendation under Article 32 of Directive 2001/83/EC
[2] Each signal refers to a substance or therapeutic class. The route of marketing authorisation is indicated in brackets (CAP for Centrally Authorised Products; NAP for Nationally Authorised Products including products authorised via Mutual Recognition Procedures and Decentralised Procedure). Product names are listed for reference Centrally Authorised Products (CAP) only. PRAC recommendations will specify the products concerned in case of any regulatory action required

Scope: Signal of insomnia

Action: For adoption of PRAC recommendation

EPITT 20077 – New signal

Lead Member State(s): FI, NL

4.1.3. Erenumab – AIMOVIG (CAP)

Applicant: Novartis Europharm Limited

PRAC Rapporteur: Kirsti Villikka

Scope: Signal of hypertension

Action: For adoption of PRAC recommendation

EPITT 20081– New signal

Lead Member State(s): FI

4.2. Signals follow-up and prioritisation

4.2.1. Aflibercept – EYLEA (CAP) - EMEA/H/C/002392/SDA/023, YESAFILI (CAP) - EMEA/H/C/006022/SDA/002

Applicant(s): Bayer AG (Eylea), Biosimilar Collaborations Ireland Limited (Yesafili)

PRAC Rapporteur: Nathalie Gault

Scope: Signal of nephropathy toxic after intravitreal administration

Action: For adoption of PRAC recommendation

EPITT 20024 – Follow-up to January 2024

4.2.2. Baricitinib – OLUMIANT (CAP) - EMEA/H/C/004085/SDA/018

Applicant: Eli Lilly Nederland B.V.

PRAC Rapporteur: Adam Przybylkowski

Scope: Signal of hypoglycaemia in diabetic patients

Action: For adoption of PRAC recommendation

EPITT 20038 – Follow-up to January 2024

4.2.3. Clobazam (NAP)

Applicant(s): various

PRAC Rapporteur: Kimmo Jaakkola

Scope: Signal of drug rash with eosinophilia and systemic symptoms (DRESS)

Action: For adoption of PRAC recommendation

EPITT 20041 – Follow-up to January 2024

4.2.4. Dabrafenib – TAFINLAR (CAP) - EMEA/H/C/002604/SDA/024, FINLEE (CAP) - EMEA/H/C/005885/SDA/003; Trametinib – MEKINIST (CAP) - EMEA/H/C/002643/SDA/019, SPEXOTRAS (CAP) - EMEA/H/C/005886/SDA/002

Applicant: Novartis Europharm Limited

PRAC Rapporteur: David Olsen

Scope: Signal of acute febrile neutrophilic dermatosis

Action: For adoption of PRAC recommendation

EPITT 20022 – Follow-up to January 2024

4.2.5. Ivacaftor, tezacaftor, elexacaftor - KAFTRIO (CAP) – EMEA/H/C/005269/SDA/017; Ivacaftor - KALYDECO (CAP) - EMEA/H/C/002494/SDA/034; Lumacaftor, ivacaftor - ORKAMBI (CAP) - EMEA/H/C/003954/SDA/018; Tezacaftor, ivacaftor - SYMKEVI (CAP) - EMEA/H/C/004682/SDA/009

Applicant: Vertex Pharmaceuticals (Ireland) Limited

PRAC Rapporteur: Martin Huber

Scope: Signal of intracranial pressure increased

Action: For adoption of PRAC recommendation

EPITT 20000 – Follow-up to December 2023[3]

4.2.6. Manidipine (NAP)

Applicant(s): various

PRAC Rapporteur: Amelia Cupelli

Scope: Signal of ascites

Action: For adoption of PRAC recommendation

EPITT 20026 – Follow-up to January 2024

4.2.7. Propofol (NAP)

Applicant(s): various

PRAC Rapporteur: Karen Pernille Harg

Scope: Signal of hepatic failure

Action: For adoption of PRAC recommendation

EPITT 20020 – Follow-up to January 2024

[3] Held 27-30 November 2023

4.3. Variation procedure(s) resulting from signal evaluation

4.3.1. Brolucizumab - BEOVU (CAP) - EMEA/H/C/004913/II/0028

Applicant: Novartis Europharm Limited

PRAC Rapporteur: Gabriele Maurer

Scope: Update of section 4.8 of the SmPC in order to add 'scleritis' to the list of adverse drug reactions (ADRs) with frequency 'not known', following the recommendation by PRAC in the outcome for the signal assessment of scleritis. The package leaflet is updated accordingly

Action: For adoption of PRAC Assessment Report

5. Risk management plans (RMPs)

5.1. Medicines in the pre-authorisation phase

5.1.1. Avacincaptad pegol (CAP MAA) - EMEA/H/C/006153

Scope (pre D-180 phase): Treatment of adults with geographic atrophy (GA) secondary to age-related macular degeneration (AMD)

Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.1.2. Ciclosporin (CAP MAA) - EMEA/H/C/006250

Scope (pre D-180 phase): Treatment of dry eye disease in adult patients

Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.1.3. Delgocitinib (CAP MAA) - EMEA/H/C/006109

Scope (pre D-180 phase): Treatment of moderate to severe chronic hand eczema (CHE)

Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.1.4. Elafibranor (CAP MAA) - EMEA/H/C/006231, Orphan

Applicant: Ipsen Pharma

PRAC Rapporteur: Rugile Pilviniene; PRAC Co-rapporteur: Liana Martirosyan

Scope (pre D-180 phase): Treatment of primary biliary cholangitis (PBC)

Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.1.5. Influenza vaccine (live attenuated, nasal) (CAP MAA) - EMEA/H/C/006514

Scope (pre-opinion phase): Active immunisation to prevent influenza disease

Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.1.6. Leriglitazone - NEZGLYAL (CAP MAA) - EMEA/H/C/005757, Orphan

Applicant: Minoryx Therapeutics S.L.

Scope (under re-examination): Treatment of cerebral progression and myelopathy in male patients with adrenoleukodystrophy (ALD)

Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.1.7. Lutetium (177Lu) chloride (CAP MAA) - EMEA/H/C/005882

Scope (pre D-180 phase): Radiolabelling of carrier molecules, which have been specifically developed for radiolabelling with this radionuclide

Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.1.8. Trastuzumab (CAP MAA) - EMEA/H/C/006252

Scope (pre D-180 phase): Treatment of adult patients with HER2 positive metastatic breast cancer (MBC) and HER2 positive early breast cancer (EBC)

Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.1.9. Ustekinumab (CAP MAA) - EMEA/H/C/005805

Scope (pre D-180 phase): Treatment of moderate to severe plaque psoriasis, active psoriatic arthritis, Crohn’s disease and ulcerative colitis

Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.1.10. Ustekinumab (CAP MAA) - EMEA/H/C/006544

Scope : Treatment of Crohn’s disease and ulcerative colitis, moderate to severe plaque psoriasis, active and psoriatic arthritis

Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.1.11. Zapomeran (CAP MAA) - EMEA/H/C/006207

Scope (pre D-180 phase): Active immunisation to prevent COVID-19

Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.2. Medicines in the post-authorisation phase – PRAC-led procedures

5.2.1. Amlodipine, valsartan - AMLODIPINE-VALSARTAN MYLAN (CAP) - EMEA/H/C/004037/II/0021

Applicant: Mylan Pharmaceuticals Limited

PRAC Rapporteur: Karin Erneholm

Scope: Submission of an updated RMP version 4.0 in order to align the safety concerns with the latest version of RMP for amlodipine/valsartan available in the public domain and to bring the RMP in line with the latest RMP template

Action: For adoption of PRAC Assessment Report

5.2.2. Colesevelam - CHOLESTAGEL (CAP) - EMEA/H/C/000512/II/0053

Applicant: CHEPLAPHARM Arzneimittel GmbH

PRAC Rapporteur: Bianca Mulder

Scope: Submission of an updated RMP version 2.0 in order to remove important identified and potential risks, as well as missing information to bring it in line with GVP module V. Additionally, epidemiological data on indication and target population, clinical data and postmarking exposure data was updated

Action: For adoption of PRAC Assessment Report

5.2.3. Fluticasone furoate - AVAMYS (CAP) - EMEA/H/C/000770/II/0051/G

Applicant: GlaxoSmithKline (Ireland) Limited

PRAC Rapporteur: Adam Przybylkowski

Scope: Grouped application comprising two type II variations as follows:
C.I.11.b – Submission of an updated RMP version 12 in order to remove headache, nasal events (including: epistaxis, nasal ulceration, nasal septum perforation and other nasal events), hypersensitivity, cataract and glaucoma as important identified risks; to remove taste and smell disorders, pyrexia, systemic corticosteroids effect: adrenal suppression, Systemic corticosteroid effect: growth retardation, psychiatric effects as important potential risks and to remove use in pregnancy and lactation, off-label use (sinusitis and children < 6 years of age) as missing information.
C.I.11.b – Submission of an updated RMP version 12 in order to remove targeted follow up questionnaires.
In addition, the MAH took this opportunity to align the RMP template with GVP Module V Revision 2

Action: For adoption of PRAC Assessment Report

5.2.4. Infliximab - REMICADE (CAP) - EMEA/H/C/000240/II/0247

Applicant: Janssen Biologics B.V.

PRAC Rapporteur: Mari Thorn

Scope: Submission of an updated RMP version 22.1 in order to remove reference to the immunogenicity substudy as part of protocol REMICADEPIB4002 in Part III. The MAH proposes to discontinue the Dutch portion of the immunogenicity substudy, which is part of protocol REMICADEPIB4002

Action: For adoption of PRAC Assessment Report

5.2.5. Infliximab - ZESSLY (CAP) - EMEA/H/C/004647/II/0033

Applicant: Sandoz GmbH

PRAC Rapporteur: Mari Thorn

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

Action: For adoption of PRAC Assessment Report

5.2.6. Pegcetacoplan - ASPAVELI (CAP) - EMEA/H/C/005553/II/0018, Orphan

Applicant: Swedish Orphan Biovitrum AB (publ)

PRAC Rapporteur: Kimmo Jaakkola

Scope: Submission of an updated RMP version 2.1 in order to revise the category 3 PASS Sobi.PEGCET-301 and Sobi.PEGCET-302

Action: For adoption of PRAC Assessment Report

5.2.7. Prasugrel - EFIENT (CAP) - EMEA/H/C/000984/II/0037

Applicant: Substipharm

PRAC Rapporteur: Marie Louise Schougaard Christiansen

Scope: Submission of an updated RMP version 13 in order to remove of a region-specific additional risk-minimisation activity following previous PSUSA procedure (EMEA/H/C/PSUSA/00002499/202102), as well as to align content and format with new requirements according to GVP Module V Rev. 2. In addition, the MAH took the opportunity to update Annex II of the product information and to update the list of local representatives in the package leaflet

Action: For adoption of PRAC Assessment Report

5.2.8. Sacituzumab govitecan - TRODELVY (CAP) - EMEA/H/C/005182/II/0031

Applicant: Gilead Sciences Ireland UC

PRAC Rapporteur: Bianca Mulder

Scope: Submission of an updated RMP version 3.1 in order to propose the removal of some safety concerns and the extension of remaining study milestones to date for category 3 study IMMU-132-15

Action: For adoption of PRAC Assessment Report

5.2.9. Telmisartan - KINZALMONO (CAP) - EMEA/H/C/000211/WS2577/0120; MICARDIS (CAP) - EMEA/H/C/000209/WS2577/0129; PRITOR (CAP) - EMEA/H/C/000210/WS2577/0133

Applicant: Boehringer Ingelheim International GmbH

PRAC Rapporteur: Amelia Cupelli

Scope: Submission of an updated RMP version 6.1 in order to implement an overall update regarding safety concerns based on literature and post marketing data; and to adapt the RMP to the current RMP format (Rev 2.0.1), in line with GVP Module V, Revision 2

Action: For adoption of PRAC Assessment Report

5.2.10. Telmisartan, hydrochlorothiazide - KINZALKOMB (CAP) - EMEA/H/C/000415/WS2611/0123; MICARDISPLUS (CAP) - EMEA/H/C/000413/WS2611/0130; PRITORPLUS (CAP) - EMEA/H/C/000414/WS2611/0133

Applicant: Boehringer Ingelheim International GmbH

PRAC Rapporteur: Amelia Cupelli

Scope: Submission of an updated RMP version 9.1 for MicardisPlus, PritorPlus and Kinzalkomb in order to remove all important identified and potential risks from the list of safety concerns and to adapt the RMP to the current RMP format (Rev 2.0.1), in line with GVP Module V, Revision 2

Action: For adoption of PRAC Assessment Report

5.3. Medicines in the post-authorisation phase – CHMP-led procedures

5.3.1. Afamelanotide - SCENESSE (CAP) - EMEA/H/C/002548/II/0052, Orphan

Applicant: Clinuvel Europe Limited

PRAC Rapporteur: Martin Huber

Scope: Update of section 4.2 of the SmPC in order to update the posology recommendations by removing the current recommendation of a maximum of four implants per year, based on a literature review and analysis of safety data. The package leaflet is updated accordingly. The RMP version 9.8 has also been submitted. In addition, the MAH took the opportunity to introduce a minor editorial change to the product information

Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.2. Aflibercept - EYLEA (CAP) - EMEA/H/C/002392/II/0090

Applicant: Bayer AG

PRAC Rapporteur: Nathalie Gault

Scope: Update of sections 4.8 and 5.1 of the SmPC in order to update safety and clinical information based on results from studies PULSAR (20968) and PHOTON (21091). PULSAR (20968) is an ongoing pivotal Phase 3 study to investigate the efficacy and safety

of HD aflibercept at treatment intervals of 12 weeks and longer for indication neovascular age-related macular degeneration (nAMD).
PHOTON (21091) is an ongoing pivotal phase 2/3 study to investigate the efficacy and safety of HD aflibercept at treatment intervals of 12 weeks and longer for indication diabetic macular oedema. The package leaflet is updated accordingly. The RMP version 34.1 has also been submitted. In addition, the MAH took the opportunity to implement an editorial update in section 6.6 of the SmPC to align the text with other similar products

Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.3. Alpelisib - PIQRAY (CAP) - EMEA/H/C/004804/II/0022/G

Applicant: Novartis Europharm Limited

PRAC Rapporteur: Bianca Mulder

Scope: Grouped application comprising two type II variations (C.I.4) as follows:
- Update of sections 4.2, 4.4, 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, 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

Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.4. Amivantamab - RYBREVANT (CAP) - EMEA/H/C/005454/II/0013

Applicant: Janssen-Cilag International N.V.

PRAC Rapporteur: Gabriele Maurer

Scope: Extension of indication to include amivantamab in combination with lazertinib for the first-line treatment of adult patients with advanced non-small cell lung cancer (NSCLC) with EGFR exon 19 deletions or exon 21 L858R substitution mutations (EGFRm NSCLC), based on results from study 73841937NSC3003 (MARIPOSA). This is a randomised, open-label, phase 3 study that compares the efficacy and safety of the combination of amivantamab and lazertinib (Arm A) versus osimertinib monotherapy (Arm B) and lazertinib monotherapy (Arm C) in participants with EGFRm NSCLC. The primary objective of the MARIPOSA study was to assess the efficacy of the combination of amivantamab and lazertinib (Arm A), compared with osimertinib (Arm B), as measured by PFS assessed by BICR in adult participants with EGFRm NSCLC. As a consequence, sections 4.1, 4.2, 4.4, 4.8, 4.9, 5.1, 5.2, 6.6 and 9 of the SmPC are updated. The package leaflet is updated in accordance. Version 3.3 of the EU RMP has also been submitted. As part of the application the MAH is requesting a 1-year extension of the market protection

Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.5. Apixaban - ELIQUIS (CAP) - EMEA/H/C/002148/X/0089/G

Applicant: Bristol-Myers Squibb / Pfizer EEIG

PRAC Rapporteur: Bianca Mulder

Scope: Extension application to:
1) Introduce a new pharmaceutical form (granules in single-dose container) associated with a new strength (0.15 mg).
2) Introduce a new pharmaceutical form (coated granules in sachet) associated with 3 new strengths (0.5 mg, 1.5 mg and 2 mg);
The above two line extensions are grouped with a type II - C.I.6.a variation:
Extension of indication to include the treatment of venous thromboembolism (VTE) and prevention of recurrent VTE in paediatric patients from 28 days to less than 18 years of age for Eliquis (all strengths), based on a pre-specified interim analysis from study CV185325; this is an open-label, multi-centre, randomised, active controlled trial to provide pharmacokinetic (PK) data and data on anti-Xa activity to support the extrapolation of efficacy to children, to evaluate safety and efficacy of apixaban in children who require anticoagulation for a venous thromboembolism; as a consequence, sections 4.1, 4.2, 4.4, 4.5, 4.8, 4.9, 5.1 and 5.2 of the SmPCs are updated. The package leaflet and Annex II are updated in accordance. Version 21.0 of the RMP has also been submitted

Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.6. Blinatumomab - BLINCYTO (CAP) - EMEA/H/C/003731/II/0056, Orphan

Applicant: Amgen Europe B.V.

PRAC Rapporteur: Jana Lukacisinova

Scope: Extension of indication to include treatment as part of consolidation therapy for the treatment of patients with Philadelphia chromosome negative CD19 positive B-cell precursor acute lymphoblastic leukemia (ALL) for BLINCYTO. The proposed indication is supported by efficacy data from Studies E1910, 20120215, and AALL1331, safety data for Studies E1910, 20120215, AALL1331, MT103-202, and MT103-203, and pharmacokinetic data for studies 20120215, AALL1331, MT103-202, MT103-203, and 20190360. 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 18.0 of the RMP has also been submitted

Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.7. Brolucizumab - BEOVU (CAP) - EMEA/H/C/004913/II/0029

Applicant: Novartis Europharm Limited

PRAC Rapporteur: Gabriele Maurer

Scope: Update of sections 4.2 and 5.1 of the SmPC in order to include information on maintenance treatment and to update efficacy and safety information based on final results from studies CRTH258A2303 (TALON) and CRTH258A2303E1 (TALON Extension). TALON is a 64-week, two-arm, randomised, double-masked, phase IIIb study assessing the efficacy and safety of brolucizumab 6 mg compared to aflibercept 2 mg in a treat-to-control regimen in patients with neovascular age-related macular degeneration. TALON Extension is a 56-week phase IIIb/IV, open-label, one-arm extension study to assess the efficacy and safety

of brolucizumab 6 mg in a treat-to-control regimen with maximum treatment intervals up to 20 weeks for the treatment of subjects with neovascular age-related macular degeneration who have completed the CRTH258A2303 (TALON) study.
The package leaflet is updated accordingly. The RMP version 12.0 has also been submitted

Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.8. Casirivimab, imdevimab - RONAPREVE (CAP) - EMEA/H/C/005814/II/0017

Applicant: Roche Registration GmbH

PRAC Rapporteur: Ulla Wändel Liminga

Scope: Extension of indication to include treatment of paediatric patients from 2 to less than 12 years old, weighing at least 10kg, who do not require supplemental oxygen and who are at increased risk of progression to severe COVID-19 for Ronapreve, based on final results from study COV-2067; this was a seamless, adaptive, phase 3, randomised, double-blinded, placebo-controlled, multi-centre study to evaluate the efficacy, safety, and tolerability of casirivimab+imdevimab combination therapy in paediatric and adult outpatients with mild to moderate COVID-19. As a consequence, sections 4.1, 4.2, 4.8, 5.1, 5.2 and 6.6 of the SmPC are updated. The package leaflet is updated in accordance. Version 4.0 of the RMP has also been submitted

Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.9. Defatted powder of Arachis hypogaea L., semen (peanuts) - PALFORZIA (CAP) - EMEA/H/C/004917/II/0014/G

Applicant: Aimmune Therapeutics Ireland Limited

PRAC Rapporteur: Kirsti Villikka

Scope: Grouped variation consisting of:
C.I.6.a: Extension of indication to include treatment of patients 1 to 3 years old for PALFORZIA, based on final results from study ARC005; this is a phase 3 randomised, double-blind, placebo-controlled Peanut Oral Immunotherapy Study of Early Intervention for Desensitization (POSEIDON) to evaluate the safety and efficacy of peanut powder in terms of superiority of placebo in children of 1 year to less than 4 years of age with peanut allergy. As a consequence, sections 4.1, 4.2, 4.4, 4.8, 5.1, 6.5 and 8 of the SmPC are updated. The package leaflet and labelling were updated accordingly. Version 1.1 of the RMP has also been submitted. In addition, the marketing authorisation holder (MAH) took the opportunity to implement editorial changes to the SmPC 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.

B.II.e.5.a: Introduction of a new pack-size of 16 capsules of 1 mg (Level 0) in blisters for PALFORZIA, 1 mg, oral powder in capsules for opening.
Due to the lack of a suitable pack-size for the up-dosing phase for patients 1 to 3 years old, a new pack size Level 0 for the up-dosing phase will be introduced. Labelling was updated accordingly. In addition, the MAH took the opportunity to update module 3.2.P.3.1 to take out the EU importation site (editorial change)

Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.10. Dupilumab - DUPIXENT (CAP) - EMEA/H/C/004390/II/0079

Applicant: Sanofi Winthrop Industrie

PRAC Rapporteur: Kimmo Jaakkola

Scope: Extension of indication for DUPIXENT to include treatment of adults as add-on maintenance treatment for uncontrolled chronic obstructive pulmonary disease (COPD) with type 2 inflammation on triple therapy or double therapy if inhaled corticosteroids (ICS) are contraindicated, based on final results from study EFC15804 (BOREAS); this is a phase 3, randomised, double blind, placebo-controlled, multi-centre, parallel group, 52-week study to assess the efficacy, safety and tolerability of dupilumab in patients with moderate-to-severe COPD with type 2 inflammation. 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 10.0 of the RMP has also been submitted

Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.11. Dupilumab - DUPIXENT (CAP) - EMEA/H/C/004390/II/0083

Applicant: Sanofi Winthrop Industrie

PRAC Rapporteur: Kimmo Jaakkola

Scope: Extension of indication to include treatment of moderate to severe chronic spontaneous urticaria (CSU) in adults and adolescents 12 years and older, who are symptomatic despite treatment with H1 antihistamines and who are intolerant to or inadequately controlled by anti-IgE therapy for Dupixent, based on the results from studies EFC16461 (CUPID) study B (pivotal) and study A (supportive); EFC16461 Study B was a 24-week, double-blind, randomised, placebo-controlled study to evaluate the efficacy and safety of dupilumab in adult and adolescent participants with CSU who remained symptomatic despite the use of H1-antihistamine and who were intolerant or incomplete responders to omalizumab and EFC16461 Study A was a 24-week, double-blind, randomised, placebo-controlled study to evaluate the efficacy and safety of dupilumab in participants with CSU who remained symptomatic despite the use of H1-antihistamine and who were naïve to omalizumab. 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 11.0 of the RMP has also been submitted

Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.12. Encorafenib - BRAFTOVI (CAP) - EMEA/H/C/004580/WS2538/0034; Binimetinib - MEKTOVI (CAP) - EMEA/H/C/004579/WS2538/0030

Applicant: Pierre Fabre Medicament

PRAC Rapporteur: Rugile Pilviniene

Scope: Extension of indication to include binimetinib in combination with encorafenib for the treatment of adult patients with advanced non-small cell lung cancer (NSCLC) with a BRAF V600 mutation for MEKTOVI and BRAFTOVI based on results from study PHAROS (study ARRAY-818-202) at the primary completion date; this is a phase II, open-label, multicentre, non-comparative study (interventional). As a consequence, sections 4.1, 4.4, 4.8, 5.1, 5.2, 9 and 10 of the SmPC are updated. The package leaflet is updated in accordance. Version

2.1 of the RMP has also been submitted. As part of the application, the MAH is requesting a 1-year extension of the market protection for MEKTOVI

Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.13. Empagliflozin, metformin - SYNJARDY (CAP) - EMEA/H/C/003770/II/0078

Applicant: Boehringer Ingelheim International GmbH

PRAC Rapporteur: Maria del Pilar Rayon

Scope: Extension of indication to include the treatment of children aged 10 years and above with type 2 diabetes for Synjardy, based on the final results from study 1218-0091 (DINAMO) - 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. As a consequence, sections 4.1, 4.2, 4.4, 4.5, 4.8, 5.1, and 5.2 of the SmPC are updated. The package leaflet is updated in accordance. Version 16.0 of the RMP has also been submitted. In addition, the marketing authorisation holder (MAH) took the opportunity to update the list of local representatives in the package leaflet

Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.14. Epcoritamab - TEPKINLY (CAP) - EMEA/H/C/005985/II/0001, Orphan

Applicant: AbbVie Deutschland GmbH & Co. KG

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 product information

Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.15. Ganaxolone - ZTALMY (CAP) - EMEA/H/C/005825/II/0005, Orphan

Applicant: Marinus Pharmaceuticals Emerald Limited

PRAC Rapporteur: Adam Przybylkowski

Scope: Update of section 4.2 of the SmPC in order to update dosing instructions in severe hepatic impairment based on data from phase I study 1042-IHF-1001. The RMP version 1.3 has also been submitted

Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.16. Glycopyrronium - SIALANAR (CAP) - EMEA/H/C/003883/II/0029

Applicant: Proveca Pharma Limited

PRAC Rapporteur: Zane Neikena

Scope: Extension of indication to include treatment of children aged from 2 years and older for SIALANAR, based on the interim results from study PRO/GLY/005. This is a retrospective analysis of real-world data from children aged under 3 years treated with glycopyrronium for severe drooling. As a consequence, sections 4.1, 4.2, and 4.4 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 implement editorial changes to the SmPC. As part of the application the MAH is requesting a 1-year extension of the market protection

Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.17. Imlifidase - IDEFIRIX (CAP) - EMEA/H/C/004849/II/0019, Orphan

Applicant: Hansa Biopharma AB

PRAC Rapporteur: Bianca Mulder

Scope: Update of section 5.1 of the SmPC in order to include the description of the final results from post-authorisation efficacy study (PAES) study 17-HMedIdeS-14 listed as a specific obligation in the Annex II (SOB/002); this is a prospective, observational long-term follow-up study of patients treated with imlifidase (IdeS) prior to kidney transplantation. The primary objective of this trial was to evaluate graft survival in patients who have undergone kidney transplantation after imlifidase administration in earlier trials and relates to both safety and efficacy. The RMP version 1.2 has also been submitted. In addition, the MAH took the opportunity to update section E of Annex II and to implement editorial changes to sections 4.4, 4.6 and 9 of the SmPC. Furthermore, the MAH took the opportunity to bring the product information in line with the latest QRD template version 10.3

Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.18. Ivacaftor, tezacaftor, elexacaftor - KAFTRIO (CAP) - EMEA/H/C/005269/WS2551/0043; Ivacaftor - KALYDECO (CAP) - EMEA/H/C/002494/WS2551/0121

Applicant: Vertex Pharmaceuticals (Ireland) Limited

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 product information

Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.19. Linzagolix choline - YSELTY (CAP) - EMEA/H/C/005442/II/0013

Applicant: Theramex Ireland Limited

PRAC Rapporteur: Martin Huber

Scope: Extension of indication to include treatment of endometriosis-associated pain in adult women of reproductive age for YSELTY, based on final results from studies Edelweiss 3 (18-OBE2109-003) and Edelweiss 6 (19-OBE2109-006) as well as additional supporting studies. Edelweiss 3 is a pivotal phase 3, randomised, double-blind, placebo-controlled, safety and efficacy study to evaluate linzagolix with add-back therapy as a therapy for pain associated with endometriosis, while Edelweiss 6 is an open-label extension study including patients who completed Edelweiss 3 pivotal study regardless of their previous treatment assignment and met the eligibility criteria. 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 1.1 of the RMP has also been submitted. As part of the application, the MAH is requesting a 1-year extension of the market protection

Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.20. Lisocabtagene maraleucel - BREYANZI (CAP) - EMEA/H/C/004731/II/0036/G

Applicant: Bristol-Myers Squibb Pharma EEIG, ATMP

PRAC Rapporteur: Gabriele Maurer

Scope: 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. RMP version 4.0 has also been submitted

Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CAT and CHMP

5.3.21. Lumacaftor, ivacaftor - ORKAMBI (CAP) - EMEA/H/C/003954/II/0088

Applicant: Vertex Pharmaceuticals (Ireland) Limited

PRAC Rapporteur: Eamon O'Murchu

Scope: Submission of the final report from study VX19-809-124 (Study 124), listed as a category 3 study in the RMP. This is a phase 3, open-label, rollover study to evaluate the long-term safety and tolerability of lumacaftor/ivacaftor in cystic fibrosis subjects homozygous for F508del who were 1 to <2 years of age at treatment initiation and who completed the safety follow-up (SFU) visit in Study 122 (Part B) or were lumacaftor/ivacaftor naïve. The RMP version 11.5 has also been submitted

Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.22. Niraparib, abiraterone acetate - AKEEGA (CAP) - EMEA/H/C/005932/II/0003

Applicant: Janssen-Cilag International N.V.

PRAC Rapporteur: Jan Neuhauser

Scope: Update of sections 4.8 and 5.1 of the SmPC in order to update the frequency of adverse drug reactions and to update information from MAGNITUDE study based on final results from study 64091742PCR3001 (MAGNITUDE) listed as a post-authorisation efficacy study (PAES) in the Annex II. This is a phase 3 randomised, placebo-controlled, double-blind, multicenter study which assessed the efficacy and safety of niraparib 200 mg in combination with AA 1,000 mg once daily plus prednisone or prednisolone 10 mg daily (AAP), compared with placebo plus AAP in men with mCRPC and HRR gene alterations, approximately half of whom had BRCA gene alterations and comprised the prespecified BRCA subgroup.
The Annex II and package leaflet are updated accordingly. The RMP version 2.1 has also been submitted. In addition, the MAH took this opportunity to update the list of local representatives in the package leaflet and to introduce editorial changes to the product information

Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.23. Ozanimod - ZEPOSIA (CAP) - EMEA/H/C/004835/II/0023

Applicant: Bristol-Myers Squibb Pharma EEIG

PRAC Rapporteur: Maria del Pilar Rayon

Scope: Update of sections 4.4 and 5.1 of the SmPC in order to update efficacy and safety information based on the final results from study RPC01-3001, listed as a category 3 study in the RMP. This is a multi-site, open label extension trial of RPC1063 in relapsing multiple sclerosis. The study's main objectives were to characterise the long-term safety and tolerability, and the long-term efficacy of ozanimod in patients with relapsing multiple sclerosis. The RMP version 7.0 has also been submitted

Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.24. Peginterferon alfa-2a - PEGASYS (CAP) - EMEA/H/C/000395/II/0119/G

Applicant: Pharmaand GmbH

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 product information is brought in line with the latest QRD template version 10.3

Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.25. Pregabalin - LYRICA (CAP) - EMEA/H/C/000546/X/0127

Applicant: Upjohn EESV

PRAC Rapporteur: Liana Martirosyan

Scope: Extension application to introduce a new pharmaceutical form (orodispersible tablet)

Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.26. Pretomanid - DOVPRELA (CAP) - EMEA/H/C/005167/II/0019/G, Orphan

Applicant: Mylan IRE Healthcare Limited

PRAC Rapporteur: Liana Martirosyan

Scope: Grouped application comprising two variations as follows:
Type II (C.I.4) – Update of sections 4.1 and 5.1 of the SmPC in order to rephrase the indication wording to align with the current WHO definitions. The package leaflet is updated accordingly. In addition, the MAH took the opportunity to update the list of local representatives in the package leaflet.
Type IB (C.I.11.z) - Submission of an updated RMP version 2.0 in order to align the safety concerns following the assessment of procedure EMEA/H/C/005167/11/0013

Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.27. Rilpivirine - EDURANT (CAP) - EMEA/H/C/002264/X/0042/G

Applicant: Janssen-Cilag International N.V.

PRAC Rapporteur: Liana Martirosyan

Scope: Grouped application consisting of: 1) Extension application to introduce a new pharmaceutical form associated with new strength (2.5 mg dispersible tablets). The new presentation is indicated, in combination with other antiretroviral medicinal products, for the treatment of HIV-1 infection in patients ≥2 to <18 years of age and weighing at least 10 kg to less than 25 kg. The product information and RMP have been updated in accordance. 2) Type II variation (C.I.6.a) to modify the approved therapeutic indication of the already authorised 25 mg film-coated tablets presentation to include, in combination with other antiretroviral medicinal products, treatment of human immunodeficiency virus type 1 (HIV-1) infection in antiretroviral treatment-naïve and virologically suppressed (HIV-1 RNA less than 50 copies per ml) paediatric patients from 2 to less than 12 years weighing at least 25 kg, based on final results from study studies TMC278-TiDP38-C213 Cohort 2. As a consequence, sections 4.1, 4.2, 4.4, 4.5, 4.8, 5.1, 5.2 and 5.3 of the SmPC are updated. The package leaflet and labelling are updated in accordance. The updated RMP version 10.1 has also been submitted. In addition, the MAH took the opportunity to implement editorial changes to Annex II and to update the list of local representatives in the package leaflet

Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.28. Smallpox vaccine (live modified vaccinia virus Ankara) - IMVANEX (CAP) - EMEA/H/C/002596/II/0100

Applicant: Bavarian Nordic A/S

PRAC Rapporteur: Gabriele Maurer

Scope: Update of section 5.1 of the SmPC in order to add vaccine effectiveness data, and the removal of the two open specific obligations (POX-MVA-039 (SOB02) and SEMVAc (SOB03)), based on the IMVANEX vaccine effectiveness data in real-world use during the 2022 monkeypox outbreak. Consequently, the MAH proposes a switch from exceptional marketing authorisation to full marketing authorisation. The Annex II and package leaflet are updated accordingly. The RMP version 10.0 has also been submitted. In addition, the MAH took the opportunity to introduce minor editorial changes to the product information

Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.29. Teclistamab - TECVAYLI (CAP) - EMEA/H/C/005865/II/0012

Applicant: Janssen-Cilag International N.V.

PRAC Rapporteur: Jana Lukacisinova

Scope: Update of sections 4.2, 4.8, 5.1 of the SmPC in order to amend the recommendations for dose delays, as well as, to update safety and efficacy information based on final results from study 64007957MMY1001 listed as a specific obligation in the Annex II (SOB/005); this is a phase 1/2, first in human, open label, dose escalation study of teclistamab in subjects with relapsed or refractory multiple myeloma. The package leaflet is updated accordingly. The RMP version 4.2 has also been submitted. In addition, the MAH took the opportunity to update Annex II and Annex IV of the product information

Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.30. Tildrakizumab - ILUMETRI (CAP) - EMEA/H/C/004514/II/0054

Applicant: Almirall S.A

PRAC Rapporteur: Adam Przybylkowski

Scope: Update of section 5.1 of the SmPC in order to update clinical and safety information based on long-term results from the extension periods of the pivotal clinical studies MK-3222-010 (a 64-week, phase 3, randomised, placebo-controlled, parallel design study to evaluate the efficacy and safety/tolerability of subcutaneous tildrakizumab (SCH 900222/MK-3222), followed by an optional long-term safety extension study, in subjects with moderate-to-severe chronic plaque psoriasis (Protocol No. MK-3222-010)) and MK-3222-011 (A 52-Week, Phase 3, Randomised, Active Comparator and Placebo-Controlled, Parallel Design Study to Evaluate the Efficacy and Safety/Tolerability of Subcutaneous Tildrakizumab (SCH 900222 / MK-3222), Followed by an Optional Long-Term Safety Extension Study, in Subjects With Moderate-to-Severe Chronic Plaque Psoriasis). The RMP version 1.4 has also been submitted

Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.31. Tislelizumab - TEVIMBRA (CAP) - EMEA/H/C/005919/II/0006

Applicant: Beigene Ireland Limited

PRAC Rapporteur: Bianca Mulder

Scope: Extension of indication to include in combination with platinum and fluoropyrimidine-based chemotherapy the first-line treatment of adult patients with human epidermal growth factor receptor-2 (HER-2)-negative locally advanced unresectable or metastatic gastric or gastroesophageal junction (G/GEJ) adenocarcinoma for TEVIMBRA, based on results from the phase 3 study BGB-A317-305 (study 305); this is a global, randomised, double-blind, placebo-controlled study at the approved registrational dosing regimen for Tevimbra (200 mg administered IV Q3W), in combination with platinum and fluoropyrimidine-based chemotherapy, in adult patients with HER-2 negative locally advanced unresectable or metastatic gastric or gastroesophageal junction (G/GEJ) adenocarcinoma. 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 1.2 of the RMP has also been submitted. In addition, the MAH took the opportunity to introduce minor editorial changes to the product information

Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.32. Tislelizumab - TEVIMBRA (CAP) - EMEA/H/C/005919/II/0008

Applicant: Beigene Ireland Limited

PRAC Rapporteur: Bianca Mulder

Scope: Extension of indication to include treatment of adult patients with non-small cell lung cancer (NSCLC) in combination and as monotherapy for TEVIMBRA, based on results from studies BGB-A317-303, BGB-A317-304, BGB-A317-307 and BGB A317-206. As a consequence, sections 4.1, 4.2, 4.4, 4.5, 4.8, 5.1 and 5.2 of the SmPC are updated. The package leaflet is updated in accordance. Version 2.0 of the RMP has also been submitted. In addition, the MAH took the opportunity to introduce minor editorial changes to the product information

Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.33. Upadacitinib - RINVOQ (CAP) - EMEA/H/C/004760/II/0052

Applicant: AbbVie Deutschland GmbH & Co. KG

PRAC Rapporteur: Petar Mas

Scope: Update of sections 4.2, 4.8 and 5.1 of the SmPC in order to change posology recommendations in adolescents with atopic dermatitis to include the 30mg dose option based on results from studies M16-045, M16-047 and M18-891 (pivotal phase 3 studies with adolescent substudies). The package leaflet is updated accordingly. The RMP version 14.0 has also been submitted

Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

6. Periodic safety update reports (PSURs)

6.1. PSUR single assessment (PSUSA) procedures including centrally authorised products (CAPs) only

6.1.1. Abaloparatide - ELADYNOS (CAP) - PSUSA/00011029/202310

Applicant: Theramex Ireland Limited

PRAC Rapporteur: Karin Erneholm

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

6.1.2. Acalabrutinib - CALQUENCE (CAP) - PSUSA/00010887/202310 (with RMP)

Applicant: AstraZeneca AB

PRAC Rapporteur: Barbara Kovacic Bytyqi

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

6.1.3. Afatinib - GIOTRIF (CAP) - PSUSA/00010054/202309

Applicant: Boehringer Ingelheim International GmbH

PRAC Rapporteur: Ulla Wändel Liminga

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

6.1.4. Andexanet alfa - ONDEXXYA (CAP) - PSUSA/00010764/202310

Applicant: AstraZeneca AB

PRAC Rapporteur: Bianca Mulder

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

6.1.5. Arsenic trioxide - TRISENOX (CAP) - PSUSA/00000235/202309

Applicant: Teva B.V.

PRAC Rapporteur: Tiphaine Vaillant

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

6.1.6. Asciminib - SCEMBLIX (CAP) - PSUSA/00011008/202310

Applicant: Novartis Europharm Limited

PRAC Rapporteur: Eva Jirsová

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

6.1.7. Axicabtagene ciloleucel - YESCARTA (CAP) - PSUSA/00010703/202310

Applicant: Kite Pharma EU B.V.

PRAC Rapporteur: Karin Erneholm

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

6.1.8. Aztreonam[4] - CAYSTON (CAP) - PSUSA/00000283/202309

Applicant: Gilead Sciences Ireland UC

PRAC Rapporteur: Liana Martirosyan

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

6.1.9. Bexarotene - TARGRETIN (CAP) - PSUSA/00000404/202309

Applicant: Eisai GmbH

PRAC Rapporteur: Tiphaine Vaillant

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

6.1.10. Brolucizumab - BEOVU (CAP) - PSUSA/00010829/202310

Applicant: Novartis Europharm Limited

PRAC Rapporteur: Gabriele Maurer

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

6.1.11. Bupivacaine[5] - EXPAREL LIPOSOMAL (CAP) - PSUSA/00010889/202310

Applicant: Pacira Ireland Limited

PRAC Rapporteur: Eamon O'Murchu

[4] For inhalation use only
[5] Liposomal formulations only

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

6.1.12. Cetuximab - ERBITUX (CAP) - PSUSA/00000635/202309

Applicant: Merck Europe B.V.

PRAC Rapporteur: Ulla Wändel Liminga

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

6.1.13. Chenodeoxycholic acid[6] - CHENODEOXYCHOLIC ACID LEADIANT (CAP) - PSUSA/00010590/202310

Applicant: Leadiant GmbH

PRAC Rapporteur: Adam Przybylkowski

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

6.1.14. Conestat alfa - RUCONEST (CAP) - PSUSA/00000873/202310

Applicant: Pharming Group N.V

PRAC Rapporteur: Jan Neuhauser

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

6.1.15. Coronavirus (COVID-19) vaccine (recombinant protein receptor binding domain fusion heterodimer) - BIMERVAX (CAP) - PSUSA/00011045/202309

Applicant: Hipra Human Health S.L.

PRAC Rapporteur: Zane Neikena

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

6.1.16. Dapagliflozin - EDISTRIDE (CAP); FORXIGA (CAP) - PSUSA/00010029/202310

Applicant: AstraZeneca AB

PRAC Rapporteur: Mari Thorn

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

[6] Inborn error in primary bile acid synthesis, xanthomatosi - centrally authorised products only

6.1.17. Delamanid - DELTYBA (CAP) - PSUSA/00010213/202310

Applicant: Otsuka Novel Products GmbH

PRAC Rapporteur: Jo Robays

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

6.1.18. Denosumab[7] - XGEVA (CAP) - PSUSA/00009119/202309

Applicant: Amgen Europe B.V.

PRAC Rapporteur: Mari Thorn

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

6.1.19. Dexamethasone[8] - NEOFORDEX (CAP) - PSUSA/00010480/202309

Applicant: Theravia

PRAC Rapporteur: Tiphaine Vaillant

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

6.1.20. Dostarlimab - JEMPERLI (CAP) - PSUSA/00010931/202310

Applicant: GlaxoSmithKline (Ireland) Limited

PRAC Rapporteur: Carla Torre

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

6.1.21. Eculizumab - BEKEMV (CAP); EPYSQLI (CAP); SOLIRIS (CAP) - PSUSA/00001198/202310

Applicant(s): Alexion Europe SAS (Soliris), Amgen Technology (Ireland) Unlimited Company (Bekemv), Samsung Bioepis NL B.V. (Epysqli)

PRAC Rapporteur: Monica Martinez Redondo

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

[7] Indicated for skeletal related events associated with bone metastases and for giant cell tumour of bone only
[8] Centrally authorised product indicated in symptomatic multiple myeloma

6.1.22. Edoxaban - LIXIANA (CAP); ROTEAS (CAP) - PSUSA/00010387/202310

Applicant(s): Berlin Chemie AG (Roteas), Daiichi Sankyo Europe GmbH (Lixiana)

PRAC Rapporteur: Nathalie Gault

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

6.1.23. Etravirine - INTELENCE (CAP) - PSUSA/00001335/202309

Applicant: Janssen-Cilag International N.V.

PRAC Rapporteur: Nathalie Gault

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

6.1.24. Futibatinib - LYTGOBI (CAP) - PSUSA/00000068/202309

Applicant: Taiho Pharma Netherlands B.V.

PRAC Rapporteur: Ulla Wändel Liminga

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

6.1.25. Herpes zoster vaccine (recombinant, adjuvanted) - SHINGRIX (CAP) - PSUSA/00010678/202310

Applicant: GlaxoSmithkline Biologicals SA

PRAC Rapporteur: Sonja Hrabcik

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

6.1.26. Histamine[9] - CEPLENE (CAP) - PSUSA/00001610/202310

Applicant: Laboratoires Delbert

PRAC Rapporteur: Eamon O'Murchu

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

6.1.27. Indacaterol, glycopyrronium bromide - ULTIBRO BREEZHALER (CAP); ULUNAR BREEZHALER (CAP); XOTERNA BREEZHALER (CAP) - PSUSA/00010105/202309

Applicant: Novartis Europharm Limited

[9] Indicated for acute myeloid leukemia only

PRAC Rapporteur: Marie Louise Schougaard Christiansen

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

6.1.28. Insulin aspart - FIASP (CAP); INSULIN ASPART SANOFI (CAP); KIRSTY (CAP); NOVOMIX (CAP); NOVORAPID (CAP); TRUVELOG MIX 30 (SRD) (CAP) - PSUSA/00001749/202309

Applicant(s): Biosimilar Collaborations Ireland Limited (Kirsty), Novo Nordisk A/S (Fiasp, NovoMix, NovoRapid), Sanofi Winthrop Industrie (Insulin aspart Sanofi, Truvelog Mix 30 (SRD))

PRAC Rapporteur: Mari Thorn

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

6.1.29. Insulin degludec – TRESIBA; insulin degludec, insulin aspart - RYZODEG (CAP); (CAP) - PSUSA/00010036/202309

Applicant: Novo Nordisk A/S

PRAC Rapporteur: Mari Thorn

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

6.1.30. Insulin human[10] - INSUMAN (CAP) - PSUSA/00010107/202309

Applicant: Sanofi-Aventis Deutschland GmbH

PRAC Rapporteur: Jean-Michel Dogné

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

6.1.31. Ivacaftor, tezacaftor, elexacaftor - KAFTRIO (CAP) - PSUSA/00010868/202310

Applicant: Vertex Pharmaceuticals (Ireland) Limited

PRAC Rapporteur: Martin Huber

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

6.1.32. Lasmiditan - RAYVOW (CAP) - PSUSA/00011011/202310

Applicant: Eli Lilly Nederland B.V.

[10] For intraperitoneal use only

PRAC Rapporteur: Anna Mareková

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

6.1.33. Loncastuximab tesirine - ZYNLONTA (CAP) - PSUSA/00011027/202310

Applicant: Swedish Orphan Biovitrum AB (publ)

PRAC Rapporteur: Eva Jirsová

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

6.1.34. Mavacamten - CAMZYOS (CAP) - PSUSA/00000074/202310

Applicant: Bristol-Myers Squibb Pharma EEIG

PRAC Rapporteur: Kimmo Jaakkola

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

6.1.35. Nintedanib[11] - OFEV (CAP) - PSUSA/00010319/202310

Applicant: Boehringer Ingelheim International GmbH

PRAC Rapporteur: Barbara Kovacic Bytyqi

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

6.1.36. Niraparib, abiraterone acetate - AKEEGA (CAP) - PSUSA/00011051/202310

Applicant: Janssen-Cilag International N.V.

PRAC Rapporteur: Jan Neuhauser

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

6.1.37. Nirsevimab - BEYFORTUS (CAP) - PSUSA/00011026/202310

Applicant: Sanofi Winthrop Industrie

PRAC Rapporteur: Kimmo Jaakkola

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

[11] Respiratory indication only

6.1.38. Parathyroid hormone - NATPAR (CAP) - PSUSA/00010591/202310

Applicant: Takeda Pharmaceuticals International AG Ireland Branch

PRAC Rapporteur: Rhea Fitzgerald

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

6.1.39. Pemigatinib - PEMAZYRE (CAP) - PSUSA/00010923/202310

Applicant: Incyte Biosciences Distribution B.V.

PRAC Rapporteur: Bianca Mulder

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

6.1.40. Pitolisant - OZAWADE (CAP); WAKIX (CAP) - PSUSA/00010490/202309

Applicant: Bioprojet Pharma

PRAC Rapporteur: Kirsti Villikka

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

6.1.41. Potassium citrate, potassium hydrogen carbonate - SIBNAYAL (CAP) - PSUSA/00010932/202310

Applicant: Advicenne

PRAC Rapporteur: Adam Przybylkowski

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

6.1.42. Regorafenib - STIVARGA (CAP) - PSUSA/00010133/202309

Applicant: Bayer AG

PRAC Rapporteur: Bianca Mulder

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

6.1.43. Recombinant respiratory syncytial virus pre-fusion F protein, adjuvanted with AS01E - AREXVY (CAP) - PSUSA/00000031/202311

Applicant: GlaxoSmithkline Biologicals S.A.

PRAC Rapporteur: Maria del Pilar Rayon

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

6.1.44. Selumetinib - KOSELUGO (CAP) - PSUSA/00010936/202310

Applicant: AstraZeneca AB

PRAC Rapporteur: Ulla Wändel Liminga

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

6.1.45. Sirolimus[12] - RAPAMUNE (CAP) - PSUSA/00002710/202309

Applicant: Pfizer Europe MA EEIG

PRAC Rapporteur: Mari Thorn

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

6.1.46. Somatrogon - NGENLA (CAP) - PSUSA/00010982/202310

Applicant: Pfizer Europe MA EEIG

PRAC Rapporteur: Liana Martirosyan

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

6.1.47. Sulfur hexafluoride - SONOVUE (CAP) - PSUSA/00002822/202309

Applicant: Bracco International B.V.

PRAC Rapporteur: Tiphaine Vaillant

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

6.1.48. Talazoparib - TALZENNA (CAP) - PSUSA/00010781/202310

Applicant: Pfizer Europe MA EEIG

PRAC Rapporteur: Carla Torre

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

[12] For prophylaxis of organ rejection indication only

6.1.49. Talimogene laherparepvec - IMLYGIC (CAP) - PSUSA/00010459/202310

Applicant: Amgen Europe B.V., ATMP

PRAC Rapporteur: Gabriele Maurer

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CAT and CHMP

6.1.50. Toremifene - FARESTON (CAP) - PSUSA/00002999/202309

Applicant: Orion Corporation

PRAC Rapporteur: Tiphaine Vaillant

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

6.1.51. Tremelimumab - IMJUDO (CAP) - PSUSA/00011038/202310

Applicant: AstraZeneca AB

PRAC Rapporteur: David Olsen

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

6.1.52. Tucatinib - TUKYSA (CAP) - PSUSA/00010918/202310

Applicant: Seagen B.V.

PRAC Rapporteur: Jean-Michel Dogné

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

6.1.53. Volanesorsen - WAYLIVRA (CAP) - PSUSA/00010762/202311

Applicant: Akcea Therapeutics Ireland Limited

PRAC Rapporteur: Martin Huber

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

6.1.54. Zoonotic influenza vaccine (H5N1)[13] - FOCLIVIA (CAP); prepandemic influenza vaccine (H5N1)[13] - AFLUNOV (CAP); ZOONOTIC INFLUENZA VACCINE SEQIRUS (CAP) - PSUSA/00010008/202310

Applicant: Seqirus S.r.l.

[13] Surface antigen, inactivated, adjuvanted

PRAC Rapporteur: Amelia Cupelli

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

6.2. PSUR single assessment (PSUSA) procedures including centrally authorised products (CAPs) and nationally authorised products (NAPs)

6.2.1. Buprenorphine[14] - BUVIDAL (CAP); NAP - PSUSA/00000459/202309

Applicant: Camurus AB (Buvidal), various

PRAC Rapporteur: Tiphaine Vaillant

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

6.2.2. Buprenorphine, naloxone - SUBOXONE (CAP); ZUBSOLV (CAP); NAP - PSUSA/00002113/202309

Applicant(s): Accord Healthcare S.L.U. (Zubsolv), Indivior Europe Limited (Suboxone), various

PRAC Rapporteur: Martin Huber

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

6.2.3. Iloprost[15] - VENTAVIS (CAP); NAP - PSUSA/00001724/202309

Applicant: Bayer AG (Ventavis), various

PRAC Rapporteur: Nathalie Gault

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

6.2.4. Leflunomide - ARAVA (CAP); LEFLUNOMIDE MEDAC (CAP); LEFLUNOMIDE ZENTIVA (CAP); NAP - PSUSA/00001837/202309

Applicant(s): Medac Gesellschaft fur klinische Spezialpraparate mbH (Leflunomide medac), Sanofi-Aventis Deutschland GmbH (Arava), Zentiva, k.s. (Leflunomide Zentiva), various

PRAC Rapporteur: Liana Martirosyan

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

[14] All formulations except implants
[15] Nebuliser solution only

6.2.5. Melatonin - CIRCADIN (CAP); MELATONIN NEURIM (CAP); SLENYTO (CAP); NAP - PSUSA/00001963/202309

Applicant: RAD Neurim Pharmaceuticals EEC SARL (Circadin, Melatonin Neurim, Slenyto), various

PRAC Rapporteur: Ana Sofia Diniz Martins

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

6.2.6. Memantine - AXURA (CAP); EBIXA (CAP); MEMANTINE MERZ (CAP); NAP - PSUSA/00001967/202309

Applicant(s): H. Lundbeck A/S (Ebixa), Merz Pharmaceuticals GmbH (Axura, Memantine Merz), various

PRAC Rapporteur: Maria del Pilar Rayon

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

6.2.7. Posaconazole - NOXAFIL (CAP); NAP - PSUSA/00002480/202310

Applicant: Merck Sharp & Dohme B.V. (Noxafil), various

PRAC Rapporteur: Nathalie Gault

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

6.2.8. Sodium oxybate[16] - XYREM (CAP); NAP - PSUSA/00010612/202310

Applicant: UCB Pharma S.A. (Xyrem), various

PRAC Rapporteur: Ana Sofia Diniz Martins

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

6.2.9. Teriflunomide - AUBAGIO (CAP); TERIFLUNOMIDE ACCORD (CAP); TERIFLUNOMIDE MYLAN (CAP); NAP - PSUSA/00010135/202309

Applicant(s): Accord Healthcare S.L.U. (Teriflunomide Accord), Mylan Pharmaceuticals Limited (Teriflunomide Mylan), Sanofi Winthrop Industrie (AUBAGIO), various

PRAC Rapporteur: Martin Huber

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

[16] Oral use

6.2.10. Vigabatrin - KIGABEQ (CAP); NAP - PSUSA/00003112/202309

Applicant: ORPHELIA Pharma SAS (Kigabeq), various

PRAC Rapporteur: Kirsti Villikka

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CHMP

6.3. PSUR single assessment (PSUSA) procedures including nationally authorised products (NAPs) only

6.3.1. Acetylcysteine (NAP) - PSUSA/00000034/202309

Applicant(s): various

PRAC Lead: Adam Przybylkowski

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CMDh

6.3.2. Acitretin (NAP) - PSUSA/00000051/202310

Applicant(s): various

PRAC Lead: Marie Louise Schougaard Christiansen

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CMDh

6.3.3. Adapalene, benzoyl peroxide (NAP) - PSUSA/00000059/202309

Applicant(s): various

PRAC Lead: Mari Thorn

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CMDh

6.3.4. Baclofen[17] (NAP) - PSUSA/00000294/202309

Applicant(s): various

PRAC Lead: Eamon O'Murchu

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CMDh

[17] Oral use, for muscle spasticity indication only

6.3.5. Calcium carbonate, famotidine, magnesium hydroxide (NAP) - PSUSA/00001351/202309

Applicant(s): various

PRAC Lead: Nathalie Gault

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CMDh

6.3.6. Desflurane (NAP) - PSUSA/00000958/202309

Applicant(s): various

PRAC Lead: Melinda Palfi

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CMDh

6.3.7. Diclofenac[18] (NAP) - PSUSA/00010342/202309

Applicant(s): various

PRAC Lead: Karin Erneholm

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CMDh

6.3.8. Felodipine, ramipril (NAP) - PSUSA/00001358/202309

Applicant(s): various

PRAC Lead: Martin Huber

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CMDh

6.3.9. Human von Willebrand factor (NAP) - PSUSA/00001642/202309

Applicant(s): various

PRAC Lead: Gabriele Maurer

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CMDh

6.3.10. Levofloxacin[19] (NAP) - PSUSA/00010767/202310

Applicant(s): various

[18] Topical formulations only
[19] Intravenous and oral use only

PRAC Lead: Martin Huber

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CMDh

6.3.11. Levofloxacin[20] (NAP) - PSUSA/00010768/202310

Applicant(s): various

PRAC Lead: Martin Huber

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CMDh

6.3.12. Prulifloxacin (NAP) - PSUSA/00002569/202309

Applicant(s): various

PRAC Lead: Amelia Cupelli

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CMDh

6.3.13. Rubidium (^{82}Rb) chloride (NAP) - PSUSA/00010806/202310

Applicant(s): various

PRAC Lead: Martin Huber

Scope: Evaluation of a PSUSA procedure

Action: For adoption of recommendation to CMDh

6.4. Follow-up to PSUR/PSUSA procedures

6.4.1. Tenofovir disoproxil - VIREAD (CAP) - EMEA/H/C/000419/LEG 279

Applicant: Gilead Sciences Ireland UC

PRAC Rapporteur: Nathalie Gault

Scope: From PSUSA/00002892/202303: Cumulative safety review on dental disorders, increased parathyroid hormone and congenital anomalies

Action: For adoption of advice to CHMP

[20] Ocular use only

6.5. Variation procedure(s) resulting from PSUSA evaluation

6.5.1. Dabrafenib - FINLEE (CAP) - EMEA/H/C/005885/WS2671/0005; Trametinib - SPEXOTRAS (CAP) - EMEA/H/C/005886/WS2671/0004; Dabrafenib - TAFINLAR (CAP) - EMEA/H/C/002604/WS2671/0067

Applicant: Novartis Europharm Limited

PRAC Rapporteur: Ulla Wändel Liminga

Scope: Update of section 4.8 of the SmPC for Tafinlar, Finlee and Spexotras in order to add 'Atrioventricular (AV) block' and 'Bundle branch block' to the list of adverse drug reactions (ADRs), following the PRAC recommendation in the PSUR for Mekinist (PSUSA/00010262/202305). The package leaflet is updated accordingly

Action: For adoption of PRAC Assessment Report

6.5.2. Ioflupane (^{123}I) - DATSCAN (CAP) - EMEA/H/C/000266/II/0067

Applicant: GE Healthcare B.V.

PRAC Rapporteur: Tiphaine Vaillant

Scope: To update sections 4.4 and 4.5 of the SmPC and section 2 of the Package Leaflet to implement the recommendation of PRAC following the PSUSA procedure (EMEA/H/C/PSUSA/00001767/202207). In addition, the MAH took the opportunity to update the list of local representatives in the package leaflet.

Action: For adoption of PRAC Assessment Report

6.5.3. Mitotane - LYSODREN (CAP) - EMEA/H/C/000521/II/0030

Applicant: HRA Pharma Rare Diseases

PRAC Rapporteur: Maria del Pilar Rayon

Scope: Update of section 4.4 of the SmPC in order amend an existing warning on hepatic impairment based on a cumulative review of cases with increase of transaminases >5 ULN and the outcome of these elevations after mitotane discontinuation, following the request by PRAC in the PSUSA/00002075/202304

Action: For adoption of PRAC Assessment Report

6.5.4. Naltrexone hydrochloride, Bupropion hydrochloride - MYSIMBA (CAP) - EMEA/H/C/003687/II/0063

Applicant: Orexigen Therapeutics Ireland Limited

PRAC Rapporteur: Martin Huber

Scope: Submission of an update of sections 4.3, 4.4 and 4.5 of the SmPC to update and streamline the relevant wording on opioids as requested in the conclusions of the PSUR single assessment (PSUSA) procedure (PSUSA/00010366/202209) adopted in April 2023.

The package leaflet is updated accordingly. The RMP version 12.9 has also been submitted

Action: For adoption of PRAC Assessment Report

6.5.5. Rituximab - MABTHERA (CAP) - EMEA/H/C/000165/II/0201/G

Applicant: Roche Registration GmbH

PRAC Rapporteur: Karin Erneholm

Scope: 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 healthcare professionals (HCPs) and patients, following the request by PRAC in the assessment report 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 product information 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

Action: For adoption of PRAC Assessment Report

6.6. Expedited summary safety reviews[21]

None

7. Post-authorisation safety studies (PASS)

7.1. Protocols of PASS imposed in the marketing authorisation(s)[22]

7.1.1. Axicabtagene ciloleucel (CAP) - YESCARTA – EMEA/H/C/PSA/S/0102.4

Applicant: Kite Pharma EU B.V.

PRAC Rapporteur: Karin Erneholm

Scope: Substantial amendment to a protocol for a long-term, non-interventional study of recipients of Yescarta for treatment of relapsed or refractory diffuse large B-cell lymphoma and primary mediastinal B-cell lymphoma [MAH's response to PSA/S/0102.3]

Action: For adoption of PRAC Assessment Report, PRAC outcome letter

[21] Submission of expedited summary safety reports for review in addition to the requirements for submission of PSUR(s) falling within the pandemic period and requirements set out in the list of Union reference dates (EURD list) provided for under Article 107c(7) of Directive 2001/83/EC

[22] In accordance with Article 107n of Directive 2001/83/EC

7.1.2. Evinacumab - EVKEEZA (CAP) – EMEA/H/C/PSA/S/0112

Applicant: Regeneron Ireland DAC

PRAC Rapporteur: Mari Thorn

Scope: Substantial amendment to a protocol for an evaluation of the long-term effects of evinacumab treatment in patients with homozygous familial hypercholesterolemia (HoFH):

- safety outcomes in patients with HoFH who are ⩾12 years old
- frequency and outcomes of pregnancy in female patients with HoFH
- atherosclerosis process over time in patients with HoFH who undergo cardiovascular imaging (as data allow)
- frequency of cardiovascular imaging of patients with HoFH

Action: For adoption of PRAC Assessment Report, PRAC outcome letter

7.1.3. Methylphenidate hydrochloride (NAP) – EMEA/H/N/PSA/S/0113

Applicant: Medice Arzneimittel Pütter GmbH Co. KG (MAH for Medikinet)

PRAC Rapporteur: Martin Huber

Scope: Substantial amendments to the protocol for the post-authorisation safety study (PASS) evaluating the long-term cardiovascular and psychiatric safety profile of methylphenidate in adult patients with attention deficit/hyperactivity disorder (ADHD) in European Countries. A revised protocol version 5.2 dated 2 February 2024 was submitted. The most significant changes introduced by the amended protocol concern the study milestones and the data analysis

Action: For adoption of PRAC Assessment Report, PRAC outcome letter

7.1.4. Pegzilarginase - LOARGYS (CAP) – EMEA/H/C/PSP/S/0105

Applicant: Immedica Pharma AB

PRAC Rapporteur: Martin Huber

Scope: A European, non-interventional, multicentre, registry-based PASS to evaluate the long-term safety of Loargys treatment in arginase 1 deficiency patients in standard clinical care

Action: For adoption of PRAC Assessment Report, PRAC outcome letter

7.1.5. Valproate (CAP) – EMEA/H/N/PSP/J/0074.9

Applicant: Sanofi-Aventis Recherche & Développement (on behalf of a consortium)

PRAC Rapporteur: Jean-Michel Dogné

Scope: Responses to the 2nd request for supplementary information (RSI) of the 2nd interim report: observational study to evaluate and identify the best practices for switching of valproate in clinical practice [MAH's response to PSP/J/0074.8], non-interventional retrospective longitudinal study in the UK and France to investigate the therapeutic strategies after discontinuation of valproate (VPA) and related substances in clinical

practice: VALSE study (VALNAC09344)

- Response document to PRAC assessment report (Procedure no.: EMEA/H/N/PSP/J/0074.7)
- Sensitivity analysis performed in the SNDS database, the national healthcare data system in France, to specify the indication for the use of valproate and reduce the proportion of women with unspecified indication.

Action: For adoption of PRAC Assessment Report, PRAC outcome letter

7.2. Protocols of PASS non-imposed in the marketing authorisation(s)[23]

7.2.1. Abaloparatide - ELADYNOS (CAP) - EMEA/H/C/005928/MEA 001.2

Applicant: Theramex Ireland Limited

PRAC Rapporteur: Karin Erneholm

Scope: From initial marketing autorisation application (MAA): REVISED PASS PROTOCOL (Study number not assigned yet); Non-imposed/non-interventional; European non-interventional post-authorisation safety study (PASS) to assess serious cardiovascular events of myocardial infarction (MI), stroke, all-cause and cardiovascular mortality, and arrhythmias for abaloparatide

Action: For adoption of advice to CHMP

7.2.2. Alemtuzumab - LEMTRADA (CAP) - EMEA/H/C/003718/MEA 007.17

Applicant: Sanofi Belgium

PRAC Rapporteur: Karin Erneholm

Scope: MAH's response to MEA 007.14 [Amended Protocol OBS13434 version 4]; request for supplementary information (RSI) as adopted in October 2023. A prospective, multicentre, observational, post-authorisation safety study to evaluate the long-term safety profile of LEMTRADA (alemtuzumab) treatment in patients with relapsing forms of multiple sclerosis

Action: For adoption of advice to CHMP

7.2.3. Atogepant - AQUIPTA (CAP) - EMEA/H/C/005871/MEA 003

Applicant: AbbVie Deutschland GmbH & Co. KG

PRAC Rapporteur: Rugile Pilviniene

Scope: Study P22-392; Atogepant pregnancy exposure registry: Study P22-392 aims to prospectively evaluate maternal, foetal, and infant outcomes through 12 months of age among women exposed to atogepant during pregnancy

Action: For adoption of advice to CHMP

[23] In accordance with Article 107m of Directive 2001/83/EC, supervised by PRAC in accordance with Article 61a (6) of Regulation (EC) No 726/2004

7.2.4. Atogepant - AQUIPTA (CAP) - EMEA/H/C/005871/MEA 004

Applicant: AbbVie Deutschland GmbH & Co. KG

PRAC Rapporteur: Rugile Pilviniene

Scope: Study P22-419; Pregnancy Study; Study P22-419 aims to describe and compare the incidence of pregnancy outcomes in women with migraine who are exposed to atogepant during pregnancy

Action: For adoption of advice to CHMP

7.2.5. Dabigatran etexilate - PRADAXA (CAP) - EMEA/H/C/000829/MEA 051.2

Applicant: Boehringer Ingelheim International GmbH

PRAC Rapporteur: Marie Louise Schougaard Christiansen

Scope: From X/0122/G: REVISED PROTOCOL (3.0) for PASS 1160.307 (non-imposed); Safety of dabigatran etexilate for treatment of venous thromboembolism (VTE) and prevention of recurrent VTE in paediatric patients from birth to less than 2 years of age: A European non-interventional cohort study based on new data collection

Action: For adoption of advice to CHMP

7.2.6. Glofitamab - COLUMVI (CAP) - EMEA/H/C/005751/MEA 004

Applicant: Roche Registration GmbH

PRAC Rapporteur: Jana Lukacisinova

Scope: From initial MAA; PASS No BO43309 (non-imposed/non-interventional); Evaluation of the Effectiveness of the Additional Risk Minimisation Measures for Glofitamab - A Survey Among Healthcare Professionals in 10 Countries in the European Economic Area (EEA)

Action: For adoption of advice to CHMP

7.2.7. Neratinib - NERLYNX (CAP) - EMEA/H/C/004030/MEA 003.3

Applicant: Pierre Fabre Medicament

PRAC Rapporteur: Bianca Mulder

Scope: Amended Protocol / Study PUMA-NER-7402 (version 4.0); Safety of neratinib among breast cancer patients to characterise the incidence and duration of diarrhoea in a real-world setting, describe patient characteristics, incidence rates and duration of diarrhoea, describe use of loperamide and other concomitant antidiarrheal medication, describe adherence to neratinib therapy, assess the impact of neratinib therapy on patient self-reported, health related quality of life and their ability to perform their activities of daily living and to further assess and characterise adverse events hepatic, cardiac (LVEF decreased), pulmonary (interstitial lung disease), reproductive and developmental toxicity

Action: For adoption of advice to CHMP

7.2.8. Niraparib, Abiraterone acetate - AKEEGA (CAP) - EMEA/H/C/005932/MEA 001.2

Applicant: Janssen-Cilag International N.V.

PRAC Rapporteur: Jan Neuhauser

Scope: Revised PASS Protocol / Study no.: PCSONCA0485; Study title: Post authorisation safety study to characterise the risk of second primary malignancies (SPM) including MDS/AML among metastatic prostate cancer patients exposed to AKEEGA

Action: For adoption of advice to CHMP

7.2.9. Risankizumab - SKYRIZI (CAP) - EMEA/H/C/004759/MEA 010.3

Applicant: AbbVie Deutschland GmbH & Co. KG

PRAC Rapporteur: Liana Martirosyan

Scope: Revised PASS Protocol (v 2.1) / Study no.: P23-654 (Non-imposed/Non-interventional); Title: Long-Term Comparative Cohort Study in Patients with Crohn's Disease in a Real World Setting. Additional long-term data from the real-world experience of patients with Crohn's disease treated with risankizumab to assess product potential risks. A comparative cohort study will be conducted to estimate rates of malignancy (malignancy excluding NMSC, NMSC), serious infections, serious hypersensitivity reactions, and major adverse cardiovascular events (MACE) in risankizumab treated patients with Crohn's disease, relative to alternative systemic therapies (e.g., biologics). MAH’s responses to MEA010.2 as adopted in January 2024: The current protocol insufficiently addresses whether duration of treatment (exposure) will be taken into account in the analysis of MACE and also for the other outcomes of interest. The MAH is requested to clarify whether duration of treatment (exposure) will be considered in the analysis of the outcomes of interest or otherwise include this consideration in the protocol

Action: For adoption of advice to CHMP

7.2.10. Romosozumab - EVENITY (CAP) - EMEA/H/C/004465/MEA 002.10

Applicant: UCB Pharma S.A.

PRAC Rapporteur: Tiphaine Vaillant

Scope: MAH's response to MEA 002.9 [Protocol Amendment for study OP0004] request for supplementary information as adopted in January 2024

Action: For adoption of advice to CHMP

7.2.11. Romosozumab - EVENITY (CAP) - EMEA/H/C/004465/MEA 003.8

Applicant: UCB Pharma S.A.

PRAC Rapporteur: Tiphaine Vaillant

Scope: MAH's response to MEA 003.7 [Protocol Amendment for study OP0006] request for supplementary information as adopted in January 2024

Action: For adoption of advice to CHMP

7.2.12. Ruxolitinib - OPZELURA (CAP) - EMEA/H/C/005843/MEA 001.1

Applicant: Incyte Biosciences Distribution B.V.

PRAC Rapporteur: Adam Przybylkowski

Scope: From Initial MAA: Revised protocol for PASS INCB 88888-037 (non-imposed non-interventional, RMP, Cat. 3); Title: To evaluate the safety of long-term ruxolitinib cream use with respect to incidence of non-melanoma skin cancers

Action: For adoption of advice to CHMP

7.2.13. Somapacitan - SOGROYA (CAP) - EMEA/H/C/005030/MEA 005.1

Applicant: Novo Nordisk A/S

PRAC Rapporteur: Martin Huber

Scope: From X/0006/G: Revised protocol / Study No. NN8640-4787; Paediatric growth hormone deficiency (GHD) register-based study: A non-interventional, observational, register-based study to investigate long-term safety and clinical parameters of somapacitan treatment in paediatric patients with GHD in the setting of routine clinical practice. As a follow-up of MEA 005.1, a revised protocol for the non-imposed, non-interventional PASS should be submitted in clean and change-track format taking into account the comments adopted by PRAC

Action: For adoption of advice to CHMP

7.2.14. Tofacitinib - XELJANZ (CAP) - EMEA/H/C/004214/MEA 025.3

Applicant: Pfizer Europe MA EEIG

PRAC Rapporteur: Liana Martirosyan

Scope: From initial MAA: PASS DUS A3921403 (non-imposed RMP Cat. 3); PASS of the utilisation and prescribing patterns of Xeljanz (tofacitinib) using an administrative healthcare database in France, is a descriptive drug utilisation study using real-world data collected from routine clinical care in France. The overall goal is to determine if there is evidence that prescribers in France are compliant with the recommendations and limitations for use described in the tofacitinib additional risk minimisation measures (aRMM) materials

Action: For adoption of advice to CHMP

7.3. Results of PASS imposed in the marketing authorisation(s)[24]

7.3.1. Levofloxacin – QUINSAIR (CAP) - EMEA/H/C/PSR/S/0046

Applicant: Chiesi Farmaceutici S.p.A.

PRAC Rapporteur: Maria del Pilar Rayon

Scope: Final study report for a post-marketing, observational safety study of Quinsair (levofloxacin hemihydrate) in patients with cystic fibrosis to evaluate the long-term safety

[24] In accordance with Article 107p-q of Directive 2001/83/EC

compared to other inhaled approved antibiotic therapies in cystic fibrosis patients

Action: For adoption of recommendation to CHMP (or request for supplementary information (RSI)

7.4. Results of PASS non-imposed in the marketing authorisation(s)[25]

7.4.1. Dolutegravir, Lamivudine - DOVATO (CAP) - EMEA/H/C/004909/WS2620/0047; Dolutegravir, Rilpivirine - JULUCA (CAP) - EMEA/H/C/004427/WS2620/0056; Dolutegravir - TIVICAY (CAP) - EMEA/H/C/002753/WS2620/0092; Dolutegravir, Abacavir, Lamivudine - TRIUMEQ (CAP) - EMEA/H/C/002754/WS2620/0118

Applicant: ViiV Healthcare B.V.

PRAC Rapporteur: Martin Huber

Scope: Update of section 4.6 of the SmPC in order to update information about the use of dolutegravir-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 s a non-interventional study to Assess 'real-world' maternal and foetal outcomes following DTG use during pregnancy and to describe patterns of dolutegravir utilisation; 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

Action: For adoption of PRAC Assessment Report

7.4.2. Epoetin alfa - ABSEAMED (CAP) - EMEA/H/C/000727/WS2615/0108; Epoetin alfa - BINOCRIT (CAP) - EMEA/H/C/000725/WS2615/0108; Epoetin alfa - EPOETIN ALFA HEXAL (CAP) - EMEA/H/C/000726/WS2615/0108

Applicant: Sandoz GmbH

PRAC Rapporteur: Tiphaine Vaillant

Scope: 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 study (NIS PASS) study HX575-507 was conducted to address a post-approval requirement (MEA 13.5) to evaluate the safety profile of HX575 administered subcutaneous (s.c.) in patients with CKD-induced anaemia 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

Action: For adoption of PRAC Assessment Report

[25] In accordance with Article 61a (6) of Regulation (EC) No 726/2004, in line with the revised variations regulation for any submission as of 4 August 2013

7.4.3. Guanfacine - INTUNIV (CAP) - EMEA/H/C/003759/II/0033/G

Applicant: Takeda Pharmaceuticals International AG Ireland Branch

PRAC Rapporteur: Maria del Pilar Rayon

Scope: Submission of the final reports from the Drug Utilisation Study of Intuniv (guanfacine extended release) in European countries: a prescriber survey (EUPAS18739) and a retrospective database study (EUPAS18735), listed as category 3 studies in the RMP. The RMP version 4.0 has also been submitted

Action: For adoption of PRAC Assessment Report

7.4.4. Human normal immunoglobulin - HYQVIA (CAP) - EMEA/H/C/002491/II/0096

Applicant: Baxalta Innovations GmbH

PRAC Rapporteur: Gabriele Maurer

Scope: Update of sections 4.8 and 5.1 of the SmPC in order to update long-term safety information based on final results from studies 161406 'non-interventional post-marketing safety study on the long-term safety of HYQVIA (Global)' listed as category 3 a study in the RMP and 161302 'non-interventional post-authorisation safety study on the long-term safety of HyQvia in subjects treated with HyQvia'. Both studies were non-interventional, prospective, uncontrolled, multicentre, open-label, post-authorisation studies. The RMP version 15.0 has also been submitted. In addition, the MAH took the opportunity to bring the product information in line with the latest QRD template version 10.3, to update the list of local representatives in the package leaflet and to introduce minor editorial changes to the product information

Action: For adoption of PRAC Assessment Report

7.4.5. Naloxegol - MOVENTIG (CAP) - EMEA/H/C/002810/II/0043

Applicant: Gruenenthal GmbH

PRAC Rapporteur: Eamon O'Murchu

Scope: Submission of the final report from the PASS study D3820R0008 listed as a category 3 study in the RMP. This is a US post-marketing, comparative, observational study to evaluate the cardiovascular safety of Naloxegol in patients with non-cancer pain in comparison to other treatments for opioid induced constipation. The RMP version 9.0 has also been submitted

Action: For adoption of PRAC Assessment Report

7.4.6. Piperaquine tetraphosphate, artenimol - EURARTESIM (CAP) - EMEA/H/C/001199/II/0040/G

Applicant: Alfasigma S.p.A.

PRAC Rapporteur: Martin Huber

Scope: Grouped variation consisting of: 1) Submission of the final report from the effectiveness evaluation survey for Eurartesim (protocol no. 3366) (listed as a category 3

study in the RMP): an European multi-centre online survey to assess physician understanding of the revised edition of the educational material. Consequential changes to RMP version 16.1 have been implemented; 2) Submission of an updated RMP version 16.1 in order to delete 'severe malaria' as missing information from the list of safety specifications

Action: For adoption of PRAC Assessment Report

7.5. Interim results of imposed and non-imposed PASS submitted before the entry into force of the revised variation regulation

7.5.1. Abatacept - ORENCIA (CAP) - EMEA/H/C/000701/MEA 048.12

Applicant: Bristol-Myers Squibb Pharma EEIG

PRAC Rapporteur: Kimmo Jaakkola

Scope: From II-0024: Study number: IM101240 JIA registry; Yearly Recruitment update in the context of MEA-048; An Observational Registry of Abatacept in Patients with Juvenile Idiopathic Arthritis is ongoing. The primary objective is to describe the long-term safety of abatacept treatment for juvenile idiopathic arthritis (JIA) in routine clinical practice by quantifying the incidence rates of serious infections, autoimmune disorders, and malignancies. The data in these studies do not change the safety profile of abatacept. The MAH will supply interim reports from the above-mentioned studies according to the set schedules. Recruitment updates are provided each February, and interim reports will be submitted in 2014, 2019, and 2024. The planned date for submission of final data is 2029

Action: For adoption of advice to CHMP

7.5.2. Alemtuzumab - LEMTRADA (CAP) - EMEA/H/C/003718/ANX 009.6

Applicant: Sanofi Belgium

PRAC Rapporteur: Karin Erneholm

Scope: MAH's response to MEA 009.4 request for supplementary information (RSI) and updated protocol [feasibility analysis report for the Cat. 1 non-interventional PASS to investigate the risk of mortality in multiple sclerosis (MS) patients treated with alemtuzumab (LEMTRADA) relative to comparable MS patients using other disease modifying treatments (DMTs)] as adopted by PRAC in January 2024

Action: For adoption of advice to CHMP

7.5.3. Chenodeoxycholic acid - CHENODEOXYCHOLIC ACID LEADIANT (CAP) - EMEA/H/C/004061/SOB 003.2

Applicant: Leadiant GmbH

PRAC Rapporteur: Adam Przybylkowski

Scope: Cerebrotendinous Xanthomatosis Registry: Long term Non-Interventional Follow-up of Safety and Effectiveness of Chenodeoxycholic Acid Leadiant. (Refer also to SOB-001) In order to collect long term safety and efficacy data in patients treated with

chenodeoxycholic acid, the MAH will submit the results of a study deriving from a registry of patients with inborn errors of primary bile acid synthesis due to sterol 27 hydroxylase deficiency in infants, children and adolescents aged 1 month to 18 years and adults

Action: For adoption of advice to CHMP

7.5.4. Cinacalcet - MIMPARA (CAP) - EMEA/H/C/000570/MEA 035.9

Applicant: Amgen Europe B.V.

PRAC Rapporteur: Mari Thorn

Scope: From X-0055: Observational registry study (Study 20180204): To evaluate the risk of hypocalcaemia (e.g., clinical characteristics, laboratory variables [PTH, Ca, and P], hospitalisation due to hypocalcaemia, co-medication, cinacalcet doses) in paediatric patients treated with cinacalcet

Action: For adoption of advice to CHMP

7.5.5. Damoctocog alfa pegol - JIVI (CAP) - EMEA/H/C/004054/ANX 001.3

Applicant: Bayer AG

PRAC Rapporteur: Bianca Mulder

Scope: MAH's response to questions on ANX 001.2 [Second Interim Report: Study number 20904 (HA-SAFE)] as adopted in January 2024

Action: For adoption of advice to CHMP

7.5.6. Erenumab - AIMOVIG (CAP) - EMEA/H/C/004447/MEA 001.3

Applicant: Novartis Europharm Limited

PRAC Rapporteur: Kirsti Villikka

Scope: From Initial MAA: NIS - A Non-Interventional Study to examine patient characteristics and drug utilisation patterns in migraine patients treated with prophylactic drugs in the Nordic registries

Action: For adoption of advice to CHMP

7.5.7. Lenvatinib - LENVIMA (CAP) - EMEA/H/C/003727/MEA 014.7

Applicant: Eisai GmbH

PRAC Rapporteur: Ulla Wändel Liminga

Scope: From II-0011-G: Third annual progress report for PASS No. E7080-M000-508; HCC: To assess the severity of hepatotoxicity and factors that contribute to hepatoxicity of lenvatinib, and the influence on overall survival in the Western population.
A multicentre, observational post authorisation safety study (PASS), phase 4 study to evaluate the safety and tolerability of lenvatinib in patients with advanced or unresectable hepatocellular carcinoma (STELLAR)

Action: For adoption of advice to CHMP

7.5.8. Tozinameran - COMIRNATY (CAP) - EMEA/H/C/005735/MEA 054.2

Applicant: BioNTech Manufacturing GmbH

PRAC Rapporteur: Liana Martirosyan

Scope: Study C4591022; Pfizer-BioNTech COVID-19 Vaccine Exposure during Pregnancy: A Non-Interventional Post-Approval Safety Study of Pregnancy and Infant Outcomes in the Organization of Teratology Information Specialists (OTIS)/MotherToBaby Pregnancy Registry

Action: For adoption of advice to CHMP

7.6. Others

None

7.7. New Scientific Advice

None

7.8. Ongoing Scientific Advice

None

7.9. Final Scientific Advice (Reports and Scientific Advice letters)

None

8. Renewals of the marketing authorisation, conditional renewal and annual reassessments

8.1. Annual reassessments of the marketing authorisation

8.1.1. Glucarpidase - VORAXAZE (CAP) - EMEA/H/C/005467/S/0025 (without RMP)

Applicant: SERB S.A.S.

PRAC Rapporteur: Martin Huber

Scope: Annual reassessment of the marketing authorisation

Action: For adoption of advice to CHMP

8.1.2. Histamine dihydrochloride - CEPLENE (CAP) - EMEA/H/C/000796/S/0048 (without RMP)

Applicant: Laboratoires Delbert

PRAC Rapporteur: Eamon O'Murchu

Scope: Annual reassessment of the marketing authorisation

Action: For adoption of advice to CHMP

8.1.3. Susoctocog alfa - OBIZUR (CAP) - EMEA/H/C/002792/S/0056 (without RMP)

Applicant: Baxalta Innovations GmbH

PRAC Rapporteur: Gabriele Maurer

Scope: Annual reassessment of the marketing authorisation

Action: For adoption of advice to CHMP

8.1.4. Tabelecleucel - EBVALLO (CAP) - EMEA/H/C/004577/S/0008 (without RMP)

Applicant: Pierre Fabre Medicament, ATMP

PRAC Rapporteur: Amelia Cupelli

Scope: Annual reassessment of the marketing authorisation

Action: For adoption of advice to CAT and CHMP

8.1.5. Tagraxofusp - ELZONRIS (CAP) - EMEA/H/C/005031/S/0025 (without RMP)

Applicant: Stemline Therapeutics B.V.

PRAC Rapporteur: Bianca Mulder

Scope: Annual reassessment of the marketing authorisation

Action: For adoption of advice to CHMP

8.1.6. Tecovirimat - TECOVIRIMAT SIGA (CAP) - EMEA/H/C/005248/S/0010 (without RMP)

Applicant: SIGA Technologies Netherlands B.V.

PRAC Rapporteur: Martin Huber

Scope: Annual reassessment of the marketing authorisation

Action: For adoption of advice to CHMP

8.2. Conditional renewals of the marketing authorisation

8.2.1. Avapritinib - AYVAKYT (CAP) - EMEA/H/C/005208/R/0034 (without RMP)

Applicant: Blueprint Medicines (Netherlands) B.V.

PRAC Rapporteur: Bianca Mulder

Scope: Conditional renewal of the marketing authorisation

Action: For adoption of advice to CHMP

8.2.2. Epcoritamab - TEPKINLY (CAP) - EMEA/H/C/005985/R/0004 (without RMP)

Applicant: AbbVie Deutschland GmbH & Co. KG

PRAC Rapporteur: Monica Martinez Redondo

Scope: Conditional renewal of the marketing authorisation

Action: For adoption of advice to CHMP

8.2.3. Imlifidase - IDEFIRIX (CAP) - EMEA/H/C/004849/R/0020 (without RMP)

Applicant: Hansa Biopharma AB

PRAC Rapporteur: Bianca Mulder

Scope: Conditional renewal of the marketing authorisation

Action: For adoption of advice to CHMP

8.2.4. Larotrectinib - VITRAKVI (CAP) - EMEA/H/C/004919/R/0035 (without RMP)

Applicant: Bayer AG

PRAC Rapporteur: Rugile Pilviniene

Scope: Conditional renewal of the marketing authorisation

Action: For adoption of advice to CHMP

8.2.5. Tafasitamab - MINJUVI (CAP) - EMEA/H/C/005436/R/0015 (without RMP)

Applicant: Incyte Biosciences Distribution B.V.

PRAC Rapporteur: Ulla Wändel Liminga

Scope: Conditional renewal of the marketing authorisation

Action: For adoption of advice to CHMP

8.2.6. Valoctocogene roxaparvovec - ROCTAVIAN (CAP) - EMEA/H/C/005830/R/0011 (with RMP)

Applicant: BioMarin International Limited, ATMP

PRAC Rapporteur: Bianca Mulder

Scope: Conditional renewal of the marketing authorisation

Action: For adoption of advice to CAT and CHMP

8.3. Renewals of the marketing authorisation

8.3.1. Arsenic trioxide - ARSENIC TRIOXIDE ACCORD (CAP) - EMEA/H/C/005175/R/0009 (without RMP)

Applicant: Accord Healthcare S.L.U.

PRAC Rapporteur: Tiphaine Vaillant

Scope: 5-year renewal of the marketing authorisation

Action: For adoption of advice to CHMP

8.3.2. Bortezomib - BORTEZOMIB FRESENIUS KABI (CAP) - EMEA/H/C/005074/R/0010 (without RMP)

Applicant: Fresenius Kabi Deutschland GmbH

PRAC Rapporteur: Amelia Cupelli

Scope: 5-year renewal of the marketing authorisation

Action: For adoption of advice to CHMP

8.3.3. Cannabidiol - EPIDYOLEX (CAP) - EMEA/H/C/004675/R/0031 (with RMP)

Applicant: Jazz Pharmaceuticals Ireland Limited

PRAC Rapporteur: Ana Sofia Diniz Martins

Scope: 5-year renewal of the marketing authorisation

Action: For adoption of advice to CHMP

8.3.4. Deferasirox - DEFERASIROX MYLAN (CAP) - EMEA/H/C/005014/R/0013 (without RMP)

Applicant: Mylan Pharmaceuticals Limited

PRAC Rapporteur: Tiphaine Vaillant

Scope: 5-year renewal of the marketing authorisation

Action: For adoption of advice to CHMP

8.3.5. Upadacitinib - RINVOQ (CAP) - EMEA/H/C/004760/R/0051 (without RMP)

Applicant: AbbVie Deutschland GmbH & Co. KG

PRAC Rapporteur: Petar Mas

Scope: 5-year renewal of the marketing authorisation

Action: For adoption of advice to CHMP

9. Product related pharmacovigilance inspections

9.1. List of planned pharmacovigilance inspections

None

9.2. Ongoing or concluded pharmacovigilance inspections

Disclosure of information on results of pharmacovigilance inspections could undermine the protection of the purpose of these inspections, investigations and audits. Therefore such information is not reported in the agenda.

9.3. Others

None

10. Other safety issues for discussion requested by the CHMP or the EMA

10.1. Safety related variations of the marketing authorisation

None

10.2. Timing and message content in relation to Member States' safety announcements

None

10.3. Other requests

None

10.4. Scientific Advice

Information related to this section cannot be released at the present time as it is deemed to contain commercially confidential information.

11. Other safety issues for discussion requested by the Member States

11.1. Safety related variations of the marketing authorisation

None

11.2. Other requests

None

12. Organisational, regulatory and methodological matters

12.1. Mandate and organisation of the PRAC

12.1.1. PRAC membership

Action: For information

12.1.2. Vote by proxy

None

12.1.3. PRAC working group - Best practice guide on using PRAC plenary time efficiently and effectively – update on the implementation of quantitative goals – Q2 2023

Action: For discussion

12.2. Coordination with EMA Scientific Committees or CMDh-v

None

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

None

12.4. Cooperation within the EU regulatory network

12.4.1. Health threats and EMA Emergency Task Force (ETF) activities - update

Action: For discussion

12.5. Cooperation with International Regulators

12.5.1. International Council for Harmonisation (ICH) M14 Guideline on 'General principles on plan, design and analysis of pharmacoepidemiological studies that utilize real-world data for safety assessment of medicines'

PRAC lead: Annalisa Capuano

Action: For discussion

12.6. Contacts of PRAC with external parties and interaction with the Interested Parties to the Committee

None

12.7. PRAC work plan

None

12.8. Planning and reporting

12.8.1. EU Pharmacovigilance system - quarterly workload measures and performance indicators – Q1 2024 and predictions

Action: For discussion

12.8.2. PRAC workload statistics – Q1 2024

Action: For discussion

12.9. Pharmacovigilance audits and inspections

12.9.1. Pharmacovigilance systems and their quality systems

None

12.9.2. Pharmacovigilance inspections

None

12.9.3. Pharmacovigilance audits

None

12.10. Periodic safety update reports (PSURs) & Union reference date (EURD) list

12.10.1. Periodic safety update reports

None

12.10.2. Granularity and Periodicity Advisory Group (GPAG)

PRAC lead: Jana Lukačišinová

Action: For discussion

12.10.3. PSURs repository

None

12.10.4. Union reference date list – consultation on the draft list

Action: For adoption

12.11. Signal management

12.11.1. Signal management – feedback from Signal Management Review Technical (SMART) Working Group

PRAC lead(s): Martin Huber, Jean-Michel Dogné

Action: For discussion

12.12. Adverse drug reactions reporting and additional reporting

12.12.1. Management and reporting of adverse reactions to medicinal products

None

12.12.2. Additional monitoring

None

12.12.3. List of products under additional monitoring – consultation on the draft list

Action: For adoption

12.13. EudraVigilance database

12.13.1. Activities related to the confirmation of full functionality

None

12.14. Risk management plans and effectiveness of risk minimisations

12.14.1. Risk management systems

None

12.14.2. Tools, educational materials and effectiveness measurement of risk minimisations

None

12.15. Post-authorisation safety studies (PASS)

12.15.1. Post-authorisation Safety Studies – imposed PASS

None

12.15.2. Post-authorisation Safety Studies – non-imposed PASS

None

12.16. Community procedures

12.16.1. Referral procedures for safety reasons

None

12.17. Renewals, conditional renewals, annual reassessments

None

12.18. Risk communication and transparency

12.18.1. Public participation in pharmacovigilance

None

12.18.2. Safety communication

None

12.19. Continuous pharmacovigilance

12.19.1. Incident management

None

12.20. Impact of pharmacovigilance activities

12.21. Others

12.21.1. Antiepileptic drugs and pregnancy

Action: For discussion

13. Any other business

Next meeting on 10-13 June 2024

14. Explanatory notes

The Notes give a brief explanation of relevant agenda items and should be read in conjunction with the agenda.

EU Referral procedures for safety reasons: Urgent EU procedures and Other EU referral procedures
(Items 2 and 3 of the PRAC agenda)

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 European Union (EU). For further detailed information on safety related referrals please see: Referral procedures: human medicines | European Medicines Agency (europa.eu)

Signals assessment and prioritisation
(Item 4 of the PRAC agenda)

A safety signal is information on a new or incompletely documented adverse event that is potentially caused by a medicine and that warrants further investigation. Signals are generated from several sources such as spontaneous reports, clinical studies and the scientific literature. The evaluation of safety signals is a routine part of pharmacovigilance and is essential to ensuring that regulatory authorities have a comprehensive knowledge of a medicine's benefits and risks.
The presence of a safety signal does not mean that a medicine has caused the reported adverse event. The adverse event could be a symptom of another illness or caused by another medicine taken by the patient. The evaluation of safety signals is required to establish whether or not there is a causal relationship between the medicine and the reported adverse event.
The evaluation of safety signals may not necessarily conclude that the medicine caused the adverse event in question. In cases where a causal relationship is confirmed or considered likely, regulatory action may be necessary and this usually takes the form of an update of the summary of product characteristics and the package leaflet.

Risk Management Plans (RMPs)
(Item 5 of the PRAC agenda)

The RMP describes what is known and not known about the side effects of a medicine and states how these risks will be prevented or minimised in patients. It also includes plans for studies and other activities to gain more knowledge about the safety of the medicine and risk factors for developing side effects.
RMPs are continually modified and updated throughout the lifetime of the medicine as new information becomes available.

Assessment of Periodic Safety Update Reports (PSURs)
(Item 6 of the PRAC agenda)

A PSUR is a report providing an evaluation of the benefit-risk balance of a medicine, which is submitted by marketing authorisation holders at defined time points following a medicine's authorisation.
PSURs summarises data on the benefits and risks of a medicine and includes the results of all studies carried

out with this medicine (in the authorised and unauthorised indications).

Post-authorisation Safety Studies (PASS)
(Item 7 of the PRAC agenda)

A PASS is a study of an authorised medicinal product carried out to obtain further information on its safety, or to measure the effectiveness of risk management measures. The results of a PASS help regulatory agencies to evaluate the safety and benefit-risk profile of a medicine.

Product related pharmacovigilance inspections
(Item 9 of the PRAC agenda)

Inspections carried out by regulatory agencies to ensure that marketing authorisation holders comply with their pharmacovigilance obligations.

For a list of acronyms and abbreviations, see:
List of abbreviations used in EMA human medicines scientific committees and CMDh documents, and in Pharmacovigilance Risk Assessment Committee (PRAC)

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