06 July 2015
EMA/PRAC/455682/2015
Management and Committees Support Division

Pharmacovigilance Risk Assessment Committee (PRAC)

Draft agenda for the meeting on 06-09 July 2015

Chair: June Raine – Vice-Chair: Almath Spooner

06 July 2015, 13:00 – 19:00, room 3/A

07 July 2015, 08:30 – 19:00, room 3/A

08 July 2015, 08:30 – 19:00, room 3/A

09 July 2015, 08:30 – 16:00, room 3/A

Organisational, regulatory and methodological matters (ORGAM)

23 July 2015, 10:00-12:00, room 6/B, 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).

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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 8-11 June 2015. See (current) July 2015 PRAC minutes (to be published post September 2015 PRAC meeting).

1.2. Adoption of agenda for the meeting on 6-9 July 2015

1.3. Adoption of minutes of the previous meeting on 8-11 June 2015

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

2.4. Planned public hearings

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

None

3.4. Article 5(3) of Regulation (EC) No 726/2004 as amended: PRAC advice on CHMP request

None

3.5. Others

3.5.1. Ambroxol (NAP); bromhexine (NAP)

MAH(s): Boehringer Ingelheim, various

PRAC Rapporteur: Margarida Guimarães
PRAC Co-rapporteur: Jean-Michel Dogné, Jan Neuhauser

Scope: Review of recommendations of a referral procedure under Article 31 of Directive 2001/83/EC adopted in January 2015, at the request of the European Commission
Action: For discussion

4. Signals assessment and prioritisation[1]

4.1. New signals detected from EU spontaneous reporting systems

4.1.1. Human fibrinogen, human thrombin – TACHOSIL (CAP)

Applicant: Takeda Austria GmbH

PRAC Rapporteur: Brigitte Keller-Stanislawski

Scope: Signal of intestinal obstruction
Action: For adoption of PRAC recommendation
EPITT 18373 – New signal

4.1.2. Ipilimumab – YERVOY (CAP)

Applicant: Bristol-Myers Squibb Pharma EEIG

PRAC Rapporteur: Sabine Straus

Scope: Signal of Vogt-Koyanagi-Harada syndrome (VKH)
Action: For adoption of PRAC recommendation
EPITT 18403 – New signal

4.1.3. Palifermin – KEPIVANCE (CAP)

Applicant: Swedish Orphan Biovitrum AB

PRAC Rapporteur: Rafe Suvarna

Scope: Signal of infection
Action: For adoption of PRAC recommendation
EPITT 18401 – New signal

[1] 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

4.1.4. Saxagliptin – ONGLYZA (CAP); saxagliptin, metformin – KOMBOGLYZE (CAP)

Applicant: AstraZeneca AB

PRAC Rapporteur: Menno van der Elst

Scope: Signal of acute kidney injury
Action: For adoption of PRAC recommendation
EPITT 18379 – New signal

4.2. New signals detected from other sources

None

4.3. Signals follow-up and prioritisation

4.3.1. Adalimumab – HUMIRA (CAP) – EMEA/H/C/00000481/SDA/0242

Applicant: AbbVie Ltd.

PRAC Rapporteur: Ulla Wändel Liminga

Scope: Signal of convulsion
Action: For adoption of PRAC recommendation
EPITT 18211 – Follow-up to March 2015

4.3.2. Amiodarone (NAP)

Applicant: Sanofi, various

PRAC Rapporteur: Menno van der Elst

Scope: Signal of pancreatitis
Action: For adoption of PRAC recommendation
EPITT 18216 – Follow-up to March 2015

4.3.3. Donepezil (NAP)

Applicant: Eisai Ltd.

PRAC Rapporteur: Julie Williams

Scope: Signal of rhabdomyolysis
Action: For adoption of PRAC recommendation
EPITT 18261 – Follow-up to March 2015

4.3.4. Infliximab - INFLECTRA (CAP) - EMEA/H/C/002778/SDA/020, REMICADE (CAP) - EMEA/H/C/000240/SDA/152, REMSIMA (CAP) - EMEA/H/C/002576/SDA/019

Applicant: Hospira UK Limited (Inflectra), Janssen Biologics B.V. (Remicade), Celltrion Healthcare Hungary Kft. (Remsima)

PRAC Rapporteur: Ulla Wändel Liminga

Scope: Signal of rhabdomyolysis
Action: For adoption of PRAC recommendation
EPITT 18129– Follow-up to March 2015

4.3.5. Pantoprazole – CONTROLOC CONTROL (CAP) - EMEA/H/C/001097/SDA/014, PANTECTA CONTROL (CAP) - EMEA/H/C/001099/SDA/014, PANTOLOC CONTROL (CAP) - EMEA/H/C/001100/SDA/013, PANTOZOL CONTROL (CAP) - EMEA/H/C/001013/SDA/014, SOMAC CONTROL (CAP) - EMEA/H/C/001098/SDA/019

Applicant: Takeda GmbH

PRAC Rapporteur: Rafe Suvarna

Scope: Signal of subacute cutaneous lupus erythematosus (SCLE)
Action: For adoption of PRAC recommendation
EPITT 18119 – Follow-up to April 2015

4.3.6. Warfarin (NAP)

Applicant: various

PRAC Rapporteur: Torbjörn Callreus

Scope: Signal of bone density decrease
Action: For adoption of PRAC recommendation
EPITT 18173 – Follow-up to March 2015

5. Risk management plans (RMPs)

5.1. Medicines in the pre-authorisation phase

5.1.1. Albutrepenonacog alfa - EMEA/H/C/003955

Scope: Prophylaxis and treatment of bleeding in all patients with haemophilia B, treatment of bleeding in all patients with haemophilia B
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.1.2. Alirocumab - EMEA/H/C/003882

Scope: Reduction of low-density lipoprotein cholesterol (LDL-C) and increase high-density lipoprotein cholesterol (HDL-C)
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.1.3. Aripiprazole - EMEA/H/C/004021

Generic

Scope: Treatment of schizophrenia and treatment and prevention of manic episodes in bipolar I disorder
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.1.4. Blinatumomab - EMEA/H/C/003731 - Orphan

Applicant: Amgen Europe B.V.

Scope: Treatment of adults with Philadelphia chromosome negative relapsed or refractory B-precursor acute lymphoblastic leukaemia
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.1.5. Bortezomib - EMEA/H/C/004076

Generic

Scope: Treatment of multiple myeloma
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.1.6. Caspofungin - EMEA/H/C/004134

Generic

Scope: Treatment of invasive candidiasis and invasive aspergillosis
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.1.7. Ceftolozane, tazobactam - EMEA/H/C/003772

Scope: Treatment of intra-abdominal urinary tract infections
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.1.8. Daclizumab - EMEA/H/C/003862

Scope: Treatment of relapsing multiple sclerosis (RMS)
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.1.9. Efmoroctocog alfa - EMEA/H/C/003964 - Orphan

Applicant: Biogen Idec Ltd

Scope: Treatment of haemophilia A
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.1.10. Elvitegravir, cobicistat, emtricitabine, tenofovir alafenamide - EMEA/H/C/004042

Scope: Treatment of human immunodeficiency virus (HIV)-1
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.1.11. Enoxaparin sodium - EMEA/H/C/004264, EMEA/H/C/003795

Biosimilar

Scope: Prophylaxis of thromboembolic disorders of venous origin
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.1.12. Fentanyl - EMEA/H/C/002715

Scope: Treatment of acute moderate to severe post-operative pain
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.1.13. Ferric citrate coordination complex - EMEA/H/C/003776

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

5.1.14. Guanfacine - EMEA/H/C/003759

Scope: Treatment of attention deficit hyperactivity disorder (ADHD)

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

5.1.15. Idarucizumab - EMEA/H/C/003986

Scope: Prevention and treatment of dabigatran associated haemorrhage
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.1.16. Infliximab - EMEA/H/C/004020

Biosimilar

Scope: Treatment of rheumatoid arthritis, Crohn's disease, ankylosing spondylitis, psoriatic arthritis, psoriasis and ulcerative colitis
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.1.17. Isavuconazole - EMEA/H/C/002734 - Orphan

Applicant: Basilea Medical Ltd

Scope: Treatment of aspergillosis and mucormycosis
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.1.18. Levodopa, carbidopa - EMEA/H/C/002611

Scope: Treatment of Parkinson's disease
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.1.19. Lumacaftor, ivacaftor - EMEA/H/C/003954 - Orphan

Applicant: Vertex Pharmaceuticals (U.K.) Ltd

Scope: Treatment of cystic fibrosis
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.1.20. Mepolizumab - EMEA/H/C/003860

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

5.1.21. Methotrexate - EMEA/H/C/003756

Hybrid

Scope: Treatment of rheumatological and dermatological diseases
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.1.22. Miglustat - EMEA/H/C/004016

Generic

Scope: Treatment of Gaucher disease
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.1.23. Pandemic influenza vaccine H5N1 (live attenuated, nasal) - EMEA/H/C/003963

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

5.1.24. Pegaspargase - EMEA/H/C/003789

Scope: Treatment therapy in acute lymphoblastic leukaemia (ALL)
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.1.25. Pemetrexed - EMEA/H/C/003788

Generic

Scope: Treatment of malignant pleural mesothelioma and non-small cell lung cancer
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.1.26. Pemetrexed - EMEA/H/C/003970

Generic

Scope: Treatment of malignant pleural mesothelioma and non-small cell lung cancer
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.1.27. Pemetrexed - EMEA/H/C/004114

Generic

Scope: Treatment of malignant pleural mesothelioma and non-small cell lung cancer
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.1.28. Pemetrexed - EMEA/H/C/003905

Generic

Scope: Treatment of malignant pleural mesothelioma and non-small cell lung cancer
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.1.29. Pemetrexed - EMEA/H/C/004011

Generic

Scope: Treatment of malignant pleural mesothelioma and non-small cell lung cancer
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.1.30. Plasmodium falciparum circumsprozoite protein fused with hepatitis B surface antigen (rts) and combined with hepatitis B surface antigen(s) in the form of non-infectious virus-like particles (vlps) produced in yeast cells (saccharomyces cerevisiae) by recombinant DNA technology) - EMEA/H/W/002300

Scope: Active immunisation against malaria
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.1.31. Rasagiline - EMEA/H/C/004064

Generic

Scope: Treatment of idiopathic Parkinson's disease (PD)
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.1.32. Sufentanil - EMEA/H/C/002784

Hybrid

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

5.1.33. Susoctocog alfa - EMEA/H/C/002792 – Orphan

Applicant: Baxter AG

Scope: Treatment of haemophilia A
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.1.34. Trifluridine, tipiracil - EMEA/H/C/003897

Scope: Treatment of colorectal cancer
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. Filgrastim – FILGRASTIM HEXAL (CAP) - EMEA/H/C/000918/WS0779/0029/G; ZARZIO (CAP) - EMEA/H/C/000917/WS0779/0030/G

Applicant: Sandoz GmbH

PRAC Rapporteur: Julie Williams

Scope: Submission of long term safety and immunogenicity data in additional studies EP06-302 to address post authorisation measure MEA 005 and submission of a revised RMP (version 11) to include two important potential risks (extramedullary haematopoiesis and venous thrombotic events) following the assessment of PSUR#13
Action: For adoption of PRAC AR

5.2.2. Filgrastim – ACCOFIL (CAP) - EMEA/H/C/003956/II/0002

Applicant: Accord Healthcare Ltd

PRAC Rapporteur: Julie Williams

Scope: Update of the RMP following a product information update with regard to routine risk minimisation measures of for several safety concerns
Action: For adoption of PRAC AR

5.2.3. Micafungin – MYCAMINE (CAP) - EMEA/H/C/000734/II/0026

Applicant: Astellas Pharma Europe B.V.

PRAC Rapporteur: Martin Huber

Scope: Submission of a revised RMP in order to update the important identified risk of drug interaction; include a second survey that will be conducted in Q1 2015 to further assess the effectiveness of risk minimization measures as requested by the PRAC in May 2014
Action: For adoption of PRAC AR

5.2.4. Tadalafil – ADCIRCA (CAP) - EMEA/H/C/001021/WS0762/0021; CIALIS (CAP) – EMEA/H/C/000436/WS0762/0078

Applicant: Eli Lilly Nederland B.V.

PRAC Rapporteur: Miguel-Angel Macia

Scope: Update of the due date of the final study report of the category 3 study LVHQ from 'Q3 2015' to 'Q2 2016', which addresses the specific safety concern of non-arteritic anterior ischemic optic neuropathy (NAION), in the tadalafil EU RMP. In addition, other minor changes have been made to the RMP, mainly to update the exposure numbers in line with an updated data cut-off date
Action: For adoption of PRAC AR

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

5.3.1. Abatacept – ORENCIA (CAP) - EMEA/H/C/000701/II/0089

Applicant: Bristol-Myers Squibb Pharma EEIG

PRAC Rapporteur: Kirsti Villikka

Scope: Update of section 4.6 of the SmPC in order to update the safety information on the risk of infection associated with live vaccination in infants born to women treated with abatacept during pregnancy. The Package Leaflet is updated accordingly
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.2. Alipogene triparvovec – GLYBERA (CAP) - EMEA/H/C/002145/II/0038

Applicant: UniQure biopharma B.V.

PRAC Rapporteur: Julie Williams

Scope: Update of section 5.1 of the SmPC based on the final CSR for study CT-AMT-011-05, a retrospective clinical records review study undertaken to generate further long-term follow-up data on the incidence and severity of acute pancreatitis episodes in LPLD subjects who previously participated in clinical studies with alipogene tiparvovec or AMT-10
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.3. Ambrisentan – VOLIBRIS (CAP) - EMEA/H/C/000839/II/0041

Applicant: Glaxo Group Ltd

PRAC Rapporteur: Dolores Montero Corominas

Scope: Update of sections 4.1, 4.2, 4.4, 4.8 and 5.1 of the SmPC in order to include an expanded therapeutic indication for the treatment of pulmonary arterial hypertension (PAH) (World Health Organization [WHO] Group 1). In addition, the MAH took the opportunity to update Annex II to reflect a change in the PSUR cycle. The Package Leaflet is updated accordingly
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.4. Atazanavir – REYATAZ (CAP) - EMEA/H/C/000494/X/0094/G

Applicant: Bristol-Myers Squibb Pharma EEIG

PRAC Rapporteur: Isabelle Robine

Scope: Grouped variation of 1) Line extension covering a new pharmaceutical form (oral powder), a new strength for the oral powder presentation (50 mg), and a new paediatric indication (patients from 3 months of age and weighing at least 5 kg); 2) addition of new paediatric data for the capsules; 3) minor revisions to the RMP with regard to nephrolithiasis following PRAC's assessment of RMP version 7.3
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.5. Atazanavir – REYATAZ (CAP) - EMEA/H/C/000494/II/0096

Applicant: Bristol-Myers Squibb Pharma EEIG

PRAC Rapporteur: Isabelle Robine

Scope: Update of sections 4.2, 4.3, 4.4, 4.5, 4.6, 5.1 and 5.2 of the SmPC in order to provide important information and guidance to prescribers when they consider using unboosted atazanavir (ATV) in line with international guidelines based on study INDUMA/AI424-136. In addition, the MAH took the opportunity to make a minor change in section 4.7 of the SmPC for increased clarity, and minor editorial changes to the SmPC and Package Leaflet. The RMP version 9 has been submitted
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.6. Bevacizumab – AVASTIN (CAP) - EMEA/H/C/000582/II/0082

Applicant: Roche Registration Ltd

PRAC Rapporteur: Doris Stenver

Scope: Update of sections 4.2, 4.4 and 4.8 of the SmPC to include information regarding osteonechrosis in children. The Package Leaflet is updated accordingly. The RMP is updated accordingly
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.7. Capsaicin – QUTENZA (CAP) - EMEA/H/C/000909/II/0039

Applicant: Astellas Pharma Europe B.V.

PRAC Rapporteur: Magda Pedro

Scope: Extension of indication to include treatment of diabetic patients with peripheral neuropathic pain based on the results of studies E05-CL-3004 (STEP) and E05-CL-3002 (PACE). As a consequence, sections 4.1, 4.4 and 4.8 of the SmPC have been updated; Annex II (additional risk minimisation measures) and the package leaflet have been updated accordingly. In addition, the MAH took the opportunity to implement minor editorial changes in the SmPC, Annex II, labelling and Package Leaflet. An updated RMP (version 18) was submitted accordingly
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.8. Ceritinib – ZYKADIA (CAP) - EMEA/H/C/003819/II/0001

Applicant: Novartis Europharm Ltd

PRAC Rapporteur: Ulla Wändel Liminga

Scope: Update of sections 4.2, 4.4 and 4.8 of the SmPC in order to update the safety information after the MAH's assessment of the association between the use of ceritinib and acute pancreatitis. The Package Leaflet is updated accordingly. In addition, the updated RMP version 2.5 is submitted accordingly
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.9. Collagenase clostridium histolyticum – XIAPEX (CAP) - EMEA/H/C/002048/II/0059

Applicant: Swedish Orphan Biovitrum AB (publ)

PRAC Rapporteur: Martin Huber

Scope: Update of sections 4.2, 4.4, 4.8, 5.1, 5.2 and 6.6 of the SmPC to include information related to the treatment of Dupuytren's contracture with two concurrent injections of Xiapex. The Package Leaflet and RMP have been updated accordingly
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.10. Conestat alfa – RUCONEST (CAP) - EMEA/H/C/001223/R/0023

Applicant: Pharming Group N.V

PRAC Rapporteur: Rafe Suvarna

Scope: Evaluation of an RMP submitted in the context of a 5-year renewal of the marketing authorisation
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.11. Crizotinib – XALKORI (CAP) - EMEA/H/C/002489/II/0024

Applicant: Pfizer Limited

PRAC Rapporteur: Corinne Fechant

Scope: Extension of indication for the first-line treatment of anaplastic lymphoma kinase (ALK)-positive advanced non-small-cell lung carcinoma (NSCLC). This variation is based on results of study A8081014. As a consequence, sections 4.2, 4.4, 4.8, 5.1 and 5.2 of the SmPC have been amended. The Package Leaflet is updated accordingly
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.12. Dabrafenib - TAFINLAR (CAP) – EMEA/H/C/002604/WS0736/0011 trametinib – MEKINIST (CAP) - EMEA/H/C/002643/WS0736/0008

Applicant: Novartis Europharm Ltd

PRAC Rapporteur: Ulla Wändel Liminga

Scope: Extension of indication to add a new therapeutic indication for the use in combination of trametinib and dabrafenib for the treatment of adult patients with unresectable or metastatic melanoma with a BRAF V600 mutation. As a consequence, sections 4.1, 4.2, 4.4, 4.5, 4.6, 4.8, 5.1, 5.3 of the SmPC are updated. The Package Leaflet and RMP are updated accordingly
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.13. Daclatasvir – DAKLINZA (CAP) - EMEA/H/C/003768/II/0008/G

Applicant: Bristol-Myers Squibb Pharma EEIG

PRAC Rapporteur: Margarida Guimarães

Scope: Update of sections 4.2, 4.4, 4.5, 4.8 and 5.1 of the SmPC in order to update the safety information based on the final results of clinical study AI444043 (daclatasvir (DCV) in combination with peginterferon alfa plus ribavirin in hepatitis C virus (HCV) and human immunodeficiency virus (HIV) co-infected treatment-naive subjects with genotype 1 (GT-1) infection).The Package Leaflet is updated accordingly
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.14. Deferasirox – EXJADE (CAP) - EMEA/H/C/000670/X/0043

Applicant: Novartis Europharm Ltd

PRAC Rapporteur: Corinne Fechant

Scope: Extension application for a new pharmaceutical form and new strengths (Exjade 90, 180 mg and 360 mg film-coated tablets)
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.15. Dimethyl fumarate – TECFIDERA (CAP) - EMEA/H/C/002601/WS0689/0011/G

Applicant: Biogen Idec Ltd

PRAC Rapporteur: Martin Huber

Scope: Update of section 4.4 of the SmPC in order to add a recommendation to consider interruption of treatment in patients with low lymphocyte counts (<0.5 x 109/L) persisting for more than six months and to monitor lymphocyte counts until recovery. Update of section 4.8 of the SmPC with information on observed low lymphocyte counts in clinical studies with Tecfidera and progressive multifocal leukoencephalopathy (PML) occurrence in the setting of severe and prolonged lymphopenia. Furthermore, the due dates of two commitments as part of the RMP have been revised
Action: For discussion (Report from the SAG Neurology dated 11 June 2015)

5.3.16. Epoetin beta – NEORECORMON (CAP) - EMEA/H/C/000116/II/0083

Applicant: Roche Registration Ltd

PRAC Rapporteur: Valerie Strassmann

Scope: Update of SmPC sections 4.2 and 4.4, upon request by PRAC following the assessment of PSU 047 and MEA 052.1, to include information related to the potential risk of retinopathy of prematurity (RoP). The Package Leaflet is updated accordingly. In addition, the MAH took the opportunity to align the annexes with the QRD template v.9. An updated RMP version 3.2 was provided
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.17. Fentanyl – INSTANYL (CAP) - EMEA/H/C/000959/X/0030/G

Applicant: Takeda Pharma A/S

PRAC Rapporteur: Isabelle Robine

Scope: Grouped variations to 1) line extension to add the new strength of 400 micrograms/dose in a multi-dose nasal spray in pack size of 10's, 20's, 30's & 40 doses; 2) replace the current multi-dose nasal spray by a new improved child resistant multi-dose nasal spray; 3) add a new pack-size of 30 doses for each current strength (50 micrograms/dose, 100 micrograms/dose & 200 micrograms/dose); 4) tighten the assay release limit of the multi-dose finished product to 98.0%-102.0%; 5) reduce the shelf life of all strengths of the multi-dose finished product to 24 months
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.18. Human hepatitis b immunoglobulin – ZUTECTRA (CAP) - EMEA/H/C/001089/II/0024

Applicant: Biotest Pharma GmbH

PRAC Rapporteur: Brigitte Keller Stanislawski

Scope: Extension of indication for the prevention of hepatitis B virus (HBV) re-infection in HBsAg and HBV-DNA negative patients at least one week – instead of the approved at least 6 months - after liver transplantation for hepatitis B induced liver failure. 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
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.19. Human normal immunoglobulin – KIOVIG (CAP) - EMEA/H/C/000628/II/0065/G

Applicant: Baxter AG

PRAC Rapporteur: Brigitte Keller-Stanislawski

Scope: Update of section 4.4 of the SmPC in order to include further information to avoid thromboembolic events and renal complications and to update information on aseptic meningitis syndrome. In addition, new information is provided on possible false positive testing of assays used for diagnosis of fungal infections depending on detection of beta-D-glucans. Update of section 4.8 of the SmPC in order to update the safety information and to change frequencies of existing side effects as revealed in several clinical studies in which Kiovig was used as an investigational medicinal product. The Package Leaflet is updated accordingly. In addition, the MAH took the opportunity to update the RMP in reference to the new warning for the false positive testing for ß-d-glucan and including the changes requested during assessment of procedure EMEA/H/C/0000628/II/0056
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.20. Ivacaftor – KALYDECO (CAP) - EMEA/H/C/002494/X/0034/G

Applicant: Vertex Pharmaceuticals (U.K.) Ltd

PRAC Rapporteur: Miguel-Angel Macia

Scope: Line extension to include a new pharmaceutical form (granules) and two new strengths (50 mg and 75 mg unit doses) to enable administration of Kalydeco to patients aged 2 to less than 6 years of age. Changes to SmPC sections 4.2, 4.4, 4.5, 4.8 and 5.2 are proposed to provide clarity and relevant updates in line with the proposed paediatric extension application. The Package Leaflet is updated accordingly
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.21. Lenalidomide – REVLIMID (CAP) - EMEA/H/C/000717/II/0079

Applicant: Celgene Europe Limited

PRAC Rapporteur: Corinne Fechant

Scope: Extension of indication to add treatment of adult patients with relapsed and/or refractory mantle cell lymphoma (MCL). As a consequence, SmPC sections 4.1, 4.2, 4.5, 4.8, 5.1 and 5.2 have been updated and the Package Leaflet is updated accordingly. In addition, the MAH took the opportunity to make minor editorial changes in the SmPC and Package Leaflet. A revised version of the RMP (version 25.0) was provided
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.22. Olaparib – LYNPARZA (CAP) - EMEA/H/C/003726/II/0001/G

Applicant: AstraZeneca AB

PRAC Rapporteur: Carmela Macchiarulo

Scope: Update of sections 4.4, 4.5 and 4.6 of the SmPC in order to include further information related to pharmacokinetic interactions based on the *in vivo* interaction study D0816C00008 and 4, 3 *in vitro* interaction studies (ADME-AZS-Wave3-140714, ADME-AZS-

Wave3-140725 and 140483) and data from previously submitted interaction studies. The provision of the final CSR of study D0816C00008 addresses the post-authorisation measure MEA 004. Further, the MAH provided the study report of *in vitro* study 8305083. In addition, the MAH took the opportunity to add the published ATC code in section 5.1 of the SmPC, and to implement minor editorial changes in the SmPC, labelling and Package Leaflet. A revised RMP version 6 was provided, which includes consequential changes related to data on interactions. Further, the MAH is taking the opportunity to update the due dates for the provision of the final study reports of the category 3 studies D0816C00005 and D0816C00006, and to add the new category 3 study D0816C00010
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.23. Pneumococcal polysaccharide conjugate vaccine (adsorbed) – SYNFLORIX (CAP) - EMEA/H/C/000973/II/0096/G

Applicant: GlaxoSmithKline Biologicals

PRAC Rapporteur: Qun-Ying Yue

Scope: Grouped application including one type II variation and two type IB variations to update section 5.1 of the SmPC with effectiveness data against pneumococcal vaccine serotypes and against vaccine related serotype 19A, and update of section 4.4 of the SmPC to include information on the immune response against serotype 19A observed in infants and children. In addition, the MAH took the opportunity to make minor editorial changes in the SmPC, and to request extensions to the due dates for MEA 009: study 10PN-PD-DIT-034 (111634) and MEA 018.5: study 10PN-PD-DIT-064 (114056). A revised RMP version 12 was provided
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.24. Ruxolitinib – JAKAVI (CAP) - EMEA/H/C/002464/II/0024

Applicant: Novartis Europharm Ltd

PRAC Rapporteur: Ulla Wändel Liminga

Scope: Update of section 4.4 of the SmPC in order to add a warning on reported cases of Merkell cell carcinoma in patients treated with ruxolitinib
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.25. Thiotepa – TEPADINA (CAP) - EMEA/H/C/001046/II/0021

Applicant: Adienne S.r.l. S.U.

PRAC Rapporteur: Corinne Fechant

Scope: Update of section 4.8 of the SmPC to add the new adverse drug reaction (ADR) 'toxic skin reactions' with unknown frequency. The Package Leaflet is updated accordingly. A revised RMP version 12 was provided
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.26. Ticagrelor – BRILIQUE (CAP) - EMEA/H/C/001241X/0029/G

Applicant: AstraZeneca AB

PRAC Rapporteur: Menno van der Elst

Scope: Grouped application to 1) extension application to add a new strength of 60mg with a new indication: history of myocardial infarction; 2) update of the product information of the existing Brilique 90 mg license with important clinical information from the PEGASUS study
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.27. Tobramycin – VANTOBRA (CAP) - EMEA/H/C/002633/II/0001/G

Applicant: Pari Pharma GmbH

PRAC Rapporteur: Qun-Ying Yue
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.28. Vemurafenib – ZELBORAF (CAP) - EMEA/H/C/002409/II/0024/G

Applicant: Roche Registration Ltd

PRAC Rapporteur: Ulla Wändel Liminga

Scope: Update of sections 4.4, 4.5 and 4.8 of the SmPC in order to update information on the risk of potentiation of radiation toxicity and updating the risk of progression of cancers with RAS mutations with information on progression of pre-existing pancreatic adenocarcinoma with KRAS mutation. The Package Leaflet is updated accordingly
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.29. Voriconazole – VFEND (CAP) - EMEA/H/C/000387/II/0110/G

Applicant: Pfizer Limited

PRAC Rapporteur: Sabine Straus

Scope: Update of the SmPC sections 4.4, 4.8 and 5.1 to reflect the safety and efficacy data from studies in paediatric population. The Package Leaflet is updated accordingly. Updated RMP (version 4) is also submitted accordingly
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

6. Periodic safety update reports (PSURs)

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

6.1.1. Amifampridine – FIRDAPSE (CAP) - PSUSA/00141/201412

Applicant: BioMarin Europe Ltd

PRAC Rapporteur: Julie Williams

Scope of procedure: Evaluation of a PSUSA procedure
Action: For adoption of recommendation to CHMP

6.1.2. Avanafil – SPEDRA (CAP) - PSUSA/10066/201412

Applicant: Menarini International Operations Luxembourg S.A.

PRAC Rapporteur: Miguel-Angel Macia

Scope of procedure: Evaluation of a PSUSA procedure
Action: For adoption of recommendation to CHMP

6.1.3. Belatacept – NULOJIX (CAP) - PSUSA/00311/201412

Applicant: Bristol-Myers Squibb Pharma EEIG

PRAC Rapporteur: Ulla Wändel Liminga

Scope of procedure: Evaluation of a PSUSA procedure
Action: For adoption of recommendation to CHMP

6.1.4. Besilesomab – SCINTIMUN (CAP) - PSUSA/00385/201501

Applicant: Cis Bio International

PRAC Rapporteur: Julie Williams

Scope of procedure: Evaluation of a PSUSA procedure
Action: For adoption of recommendation to CHMP

6.1.5. Brimonidine tartrate, brinzolamide – SIMBRINZA (CAP) - PSUSA/10273/201412

Applicant: Alcon Laboratories (UK) Ltd

PRAC Rapporteur: Almath Spooner

Scope of procedure: Evaluation of a PSUSA procedure
Action: For adoption of recommendation to CHMP

6.1.6. Canakinumab – ILARIS (CAP) - PSUSA/00526/201412

Applicant: Novartis Europharm Ltd

PRAC Rapporteur: Brigitte Keller-Stanislawski

Scope of procedure: Evaluation of a PSUSA procedure
Action: For adoption of recommendation to CHMP

6.1.7. Caspofungin – CANCIDAS (CAP) - PSUSA/00576/201412

Applicant: Merck Sharp & Dohme Limited

PRAC Rapporteur: Veerle Verlinden

Scope of procedure: Evaluation of a PSUSA procedure
Action: For adoption of recommendation to CHMP

6.1.8. Clofarabine – EVOLTRA (CAP) - PSUSA/00805/201412

Applicant: Genzyme Europe BV

PRAC Rapporteur: Corinne Fechant

Scope of procedure: Evaluation of a PSUSA procedure
Action: For adoption of recommendation to CHMP

6.1.9. Concentrate of proteolytic enzymes enriched in bromelain – NEXOBRID (CAP) - PSUSA/10028/201412

Applicant: MediWound Germany GmbH

PRAC Rapporteur: Valerie Strassmann

Scope of procedure: Evaluation of a PSUSA procedure
Action: For adoption of recommendation to CHMP

6.1.10. Darunavir – PREZISTA (CAP) - PSUSA/00934/201412

Applicant: Janssen-Cilag International N.V.

PRAC Rapporteur: Menno van der Elst

Scope of procedure: Evaluation of a PSUSA procedure
Action: For adoption of recommendation to CHMP

6.1.11. Dextromethorphan hydrobromide, quinidine sulfate – NUEDEXTA (CAP) - PSUSA/10089/201412

Applicant: Jenson Pharmaceutical Services Ltd

PRAC Rapporteur: Julie Williams

Scope of procedure: Evaluation of a PSUSA procedure
Action: For adoption of recommendation to CHMP

6.1.12. Eptacog alfa (activated) – NOVOSEVEN (CAP) - PSUSA/01245/201412

Applicant: Novo Nordisk A/S

PRAC Rapporteur: Sabine Straus

Scope of procedure: Evaluation of a PSUSA procedure
Action: For adoption of recommendation to CHMP

6.1.13. Ferumoxytol – RIENSO - PSUSA/01386/201412

Applicant: Takeda Pharma A/S

PRAC Rapporteur: Martin Huber

Scope of procedure: Evaluation of a PSUSA procedure (EC decision on MA withdrawal dated 13 April 2015)
Action: For information

6.1.14. Fingolimod – GILENYA (CAP) - PSUSA/01393/201502

Applicant: Novartis Europharm Ltd

PRAC Rapporteur: Isabelle Robine

Scope of procedure: Evaluation of a PSUSA procedure
Action: For discussion (Report from the SAG Neurology dated 11 June 2015)

6.1.15. Human fibrinogen, human thrombin – EVICEL (CAP); TACHOSIL (CAP) - PSUSA/01627/201412

Applicant: Omrix Biopharmaceuticals N. V.

PRAC Rapporteur: Brigitte Keller-Stanislawski

Scope of procedure: Evaluation of a PSUSA procedure
Action: For adoption of recommendation to CHMP

6.1.16. Influenza vaccine (live attenuated, nasal) – FLUENZ TETRA (CAP) - PSUSA/01742/201412

Applicant: MedImmune LLC

PRAC Rapporteur: Jean-Michel Dogné

Scope of procedure: Evaluation of a PSUSA procedure
Action: For adoption of recommendation to CHMP

6.1.17. Lenalidomide – REVLIMID (CAP) - PSUSA/01838/201412

Applicant: Celgene Europe Limited

PRAC Rapporteur: Corinne Fechant

Scope of procedure: Evaluation of a PSUSA procedure
Action: For adoption of recommendation to CHMP

6.1.18. Matrix applied characterised autologous cultured chondrocytes – MACI (CAP) - PSUSA/10116/201412

Applicant: Aastrom Biosciences DK ApS

PRAC Rapporteur: Rafe Suvarna

Scope of procedure: Evaluation of a PSUSA procedure
Action: For adoption of recommendation to CHMP

6.1.19. Mirabegron – BETMIGA (CAP) - PSUSA/10031/201412

Applicant: Astellas Pharma Europe B.V.

PRAC Rapporteur: Miguel-Angel Macia

Scope of procedure: Evaluation of a PSUSA procedure
Action: For adoption of recommendation to CHMP

6.1.20. Omalizumab – XOLAIR (CAP) - PSUSA/02214/201412

Applicant: Novartis Europharm Ltd

PRAC Rapporteur: Qun-Ying Yue

Scope of procedure: Evaluation of a PSUSA procedure
Action: For adoption of recommendation to CHMP

6.1.21. Paclitaxel albumin – ABRAXANE (CAP) - PSUSA/10123/201501

Applicant: Celgene Europe Limited

PRAC Rapporteur: Sabine Straus

Scope of procedure: Evaluation of a PSUSA procedure
Action: For adoption of recommendation to CHMP

6.1.22. Pertuzumab – PERJETA (CAP) - PSUSA/10125/201412

Applicant: Roche Registration Ltd

PRAC Rapporteur: Doris Stenver

Scope of procedure: Evaluation of a PSUSA procedure
Action: For adoption of recommendation to CHMP

6.1.23. Ponatinib – ICLUSIG (CAP) - PSUSA/10128/201412

Applicant: Ariad Pharma Ltd

PRAC Rapporteur: Rafe Suvarna

Scope of procedure: Evaluation of a PSUSA procedure
Action: For adoption of recommendation to CHMP

6.1.24. Reteplase – RAPILYSIN (CAP) - PSUSA/02623/201411

Applicant: Actavis Group PTC ehf

PRAC Rapporteur: Martin Huber

Scope of procedure: Evaluation of a PSUSA procedure
Action: For adoption of recommendation to CHMP

6.1.25. Roflumilast – DALIRESP (CAP); DAXAS (CAP); LIBERTEK (CAP) - PSUSA/02658/201501

Applicant: Takeda GmbH

PRAC Rapporteur: Miguel-Angel Macia

Scope of procedure: Evaluation of a PSUSA procedure
Action: For adoption of recommendation to CHMP

6.1.26. Ticagrelor – BRILIQUE (CAP) - PSUSA/02948/201412

Applicant: AstraZeneca AB

PRAC Rapporteur: Menno van der Elst

Scope of procedure: Evaluation of a PSUSA procedure
Action: For adoption of recommendation to CHMP

6.1.27. Trametinib – MEKINIST (CAP) - PSUSA/10262/201412

Applicant: Novartis Europharm Ltd

PRAC Rapporteur: Julie Williams

Scope of procedure: Evaluation of a PSUSA procedure
Action: For adoption of recommendation to CHMP

6.1.28. Umeclidinium bromide, vilanterol – ANORO (CAP); LAVENTAIR (CAP) - PSUSA/10264/201412

Applicant: Glaxo Group Ltd

PRAC Rapporteur: Carmela Macchiarulo

Scope of procedure: Evaluation of a PSUSA procedure
Action: For adoption of recommendation to CHMP

6.1.29. Ustekinumab – STELARA (CAP) - PSUSA/03085/201412

Applicant: Janssen-Cilag International N.V.

PRAC Rapporteur: Julie Williams

Scope of procedure: Evaluation of a PSUSA procedure
Action: For adoption of recommendation to CHMP

6.1.30. Verteporfin – VISUDYNE (CAP) - PSUSA/03110/201412

Applicant: Novartis Europharm Ltd

PRAC Rapporteur: Isabelle Robine

Scope of procedure: Evaluation of a PSUSA procedure
Action: For adoption of recommendation to CHMP

6.1.31. Ziconotide – PRIALT (CAP) - PSUSA/03142/201412

Applicant: Eisai Ltd

PRAC Rapporteur: Jean-Michel Dogné

Scope of procedure: Evaluation of a PSUSA procedure
Action: For adoption of recommendation to CHMP

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

6.2.1. Human hepatitis b immunoglobulin – ZUTECTRA (CAP), NAP - PSUSA/01631/201411

Applicant: Biotest Pharma GmbH

PRAC Rapporteur: Brigitte Keller-Stanislawski

Scope of procedure: Evaluation of a PSUSA procedure
Action: adoption of recommendation to CHMP

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

6.3.1. Amiodarone (NAP) - PSUSA/00000166/201412

Applicant: various

PRAC Lead: Menno van der Elst

Scope: Evaluation of a PSUSA procedure
Action: For adoption of recommendation to CMDh

6.3.2. Atomoxetine (NAP) - PSUSA/00000262/201411

Applicant: various

PRAC Lead: Julie Williams

Scope: Evaluation of a PSUSA procedure
Action: For adoption of recommendation to CMDh

6.3.3. Caffeine, drotaverine hydrochloride, metamizol sodium (NAP) - PSUSA/00001996/201411

Applicant: various

PRAC Lead: Tatiana Magalova

Scope: Evaluation of a PSUSA procedure
Action: For adoption of recommendation to CMDh

6.3.4. Ciprofloxacin hydrochloride, hydrocortisone (NAP) - PSUSA/00000774/201411

Applicant: various

PRAC Lead: Carmela Macchiarulo

Scope: Evaluation of a PSUSA procedure
Action: For adoption of recommendation to CMDh

6.3.5. Dapoxetine (NAP) - PSUSA/00000928/201412

Applicant: various

PRAC Lead: Ulla Wändel Liminga

Scope: Evaluation of a PSUSA procedure
Action: For adoption of recommendation to CMDh

6.3.6. Diacerein (NAP) - PSUSA/00001026/201412

Applicant: various

PRAC Lead: Isabelle Robine

Scope: Evaluation of a PSUSA procedure
Action: For adoption of recommendation to CMDh

6.3.7. Diphteria, tetanus, pertussis (acellular, component), poliomyelitis (inactivated), haemophilius type b conjugate vaccine (adsorbed) (NAP) - PSUSA/00001124/201411

Applicant: various

PRAC Lead: Brigitte Keller-Stanislawski

Scope: Evaluation of a PSUSA procedure
Action: For adoption of recommendation to CMDh

6.3.8. Drospirenone, estradiol (NAP) - PSUSA/00001184/201412

Applicant: various

PRAC Lead: Menno van der Elst

Scope: Evaluation of a PSUSA procedure
Action: For adoption of recommendation to CMDh

6.3.9. Estradiol, gestodene (NAP) - PSUSA/00001273/201412

Applicant: various

PRAC Lead: Doris Stenver

Scope: Evaluation of a PSUSA procedure
Action: For adoption of recommendation to CMDh

6.3.10. Fluorine (^{18}F) fludeoxyglucose (NAP) - PSUSA/00001437/201411

Applicant: various

PRAC Lead: Isabelle Robine

Scope: Evaluation of a PSUSA procedure
Action: For adoption of recommendation to CMDh

6.3.11. Human coagulation factor VIII (antihemophilic factor A) (NAP) - PSUSA/00001620/201411

Applicant: various

PRAC Lead: Brigitte Keller-Stanislawski

Scope: Evaluation of a PSUSA procedure
Action: For adoption of recommendation to CMDh

6.3.12. Hydroxyethyl starch (NAP) - PSUSA/00001694/201501

Applicant: various

PRAC Lead: Martin Huber

Scope: Evaluation of a PSUSA procedure
Action: For adoption of recommendation to CMDh

6.3.13. Lisuride (NAP) - PSUSA/00001896/201411

Applicant: various

PRAC Lead: Jan Neuhauser

Scope: Evaluation of a PSUSA procedure
Action: For adoption of recommendation to CMDh

6.3.14. Methylprednisolone (NAP) - PSUSA/00002026/201411

Applicant: various

PRAC Lead: Jan Neuhauser

Scope: Evaluation of a PSUSA procedure
Action: For adoption of recommendation to CMDh

6.3.15. Rabbit anti-human thymocyte (concentrate for solution for infusion) (NAP) - PSUSA/00010252/201412

Applicant: various

PRAC Lead: Maia Uusküla

Scope: Evaluation of a PSUSA procedure
Action: For adoption of recommendation to CMDh

6.3.16. Rabbit anti-human thymocyte (powder for solution for infusion) (NAP) - PSUSA/00010184/201412

Applicant: various

PRAC Lead: Maia Uusküla

Scope: Evaluation of a PSUSA procedure
Action: For adoption of recommendation to CMDh

6.3.17. Testosterone undecylate (injection) (NAP) - PSUSA/00010161/201411

Applicant: various

PRAC Lead: Maia Uusküla

Scope: Evaluation of a PSUSA procedure
Action: For adoption of recommendation to CMDh

6.4. Follow-up to PSUR procedures

6.4.1. Vemurafenib – ZELBORAF (CAP) - EMEA/H/C/002409/LEG 031

Applicant: Roche Registration Ltd

PRAC Rapporteur: Ulla Wändel Liminga

Scope: MAH's responses to PSUSA/00009329/201408 as adopted in March 2015
Action: For adoption of advice to CHMP

7. Post-authorisation safety studies (PASS)

7.1. Protocols of PASS imposed in the marketing authorisation(s)[2]

7.1.1. Flupirtine (NAP) - EMEA/H/N/PSP/j/0005.4

Applicant: Meda Pharma GmbH & Co KG

PRAC Rapporteur: Valerie Strassmann

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

Scope: Evaluation of a revised protocol for a non-interventional post-authorisation safety study to evaluate the effectiveness of the risk minimisation measures for the use of flupirtine 100 mg immediate-release capsules in daily practice
Action: For adoption of PRAC Assessment Report, PRAC outcome letter

7.1.2. Hydroxyethyl starch (NAP) - EMEA/H/N/PSP/0014.2

Applicant: Fresenius Kabi Deutschland GmbH (Volulyte, Voluven Fresenius, Voluven, HyperHAES, HAES-steril), Serumwerk Bernburg AG (VitaHES, Vitafusal, Plasma Volume Redibag, PlasmaHES Redibag, Hesra, Hesra infuusioneste)

PRAC Rapporteur: Qun-Ying Yue

Scope: Evaluation of a revised PASS protocol (drug utilisation study) to assess the effectiveness of the risk minimisation taken following the European Commission decision dated 19 December 2013 for the referral procedure EMEA/H/A-107I/1376
Action: For adoption of PRAC Assessment Report, PRAC outcome letter

7.1.3. Hydroxyethyl starch (NAP) - EMEA/H/N/PSP/0024

Applicant: B. Braun Melsungen AG

PRAC Rapporteur: Qun-Ying Yue

Scope: Evaluation of a new PASS protocol (drug utilisation study) to assess the effectiveness of the risk minimisation taken following the European Commission decision dated 19 December 2013 for the referral procedure EMEA/H/A-107I/1376
Action: For adoption of PRAC Assessment Report, PRAC outcome letter

7.1.4. Tolvaptan – JINARC (CAP) - EMEA/H/C/PSP/0028

Applicant: Otsuka Pharmaceutical Europe Ltd

PRAC Rapporteur: Julie Williams

Scope: PASS protocol for a prospective study of the safety of tolvaptan in ADPKD patients with an additional retrospective component to assess for risks associated with long term use
Action: For adoption of PRAC Assessment Report, PRAC outcome letter

7.1.5. Valproate (NAP) - EMEA/H/N/PSP/j/0029

Applicant: Sanofi R&D, various

PRAC Rapporteur: To be appointed

Scope: Protocol for a drug utilisation study to assess the effectiveness of the risk minimisation measures and to further characterise the prescribing patterns for valproate
Action: For adoption of procedure timetable

7.2. Protocols of PASS non-imposed in the marketing authorisation(s)[3]

7.2.1. Collagenase clostridium histolyticum – XIAPEX (CAP) - EMEA/H/C/002048/MEA 027

Applicant: Swedish Orphan Biovitrum AB (publ)

PRAC Rapporteur: Martin Huber

[3] 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

Scope: PASS protocol for a non-interventional survey to evaluate the effectiveness of Xiapex educational material for healthcare professionals in the treatment of Peyronie's disease
Action: For adoption of advice to CHMP

7.2.2. Flutemetamol (^{18}F) – VIZAMYL (CAP) - EMEA/H/C/002557/MEA 002.1

Applicant: GE Healthcare Ltd

PRAC Rapporteur: Julie Williams

Scope: MAH's responses to MEA 002 [PASS protocol, study no. GE067-027 CPR to assess the effectiveness of the educational training programme] as adopted in February 2015
Action: For adoption of advice to CHMP

7.2.3. Linaclotide – CONSTELLA (CAP) - EMEA/H/C/002490/MEA 009.1

Applicant: Almirall S.A

PRAC Rapporteur: Valerie Strassmann

Scope: MAH's responses to MEA 009 (linaclotide safety study for the assessment of diarrhoea - complications and associated risk factors in selected European populations with IBS-C) as adopted in March 2015
Action: For adoption of advice to CHMP

7.2.4. Linaclotide – CONSTELLA (CAP) - EMEA/H/C/002490/MEA 011

Applicant: Almirall S.A

PRAC Rapporteur: Valerie Strassmann

Scope: MAH's responses to MEA 010 (drug utilisation study (DUS) protocol - linaclotide utilisation study in selected European populations study) as adopted in March 2015
Action: For adoption of advice to CHMP

7.2.5. Mixture of polynuclear iron(III)-oxyhydroxide, sucrose and starches – VELPHORO (CAP) - EMEA/H/C/002705/MEA 002.1

Applicant: Vifor Fresenius Medical Care Renal Pharma France

PRAC Rapporteur: Julie Williams

Scope: MAH's responses to MEA 002 [PASS protocol for study VFMCRP-MEAF-PA21-01-EU] as adopted in February 2015
Action: For adoption of advice to CHMP

7.2.6. Pandemic influenza vaccine (H5N1) (surface antigen, inactivated, adjuvanted) – AFLUNOV (CAP) - EMEA/H/C/002094/MEA 022; prepandemic influenza vaccine (H5N1) (surface antigen, inactivated, adjuvanted) - FOCLIVIA (CAP) - EMEA/H/C/001208/MEA 026 , PREPANDEMIC INFLUENZA VACCINE (H5N1) (SURFACE ANTIGEN, INACTIVATED, ADJUVANTED) NOVARTIS VACCINES AND DIAGNOSTIC (CAP) - EMEA/H/C/002269/MEA 021

Applicant: Novartis Vaccines Influenza Srl
PRAC Rapporteur: Carmela Macchiarulo
EMA resources: RMS: Emil Cochino; EPL: Manuela Mura

Scope: MAH's responses to MEA 020 /MEA 019 (PASS protocol synopsis V87_27 OB) request for information (RSI) following the PRAC outcome in February 2014
Action: For adoption of advice to CHMP

7.3. Results of PASS imposed in the marketing authorisation(s)[4]

None

7.4. Results of PASS non-imposed in the marketing authorisation(s)[5]

7.4.1. Aflibercept – EYLEA (CAP) - EMEA/H/C/002392/II/0023 (with RMP)

Applicant: Bayer Pharma AG

PRAC Rapporteur: Isabelle Robine

Scope: Submission of final study report for VIEW-1 extension VGFT-OD-0910 to assess the long-term safety and tolerability of VEGF Trap-Eye in patients with neovascular age-related macular degeneration (AMD), in order to fulfil MEA 003 (category 3 study included in the RMP). The RMP has been updated (version 21.0) to reflect the completion of this study
Action: For adoption of PRAC Assessment Report

7.4.2. Etanercept – ENBREL (CAP) - EMEA/H/C/000262/II/0179 (without RMP)

Applicant: Pfizer Limited

PRAC Rapporteur: Rafe Suvarna

Scope: Submission of final report from the Organisation of Teratology Information Specialists (OTIS) registry, as listed in part III of the RMP
Action: For adoption of PRAC Assessment Report

7.4.3. Fentanyl – EFFENTORA (CAP) - EMEA/H/000833/II/0037 (with RMP)

Applicant: Teva B.V.

PRAC Rapporteur: Martin Huber

Scope: Submission of the final study report of a PASS: national descriptive and longitudinal study of patients treated with Effentora in France. The RMP has been updated accordingly
Action: For adoption of PRAC Assessment Report

7.4.4. Human rotavirus, live attenuated – ROTARIX (CAP) - EMEA/H/C/000639/II/0062 (with RMP)

Applicant: GlaxoSmithKline Biologicals S.A.

PRAC Rapporteur: Jean-Michel Dogné

Scope: Submission of the final report of genetic stability study EPI-ROTA-014 VS BE – 112560 that addresses the post-approval measure MEA 005.2 in which the MAH commits to monitor for the potential occurrence of genetic drifts and shifts in the vaccine strain in post-marketing settings
Action: For adoption of PRAC Assessment Report

[4] In accordance with Article 107p-q of Directive 2001/83/EC
[5] 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

7.4.5. Indacaterol – HIROBRIZ BREEZHALER (CAP) - EMEA/H/C/001211/WS0777/0036/G ; ONBREZ BREEZHALER (CAP) - EMEA/H/C/001114/WS0777/0035/G ; OSLIF BREEZHALER (CAP) - EMEA/H/C/001210/WS0777/0035/G (with RMPs)

Applicant: Novartis Europharm Ltd

PRAC Rapporteur: Torbjorn Callreus

Scope: Submission of the final study reports of two PASS studies (UK PASS study QAB149B2433/QAB149BS641859 and EU PASS study QAB149B2431/QAB149AS232863) included in the RMP to monitor off-label use of indacaterol; an updated RMP (version 8.0) has been submitted accordingly. Moreover, the MAH updated the RMP to include changes as agreed by the CHMP in March 2015 (MEA 017/MEA 015/MEA 015 following the review of the fourth US PASS (QAB149B2432/CQAB149BS232861) interim report)
Action: For adoption of PRAC Assessment Report

7.4.6. Pneumococcal polysaccharide conjugate vaccine (13-valent, adsorbed) – PREVENAR 13 (CAP) - EMEA/H/C/001104/II/0123 (without RMP)

Applicant: Pfizer Limited

PRAC Rapporteur: Qun-Ying Yue

Scope: Submission of final clinical study report (CSR) for study 6096A1-4024 (B1851040), a PASS to assess the impact of 13vPnC on otitis media in children after the introduction of the vaccine in 2010 in the United States
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[6]

7.5.1. Adalimumab – HUMIRA (CAP) - EMEA/H/C/000481/MEA 080.3

Applicant: AbbVie Ltd

PRAC Rapporteur: Ulla Wändel Liminga

Scope: First progress report on the P11-292 registry (long-term non-interventional registry to assess safety and effectiveness of Humira (adalimumab) in paediatric patients with moderately to severely active Crohn's disease (CD)
Action: For adoption of advice to CHMP

7.5.2. Betaine anhydrous – CYSTADANE (CAP) - EMEA/H/C/000678/MEA 022

Applicant: Orphan Europe S.A.R.L.

PRAC Rapporteur: Valerie Strassmann

Scope: Submission of the second progress report on Orphan Europe Cystadane surveillance protocol in collaboration with the network and registry for homocystinurias and methylation defects (E-HOD)
Action: For adoption of advice to CHMP

7.5.3. Elosulfase alfa – VIMIZIM (CAP) - EMEA/H/C/002779/MEA 005

Applicant: BioMarin Europe Ltd

[6] In line with the revised variations regulation for any submission before 4 August 2013

PRAC Rapporteur: Julie Williams

Scope: First annual report for the Morquio registry study (MARS)
Action: For adoption of advice to CHMP

7.5.4. Mannitol – BRONCHITOL (CAP) - EMEA/H/C/001252/ANX 002.5

Applicant: Pharmaxis Pharmaceuticals Limited

PRAC Rapporteur: Julie Williams

Scope: MAH's responses to MEA 002.4 (fourth interim analysis of the cystic fibrosis (CF) study) request for supplementary information, as adopted in February 2015
Action: For adoption of advice to CHMP

7.5.5. Prucalopride – RESOLOR (CAP) - EMEA/H/C/001012/MEA 006.10

Applicant: Shire Pharmaceuticals Ireland Ltd

PRAC Rapporteur: Rafe Suvarna

Scope: Fourth annual interim safety report for the drug utilisation study to examine the characteristics of patients prescribed prucalopride (Resolor) and a pharmacoepidemiological study of the occurrence of major cardiovascular events, pregnancy, and pregnancy outcomes in the UK Clinical Practice Research Datalink (CPRD) database
Action: For adoption of advice to CHMP

7.5.6. Raltegravir – ISENTRESS (CAP) - EMEA/H/C/000860/MEA 048.5

Applicant: Merck Sharp & Dohme Limited

PRAC Rapporteur: Julie Williams

Scope: MAH's responses to MEA 048.4 [fourth and final annual report for a post-authorisation safety study in a US managed care network] request for supplementary information (RSI) as adopted in February 2015
Action: For adoption of advice to CHMP

7.5.7. Saxagliptin, metformin – KOMBOGLYZE (CAP) - EMEA/H/C/002059/MEA 010.1 saxagliptin – ONGLYZA (CAP) - EMEA/H/C/001039/MEA 033.1

Applicant: AstraZeneca AB

PRAC Rapporteur: Menno van der Elst

Scope: Second interim analysis of PASS study CV181-099ST (comparison of risk of major cardiovascular events between patients with type 2 diabetes initiating saxagliptin and those initiating other oral anti-diabetic treatments)
Action: For adoption of advice to CHMP

7.5.8. Saxagliptin, metformin – KOMBOGLYZE (CAP) - EMEA/H/C/002059/MEA 011.1 saxagliptin – ONGLYZA (CAP) - EMEA/H/C/001039/MEA 034.1

Applicant: AstraZeneca AB

PRAC Rapporteur: Menno van der Elst

Scope: Second interim analysis of PASS study CV181-100ST (comparison of risk of hospitalisation with acute liver failure between patients with type 2 diabetes initiating saxagliptin and those initiating other oral anti-diabetic treatments)
Action: For adoption of advice to CHMP

7.5.9. Saxagliptin, metformin – KOMBOGLYZE (CAP) - EMEA/H/C/002059/MEA 012.1 saxagliptin – ONGLYZA (CAP) - EMEA/H/C/001039/MEA 036.1

Applicant: AstraZeneca AB

PRAC Rapporteur: Menno van der Elst

Scope: Second interim analysis of PASS study CV181-101ST (comparison of risk of hospitalisation with infection between patients with type 2 diabetes initiating saxagliptin and those initiating other oral anti-diabetic treatments)
Action: For adoption of advice to CHMP

7.5.10. Saxagliptin, metformin – KOMBOGLYZE (CAP) - EMEA/H/C/002059/MEA 013.1 saxagliptin – ONGLYZA (CAP) - EMEA/H/C/001039/MEA 037.1

Applicant: AstraZeneca AB

PRAC Rapporteur: Menno van der Elst

Scope: Second interim analysis of PASS study CV181-157ST (comparison of risk of hospitalisation for acute kidney injury between patients with type 2 diabetes initiating saxagliptin and those initiating other oral anti-diabetic treatments)
Action: For adoption of advice to CHMP

7.5.11. Saxagliptin, metformin – KOMBOGLYZE (CAP) - EMEA/H/C/002059/MEA 014.1 saxagliptin – ONGLYZA (CAP) - EMEA/H/C/001039/MEA 035.1

Applicant: AstraZeneca AB

PRAC Rapporteur: Menno van der Elst

Scope: Second interim analysis of PASS study CV181-103ST (comparison of risk of hospitalisation for severe hypersensitivity (including severe cutaneous reactions) between patients with type 2 diabetes initiating saxagliptin and those initiating other oral anti-diabetic treatments)
Action: For adoption of advice to CHMP

7.5.12. Ulipristal – ESMYA (CAP) - EMEA/H/C/002401/MEA 003.3

Applicant: Gedeon Richter Plc.

PRAC Rapporteur: Ulla Wändel Liminga

Scope: Third annual progress report on a non interventional study in pre-operative treatment of moderate to severe symptoms of uterine fibroids (PGL10-014)
Action: For adoption of advice to CHMP

7.5.13. Ustekinumab – STELARA (CAP) - EMEA/H/C/000958/MEA 023.6

Applicant: Janssen-Cilag International N.V.

PRAC Rapporteur: Julie Williams

Scope: Fifth annual interim safety registry report on the Nordic database initiative (protocol CNTO1275PSO4005)
Action: For adoption of advice to CHMP

7.5.14. Ustekinumab – STELARA (CAP) - EMEA/H/C/000958/MEA 024.7

Applicant: Janssen-Cilag International N.V.

PRAC Rapporteur: Julie Williams

Scope: Fifth annual interim report on the pregnancy research initiative CO743T (protocol CNTO1275PSO4007)
Action: For adoption of advice to CHMP

7.5.15. Voriconazole – VFEND (CAP) - EMEA/H/C/000387/MEA 071.11

Applicant: Pfizer Limited

PRAC Rapporteur: Sabine Straus

Scope: Submission of annual status report for PASS study A1501097 (third study progress report) to assess the potential association between voriconazole use and the development of SCC of the skin in patients with lung or heart/lung transplant (LT)
Action: For adoption of advice to CHMP

7.6. Others

7.6.1. Brinavess – VERNAKALANT (CAP) - EMEA/H/C/001215/LEG 025

Applicant: Cardiome UK Limited

PRAC Rapporteur: Menno van der Elst

Scope: From II/09: A requirement to promptly inform the CHMP of any future serious cases of hypotension, with or without fatal outcome. Such case reports will be accompanied by a causality assessment. With this LEG the MAH provides the details including the causality assessment of a hypotension case related to Brinavess
Action: For adoption of advice to CHMP

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

8.1. Annual reassessments of the marketing authorisation

8.1.1. Anagrelide – XAGRID (CAP) - EMEA/H/C/000480/S/0064 (without RMP)

Applicant: Shire Pharmaceutical Contracts Ltd

PRAC Rapporteur: Corinne Fechant

Scope: Annual reassessment of the marketing authorisation
Action: For adoption of advice to CHMP

8.2. Conditional renewals of the marketing authorisation

None

8.3. Renewals of the marketing authorisation

None

9. Product related pharmacovigilance inspections

9.1. List of planned pharmacovigilance inspections

None

9.2. List of planned 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

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

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

11.1. Safety related variations of the marketing authorisation

11.1.1. Azithromycin (NAP) - FI/H/XXXX/WS/23

Applicant: Pfizer (Zithromax)

PRAC Rapporteur: Kimmo Jaakkola

Scope: PRAC consultation on a variation procedure evaluating the draft PASS protocol (A0661209) for a non-imposed non-interventional study in the Kaiser Permanente databases to examine the effects of azithromycin use on cardiovascular outcome
Action: For adoption of advice to Member States

11.1.2. Valsartan, hydrochlorothiazide (NAP) - IS/H/0126/001-003/II/021

Applicant: Egis Pharmaceuticals PLC

PRAC Lead: Hrefna Guðmundsdóttir

Scope: PRAC consultation on a variation procedure pertaining to the hydrochlorothiazide component and concomitant use with allopurinol
Action: For adoption of advice to Member States

11.2. Other requests

11.2.1. Clarithromycin (NAP) - IE/H/PSUR/0020/003

Applicant: Abbott (Klacid), various

PRAC Lead: Almath Spooner

Scope: PRAC consultation on a PSUR worksharing procedure regarding the cardiovascular safety of clarithromycin
Action: For adoption of advice to Member States

11.2.2. Diltiazem (NAP); verapamil (NAP)

Applicant: various

PRAC Lead: Menno van der Elst

Scope: Following the completion of the Article 20 for ivarbadine, CMDh's consultation on the consequences on the diltiazem or verapamil product information
Action: For adoption of advice to Member States

11.2.3. Gadolinium-containing contrast agents (GdCA): gadoversetamide – OPTIMARK (CAP) Gadobenate dimeglumine; gadobutrol; gadodiamide; gadopentetic acid dimeglumine, gadoteric acid (intra-articular formulation); gadoteric acid (intravenous and intravascular formulations); gadoteridol; gadoxetic acid disodium (NAP)

Applicant: various

Lead member: Rafe Suvarna

Scope: PRAC consultation on a post-authorisation measure resulting from the 2010 Article 20 and Article 31 referral procedures for gadolinium-containing contrast agents
Action: For adoption of advice to Member States

12. Organisational, regulatory and methodological matters

12.1. Mandate and organisation of the PRAC

12.1.1. Mandate of Chair and Vice-Chair - prolongation

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

12.3.1. Post-authorisation efficacy study (PAES) - draft scientific guidance

Action: For discussion

12.4. Cooperation within the EU regulatory network

12.4.1. EuroMediCAT: safety of medication use in pregnancy (7th Framework project) - conclusion

Action: For discussion

12.5. Cooperation with international regulators

None

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

None

12.7. PRAC work plan

12.7.1. PRAC work plan 2015 - update

Action: For discussion

12.7.2. PRAC work plan 2016 - development

Action: For discussion

12.8. Planning and reporting

12.8.1. PRAC statistics - overview

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. Granularity and Periodicity Advisory Group (GPAG)

Action: For discussion

12.10.2. Periodic safety update reports - network capacity and planning

Action: For discussion

12.10.3. Periodic safety update reports – proposal for revised criteria for plenary discussion

Action: For adoption

12.10.4. PSURs repository – conditions for identifying pre-conditions of the network from the pilot phase to switch-on

Action: For adoption

12.10.5. 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

Action: For discussion

12.12. Adverse drug reactions reporting and additional reporting

12.12.1. Additional monitoring

None

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

Action: For adoption

12.12.3. Management and reporting of adverse reactions to medicinal products – guidance on monitoring of off label use

Action: For discussion

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

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

None

12.15.2. Post-authorisation Safety Studies and additional monitoring imposed to originator products: applicability to generic products

Action: For discussion

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. Others

13. Any other business

13.1. Enhanced early dialogue to foster development and facilitate accelerated assessment

Action: For discussion

13.2. European commission report on the performance of pharmacovigilance tasks

Action: For discussion

13.3. Tender for outcome study of combined hormonal contraceptives (CHC) Article 31 referral[7]

Action: For discussion

[7] Article 31 of Directive 2001/83/EC resulting from pharmacovigilance data, EMEA/H/A-31/1356

Explanatory notes

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

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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: http://www.ema.europa.eu/ema/index.jsp?curl=pages/regulation/general/general_content_000150.jsp&mid=WC0b01ac05800240d0

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.

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

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