10 May 2016
EMA/PRAC/325452/2016
Procedure Management and Committees Support Division

Pharmacovigilance Risk Assessment Committee (PRAC)

Draft agenda for the meeting on 10-13 May 2016

Chair: June Raine – Vice-Chair: Almath Spooner

10 May 2016, 13:00 – 19:00, room 3/A

11 May 2016, 08:30 – 19:00, room 3/A

12 May 2016, 08:30 – 19:00, room 3/A

13 May 2016, 08:30 – 16:00, room 3/A

Organisational, regulatory and methodological matters (ORGAM)

26 May 2016, Time 10:00 – 12:00, room 7/B, via Adobe Connect

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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Table of contents

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 on 10-13 May 2016. See May 2016 PRAC minutes (to be published post June 2016 PRAC meeting) on the following EMA webpage: http://www.ema.europa.eu/Committees>PRAC>Agendas, minutes and highlights

1.2. Agenda of the meeting of 10-13 May 2016

Action: For adoption

1.3. Minutes of the previous meeting on 11-14 April 2016

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

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

3.2.1. Idelalisib – ZYDELIG (CAP) - EMEA/H/A-20/1439

Applicant: Gilead Sciences International Ltd

PRAC Rapporteur: Rafe Suvarna; PRAC Co-rapporteur: Ulla Wändel Liminga

Scope: Review of the benefit-risk balance following notification by the European Commission of a referral under Article 20 of Regulation (EC) No 726/2004 based on pharmacovigilance data
Action: For adoption of a list of experts for the Scientific Advisory Group (SAG)

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

None

4. Signals assessment and prioritisation[1]

4.1. New signals detected from EU spontaneous reporting systems

4.1.1. Anakinra – KINERET (CAP); canakinumab – ILARIS (CAP)

Applicant: Swedish Orphan Biovitrum AB (publ) (Kineret), Novartis Europharm Ltd (Ilaris)

PRAC Rapporteur: To be appointed

Scope: Signal of weight increase
Action: For adoption of PRAC recommendation
EPITT 18641 – New signal
Lead Member States: DE, DK

4.1.2. Metronidazole (NAP)

Applicant: various

PRAC Rapporteur: To be appointed

Scope: Signal of severe hepatic and neurologic toxicity in patients with Cockayne syndrome
Action: For adoption of PRAC recommendation
EPITT 18663 – 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

4.1.3. Regorafenib – STIVARGA (CAP)

Applicant: Bayer Pharma AG

PRAC Rapporteur: Sabine Straus

Scope: Signal of angioedema
Action: For adoption of PRAC recommendation
EPITT 18656 – New signal
Lead Member State: NL

4.1.4. Vedolizumab - ENTYVIO (CAP)

Applicant: Takeda Pharma A/S

PRAC Rapporteur: Adam Przybylkowski

Scope: Signal of hepatotoxicity
Action: For adoption of PRAC recommendation
EPITT 18646 – New signal
Lead Member State: PL

4.2. New signals detected from other sources

4.2.1. Dexlansoprazole (NAP); lansoprazole (NAP)

Applicant: various

PRAC Rapporteur: To be appointed

Scope: Signal of unexpected histopathological findings from a juvenile rat toxicity study
Action: For adoption of PRAC recommendation
EPITT 18645 – New signal
Lead Member State: FI

4.2.2. Fluconazole (NAP)

Applicant: various

PRAC Rapporteur: To be appointed

Scope: Signal of spontaneous abortion during pregnancy and stillbirth
Action: For adoption of PRAC recommendation
EPITT 18666 – New signal
Lead Member State: DK

4.2.3. Fluoroquinolones: Ciprofloxacin (NAP); enoxacin (NAP); flumequine (NAP); levofloxacin (NAP); lomefloxacin (NAP); moxifloxacin (NAP); norfloxacin (NAP); ofloxacin (NAP); pefloxacin (NAP); prulifloxacin (NAP); rufloxacin (NAP)

Applicant: Bayer, Sanofi, various

PRAC Rapporteur: To be appointed

Scope: Signal of aortic aneurysm and dissection
Action: For adoption of PRAC recommendation
EPITT 18651 – New signal
Lead Member States: DE, DK, ES, FR, IT, NO, UK

4.3. Signals follow-up and prioritisation

4.3.1. Adalimumab – HUMIRA (CAP) - EMEA/H/C/000481/SDA/090

Applicant: AbbVie Ltd.

PRAC Rapporteur: Ulla Wändel Liminga

Scope: Signal of glomerulonephritis (GN)
Action: For adoption of PRAC recommendation
EPITT 18528 – Follow-up to December 2015

4.3.2. Clozapine (NAP)

Applicant: Novartis, various

PRAC Rapporteur: Julie Williams

Scope: Signal of myocarditis
Action: For adoption of PRAC recommendation
EPITT 18414 – Follow-up to September 2015

4.3.3. Cytarabine – DEPOCYTE (CAP) - EMEA/H/C/000317/SDA/019

Applicant: Pacira Ltd

PRAC Rapporteur: Rafe Suvarna

Scope: Signal of benign intracranial hypertension
Action: For adoption of PRAC recommendation
EPITT 18533 – Follow-up to January 2016

4.3.4. Dapagliflozin – FORXIGA (CAP)- EMEA/H/C/002322/SDA/017, EDISTRIDE (CAP); dapagliflozin, metformin - XIGDUO (CAP) - EMEA/H/C/002672/SDA/003, EBYMECT (CAP)

Applicant: AstraZeneca AB

PRAC Rapporteur: Qun-Ying Yue

Scope: Signal of pancreatitis
Action: For adoption of PRAC recommendation
EPITT 18558 – Follow-up to January 2016

4.3.5. Fluoroquinolones: Ciprofloxacin (NAP); enoxacin (NAP); flumequine (NAP); levofloxacin (NAP); lomefloxacin (NAP); moxifloxacin (NAP); norfloxacin (NAP); ofloxacin (NAP); pefloxacin (NAP); prulifloxacin (NAP); rufloxacin (NAP)

Applicant: Bayer, Sanofi, various

PRAC Rapporteur: Valerie Strassmann

Scope: Signal of retinal detachment
Action: For adoption of PRAC recommendation
EPITT 15914 – Follow-up to October 2015

4.3.6. Gefitinib – IRESSA (CAP) - EMEA/H/C/001016/SDA/020

Applicant: AstraZeneca AB

PRAC Rapporteur: Ulla Wändel Liminga

Scope: Signal of pneumatosis intestinalis
Action: For adoption of PRAC recommendation
EPITT 18575 – Follow-up to January 2016

4.3.7. Infliximab – INFLECTRA (CAP), REMICADE (CAP) - EMEA/H/C/000240/SDA/154, REMSIMA (CAP)

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

PRAC Rapporteur: Ulla Wändel Liminga

Scope: Signal of thyroid gland disorders
Action: For adoption of PRAC recommendation
EPITT 18530 – Follow-up to December 2015

4.3.8. Levetiracetam (oral solution) – KEPPRA (CAP) – EMEA/H/C/000277/SDA/082, NAP

Applicant: UCB Pharma SA, various

PRAC Rapporteur: Veerle Verlinden

Scope: Signal of medication errors associated with accidental overdoses
Action: For adoption of PRAC recommendation
EPITT 10519 – Follow-up January 2016

4.3.9. Methotrexate (NAP)

Applicant: various

PRAC Rapporteur: Doris Stenver

Scope: Signal of congenital cardiovascular anomaly
Action: For adoption of PRAC recommendation
EPITT 18481 – Follow-up to November 2015

4.3.10. Natalizumab – TYSABRI (CAP) – EMEA/H/C/000603/SDA/063

Applicant: Biogen Idec Ltd

PRAC Rapporteur: Brigitte Keller-Stanislawski

Scope: Signal of necrotising retinitis
Action: For adoption of PRAC recommendation
EPITT 18605 – Follow-up to January 2016

4.3.11. Quinine (NAP)

Applicant: various

PRAC Rapporteur: Almath Spooner

Scope: Signal of an increased mortality risk in heart failure patients with/without concomitant use of beta-blockers

Action: For adoption of PRAC recommendation
EPITT 18529 – Follow-up to January 2016

4.3.12. Selective serotonin reuptake inhibitors (SSRIs): citalopram (NAP); escitalopram (NAP); fluoxetine (NAP); fluvoxamine (NAP); mirtazapine (NAP); paroxetine (NAP); sertraline (NAP)
Serotonin–noradrenaline reuptake inhibitors (SNRIs): duloxetine - ARICLAIM (CAP), DULOXETINE LILLY (CAP), DULOXETINE MYLAN (CAP), DULOXETINE ZENTIVA (CAP), CYMBALTA (CAP), XERISTAR (CAP), YENTREVE (CAP); sibutramine (NAP); venlafaxine (NAP)

Applicant: Eli Lilly Nederland B.V. (Ariclaim, Duloxetine Lilly, Xeristar, Yentreve), Generics UK Limited (Duloxetine Mylan), Zentiva (Duloxetine Zentiva), various

PRAC Rapporteur: Isabelle Robine

Scope: Signal of risk of autistic spectrum disorders (ASD) after maternal use of SSRI
Action: For adoption of PRAC recommendation
EPITT 14082 – Follow-up to November 2015

4.3.13. Warfarin (NAP)

Applicant: various

PRAC Rapporteur: Torbjorn Callreus

Scope: Signal of calciphylaxis
Action: For adoption of PRAC recommendation
EPITT 18545 – Follow-up to January 2016

5. Risk management plans (RMPs)

5.1. Medicines in the pre-authorisation phase

5.1.1. Allogeneic T cells genetically modified to express suicide gene - EMEA/H/C/002801, Orphan

Applicant: MolMed SpA, ATMP[2]

Scope: Treatment in haploidentical haematopoietic stem cell transplantation
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.1.2. Amikacin - EMEA/H/C/003936, Orphan

Applicant: Insmed Limited

Scope: Treatment of *Pseudomonas aeruginosa* lung infection and colonisation in cystic fibrosis patients
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.1.3. Cabozantinib - EMEA/H/C/004163

Scope (accelerated assessment): Treatment of advanced renal cell carcinoma (RCC)
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

[2] Advanced-therapy medicinal product

5.1.4. Darunavir - EMEA/H/C/004068

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

5.1.5. Dinutuximab beta - EMEA/H/C/003918, Orphan

Applicant: Apeiron Biologics AG

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

5.1.6. Eluxadoline - EMEA/H/C/004098

Scope: Treatment of irritable bowel syndrome with diarrhoea
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.1.7. Emtricitabine, tenofovir disoproxil - EMEA/H/C/004050

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

5.1.8. Lenvatinib - EMEA/H/C/004224

Scope: Treatment of unresectable advanced or metastatic renal cell carcinoma (RCC) in combination with everolimus
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.1.9. Tenofovir disoproxil - EMEA/H/C/004049

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

5.1.10. Tenofovir disoproxil - EMEA/H/C/004120

Scope: Treatment of human immunodeficiency virus (HIV)-1 infection and hepatitis B infection
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. Albiglutide – EPERZAN (CAP) - EMEA/H/C/002735/II/0023/G

Applicant: GlaxoSmithKline Trading Services

PRAC Rapporteur: Julie Williams

Scope: Revised RMP (version 5) in order to add a new phase IV study to evaluate the effect of albiglutide on cholecystokinin-induced gallbladder emptying in fasting healthy subjects as an additional pharmacovigilance activity, to add 'medication error' as an important potential risk, to add 'serious hypersensitivity reaction' as important identified risk and to update the description and due dates for seven studies in the RMP
Action: For adoption of PRAC AR

5.2.2. Aliskiren – RASILEZ (CAP) - EMEA/H/C/000780/WS/0771 aliskiren, hydrochlorothiazide – RASILEZ HCT (CAP) - EMEA/H/C/000964/WS/0771

Applicant: Novartis Europharm Ltd

PRAC Rapporteur: Carmela Macchiarulo

Scope: Revised RMP with regard to identified risks, missing information, concomitant use of other medicines, drug-drug interactions, removal of safety issues attributed to the withdrawn aliskiren/amlodipine (Rasilamlo) and aliskiren/amlodipine/HCTZ (Rasitrio). The variation is supported by study report SPA100A: antihypertensive effects and long-term safety of aliskiren in elderly patients
Action: For adoption of PRAC AR

5.2.3. Characterised viable autologous cartilage cells expanded ex vivo expressing specific marker proteins – CHONDROCELECT (CAP) - EMEA/H/C/000878/II/0018/G

Applicant: TiGenix NV

PRAC Rapporteur: Brigitte Keller-Stanislawski

Scope: Revised RMP (version 10) in order to add information resulting from the assessment of MEA 016 and MEA 018 in relation to the confirmatory randomized controlled trial in small lesions. Two new important potential risks 'transmission of infective agents' and 'allergic/hypersensitivity reaction' (from PSUSA/00000273/201504) are also added together with other updated information in the RMP
Action: For adoption of PRAC AR

5.2.4. Efavirenz, emtricitabine, tenofovir disoproxil – ATRIPLA (CAP) - EMEA/H/C/000797/WS/0860/G emtricitabine – EMTRIVA (CAP) - EMEA/H/C/000533/WS/0860/G emtricitabine, rilpivirine, tenofovir disoproxil – EVIPLERA (CAP) - EMEA/H/C/002312/WS/0860/G

Applicant: Bristol-Myers Squibb and Gilead Sciences Ltd. (Atripla), Gilead Sciences International Ltd (Emtriva, Eviplera)

PRAC Rapporteur: Rafe Suvarna

Scope: Revised RMP following the PRAC review on the 'comprehensive analysis of existing data on lipodystrophy (updated literature data on non-clinical and clinical aspects)' and 'comprehensive analysis of existing data on lactic acidosis (updated literature data on non-clinical and clinical aspects)'
Action: For adoption of PRAC AR

5.2.5. Posaconazole – NOXAFIL (CAP) - EMEA/H/C/000610/II/0040

Applicant: Merck Sharp & Dohme Limited

PRAC Rapporteur: Rafe Suvarna

Scope: Revised RMP (version 12.0) in order to reflect the study results showing a lack of interaction effect of OATP1B1 and OATP1B3 substrates and inhibitors
Action: For adoption of PRAC AR

5.2.6. Sonidegib – ODOMZO (CAP) - EMEA/H/C/002839/II/0004/G

Applicant: Novartis Europharm Ltd

PRAC Rapporteur: Julie Williams

Scope: Submission of the final clinical study reports for exploratory studies X2114 and X2203, whereby the MAH committed to collect cardiac events, second primary malignancies and fractures. In addition, the MAH updated the RMP (version 3.2) to reflect the completion of studies X2114 and X2203 and changes to the due dates for provision of the final study reports for the category 3 studies LDE225C2301 and LDE225X2104
Action: For adoption of PRAC AR

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

5.3.1. Abiraterone – ZYTIGA (CAP) - EMEA/H/C/002321/X/0039

Applicant: Janssen-Cilag International N.V.

PRAC Rapporteur: Dolores Montero Corominas

Scope: Extension application to introduce a new pharmaceutical form associated with a new strength (500 mg film-coated tablets)
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/0027/G

Applicant: Bayer Pharma AG

PRAC Rapporteur: Isabelle Robine

Scope: Grouped variations to include: 1) 3-year data of the pivotal trials VIVID-DME and VISTA-DME; 2) protocol T data with a consequential update to section 5.1 of the SmPC . Furthermore, the MAH took the opportunity to condense the SmPC section 4.8 text relating to antiplatelet trialists' collaboration (APTC) as recommended by EMA during II/0018 variation (diabetic macular oedema (DME) 2 year data), to shorten SmPC section 5.1 as committed by the MAH during II/0021 variation (indication myopic choroidal neovascularisation (mCNV)), to align the annexes with the latest QRD template (version 9.1) and to implement minor changes within age-related macular degeneration (AMD) and DME posology sections
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.3. Ataluren – TRANSLARNA (CAP) - EMEA/H/C/002720/II/0012

Applicant: PTC Therapeutics International Limited

PRAC Rapporteur: Sabine Straus

Scope: Extension of indication to include the treatment of cystic fibrosis resulting from a nonsense mutation in at least one allele of the cystic fibrosis transmembrane conductance regulator (CFTR) gene. Consequently, sections 4.1, 4.2, 4.3, 4.4, 4.5, 4.8, 5.1 and 5.2 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

5.3.4. Ataluren – TRANSLARNA (CAP) - EMEA/H/C/002720/II/0019

Applicant: PTC Therapeutics International Limited

PRAC Rapporteur: Sabine Straus

Scope: Submission of the final results from the non-clinical study PTC124-15055: assessment of uncoupling protein 1 (UCP1) protein levels in brown adipose tissue (BAT) in weanling rats administered ataluren via oral gavage for two weeks, in order to address MEA 007. Part II: module SII of RMP (version 4.4) was updated to reflect in tumor findings that

in-vivo exposure to ataluren and the M4 metabolite does not activate BAT. Other sections of the RMP were updated to reflect completion of the study
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.5. Capecitabine – XELODA (CAP) - EMEA/H/C/000316/II/0070

Applicant: Roche Registration Limited

PRAC Rapporteur: Martin Huber

Scope: Update of sections 4.4 and 4.8 of the SmPC in order to include a warning on fingerprint loss. The Package Leaflet and the RMP are updated accordingly. In addition, the MAH took the opportunity to bring the product information in line with the latest QRD template (version 9.1)
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.6. Carfilzomib – KYPROLIS (CAP) - EMEA/H/C/003790/II/0001/G

Applicant: Amgen Europe B.V.

PRAC Rapporteur: Marina Dimov Di Giusti

Scope: Extension of indication to include the treatment in combination with either lenalidomide and dexamethasone or dexamethasone alone, of adult patients with multiple myeloma who have received at least one prior therapy. 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 accordingly. In addition, the MAH updated section 6.6 of the SmPC to include the option to administer Kyprolis in a 100 mL intravenous bag containing 5% glucose solution for injection in line with the extension of indication part of this variation
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.7. Ceritinib – ZYKADIA (CAP) - EMEA/H/C/003819/II/0006/G

Applicant: Novartis Europharm Ltd

PRAC Rapporteur: Ulla Wändel Liminga

Scope: Update of section 4.5 of the SmPC based on the final results of the clinical pharmacology study CLDK378A2113 and results of a sub-group evaluating the impact of gastric pH-elevating agents on the steady-state pharmacokinetic (PK), efficacy, and safety of ceritinib in anaplastic lymphoma kinase (ALK)-positive non-small cell lung cancer (NSCLC) patients. The final clinical study report for study CLDK378A2113 is submitted to fulfill MEA 003. In addition, the MAH is proposing a change to the due date for the provision of the final study report for study CLDK378A2110 (MEA 001). The RMP is updated (version 3.0) accordingly
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.8. Cobimetinib – COTELLIC (CAP) - EMEA/H/C/003960/II/0004/G

Applicant: Roche Registration Limited

PRAC Rapporteur: Sabine Straus

Scope: Update of section 5.1 of the SmPC in order to update the safety and efficacy results of studies GO28141 and NO25395. The RMP has been updated accordingly. In addition, the MAH took the opportunity to make minor amendments in sections 4.6, 5.1 and 5.3 of the SmPC
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.9. Crizotinib – XALKORI (CAP) - EMEA/H/C/002489/II/0039

Applicant: Pfizer Limited

PRAC Rapporteur: Isabelle Robine

Scope: Extension of indication to include the treatment of adults with ROS1-positive advanced non-small cell lung cancer (NSCLC) based on the results of study A8081001 (a multinational, multicentre, open-label, single-arm study of the safety, pharmacokinetics, pharmacodynamics, and antitumor activity of crizotinib in patients with advanced cancer). Consequential changes are proposed to SmPC sections 4.1, 4.2, 4.4, 4.8, 5.1 and 5.2 and the Package Leaflet and RMP (version 7.0) are updated accordingly
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.10. Dolutegravir – TIVICAY (CAP) - EMEA/H/C/002753/X/0018/G

Applicant: ViiV Healthcare UK Limited

PRAC Rapporteur: Julie Williams

Scope: Extension application to add two new strengths (10 mg and 25 mg tablets) to support the extension of the indication for the treatment of paediatric patients from 6 years of age infected with human immunodeficiency virus (HIV). Data from cohort I and II A of the clinical trial ING112578 are presented in support of the new therapeutic indication
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.11. Eltrombopag, eltrombopag olamine – REVOLADE (CAP) - EMEA/H/C/001110/II/0029/G

Applicant: Novartis Europharm Ltd

PRAC Rapporteur: Dolores Montero Corominas

Scope: Update of sections 4.4 and 4.8 of the SmPC with reference to bone marrow reticulin formation and risk of bone marrow fibrosis and section 5.1 of the SmPC with updated exposure data, based on the final study reports for study TRA112940 (a longitudinal 2-year bone marrow study of eltrombopag olamine (SB-497115-GR) in previously treated adults, with chronic immune (idiopathic) thrombocytopenic purpura (ITP)) and study TRA105325 (EXTEND (Eltrombopag eXTENded Dosing study) an extension study of eltrombopag olamine (SB-497115-GR) in adults with chronic immune (idiopathic) thrombocytopenic purpura (ITP) previously enrolled in an eltrombopag study). As a consequence, Annex II is updated in order to delete 'increased bone marrow reticulin fibres' from the key elements to be included in the educational material. In addition, the MAH took the opportunity to propose an update of the due date in the RMP for the provision of the final clinical study report (CSR) for MEA 022.1 (effectiveness of educational materials for hepatitis C associated thrombocytopenia). The RMP (version 36) is updated accordingly
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.12. Empagliflozin, metformin – SYNJARDY (CAP) - EMEA/H/C/003770/II/0015

Applicant: Boehringer Ingelheim GmbH

PRAC Rapporteur: Miguel-Angel Macia

Scope: Extension of indication to include the treatment with Synjardy as adjunct to standard care therapy in adult patients with type 2 diabetes mellitus and high cardiovascular risk when the treatment with empagliflozin and metformin is appropriate and empagliflozin is needed to reduce the risk of all-cause mortality by reducing cardiovascular death and cardiovascular death or hospitalization for heart failure. As a consequence, sections 4.1, 4.4, 4.8 and 5.1 of the SmPC are updated based on the final clinical study

report of study EMPA-REG OUTCOME. The Package Leaflet and RMP (version 5.0) is updated accordingly
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.13. Erlotinib – TARCEVA (CAP) - EMEA/H/C/000618/II/0045

Applicant: Roche Registration Limited

PRAC Rapporteur: Doris Stenver

Scope: Submission of the clinical study report for study BO25460 (IUNO) 'a randomized, double-blind, placebo controlled, phase III study of first-line maintenance Tarceva *versus* Tarceva at the time of disease progression in patients with advanced non-small cell lung cancer(NSCLC) who have not progressed following 4 cycles of platinum-based chemotherapy' requested as part of variation II/0043. The RMP is updated accordingly
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.14. Evolocumab – REPATHA (CAP) - EMEA/H/C/003766/X/0002

Applicant: Amgen Europe B.V.

PRAC Rapporteur: Kimmo Jaakkola

Scope: Addition of a new strength of 420 mg (120 mg/mL) for evolocumab solution for injection in cartridge, for subcutaneous (SC) administration by an automated mini-doser device
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.15. Golimumab – SIMPONI (CAP) - EMEA/H/C/000992/II/0063

Applicant: Janssen Biologics B.V.

PRAC Rapporteur: Ulla Wändel Liminga

Scope: Update of sections 4.2 and 5.1 of the SmPC in order to reflect the data from a multicentre, placebo-controlled, double-blind, randomised-withdrawal, parallel group study (GO KIDS) in children (2 to 17 years of age) with active polyarticular juvenile idiopathic arthritis (pJIA). The Package Leaflet and the RMP are updated accordingly
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.16. Golimumab – SIMPONI (CAP) - EMEA/H/C/000992/II/0067

Applicant: Janssen Biologics B.V.

PRAC Rapporteur: Ulla Wändel Liminga

Scope: Update of sections 4.8 and 5.1 of the SmPC in order to update the safety and efficacy information with the data from the final clinical study reports of studies C0524T18 and P07642 in fulfilment of MEA 031 and MEA 032. In addition, the MAH took the opportunity to combine the SmPC for the pre-filled pen and pre-filled syringe for 50 mg strength and for the pre-filled pen and pre-filled syringe for 100 mg strength respectively, in line with the latest QRD template (version 9.1). Moreover, the RMP (version 15) is updated accordingly
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.17. Imiquimod – ALDARA (CAP) - EMEA/H/C/000179/II/0067

Applicant: Meda AB

PRAC Rapporteur: Rafe Suvarna

Scope: Update of sections 4.2 and 5.1 of the SmPC in order to add data on the results of study X-03016-3284 (LEIDA 2, a phase IV randomised active controlled study) and of a meta-analysis of studies X-03016-3271 (LEIDA, a phase IV randomized active controlled study) and X-03016-3284. The RMP is updated (version 3) accordingly
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.18. Linagliptin – TRAJENTA (CAP) - EMEA/H/C/002110/WS/0915 linagliptin, metformin – JENTADUETO (CAP) - EMEA/H/C/002279/WS/0915

Applicant: Boehringer Ingelheim International GmbH

PRAC Rapporteur: Menno van der Elst

Scope: Extension of indication to include the use of Trajenta as a combination therapy with metformin and a sodium-glucose co-transporter-2 (SGLT-2) inhibitor as well as the use of Jentadueto with a SGLT-2 inhibitor. As a consequence, sections 4.1, 4.8 and 5.1 of the SmPC are updated based on studies 1245.30, 1275.10 and 1275.1. The Package Leaflet is updated accordingly. In addition, the MAH took the opportunity to introduce minor editorial changes in the SmPC for Jentadueto only. Moreover, the RMPs for Trajenta (version 10) and for Jentadueto (version 12) are updated accordingly
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.19. Lipegfilgrastim – LONQUEX (CAP) - EMEA/H/C/002556/II/0023

Applicant: Sicor Biotech UAB

PRAC Rapporteur: Julie Williams

Scope: Update of sections 4.4 and 4.8 of the SmPC in order to update the safety information in relation to splenomegaly. The Package Leaflet and Labelling as well as the RMP (version 9) are updated accordingly. In addition, the MAH took the opportunity to bring the product information in line with the latest QRD template (versions 9.1 and 10). Furthermore, minor editorial changes are introduced in the Package Leaflet
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.20. Meningococcal group a, c, w135 and y conjugate vaccine – MENVEO (CAP) - EMEA/H/C/001095/II/0056

Applicant: GSK Vaccines S.r.l

PRAC Rapporteur: Menno van der Elst

Scope: Update of section 4.8 of the SmPC in order to add facial paresis as a new adverse drug reaction and to provide further safety information based on the final clinical study report for study V59_34OB in order to fulfil MEA 023. The Package Leaflet and the RMP (version 8.2) are updated accordingly
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.21. Nintedanib – OFEV (CAP) - EMEA/H/C/003821/II/0006

Applicant: Boehringer Ingelheim International GmbH

PRAC Rapporteur: Marina Dimov Di Giusti

Scope: Update of sections 4.2 and 4.4 of the SmPC in order to revise the dose recommendations for patients with mild hepatic impairment, based on PK/PD modelling data. In addition, the MAH took the opportunity to bring the Annex II in line with the latest QRD template (version 10)
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.22. Ocriplasmin – JETREA (CAP) - EMEA/H/C/002381/II/0026

Applicant: ThromboGenics NV

PRAC Rapporteur: Julie Williams

Scope: Update of sections 4.4, 4.8 and 5.1 of the SmPC to reflect new long-term safety and efficacy data based on the final clinical study report for study TG-MV-014 in fulfilment of MEA 002. The Package Leaflet is updated accordingly. In addition, the MAH took the opportunity to implement editorial changes in the annexes, to align the annexes with the latest QRD templates (versions 9.1 and 10). The RMP (version 7) is updated accordingly
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.23. Perampanel – FYCOMPA (CAP) - EMEA/H/C/002434/X/0025

Applicant: Eisai Europe Ltd.

PRAC Rapporteur: Julie Williams

Scope: Line extension to add a new strength of 0.5 mg/ml and to add a new pharmaceutical form, oral solution
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.24. Ponatinib – ICLUSIG (CAP) - EMEA/H/C/002695/II/0029/G

Applicant: Ariad Pharma Ltd

PRAC Rapporteur: Rafe Suvarna

Scope: Update of section 5.3 of the SmPC in order to add pre-clinical information on fertility and early embryonic development to implantation (study 2424-001) and on carcinogenity (study 805826). In addition, the MAH has submitted final study results for pre-clinical studies ARP590, ARP591, ARP592, ARP593, ARP593 on vascular occlusion mechanism and study ARP598 on effects of ponatinib and its metabolites on in vitro kinase activity and cellular viability following commitments taken during the Article 20 referral procedure (EMEA/H/C/002695/A-20/0003, EC decision on 15 January 2015). No impact in the Product information is proposed for these 6 studies. The RMP has been updated accordingly
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.25. Ranibizumab – LUCENTIS (CAP) - EMEA/H/C/000715/II/0061

Applicant: Novartis Europharm Ltd

PRAC Rapporteur: Ulla Wändel Liminga

Scope: Extension of indication to include treatment of visual impairment due to choroidal neovascularization (CNV) based on 6-month data from the pivotal study CRFB002G2301 (MINERVA). Consequential changes are proposed to SmPC sections 4.1, 4.2, 4.8 and 5.1. The Package Leaflet and the RMP (version 16.0) are updated accordingly
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.26. Rivaroxaban – XARELTO (CAP) - EMEA/H/C/000944/II/0042/G

Applicant: Bayer Pharma AG

PRAC Rapporteur: Qun-Ying Yue

Scope: Update of section 5.1 of the SmPC following the submission of a prospective, single-arm, non-interventional, open-label cohort study conducted to investigate the safety and effectiveness in a real-world setting, study XANTUS (SN 15914) in order to fulfill MEA 025.

In addition, update of section 5.1 of the SmPC following the submission of a prospective, non-interventional, open-label cohort study that was conducted in patient with acute deep vein thrombosis (DVT) to investigate the safety and effectiveness in a real-world setting, study XALIA (SN 15915) in order to fulfill MEA 027. The RMP (version 9.0) is updated accordingly. Additionally the final clinical study reports for studies X-TRA (SN 16320, phase IIIb) and VENTURE-AF (SN 15694, phase IIIb) were also included in the RMP. Finally, the MAH took the opportunity to introduce a minor editorial change in the list of representatives in the package leaflets of all strengths
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.27. Simeprevir – OLYSIO (CAP) - EMEA/H/C/002777/II/0015

Applicant: Janssen-Cilag International N.V.

PRAC Rapporteur: Rafe Suvarna

Scope: Update of sections 4.2, 4.4, 4.5, 4.8 and 5.1 of the SmPC in order to amend the safety information regarding the use of Olysio in interferon-free regimens, based on the primary analysis (SVR12) of studies HPC3017 and HPC3018. The Package Leaflet and Labelling are updated accordingly
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.28. Tedizolid phosphate – SIVEXTRO (CAP) - EMEA/H/C/002846/II/0009

Applicant: Merck Sharp & Dohme Limited

PRAC Rapporteur: Miguel-Angel Macia

Scope: Update of sections 4.4, 4.5 and 5.2 of the SmPC based on the completed drug-drug interaction study MK-1986-004. The Package Leaflet is updated accordingly. In addition the MAH took the opportunity to implement editorial changes in the annexes and to update the annexes in line with the latest QRD template (version 10). The RMP (version 2.0) is updated by removing the missing information for potential risks for drug-drug interactions mediated by CYP3A4, as well as addressing the identified risk for drug-drug interactions mediated via inhibition of breast cancer resistance protein (BCRP), adding updates made to timelines for ongoing and planned studies for long term safety and Asian population experience
Action: For adoption of PRAC RMP AR, PRAC RMP assessment overview and advice to CHMP

5.3.29. Teduglutide – REVESTIVE (CAP) - EMEA/H/C/002345/II/0020

Applicant: NPS Pharma Holdings Limited

PRAC Rapporteur: Torbjorn Callreus

Scope: Extension of indication to include the treatment of the paediatric population. As a consequence, sections 4.1, 4.2, 4.4, 4.8, 5.1, 5.2 and 6.6 of the SmPC are updated in order to update the safety information. The Package Leaflet is updated 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. Adefovir – HEPSERA (CAP) - PSUSA/00000060/201509 (with RMP)

Applicant: Gilead Sciences International Ltd

PRAC Rapporteur: Isabelle Robine

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

6.1.2. Alipogene tiparvovec – GLYBERA (CAP) - PSUSA/00010056/201510

Applicant: UniQure biopharma B.V.

PRAC Rapporteur: Julie Williams

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

6.1.3. Bazedoxifene – CONBRIZA (CAP) - PSUSA/00000302/201510

Applicant: Pfizer Limited

PRAC Rapporteur: Martin Huber

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

6.1.4. Budesonide, formoterol – BIRESP SPIROMAX (CAP); BUDESONIDE/FORMOTEROL TEVA (CAP); BUDESONIDE/FORMOTEROL TEVA PHARMA B.V. (CAP); DUORESP SPIROMAX (CAP); VYLAER SPIROMAX (CAP) - PSUSA/00010202/201510

Applicant: Teva Pharma B.V.

PRAC Rapporteur: Torbjorn Callreus

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

6.1.5. Carbidopa, entacapone, levodopa – CORBILTA (CAP); LEVODOPA/CARBIDOPA/ENTACAPONE ORION (CAP); STALEVO (CAP) - PSUSA/00000547/201510

Applicant: Orion Corporation

PRAC Rapporteur: Kirsti Villikka

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

6.1.6. Ceftaroline fosamil – ZINFORO (CAP) - PSUSA/00010013/201510

Applicant: AstraZeneca AB

PRAC Rapporteur: Julie Williams

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

6.1.7. Ceritinib – ZYKADIA (CAP) - PSUSA/00010372/201510

Applicant: Novartis Europharm Ltd

PRAC Rapporteur: Ulla Wändel Liminga

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

6.1.8. Deferasirox – EXJADE (CAP) - PSUSA/00000939/201510 (with RMP)

Applicant: Novartis Europharm Ltd

PRAC Rapporteur: Isabelle Robine

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

6.1.9. Defibrotide – DEFITELIO (CAP) - PSUSA/00010086/201510

Applicant: Gentium S.r.l.

PRAC Rapporteur: Julie Williams

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

6.1.10. Delamanid – DELTYBA (CAP) - PSUSA/00010213/201510 (with RMP)

Applicant: Otsuka Novel Products GmbH

PRAC Rapporteur: Rafe Suvarna

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

6.1.11. Empagliflozin – JARDIANCE (CAP); empagliflozin, metformin – SYNJARDY (CAP) - PSUSA/00010388/201510

Applicant: Boehringer Ingelheim International GmbH, Boehringer Ingelheim GmbH

PRAC Rapporteur: Miguel-Angel Macia

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

6.1.12. Eptotermin alfa – OPGENRA (CAP); OSIGRAFT (CAP) - PSUSA/00001247/201509

Applicant: Olympus Biotech International Limited

PRAC Rapporteur: Kirsti Villikka

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

6.1.13. Eslicarbazepine acetate – ZEBINIX (CAP) - PSUSA/00001267/201510

Applicant: Bial - Portela & Cª, S.A.

PRAC Rapporteur: Martin Huber

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

6.1.14. Flutemetamol (^{18}F) – VIZAMYL (CAP) - PSUSA/00010293/201510

Applicant: GE Healthcare Ltd

PRAC Rapporteur: Julie Williams

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

6.1.15. Granisetron – SANCUSO (CAP) - PSUSA/00010101/201510

Applicant: ProStrakan Limited

PRAC Rapporteur: Jolanta Gulbinovic

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

6.1.16. Hydrocortisone – PLENADREN (CAP) - PSUSA/00009176/201511

Applicant: Shire Services BVBA

PRAC Rapporteur: Qun-Ying Yue

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

6.1.17. Insulin detemir – LEVEMIR (CAP) - PSUSA/00001750/201510

Applicant: Novo Nordisk A/S

PRAC Rapporteur: Doris Stenver

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

6.1.18. Ledipasvir, sofosbuvir – HARVONI (CAP) - PSUSA/00010306/201510

Applicant: Gilead Sciences International Ltd

PRAC Rapporteur: Margarida Guimarães

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

6.1.19. Levofloxacin – QUINSAIR (CAP) - PSUSA/00010429/201509

Applicant: Raptor Pharmaceuticals Europe BV

PRAC Rapporteur: Miguel-Angel Macia

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

6.1.20. Lopinavir, ritonavir – ALUVIA (Art 58[3]); KALETRA (CAP) - PSUSA/00001905/201509

Applicant: AbbVie Ltd

PRAC Rapporteur: Isabelle Robine

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

6.1.21. Lurasidone – LATUDA (CAP) - PSUSA/00010114/201510 (with RMP)

Applicant: Sunovion Pharmaceuticals Europe Ltd

PRAC Rapporteur: Qun-Ying Yue

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

6.1.22. Macitentan – OPSUMIT (CAP) - PSUSA/00010115/201510

Applicant: Actelion Registration Ltd.

PRAC Rapporteur: Dolores Montero Corominas

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

6.1.23. Meningococcal group a, c, w135 and y conjugate vaccine – NIMENRIX (CAP) - PSUSA/00010044/201510

Applicant: Pfizer Limited

PRAC Rapporteur: Rafe Suvarna

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

6.1.24. Micafungin – MYCAMINE (CAP) - PSUSA/00002051/201510

Applicant: Astellas Pharma Europe B.V.

PRAC Rapporteur: Martin Huber

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

6.1.25. Miglustat – ZAVESCA (CAP) - PSUSA/00002062/201510

Applicant: Actelion Registration Ltd.

PRAC Rapporteur: Qun-Ying Yue

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

[3] Article 58 of Regulation (EC) No 726/2004 allows the Agency's Committee for Medicinal Products for Human Use (CHMP) to give opinions, in co-operation with the World Health Organisation (WHO), on medicinal products for human use that are intended exclusively for markets outside of the European Union (EU)

6.1.26. Netupitant, palonosetron – AKYNZEO (CAP) - PSUSA/00010393/201510

Applicant: Helsinn Birex Pharmaceuticals Ltd

PRAC Rapporteur: Carmela Macchiarulo

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

6.1.27. Nintedanib – OFEV (CAP) - PSUSA/00010319/201510

Applicant: Boehringer Ingelheim International GmbH

PRAC Rapporteur: Marina Dimov Di Giusti

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

6.1.28. Obinutuzumab – GAZYVARO (CAP) - PSUSA/00010279/201510

Applicant: Roche Registration Limited

PRAC Rapporteur: Julie Williams

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

6.1.29. Ocriplasmin – JETREA (CAP) - PSUSA/00010122/201510

Applicant: ThromboGenics NV

PRAC Rapporteur: Julie Williams

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

6.1.30. Ofatumumab – ARZERRA (CAP) - PSUSA/00002202/201510

Applicant: Novartis Europharm Ltd

PRAC Rapporteur: Doris Stenver

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

6.1.31. Oseltamivir – TAMIFLU (CAP) - PSUSA/00002225/201509

Applicant: Roche Registration Limited

PRAC Rapporteur: Kirsti Villikka

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

6.1.32. Pandemic influenza vaccine (H5N1) (surface antigen, inactivated, adjuvanted) – FOCLIVIA (CAP) - Prepandemic influenza vaccine (H5N1) (surface antigen, inactivated, adjuvanted) - AFLUNOV (CAP) - PSUSA/00010008/201510

Applicant: Novartis Vaccines Influenza Srl

PRAC Rapporteur: Carmela Macchiarulo

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

6.1.33. Para-aminosalicylic acid – GRANUPAS (CAP) - PSUSA/00010171/201510

Applicant: Lucane Pharma

PRAC Rapporteur: Julie Williams

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

6.1.34. Pasireotide – SIGNIFOR (CAP) - PSUSA/00009253/201510

Applicant: Novartis Europharm Ltd

PRAC Rapporteur: Qun-Ying Yue

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

6.1.35. Pazopanib – VOTRIENT (CAP) - PSUSA/00002321/201510

Applicant: Novartis Europharm Ltd

PRAC Rapporteur: Doris Stenver

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

6.1.36. Posaconazole – NOXAFIL (CAP) - PSUSA/00002480/201510

Applicant: Merck Sharp & Dohme Limited

PRAC Rapporteur: Rafe Suvarna

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

6.1.37. Propranolol – HEMANGIOL (CAP) - PSUSA/00010250/201510

Applicant: Pierre Fabre Dermatologie

PRAC Rapporteur: Dolores Montero Corominas

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

6.1.38. Prucalopride – RESOLOR (CAP) - PSUSA/00002568/201510 (with RMP)

Applicant: Shire Pharmaceuticals Ireland Ltd.

PRAC Rapporteur: Rafe Suvarna

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

6.1.39. Ramucirumab – CYRAMZA (CAP) - PSUSA/00010323/201510 (with RMP)

Applicant: Eli Lilly Nederland B.V.

PRAC Rapporteur: Brigitte Keller-Stanislawski

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

6.1.40. Siltuximab – SYLVANT (CAP) - PSUSA/00010254/201510

Applicant: Janssen-Cilag International NV

PRAC Rapporteur: Brigitte Keller-Stanislawski

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

6.1.41. Thalidomide – THALIDOMIDE CELGENE (CAP) - PSUSA/00002919/201510

Applicant: Celgene Europe Limited

PRAC Rapporteur: Isabelle Robine

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

6.1.42. Tocilizumab – ROACTEMRA (CAP) - PSUSA/00002980/201510

Applicant: Roche Registration Limited

PRAC Rapporteur: Brigitte Keller-Stanislawski

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

6.1.43. Turoctocog alfa – NOVOEIGHT (CAP) - PSUSA/00010138/201510

Applicant: Novo Nordisk A/S

PRAC Rapporteur: Brigitte Keller-Stanislawski

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

6.1.44. Umeclidinium bromide – INCRUSE (CAP) - PSUSA/00010263/201510

Applicant: Glaxo Group Ltd

PRAC Rapporteur: Carmela Macchiarulo

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

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

6.2.1. Filgrastim – ACCOFIL (CAP); FILGRASTIM HEXAL (CAP); GRASTOFIL (CAP); NIVESTIM (CAP); RATIOGRASTIM (CAP) ; TEVAGRASTIM (CAP); ZARZIO (CAP), NAP - PSUSA/00001391/201509

Applicant: Accord Healthcare Ltd (Accofil), Hexal AG (Filgrastim Hexal), Apotex Europe BV (Gastrofil), Hospira UK Limited (Nivestim), Ratiopharm GmbH (Ratiograstim), Teva GmbH (Tevagrastim), Sandoz GmbH (Zarzio), various

PRAC Rapporteur: Kirsti Villikka

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

6.2.2. Influenza vaccine (H1N1)v (whole virion, vero cell derived, inactivated) – CELVAPAN (CAP), NAP - PSUSA/00002280/201510

Applicant: Nanotherapeutics Bohumil Sro, various

PRAC Rapporteur: Brigitte Keller-Stanislawski

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

6.2.3. Sodium oxybate – XYREM (CAP); NAP - PSUSA/00002757/201510

Applicant: UCB Pharma Ltd., various

PRAC Rapporteur: Leonor Chambel

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

6.2.4. Somatropin – NUTROPINAQ (CAP); OMNITROPE (CAP); SOMATROPIN BIOPARTNERS (CAP); NAP - PSUSA/00002772/201509

Applicant: Ipsen Pharma (NutropinAq), Sandoz GmbH (Omnitrope), BioPartners GmbH (Somatropin Biopartners), various

PRAC Rapporteur: Torbjorn Callreus

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

6.2.5. Tadalafil – ADCIRCA (CAP); CIALIS (CAP); NAP - PSUSA/00002841/201510

Applicant: Eli Lilly Nederland B.V.(Adcirca, Cialis), various

PRAC Rapporteur: Miguel-Angel Macia

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

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

6.3.1. Acetylcysteine (NAP) - PSUSA/00000034/201509

Applicant: various

PRAC Lead: Adam Przybylkowski

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

6.3.2. Alfentanil (NAP) - PSUSA/00000082/201509

Applicant: various

PRAC Lead: Almath Spooner

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

6.3.3. Beractant (NAP) - PSUSA/00000384/201510

Applicant: various

PRAC Lead: Julie Williams

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

6.3.4. Bisoprolol (NAP) - PSUSA/00000419/201509

Applicant: various

PRAC Lead: Kimmo Jaakkola

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

6.3.5. Carmustine (implant) (NAP) - PSUSA/00010348/201509

Applicant: various

PRAC Lead: Isabelle Robine

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

6.3.6. Chlorquinaldol (vaginal tablet), promestriene (NAP) - PSUSA/00009272/201509

Applicant: various

PRAC Lead: Roxana Stefania Stroe

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

6.3.7. Desogestrel, ethinylestradiol (NAP) - PSUSA/00000967/201509

Applicant: various

PRAC Lead: Kimmo Jaakkola

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

6.3.8. Diclofenac (systemic formulations) (NAP) - PSUSA/00001048/201509

Applicant: various

PRAC Lead: Doris Stenver

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

6.3.9. Diclofenac (topical formulations) (NAP) - PSUSA/00010342/201509

Applicant: various

PRAC Lead: Doris Stenver

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

6.3.10. Fenoterol (obstetric indications) (NAP) - PSUSA/00010001/201509

Applicant: various

PRAC Lead: Marina Dimov Di Giusti

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

6.3.11. Glycopyrronium (all indications except for chronic obstructive pulmonary disease) (NAP) - PSUSA/00001556/201509

Applicant: various

PRAC Lead: Almath Spooner

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

6.3.12. Glycopyrronium, neostigmine (NAP) - PSUSA/00001557/201509

Applicant: various

PRAC Lead: Almath Spooner

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

6.3.13. Ketoprofen (topical use only) (NAP) - PSUSA/00009205/201509

Applicant: various

PRAC Lead: Ulla Wändel Liminga

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

6.3.14. Lactitol (NAP) - PSUSA/00001819/201509

Applicant: various

PRAC Lead: Jan Neuhauser

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

6.3.15. Latanoprost (paediatric indication only) (NAP) - PSUSA/00001834/201510

Applicant: various

PRAC Lead: Julie Williams

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

6.3.16. Levofloxacin (NAP) - PSUSA/00001854/201510

Applicant: various

PRAC Lead: Julie Williams

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

6.3.17. Lisinopril (NAP) - PSUSA/00001894/201509

Applicant: various

PRAC Lead: Margarida Guimarães

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

6.3.18. Phloroglucinol, trimethylphloroglucinol (NAP) - PSUSA/00010355/201509

Applicant: various

PRAC Lead: Isabelle Robine

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

6.4. Follow-up to PSUR/PSUSA procedures

6.4.1. Omalizumab – XOLAIR (CAP) - EMEA/H/C/000606/LEG 050.1

Applicant: Novartis Europharm Ltd

PRAC Rapporteur: Qun-Ying Yue

Scope: MAH's response to LEG 050 [venous thromboembolism cumulative review as requested in the conclusions of EMEA/H/C/PSUSA/00002214/201412 adopted by the PRAC in July 2015] as per the request for supplementary information adopted in January 2016
Action: For adoption of advice to CHMP

6.4.2. Pemetrexed – ALIMTA (CAP) - EMEA/H/C/000564/LEG 025

Applicant: Eli Lilly Nederland B.V.

PRAC Rapporteur: Isabelle Robine

Scope: Evaluation of a cumulative review and detailed analysis of all cases pertaining to local and systemic scleroderma submitted by the MAH following the recommendation of the PSUSA/00002330/201502 procedure adopted in September 2015
Action: For adoption of advice to CHMP

6.4.3. Pemetrexed – ALIMTA (CAP) - EMEA/H/C/000564/LEG 026

Applicant: Eli Lilly Nederland B.V.

PRAC Rapporteur: Isabelle Robine

Scope: Evaluation of a cumulative analysis of cases of acute myeloid leukaemia and rhabdomyolysis, cumulative review of cases of atrial fibrillation, cumulative review of cases of posterior reversible encephalopathy syndrome (PRES) and leukoencephalopathy and a cumulative review of cases of palmar-plantar erythrodysaesthesia submitted by the MAH following the recommendation of the PSUSA/00002330/201502 procedure adopted in September 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)[4]

7.1.1. Asfotase alfa – STRENSIQ (CAP) - EMEA/H/C/PSP/0032.1

Applicant: Alexion Europe SAS

PRAC Rapporteur: Almath Spooner

Scope: Revised PASS protocol for study ALX-HPP-501: an observational, longitudinal, prospective, long-term registry of patients with hypophosphatasia to collect information on the epidemiology of the disease, including clinical outcomes and quality of life, and to evaluate safety and effectiveness data in patients treated with Strensiq
Action: For adoption of PRAC Assessment Report, PRAC outcome letter

7.1.2. Imatinib – GLIVEC (CAP) - EMEA/H/C/PSP/0042.A.1

Applicant: Novartis Europharm Ltd.

PRAC Rapporteur: Dolores Montero Corominas

Scope: Revised PASS protocol for study CSTI571I2201: a European observational registry collecting efficacy and safety data in newly diagnosed paediatric Philadelphia positive (Ph+) acute lymphoblastic leukaemia (ALL) patients treated with chemotherapy + imatinib ± hematopoietic stem cell treatment (±HSCT)
Action: For adoption of PRAC Assessment Report, PRAC outcome letter

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

7.1.3. Ivabradine – PROCORALAN (CAP); CORLENTOR (CAP); IVABRADINE ANPHARM (CAP) - EMEA/H/C/PSP/j/0019.1.A.1

Applicant: Les Laboratoires Servier (Corlentor, Procorolan), Anpharm Przedsiębiorstwo Farmaceutyczne S.A. (Ivabradine Anpharm)

PRAC Rapporteur: Menno van der Elst

Scope: Revised protocol for a drug utilisation study (DUS) for a multinational, retrospective, observational study to assess the effectiveness of risk-minimisation measures
Action: For adoption of PRAC Assessment Report, PRAC outcome letter

7.1.4. Sebelipase alfa – KANUMA (CAP) - EMEA/H/C/PSP/0036.1

Applicant: Alexion Europe SAS

PRAC Rapporteur: Qun-Ying Yue

Scope: Revised protocol for a PASS: a non-interventional, multicentre, prospective disease and clinical outcome registry of patients with lysosomal acid lipase deficiency (LAL-D) to further understand the disease, its progression and any associated complication, and to evaluate the long-term efficacy (normalisation of hepatic function) and safety of Kanuma (in particular hypersensitivity reactions, including anaphylaxis, and anti-drug antibodies development potentially impacting response to drug)
Action: For adoption of PRAC Assessment Report, PRAC outcome letter

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

7.2.1. Alemtuzumab – LEMTRADA (CAP) - EMEA/H/C/003718/MEA/007

Applicant: Genzyme Therapeutics Ltd

PRAC Rapporteur: Torbjorn Callreus

Scope: PASS protocol for study No. OBS13434: a prospective, multicentre, observational, post-authorisation safety study (PASS) to evaluate the long term safety profile of alemtuzumab treatment in patients with relapsing forms of multiple sclerosis (RMS)
Action: For adoption of advice to CHMP

7.2.2. Bromelain enriched proteolytic enzyme preparation from ananas comosus – NEXOBRID (CAP) - EMEA/H/C/002246/MEA/003.4

Applicant: MediWound Germany GmbH

PRAC Rapporteur: Valerie Strassmann

Scope: MAH's responses to MEA 003.3 [revised PASS protocol for study MW2013-06-01: drug utilisation study (DUS) to further evaluate the effectiveness of the risk minimisation activities (including evaluation of educational and training materials)] as per request for supplementary information adopted in January 2016
Action: For adoption of advice to CHMP

7.2.3. Collagenase clostridium histolyticum – XIAPEX (CAP) - EMEA/H/C/002048/MEA/027.2

Applicant: Swedish Orphan Biovitrum AB (publ)

[5] 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

PRAC Rapporteur: Martin Huber

Scope: MAH's responses to MEA 027.1 [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] as per request for supplementary information adopted in January 2016
Action: For adoption of advice to CHMP

7.2.4. Desloratadine – AERIUS (CAP) - EMEA/H/C/000313/MEA/065.1; AZOMYR (CAP) - EMEA/H/C/000310/MEA/065.1; NEOCLARITYN (CAP) - EMEA/H/C/000314/MEA/065.1

Applicant: Merck Sharp & Dohme Limited

PRAC Rapporteur: Jean-Michel Dogné

Scope: MAH's responses to MEA 065 [PASS investigating the association between the use of desloratadine and the risk of seizures, supraventricular tachycardia, and atrial fibrillation or flutter: a nordic register-based study] as per request for supplementary information adopted in October 2015
Action: For adoption of advice to CHMP

7.2.5. Empagliflozin – JARDIANCE (CAP) - EMEA/H/C/002677/MEA/004.1

Applicant: Boehringer Ingelheim International GmbH

PRAC Rapporteur: Miguel-Angel Macia

Scope: MAH's responses to MEA 004 [PASS study1245.97 to assess the risk of urinary tract malignancies in relation to empagliflozin exposure in patients with type 2 mellitus diabetes: a multi-database European study, preceded by feasibility assessment] as per request for supplementary information adopted in December 2015
Action: For adoption of advice to CHMP

7.2.6. Estrogens conjugated, bazedoxifene – DUAVIVE (CAP) - EMEA/H/C/002314/MEA/003.2

Applicant: Pfizer Limited

PRAC Rapporteur: Martin Huber

Scope: MAH's responses to MEA 003.1 [revised protocol for drug utilisation study (DUS) No. B2311061] as per request for supplementary information adopted in October 2015
Action: For adoption of advice to CHMP

7.2.7. Human normal immunoglobulin – PRIVIGEN (CAP) - EMEA/H/C/000831/MEA/022.4

Applicant: CSL Behring GmbH

PRAC Rapporteur: Brigitte Keller-Stanislawski

Scope: MAH's responses to MEA 022.3 [revised protocol for study IgPro10_5003: an observational hospital-based cohort study in the US: Privigen use and haemolytic anaemia in adults and children and the Privigen safety profile in children with chronic inflammatory demyelinating polyneuropathy (CIDP)] as per request for supplementary information adopted in January 2016
Action: For adoption of advice to CHMP

7.2.8. Lumacaftor, ivacaftor – ORKAMBI (CAP) - EMEA/H/C/003954/MEA/003

Applicant: Vertex Pharmaceuticals (Europe) Ltd.

PRAC Rapporteur: Almath Spooner

Scope: PASS protocol for study VX14 809 108: an observational study to evaluate the utilisation patterns and long-term effects of lumacaftor and ivacaftor combination therapy in patients with cystic fibrosis
Action: For adoption of advice to CHMP

7.2.9. Panobinostat – FARYDAK (CAP) - EMEA/H/C/003725/MEA/002.1

Applicant: Novartis Europharm Ltd

PRAC Rapporteur: Julie Williams

Scope: MAH's responses to MEA 002 [PASS study LBH589D2408 of panobinostat use in relapsed or relapsed/refractory multiple myeloma patients who have received at least two prior regimens including bortezomib and an immunomodulatory agent in a real-world setting according to the current EU prescribing information and document adherence to dosing regimen (including the dosing card, blister pack) by describing clinical characteristics, frequency and severity of the medication error events] as per request for supplementary information adopted in December 2015
Action: For adoption of advice to CHMP

7.2.10. Rituximab – MABTHERA (CAP) - EMEA/H/C/000165/MEA/093.2

Applicant: Roche Registration Limited

PRAC Rapporteur: Doris Stenver

Scope: MAH's responses to MEA 093.1 [revised PASS registry protocol for a long-term surveillance study of rituximab (Mabthera)-treated patients with granulomatosis with polyangiitis (GPA) or microscopic polyangiitis (MPA)] as per request for supplementary information adopted in February 2016
Action: For adoption of advice to CHMP

7.2.11. Sacubitri, valsartan – ENTRESTO (CAP) - EMEA/H/C/004062/MEA/002

Applicant: Novartis Europharm Ltd

PRAC Rapporteur: Rafe Suvarna

Scope: PASS protocol for study No. CLCZ696B2014: a non-interventional post-authorisation European database safety study (category3) to characterize the risk of angioedema and other specific safety events of interest in association with use of Entresto (sacubitril/valsartan) in adult patients with heart failure
Action: For adoption of advice to CHMP

7.2.12. Sacubitril, valsartan – ENTRESTO (CAP) - EMEA/H/C/004062/MEA/004

Applicant: Novartis Europharm Ltd

PRAC Rapporteur: Rafe Suvarna

Scope: PASS protocol for study No. CLCZ696B2015: a non-interventional post-authorisation European database safety study (category 3) to assess the risk of myotoxicity, hepatotoxicity and acute pancreatitis in statin-exposed heart failure patients with or without concomitant use of sacubitril/valsartan

Action: For adoption of advice to CHMP

7.3. Results of PASS imposed in the marketing authorisation(s)[6]

None

7.4. Results of PASS non-imposed in the marketing authorisation(s)[7]

7.4.1. Dabigatran etexilate – PRADAXA (CAP) - EMEA/H/C/000829/II/0093

Applicant: Boehringer Ingelheim International GmbH

PRAC Rapporteur: Torbjorn Callreus

Scope: Submission of the final clinical trial report of study 1160.149: a post-authorisation safety study to evaluate the effectiveness of the risk minimisation activities in the treatment of stroke prevention in atrial fibrillation in order to address part of follow-up measure MEA 026. The RMP (version 31.6) is updated with results from clinical study 1160.149
Action: For adoption of PRAC Assessment Report

7.4.2. Eptacog alfa (activated) – NOVOSEVEN (CAP) - EMEA/H/C/000074/II/0089

Applicant: Novo Nordisk A/S

PRAC Rapporteur: Sabine Straus

Scope: Submission of the final study teport for study NN7025-3601 : a prospective observational study on NovoSeven room temperature (VII25) in patients with haemophilia A and B. The submission of this study report addresses MEA 046.4. The RMP (version 6.1) is updated accordingly
Action: For adoption of PRAC Assessment Report

7.4.3. Golimumab – SIMPONI (CAP) - EMEA/H/C/000992/II/0068/G

Applicant: Janssen Biologics B.V.

PRAC Rapporteur: Ulla Wändel Liminga

Scope: Submission of the final results of a non-interventional, prospective registry study CNTO148ART4003 in order to fulfill the post-authorisation commitment MEA 006: a registry study Swedish database initiative for exposure to golimumab: a review and analysis of adverse events from the Swedish national registry system. The RMP is updated accordingly to update the due date of the completion and final report of studies: - P04480 (MEA 005.4) 'long-term observation of treatment with biologics in rheumatoid arthritis' from December 2017 to December 2021 and December 2018 to December 2022 respectively. - CNTO148ART4001 (MEA 007.1) 'exposure to golimumab during pregnancy in patients with rheumatoid arthritis, psoriatic arthritis, and ankylosing spondylitis: a review and analysis of birth outcomes from the Swedish, Danish, and Finnish medical birth registers' from February 2016 to December 2021 and February 2017 to December 2022 respectively. -MK-8259-042 (MEA0027.3) 'safety study of golimumab in ulcerative colitis using the Spanish ENEIDA registry' from July 2015 to March 2022 and December 2018 to December 2022 respectively and to introduce the date of provision of the final report (March 2023). In addition, the need to submit interim report for study CNTO148ART4002 was removed following the completion of MEA 008.2
Action: For adoption of PRAC Assessment Report

[6] In accordance with Article 107p-q of Directive 2001/83/EC
[7] 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

7.4.4. Imatinib – GLIVEC (CAP) - EMEA/H/C/000406/II/0100

Applicant: Novartis Europharm Ltd

PRAC Rapporteur: Dolores Montero Corominas

Scope: Submission of the final clinical study report for study CSTI571A2403: 'a global Gleevec/Glivec and Tasigna pregnancy exposure registry' (category 3 study)
Action: For adoption of PRAC Assessment Report

7.4.5. Nilotinib – TASIGNA (CAP) - EMEA/H/C/000798/II/0080

Applicant: Novartis Europharm Ltd

PRAC Rapporteur: Doris Stenver

Scope: Submission of the final clinical study report for study CSTI571A2403: 'a global Gleevec/Glivec and Tasigna pregnancy exposure registry' (category 3) in fulfilment of MEA 038
Action: For adoption of PRAC Assessment Report

7.4.6. Voriconazole – VFEND (CAP) - EMEA/H/C/000387/II/0115

Applicant: Pfizer Limited

PRAC Rapporteur: Sabine Straus

Scope: Submission of the final study report for a non-interventional post authorisation safety study A1501097: evaluation of the potential association between voriconazole use and squamous cell carcinoma (SCC) of the skin among patients with lung or lung/heart transplants in order to fulfil MEA 071.11. The RMP (version 4.0) is updated accordingly
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[8]

7.5.1. Abatacept – ORENCIA (CAP) - EMEA/H/C/000701/MEA/046.3

Applicant: Bristol-Myers Squibb Pharma EEIG

PRAC Rapporteur: Kirsti Villikka

Scope: Annual update for study IM101240: observational registry of abatacept in patients with juvenile idiopathic arthritis (JIA registry)
Action: For adoption of advice to CHMP

7.5.2. Abatacept – ORENCIA (CAP) - EMEA/H/C/000701/MEA/048.4

Applicant: Bristol-Myers Squibb Pharma EEIG

PRAC Rapporteur: Kirsti Villikka

Scope: Annual report on the juvenile idiopathic arthritis (JIA) registry, an observational registry of abatacept in patients with juvenile idiopathic arthritis
Action: For adoption of advice to CHMP

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

7.5.3. Dolutegravir – TIVICAY (CAP) - EMEA/H/C/002753/MEA/001

Applicant: ViiV Healthcare UK Limited

PRAC Rapporteur: Julie Williams

Scope: Interim annual report from EuroSIDA PASS study No. 201177: a prospective observational cohort study in patients receiving dolutegravir (category 3) to investigate the risk of hypersensitivity reactions, hepatotoxicity and serious rash (division of acquired immune deficiency syndrome (AIDS) (category 3 or 4)
Action: For adoption of advice to CHMP

7.5.4. Dolutegravir, abacavir, lamivudine – TRIUMEQ (CAP) - EMEA/H/C/002754/MEA/007

Applicant: ViiV Healthcare UK Limited

PRAC Rapporteur: Julie Williams

Scope: Interim annual report form a prospective observational cohort study to monitor occurrence of hypersensitivity reactions and hepatotoxicity in patients receiving dolutegravir (category 3)
Action: For adoption of advice to CHMP

7.5.5. Infliximab – INFLECTRA (CAP) – EMEA/H/C/002778/MEA 008.2, REMSIMA (CAP) - EMEA/H/C/002576/MEA/008.2

Applicant: Celltrion Healthcare Hungary Kft.

PRAC Rapporteur: Rafe Suvarna

Scope: Evaluation of the MAH's responses to MEA 008.1 [Fifth periodic report for post marketing surveillance of Remsima 100 mg to evaluate safety and efficacy in Korea] as per the request for supplementary information adopted in December 2015
Action: For adoption of advice to CHMP

7.6. Others

None

7.7. New Scientific Advice

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

7.8. Ongoing Scientific Advice

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

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

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

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

8.1. Annual reassessments of the marketing authorisation

8.1.1. Afamelanotide – SCENESSE (CAP) - EMEA/H/C/002548/S/0007 (without RMP)

Applicant: Clinuvel (UK) Limited

PRAC Rapporteur: Valerie Strassmann

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

8.1.2. Antithrombin alfa – ATRYN (CAP) - EMEA/H/C/000587/S/0026 (without RMP)

Applicant: GTC Biotherapeutics UK Limited

PRAC Rapporteur: Isabelle Robine

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. Crizotinib – XALKORI (CAP) - EMEA/H/C/0002489/R/0041 (without RMP)

Applicant: Pfizer Limited

PRAC Rapporteur: Isabelle Robine

Scope: Conditional renewal of the marketing authorisation
Action: For adoption of advice to CHMP

8.3. Renewals of the marketing authorisation

8.3.1. Antithrombin alfa – ATRYN (CAP) - EMEA/H/C/000587/R/0024 (without RMP)

Applicant: GTC Biotherapeutics UK Limited

PRAC Rapporteur: Isabelle Robine

Scope: 5-year renewal of the marketing authorisation
Action: For adoption of advice to CHMP

8.3.2. Dasatinib – SPRYCEL (CAP) - EMEA/H/C/000709/R/0050 (without RMP)

Applicant: Bristol-Myers Squibb Pharma EEIG

PRAC Rapporteur: Doris Stenver

Scope: 5-year renewal of the marketing authorisation
Action: For adoption of advice to CHMP

8.3.3. Emtricitabine, rilpivirine, tenofovir disoproxil – EVIPLERA (CAP) - EMEA/H/C/002312/R/0074 (without RMP)

Applicant: Gilead Sciences International Ltd

PRAC Rapporteur: Menno van der Elst

Scope: 5-year renewal of the marketing authorisation
Action: For adoption of advice to CHMP

8.3.4. Exenatide – BYETTA (CAP) - EMEA/H/C/000698/R/0053 (with RMP)

Applicant: AstraZeneca AB

PRAC Rapporteur: Qun-Ying Yue

Scope: 5-year renewal of the marketing authorisation
Action: For adoption of advice to CHMP

8.3.5. Fidaxomicin – DIFICLIR (CAP) - EMEA/H/C/002087/R/0026 (with RMP)

Applicant: Astellas Pharma Europe B.V.

PRAC Rapporteur: Qun-Ying Yue

Scope: 5-year renewal of the marketing authorisation
Action: For adoption of advice to CHMP

8.3.6. Hydrocortisone – PLENADREN (CAP) - EMEA/H/C/002185/R/0020 (without RMP)

Applicant: Shire Services BVBA

PRAC Rapporteur: Qun-Ying Yue

Scope: 5-year renewal of the marketing authorisation
Action: For adoption of advice to CHMP

8.3.7. Levetiracetam – LEVETIRACETAM ACCORD (CAP) - EMEA/H/C/002290/R/0012 (with RMP)

Applicant: Accord Healthcare Ltd

PRAC Rapporteur: Veerle Verlinden

Scope: 5-year renewal of the marketing authorisation
Action: For adoption of advice to CHMP

8.3.8. Levetiracetam – LEVETIRACETAM ACTAVIS (CAP) - EMEA/H/C/002355/R/0013 (without RMP)

Applicant: Actavis Group PTC ehf

PRAC Rapporteur: Veerle Verlinden

Scope: 5-year renewal of the marketing authorisation
Action: For adoption of advice to CHMP

8.3.9. Levetiracetam – LEVETIRACETAM TEVA (CAP) - EMEA/H/C/002316/R/0021 (without RMP)

Applicant: Teva B.V.

PRAC Rapporteur: Veerle Verlinden

Scope: 5-year renewal of the marketing authorisation
Action: For adoption of advice to CHMP

8.3.10. Perflutren – LUMINITY (CAP) - EMEA/H/C/000654/R/0021 (without RMP)

Applicant: Lantheus MI UK Ltd

PRAC Rapporteur: Almath Spooner

Scope: 5-year renewal of the marketing authorisation
Action: For adoption of advice to CHMP

8.3.11. Piperaquine tetraphosphate, artenimol – EURARTESIM (CAP) - EMEA/H/C/001199/R/0023 (without RMP)

Applicant: Sigma-Tau Industrie Farmaceutiche Riunite S.p.A.

PRAC Rapporteur: Julie Williams

Scope: 5-year renewal of the marketing authorisation
Action: For adoption of advice to CHMP

8.3.12. Pramipexole – PRAMIPEXOLE ACCORD (CAP) - EMEA/H/C/002291/R/0010 (without RMP)

Applicant: Accord Healthcare Ltd

PRAC Rapporteur: Doris Stenver

Scope: 5-year renewal of the marketing authorisation
Action: For adoption of advice to CHMP

8.3.13. Rilpivirine – EDURANT (CAP) - EMEA/H/C/002264/R/0022 (with RMP)

Applicant: Janssen-Cilag International N.V.

PRAC Rapporteur: Sabine Straus

Scope: 5-year renewal of the marketing authorisation
Action: For adoption of advice to CHMP

8.3.14. Saxagliptin, metformin hydrochloride – KOMBOGLYZE (CAP) - EMEA/H/C/002059/R/0032 (without RMP)

Applicant: AstraZeneca AB

PRAC Rapporteur: Menno van der Elst

Scope: 5-year renewal of the marketing authorisation
Action: For adoption of advice to CHMP

8.3.15. Tafamidis – VYNDAQEL (CAP) - EMEA/H/C/0002294/R/0032 (without RMP)

Applicant: Pfizer Limited

PRAC Rapporteur: Isabelle Robine

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

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

Disclosure of 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 working group - Recommendations on efficiency of plenary meetings – best practice guide

PRAC lead: Martin Huber, Rafe Suvarna, Ulla Wändel Liminga
Action: For discussion

12.2. Coordination with EMA Scientific Committees or CMDh

12.2.1. Advancing the Development of Paediatric Therapeutics (ADEPT): successes and challenges of performing long-term paediatric safety studies – report from the FDA public workshop held in April 2016

Action: For discussion

12.2.2. Joint Paediatric Committee (PDCO)-PRAC Working Group - guideline on conduct of pharmacovigilance for medicines used by the paediatric population - proposal for creation of new GVP chapter for special populations

Action: For discussion

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

12.3.1. Advisory Group on Summary of Product Characteristics (SmPC) - 2010-2015 activity report

Action: For discussion

12.3.2. Vaccines Working Party (VWP) / PRAC: Stakeholders proposal for the implementation of the principles of passive enhanced safety surveillance (ESS) for the upcoming pilot seasons

Action: For adoption

12.4. Cooperation within the EU regulatory network

None

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

None

12.8. Planning and reporting

None

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)

PRAC lead: Menno van der Elst; Margarida Guimarães
Action: For discussion

12.10.2. Periodic safety update reports

None

12.10.3. PSUR action group – roadmap for PSUR issues: Joint PRAC/CMDh recommendation paper on common understanding - finalisation

PRAC lead: Margarida Guimarães; Menno van der Elst; Jolanta Gulbinovic
Action: For adoption

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

Action: For adoption of the revised list

12.11. Signal management

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

PRAC lead: Sabine Straus

Action: For discussion

12.12. Adverse drug reactions reporting and additional reporting

12.12.1. Good Pharmacovigilance Practice (GVP) module VI on Management and reporting of adverse reactions to medicinal products - revision 2

Action: For discussion

12.12.2. Management and reporting of adverse reactions to medicinal products

None

12.12.3. Additional monitoring

None

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

Action: For adoption of the list

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 hearings - Plan for a 'mock-up' public hearing

Action: For discussion

12.18.2. Safety communication

None

12.19. Continuous pharmacovigilance

12.19.1. Incident management

None

12.20. Others

12.20.1. Strategy on measuring the impact of pharmacovigilance - update

PRAC lead: Marieke De Bruin
Action: For discussion

13. Any other business

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:
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/