17 November 2014
EMA/CHMP/659045/2014
Procedure Management and Business Support Division

Committee for medicinal products for human use (CHMP)

Agenda of meeting to be held on 17-20 November 2014

Chair: Tomas Salmonson – Vice-chair: Pierre Demolis

17 November 2014, 13:00 – 19:30, room 2A

18 November 2014, 08:30 – 19:30, room 2A

19 November 2014, 08:30 – 19:30, room 2A

20 November 2014, 08:30 – 16:00, room 2A

Note on access to documents

Some documents mentioned in the agenda/minutes cannot be released at present following a request for access to documents under 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).

Health & 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 and therefore not disclosed. With regards to therapeutic indications listed against products it must be noted that these may not reflect the full wording proposed by applicants and may also vary during the course of the review. The procedures discussed by the CHMP are on-going and therefore are considered confidential. Additional details on some of these procedures will be published in the CHMP meeting highlights once the procedures are finalised and start of referrals will also be available. For orphan medicinal products the applicant details are published as this information is already publicly available.

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

Further information with relevant explanatory notes can be found at the end of this document.

For adoption

Agenda (EMA/CHMP/659045/2014 rev.4) and Annex to CHMP agenda of the CHMP plenary session to be held 17-20 November 2014	
Timeschedule (EMA/CHMP/686385/2014 rev.3) of the CHMP plenary session to be held 17-20 November 2014	
Minutes (EMA/CHMP/679758/2014 rev.0) of the CHMP plenary session held 20-23 October 2014	
TOD / Minutes (EMA/CHMP/695849/2014) of the ORGAM meeting held on 10 November 2014	

For information

Pre-meeting list of participants and restrictions in relation to declarations of interests applicable to the items of the agenda for the CHMP plenary session to be held 17-20 November 2014	*See November 2014 Minutes (to be published post December 2014 CHMP meeting)*
Draft Agenda of CHMP meeting to be held on 15-18 December 2014.	

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1. Oral explanations

1.1. Pre-authorisation procedure oral explanations

VANTOBRA (EMEA/H/C/002633) (Tobramycin), Applicant: PARI Pharma GmbH, Hybrid application (Article 10(3) of Directive No 2001/83/EC)	Possible Oral explanation to be held on Tuesday 18 November 2014 at 14.00. See also 2.5 Products in the Decision Making Phase
(EMEA/H/C/002396), (safinamide), (treatment of Parkinson's disease (PD)) List of Outstanding Issues adopted on 25.09.2014. List of Questions adopted on 25.04.2014.	Possible Oral explanation to be held on Wednesday 19 November at 9.00.

1.2. Re-examination procedure oral explanation

No items

1.3. Post-authorisation procedure oral explanation

ellaOne (EMEA/H/C/001027/II/0021), (ulipristal acetate), MAH: Laboratoire HRA Pharma, SA, Rapporteur: Pieter de Graeff, Co-Rapporteur: Kristina Dunder, PRAC Rapporteur: Menno van der Elst "The marketing authorisation holder proposed a change in the classification for supply of ellaOne from "medicinal product subject to medical prescription" to " medicinal product not subject to medical prescription" in the EU. Update of the Product information in line with a non-prescription setting is proposed. The MAH also proposed updates of SmPC sections 4.2, 4.4 and 5.1 based on Repeated use study (Protocol 091015-001/CSR HRA2914-554 - SmPC sections 4.4 and 5.1) and on interim data from the STEella study in postmenarcheal girls and adult women (Protocol 2914-010/ EUDRACT nr 2009-017771-21/ CSR HRA 2914-515 - SmPC sections 4.2 and 5.1). Updates to the labelling and package leaflet have been proposed accordingly. Furthermore the MAH proposed this opportunity to bring the PI in line with the QRD template version 8 Rev2." Request for Supplementary Information adopted on 23.01.2014, 21.11.2013, 25.04.2013. Request for 1 year of data exclusivity for a change in classification (Article 74(a) of Directive 2001/83/EC)	Possible Oral explanation to be held on Tuesday 18 November 2014 at 11.00. See also 11. Post-authorisation Issues

1.4. Referral procedure oral explanation

No items

2. New full applications

2.1. New full applications; Opinions

(EMEA/H/C/003724), Orphan, (eliglustat), Applicant: Genzyme Europe BV, (treatment of Gaucher disease type 1) List of Outstanding Issues adopted on 25.09.2014., 24.07.2014. List of Questions adopted on 20.02.2014.	

(EMEA/H/C/003729), (secukinumab), (treatment of plaque psoriasis)
List of Outstanding Issues adopted on 25.09.2014.
List of Questions adopted on 20.03.2014.

- BWP Report: **For adoption**

(EMEA/H/C/003837), (dasabuvir), (treatment of chronic hepatitis C)
List of Questions adopted on 25.09.2014.

(EMEA/H/C/003821), Orphan, (nintedanib), Applicant: Boehringer Ingelheim International GmbH, (treatment of Idiopathic Pulmonary Fibrosis (IPF))
List of Questions adopted on 25.09.2014.

(EMEA/H/C/003746), (apremilast), (treatment of psoriatic arthritis, psoriasis)
List of Outstanding Issues adopted on 25.09.2014.
List of Questions adopted on 25.04.2014.

(EMEA/H/C/002780), (ospemifene), (treatment of vulvar and vaginal atrophy (VVA))
List of Outstanding Issues adopted on 24.07.2014, 20.03.2014.
List of Questions adopted on 25.07.2013.

(EMEA/H/C/003971), (sevelamer), (control of hyperphosphataemia in adult patients receiving haemodialysis or peritoneal dialysis)
List of Questions adopted on 23.10.2014.

(EMEA/H/C/003957), (rasagiline), (treatment of Parkinson's disease)

(EMEA/H/C/003839), (ombitasvir / paritaprevir / ritonavir), (treatment of chronic hepatitis C)
List of Questions adopted on 25.09.2014.

(EMEA/H/C/002814), (vorapaxar), (indicated for the reduction of atherothrombotic events)
List of Outstanding Issues adopted on 25.09.2014.
List of Questions adopted on 25.04.2014.

- List of expert to SAG CVS meeting: **Adopted by written procedure**
- Report from SAG CVS meeting: **For discussion**

2.2. New full applications; Day 180 List of outstanding issues

(EMEA/H/C/004006), (clopidogrel), (prevention of myocardial infarction and acute coronary syndrome
Myocardial infarction, ischaemic stroke, peripheral arterial disease, acute coronary syndrome, prevention of atherothrombotic and thromboembolic events in atrial fibrillation
Myocardial infarction, ischaemic stroke, peripheral arterial disease, acute coronary syndrome, prevention of atherothrombotic and thromboembolic events in atrial fibrillation
Myocardial infarction, ischaemic stroke, peripheral arterial disease, acute coronary syndrome, prevention of atherothrombotic and thromboembolic events in atrial fibrillation.)
Request for Supplementary Information adopted on 25.09.2014.

(EMEA/H/C/002830), Orphan, (mifepristone), Applicant: FGK Representative Service GmbH, (treatment of Cushing's syndrome)
List of Questions adopted on 20.03.2014.

(EMEA/H/C/002788), Orphan, (tolvaptan), Applicant: Otsuka Pharmaceutical Europe Ltd, (treatment of kidney disease (ADPKD)
List of Questions adopted on 25.04.2014.

(EMEA/H/C/003823), (lamivudine / raltegravir), (treatment of human immunodeficiency virus (HIV-1))
List of Questions adopted on 24.07.2014.

(EMEA/H/C/002629), (edoxaban), (prevention of stroke; embolism and treatment of venous thromboembolism)
List of Questions adopted on 26.06.2014.

(EMEA/H/C/003785), (oritavancin), (treatment of complicated skin and soft tissue infections (cSSTI))
List of Questions adopted on 26.06.2014.

(EMEA/H/C/002789), Orphan, (levofloxacin), Applicant: Aptalis Pharma SAS, (indicated for chronic pulmonary infections)
List of Outstanding Issues adopted on 25.09.2014.
List of Questions adopted on 25.04.2014.

(EMEA/H/C/002807), (human fibrinogen / human thrombin), ((human plasma-derived fibrinogen and thrombin) used as an adjunct to haemostasis)
List of Questions adopted on 20.03.2014.

(EMEA/H/C/002739), ((substance to be reviewed) human alpha1-proteinase inhibitor), (treatment of lung tissue)
List of Questions adopted on 25.04.2014.

- BWP Report: **For adoption**

(EMEA/H/C/002846), (tedizolid phosphate), (treatment of tissue infections (cSSTI))
List of Questions adopted on 26.06.2014.

(EMEA/H/C/003737), (voriconazole), (treatment of fungal infections)
List of Questions adopted on 24.07.2014.

2.3. New full applications; Day 120 List of Questions –

(EMEA/H/C/003899), (aripiprazole), (treatment of schizophrenia and prevention of manic episodes in bipolar I disorder)

(EMEA/H/C/003926), (aripiprazole), (treatment of schizophrenia)

(EMEA/H/C/003803), (aripiprazole), (treatment of schizophrenia)

(EMEA/H/C/004008), (aripiprazole), (treatment of schizophrenia, treatment and prevention of bipolar disorder (manic episodes))

(EMEA/H/C/003904), (atazanavir / cobicistat), (treatment of HIV-1 infected, combination with other antiretroviral medicinal products.)

(EMEA/H/W/002300), (p. falciparum circumsporozoite 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), , (indicated for active immunisation against malaria)

- BWP Report: **For adoption**

(EMEA/H/C/002792), Orphan, (susoctocog alfa), Applicant: Baxter AG, (treatment of acquired haemophilia A)

- BWP Report: **For adoption**

(EMEA/H/C/003789), (pegaspargase), (indicated as combination therapy in acute lymphoblastic leukaemia (ALL))

- BWP Report: **For adoption**

(EMEA/H/C/003910), (pegfilgrastim), (treatment of neutropenia)

(EMEA/H/C/003794), Orphan, (asfotase alfa), Applicant: Alexion Europe SAS, (treatment of paediatric-onset hypophosphatasia)

- BWP Report: **For adoption**

(EMEA/H/C/003770), (empagliflozin / metformin hydrochloride), (treatment of type II diabetes)

(EMEA/H/C/002784), (sufentanil), (indicated for the management pain)

2.4. Update on on-going new full applications for Centralised procedure

(EMEA/H/C/003725), Orphan, (panobinostat), Applicant: Novartis Pharmaceuticals UK Limited, (treatment of multiple myeloma)

- Letter from the applicant dated 31 October 2014 requesting an extension of clock stop to submit the responses to the Day 120 List of Questions adopted in September 2014 : **For information**

(EMEA/H/C/003773), (cangrelor), (inhibitor indicated for the reduction of thrombotic cardiovascular events)
List of Outstanding Issues adopted on 25.09.2014.
List of Questions adopted on 25.04.2014.

- List of experts to SAG CVS meeting: For adoption by written procedure

2.5. Products in the Decision Making Phase

VANTOBRA (EMEA/H/C/002633) (Tobramycin), Applicant: PARI Pharma GmbH, Hybrid application (Article 10(3) of Directive No 2001/83/EC)	Possible Oral explanation to be held on Tuesday 18 November 2014 at 14.00. See also 1.1 Pre-authorisation oral explanations

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

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

No items

3.2. Extension of marketing authorisation according to Annex I of Commission Regulation (EC) No 1234/2008; Day 180 List of outstanding issues

Ibandronic acid Accord (EMEA/H/C/002638/X/0006), (ibandronic acid), MAH: Accord Healthcare Ltd, Rapporteur: Alar Irs, "To add a new strength/potency and a new pharmaceutical form 3 mg solution for injection."
List of Questions adopted on 26.06.2014.

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

JETREA (EMEA/H/C/002381/X/0013), (ocriplasmin), MAH: ThromboGenics NV, Rapporteur: Greg Markey, PRAC Rapporteur: Julie Williams, "to introduce a ready-to-use (RTU) formulation with adjusted fill volume for Jetrea 0.375 mg/0.3 mL"

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

No items

4. Type II variations - Extension of indication procedures according to Annex I of Commission Regulation (EC) No 1234/2008

4.1. Type II variation; Extension of indication- Opinions or Requests for supplementary information

Adenuric (EMEA/H/C/000777/II/0037), (febuxostat), MAH: Menarini International Operations Luxembourg S.A., Rapporteur: Andrea Laslop, PRAC Rapporteur: Jan Neuhauser, "Update of sections 4.1, 4.2, 4.4, 4.5, 4.8 and 5.1 of the SmPC for the 120 mg strength further to the introduction of a new indication for prevention and treatment of hyperuricaemia in adult patients undergoing chemotherapy for haematologic malignancies at intermediate to high risk of Tumor Lysis Syndrome (TLS). The Package Leaflet is updated accordingly."
Esmya (EMEA/H/C/002041/II/0028), (ulipristal), MAH: Gedeon Richter Plc., Rapporteur: Kristina Dunder, PRAC Rapporteur: Ulla Wändel Liminga, "Update of section 4.1 of the SmPC with subsequent updates to sections 4.2, 4.4, 4.8 and 5.1 in order to extend the current indication to long term (repeated intermittent) treatment of moderate to severe symptoms of uterine fibroids in adult women of reproductive age. The Package Leaflet is updated accordingly."
InductOs (EMEA/H/C/000408/II/0071), (dibotermin alfa), MAH: Medtronic BioPharma B.V., Rapporteur: Pieter de Graeff, Co-Rapporteur: Janne Komi, PRAC Rapporteur: Menno van der Elst, "Extension of indication to broaden the use of Inductos in interbody lumbar spine fusion." Request for Supplementary Information adopted on 24.07.2014.

Invega (EMEA/H/C/000746/II/0043), (paliperidone), MAH: Janssen-Cilag International N.V., Rapporteur: Filip Josephson, PRAC Rapporteur: Qun-Ying Yue, “Update of sections 4.1 of the SmPC in order to extend the Invega indication to include depressive symptom domain of schizoaffective disorder. Additionally section 5.1 has been updated to reflect the data from the study SCA-3004 on paliperidone palmitate effects in the maintenance of symptom control. Minor editorial changes have been introduced throughout the PI. In addition, the MAH took the opportunity to update the list of local representatives in the Package Leaflet.”	
Relistor (EMEA/H/C/000870/II/0030), (methylnaltrexone bromide), MAH: TMC Pharma Services Ltd, Rapporteur: Harald Enzmann, Co-Rapporteur: Greg Markey, PRAC Rapporteur: Valerie Strassmann, “The MAH applied for an extension of the indication for the treatment of opioid induced constipation in adult non cancer pain patients. Consequently, the MAH proposed the update of sections 4.1, 4.2, 4.4 and 5.1 of the SmPC. The Package Leaflet was proposed to be updated in accordance.” Request for Supplementary Information adopted on 26.06.2014. Request for 1 year of market protection for a new indication (Article 14(11) of Regulation (EC) 726/2004)	

Rienso (EMEA/H/C/002215/II/0008), (ferumoxytol), MAH: Takeda Pharma A/S, Rapporteur: Harald Enzmann, Co-Rapporteur: Romaldas Mačiulaitis, PRAC Rapporteur: Martin Huber, "Extension of indication: all cause iron deficiency anaemia when oral therapy is ineffective or inappropriate or where there is a need for rapid iron repletion
As a consequence, sections 4.1, 4.2 , 4.4, 4.8 and 5.1 of the SmPC were proposed to be updated. The Package Leaflet was proposed to be updated accordingly. The MAH took the opportunity to propose minor editorial changes to the SmPC and to propose the update of the Product Information in line with the latest version of the QRD template (9.0)"
Request for Supplementary Information adopted on 26.06.2014, 25.04.2014, 24.10.2013.

Soliris (EMEA/H/C/000791/II/0066), Orphan, (eculizumab), MAH: Alexion Europe SAS, Rapporteur: Arantxa Sancho-Lopez, Co-Rapporteur: Pierre Demolis, "Update of sections 4.1 and 5.1 of the SmPC with an extension of the indication in patients with Paroxysmal nocturnal haemoglobinuria (PNH) regardless of their history of transfusion. The PL has been updated accordingly. In addition, some minor corrections are proposed in Section 5.1 of the SmPC and in the PL."
Request for Supplementary Information adopted on 24.07.2014.

Tracleer (EMEA/H/C/000401/II/0066), (bosentan), MAH: Actelion Registration Ltd., Rapporteur: Pierre Demolis, Co-Rapporteur: Kristina Dunder, PRAC Rapporteur: Arnaud Batz, "Extension of Indication to include treatment of symptomatic pulmonary arterial hypertension in paediatric patients aged from 3 months to 18 years. The SmPC has been updated in order to include the data generated in studies conducted according to the agreed Paediatric Investigationa Plan for bosentan (EMEA-000425-PIP02-10-M04). As a consequence, sections 4.1, 4.2, 4.5, 4.8, 5.1, 5.2, 5.3 and 6.6 of the SmPC have been updated. The Package Leaflet has been updated accordingly. In addition, taking into account the new data in the paediatric population, an updated version of the RMP (version 5) has been provided."
Request for Supplementary Information adopted on 24.07.2014.

Travatan (EMEA/H/C/000390/II/0046), (travoprost), MAH: Alcon Laboratories (UK) Ltd, Rapporteur: Concepcion Prieto Yerro, Co-Rapporteur: Greg Markey, PRAC Rapporteur: Dolores Montero Corominas, "Extension of the therapeutic indication for decrease of elevated intraocular pressure in paediatric patients with ocular hypertension or paediatric glaucoma."
Request for Supplementary Information adopted on 25.09.2014.

4.2. Update on on-going type II variation; extension of indications

No items

5. Ancillary medicinal substances in medical devices

5.1. Ancillary medicinal substances in medical devices - Opinions / List of outstanding issues / List of Questions

No items

6. Re-examination procedure (new applications) under Article 9(2) of Regulation no 726/2004

No items

7. Re-examination procedure (Type II variations) under Article Article 16 of Commission Regulation (EC) No 1234/2008 and 9(2) of Regulation (EC) No 726/2004

No items

8. Withdrawal of full new application

Egranli (EMEA/H/C/002637), (balugrastim), Applicant: Teva Pharma B.V., (treatment of neutropenia)
New active substance (Article 8(3) of Directive No 2001/83/EC). Positive Opinion adopted on 25 September 2014.

- Letter from the applicant dated 6 November 2014 informing of a decision to withdraw the Marketing Authorisation Application: **For information**

9. Procedure under Article 83(1) of Regulation (EC) 726/2004 (Compassionate Use)

No items

10. Pre-submission issues

Some items in this section are considered commercially confidential or sensitive and therefore not disclosed.

(H0003882) (Alirocumab), (Alirocumab is indicated, as adjunct therapy to diet, for long-term treatment of adult patients with primary hypercholesterolaemia (non-familial and heterozygous familial) or mixed dyslipidaemia to reduce low-density lipoprotein cholesterol (LDL-C).
Combination therapy with a statin
Alirocumab is indicated in combination with a statin (HMG-CoA reductase inhibitor), with or without other lipid modifying therapy (LMT), in patients not appropriately controlled with a statin alone.
Monotherapy
Alirocumab is indicated as monotherapy, or as add-on to other non-statin LMT, in patients who cannot tolerate statins),

- Request for accelerated assessment dated 2 October 2014: **For discussion**
- Rapporteurs' accelerated assessment briefing note: **For adoption**

(H0004062) (Sacubitril/Valsartan), (indicated for the treatment of heart failure (NYHA class II-IV) to reduce the rate of cardiovascular death and heart failure hospitalization. TRADENAME was also shown to reduce all-cause mortality).

- Request for accelerated assessment dated 31 October 2014: **For discussion**
- Rapporteurs' accelerated assessment briefing note: **For adoption**

(H0004004), Orphan
(Sebelipase Alfa), Applicant: Synageva BioPharma Ltd, (indicated for long-term enzyme replacement therapy (ERT) for patients with Lysosomal Acid Lipase Deficiency (LAL D).),

- Request for accelerated assessment dated 3 November 2014: **For discussion**
- Rapporteurs' accelerated assessment briefing note: **For adoption**

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11. Post-authorisation issues

Some items in this section are considered commercially confidential or sensitive and therefore not disclosed.

ellaOne (EMEA/H/C/001027/II/0021), (ulipristal acetate), MAH: Laboratoire HRA Pharma, SA, Rapporteur: Pieter de Graeff, Co-Rapporteur: Kristina Dunder, PRAC Rapporteur: Menno van der Elst "The marketing authorisation holder proposed a change in the classification for supply of ellaOne from "medicinal product subject to medical prescription" to " medicinal product not subject to medical prescription" in the EU. Update of the Product information in line with a non-prescription setting is proposed. The MAH also proposed updates of SmPC sections 4.2, 4.4 and 5.1 based on Repeated use study (Protocol 091015-001/CSR HRA2914-554 - SmPC sections 4.4 and 5.1) and on interim data from the STEella study in postmenarcheal girls and adult women (Protocol 2914-010/ EUDRACT nr 2009-017771-21/ CSR HRA 2914-515 - SmPC sections 4.2 and 5.1). Updates to the labelling and package leaflet have been proposed accordingly. Furthermore the MAH proposed this opportunity to bring the PI in line with the QRD template version 8 Rev2 ." Request for Supplementary Information adopted on 23.01.2014, 21.11.2013, 25.04.2013. Request for 1 year of data exclusivity for a change in classification (Article 74(a) of Directive 2001/83/EC) • Opinion: **For adoption**	Possible Oral explanation to be held on Tuesday 18 November 2014 at 11.00. See also 1.3 Post-authorisation procedure oral explanation

Ferriprox (EMEA/H/C/000236/II/0089/G), (deferiprone), MAH: Apotex Europe BV, Rapporteur: Pierre Demolis, PRAC Rapporteur: Isabelle Robine, "Update of section 4.5 of the SmPC regarding combination of deferiprone with other iron chelators further to request of the PRAC in the assessment of the PSUR (PSUV/083). Update of section 5.1 of the SmPC and the RMP with the results of Study LA37-111 conducted to evaluate the effect of deferiprone on cardiac QT and QTC interval duration. The Package leaflet is updated accordingly. The MAH also takes the opportunity to align the product information with QRD template (version 9) and to make minor editorial corrections. The package leaflet is also updated to add the local representatives in Croatia."
Request for Supplementary Information adopted on 24.07.2014.

- Request for Supplementary information or Opinion: **For adoption**
- Letter from patient and UK thalassaemia patients' representative on use of deferiprone administered in combination with other iron chelators: **For information**

Brintellix (EMEA/H/C/002717/II/0004), (vortioxetine), MAH: H. Lundbeck A/S, Rapporteur: Bart Van der Schueren "Update of section 5.1 of the SmPC with information on the effect of vortioxetine on cognitive dysfunction in Major Depressive Disorder."

- Request for Supplementary information: **For adoption**

Herceptin (EMEA/H/C/000278/II/0084/G), (trastuzumab), MAH: Roche Registration Ltd, Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Brigitte Keller-Stanislawski, "Update of sections 4.2 and 4.8 of the SmPC with information on switching between intravenous (IV) and subcutaneous (SC) formulations further to safety data from study MO22982. The Package Leaflet is updated accordingly. Update of section 4.2 with a statement regarding switching between Herceptin and biosimilars. In addition, the MAH took the opportunity make corrections to the SmPC and Package leaflet." • Request for Supplementary information: **For adoption**	
Revlimid (EMEA/H/C/000717/II/0076), Orphan, (lenalidomide), MAH: Celgene Europe Limited, Rapporteur: Pierre Demolis "Update of section 4.4 of the SmPC with a new warning regarding an increased risk of mortality with the use of Revlimid in patients with chronic lymphocytic leukemia (CLL). The package leaflet is updated accordingly." Request for Supplementary Information adopted on 26.06.2014. • Request for Supplementary information or Opinion: **For adoption**	
Vectibix (EMEA/H/C/000741/R/0064), (panitumumab), MAH: Amgen Europe B.V., Rapporteur: Robert James Hemmings, PRAC Rapporteur: Julie Williams, • Opinion: **For adoption**	Renewal procedure.

Hexacima (EMEA/H/C/002702/ PSUV 011)

Hexyon (EMEA/H/C/002796/ PSUV 013)

Hexaxim (EMEA/H/W/002495/ PSUV 019)

(Diphtheria Toxoid,Filamentous Haemagglutinin,Hepatitis B Surface Antigen,Pertussis Toxoid,Tetanus Toxoid,Haemophilus Influenzae Type B Polysaccharide,Polyribosylribitol Phosphate Conjugated To Tetanus Protein,Type 1 (Mahoney),Type 2 (Mef-1),Type 3 (Saukett)), MAH: Sanofi Pasteur MSD SNC, Rapporteur: Jan Mueller-Berghaus, Co-Rapporteur: Daniel Brasseur, (treatment of diphtheria, tetanus, pertussis, hepatitis B, poliomyelitis and invasive diseases caused by Haemophilus influenzae type b.)

- Outcome of PRAC discussion : **For discussion**

12. Referral procedures

12.1. Procedure for Centrally Authorised products under Article 20 Council Regulation (EC) No 726/2004

Procoralan (EMEA/H/C/000597)
Corlentor (EMEA/H/C/000598)
Procedure number: EMEA/H/A20/1404/C/000597-598/0031-0032
(ivabradine hydrochloride), MAH: Les Laboratoires Servier, Rapporteur: Pieter de Graeff, Co-Rapporteur: Janne Komi,
Article 20 procedure triggered by the EC to assess how the results of the SIGNIFY study, which showed a statistically significant increase in a composite endpoint of cardiovascular death and non-fatal MI in a pre-defined subgroup of symptomatic angina patients, impact on the benefit-risk balance of Corlentor and Procoralan.

- PRAC recommendation: **For discussion**
- Opinion: **For adoption**

12.2. Requests for CHMP Opinion under Article 5(3) and 57 (1)p of Regulation (EC) No 726/2004

Medicinal products under development for

the treatment of Ebola (EMEA/H/A-5(3)/1410)

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

No items

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

Seasonique film coated tablets (EMEA/H/A-29(4)/1392)
(levonogestrel 150 µg and ethinylestradiol 30 µg / 10 µg), MAH: Teva Pharma B.V (NL), Rapporteur: Joseph Emmerich, Co-Rapporteur: Martina Weise, , RMS: FR, CMS: AT, BE, DE, IT, PL, RO, SI ,SK, Procedure number: FR/H/0516/001/DC,
Article 29(4) triggered due to disagreement with regard to the demonstration of contraceptive effectiveness and treatment compliance. Opinion adopted on 26.06.2014.

Nasonex (EMEA/H/A-30/1374)
(mometasone), nasal spray suspension, MAH: Merck Sharp & Dohme, Rapporteur: Kristina Dunder, Co-Rapporteur: David Lyons,
Nasonex was included in the list of products for SmPC harmonisation, drawn up by the CMDh, in accordance with Article 30(2) of Directive 2001/83/EC. List of Questions adopted on 19.09.2013. List of Outstanding Issues adopted on23.10.2014, 25.09.2014, 20.02.2014.

- Opinion: **For adoption**

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

No items

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

GVK Biosciences (EMEA/H/A-31/1408)
Rapporteur: Harald Enzmann, Co-Rapporteur: Christian Schneider,

Article 31 procedure triggered by the European Commission concerning GVK Biosciences Private Limited (GVK Bio), Swarna Jayanthi commercial complex, Ameerpet, Hyderabad 500 038, India

- List of outstanding issues or Opinion: **For adoption**

Gadolinium containing contrast agents, Gd-Cas (EMEA/H/A-31/1097),
Rapporteur: Rafe Suvarna, Co-Rapporteur: Pieter de Graeff,
FUM related to the updated 3rd annual cumulative safety reviews on nephrogenic systemic fibrosis (NSF) with regards to Omniscan: GE HealthCare

- Assessment report: **For adoption**

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

No items

12.8. Procedure under Article 107(2) of Directive 2001/83/EC

No items

12.9. Disagreement between Member States on Type II variation– Arbitration procedure initiated by MAH under Article 6(13) (EC) No 1084/2003)

No items

12.10. Procedure under Article 29 Regulation (EC) 1901/2006

No items

12.11. Referral under Article 13 Disagreement between Member States on Type II variation– Arbitration procedure initiated by Member State under Article 13 (EC) No 1234/2008)

No items

13. Pharmacovigilance issues

Summary of recommendations and advice of PRAC meeting held on 3-6 November 2014: **For information**

List of Union Reference Dates and frequency of submission of Periodic Safety Update Reports (EURD list) for November 2014: **For adoption**

Early Notification System:

November 2014 Early Notification System on Envisaged CHMP Recommendations for Regulatory Action (based on Identified Safety Concerns) Accompanied by Communication to the General Public: **for information**

Signal detection

PRAC recommendations on signals adopted at the 3-6 November 2014 meeting: **For adoption**

TECFIDERA (EMEA/H/C/002601)
(Dimethyl Fumarate), MAH: Biogen Idec Ltd, Rapporteur: Martina Weise, Co-Rapporteur: Robert James Hemmings, (treatment of multiple sclerosis), Known active substance (Article 8(3) of Directive No 2001/83/EC)

Signal of Progressive multifocal leukoencephalopathy (PML)

- DHPC: **For information**

14. Inspections

14.1. GMP inspections

Request for GMP inspections: **For adoption**	*Disclosure of information related to GMP inspections will not be published as it undermines the purpose of such inspections.*

14.2. GCP inspections

Request for GCP inspections: **For adoption**	*Disclosure of information related to GCP inspections will not be published as it undermines the purpose of such inspections.*

14.3. Pharmacovigilance inspections

Request for Pharmacovigilance inspections: **For adoption**	*Disclosure of information related to Pharmacovigilance inspections will not be published as it undermines the purpose of such inspections.*

14.4. GLP inspections

Request for GLP inspections: **For adoption**	*Disclosure of information related to GLP inspections will not be published as it undermines the purpose of such inspections.*

15. Innovation Task Force

15.1. Minutes of Innovation Task Force: For information

15.2. Briefing meetings (Innovation Task Force)

Disclosure of information related to briefing meetings taking place with applicants cannot be released at present time as deemed to contain commercially confidential information.

15.3. Eligibility to EMA scientific services

No items

15.4. Requests for CHMP Opinion under Article 57 (1)P of Regulation (EC) No 726/2004

No items

15.5. Nanomedicine activities

No items

16. Scientific Advice Working Party (SAWP)

Report from the SAWP meeting held on 3-6 November 2014. Table of conclusions: **For information**	
Scientific advice letters:	*Disclosure of information related to scientific advice letters cannot be released at present time as these contain commercially confidential information.*

17. Satellite Groups

17.1. Coordination Group for Mutual Recognition and Decentralised Procedures

Report from the Coordination Group for Mutual Recognition and Decentralised Procedures – Human (CMDh) on the meeting held on 17-19	

November 2014: **For information**

18. Other Committees

18.1. Committee for Orphan Medicinal Products (COMP)

Press release of the COMP meeting held on 12-23 November 2014: **For information**

18.2. Committee for Herbal Medicinal Products (HMPC)

Not applicable this month.

18.3. Paediatric Committee (PDCO)

PIPs reaching D30 November 2014 PDCO: **For information**

Report from the PDCO meeting held on 12-14 November 2014: **For information**

18.4. Committee for Advanced Therapies (CAT)

Table of Decisions of CAT meeting held on 13-14 November 2014: **For information**

19. Invented name issues

Table of Decisions of the NRG meeting held on: **For adoption**

20. Any other business

Initiative on registries: tools to improve the quality, consistency and utility of data and information from patient registries: **For discussion**

Presentations given at the CHMP, CAT, COMP joint Presidency meeting, 29 – 30 October 2014 in Rome: **For information**

Participation of the centralised procedure in the IGDRP-Information sharing pilot The centralised procedure will participate in the "IGDRP-Information sharing pilot". Under this pilot all assessment reports (from D80 to D210) will be shared with participating non-EEA agencies, for selected applications of generic medicinal products. The anticipated start date is January 2015, first assessment reports are expected to be shared by the agency from May 2015.	
Questions and answers on propylene glycol in the context of the revision of the guideline on 'Excipients in the label and package leaflet of medicinal products for human use' (EMA/CHMP/704195/2013): **For adoption for 3-month public consultation** Draft background report (EMA/CHMP/334655/2013): **For information**	
Questions and answers on cyclodextrins in the context of the revision of the guideline on 'Excipients in the label and package leaflet of medicinal products for human use' (EMA/CHMP/495747/2013): **For adoption for 3-month public consultation** Draft background report (EMA/CHMP/333892/2013): **For information**	
CHMP Safety Working Party's response to the CHMP query regarding the contra-indication for pregnancy is justifiable on the basis of non-clinical data of apremilast (EMA/CHMP/SWP/680234/2014): **For adoption**	
Guideline on similar biological medicinal products containing biotechnology-derived proteins as active substance: non-clinical and clinical issues (Doc Ref: EMEA/CHMP/BMWP/42832/2005 Rev1): **For adoption**	
Guideline on the pharmacokinetic and clinical evaluation of modified release dosage forms - EMA/CHMP/EWP/280/96 Corr1: **For adoption**	
Update to Committees about changes to the MAA process: **For information**	See also topic below

Draft Guidance with applicants on responses to questions received from EMA Scientific Committees during the evaluation within the centralised procedure: **For comments by 5 December 2014**
Adjustment of timetable for procedures in phase outcome in December 2014/January 2015

List of participants

List of participants including any restrictions with respect to involvement of members / alternates / experts following evaluation of declared interests for the 17-20 November 2014 meeting.

Explanatory notes

The notes below give a brief explanation of the main sections and headings in the CHMP agenda and should be read in conjunction with the agenda or the minutes.

Oral explanations *(section 1)*

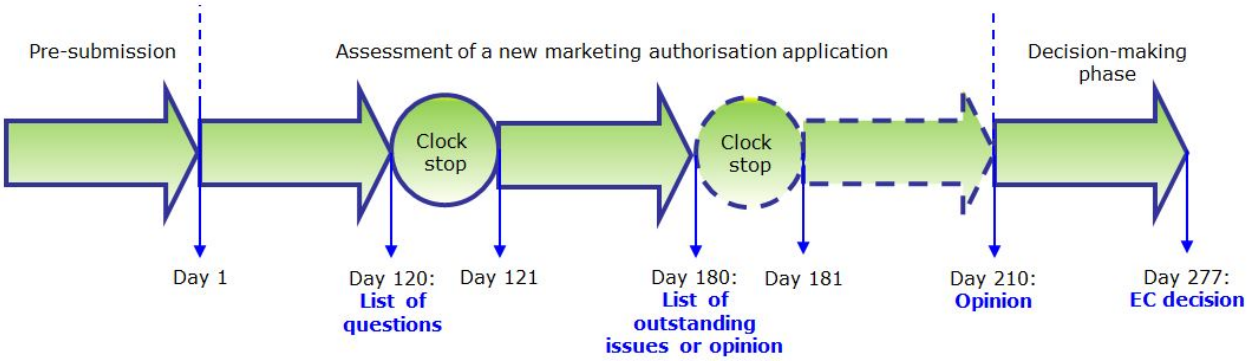
The items listed in this section are those for which marketing authorisation holders (MAHs) or applicants have been invited to the CHMP plenary meeting to address questions raised by the Committee. Oral explanations normally relate to on-going applications (section 2 and 3) or referral procedures (section 12) but can relate to any other issue for which the CHMP would like to discuss with company representatives in person.

New applications *(section 2)*

This section lists applications for marketing authorisations of new medicines that are to be discussed by the Committee.

Section 2.1 is for medicinal products nearing the end of the evaluation and for which the CHMP is expected to adopt an **opinion** at this meeting on whether marketing authorisation should be granted. Once adopted, the CHMP opinion will be forwarded to the European Commission for a final legally binding decision valid throughout the EU.

The other items in the section are listed depending on the stage of the evaluation, which is shown graphically below:



The assessment of an application for a new medicine takes up to 210 'active' days. This active evaluation time is interrupted by at least one 'clock-stop' during which time the applicant prepares the answers to questions from the CHMP. The clock stop happens after day 120 and may also happen after day 180, when the CHMP has adopted a list of questions or outstanding issues to be addressed by the company. Related discussions are listed in the agenda under sections 2.2 (**Day 180 List of outstanding issues**) and 2.3 (**Day 120 list of questions**).

CHMP discussions may also occur at any other stage of the evaluation, and these are listed under section 2.4, **update on ongoing new applications for centralised procedures**.

The assessment leads to an opinion from the CHMP by day 210. Following a CHMP opinion the European Commission takes usually 67 days to issue a legally binding decision (i.e. by day 277 of the procedure). CHMP discussions on products that have received a CHMP opinion and are awaiting a decision are listed under section 2.5, **products in the decision making phase**.

Extension of marketing authorisations according to Annex I of Reg. 1234/2008 *(section 3)*

Extensions of marketing authorisations are applications for the change or addition of new strengths, formulations or routes of administration to existing marketing authorisations. Extension applications follow a 210-day evaluation process, similarly to applications for new medicines (see figure above).

Type II variations - Extension of indication procedures *(section 4)*

Type II variations are applications for a change to the marketing authorisation which requires an update of the product information and which is not covered in section 3. Type II variations include applications for a new use of the medicine (extension of indication), for which the assessment takes up to 90 days. For the applications listed in this section, the CHMP may adopt an opinion or request supplementary information from the applicant.

Ancillary medicinal substances in medical devices *(section 5)*

Although the EMA does not regulate medical devices it can be asked by the relevant authorities (the so-called Notified Bodies) that are responsible for regulating these devices to give a scientific opinion on a medicinal substance contained in a medical device.

Re-examination procedures (new applications) under article 9(2) of regulation no 726/2004 *(section 6)*

This section lists applications for new marketing authorisation for which the applicant has requested a re-examination of the opinion previously issued by the CHMP.

Re-examination procedures *(section 7)*

This section lists applications for type II variations (including extension of indication applications) for which the applicant has requested re-examination of the opinion previously issued by the CHMP.

Withdrawal of application *(section 8)*

Applicants may decide to withdraw applications at any stage during the assessment and a CHMP opinion will therefore not be issued. Withdrawals are included in the agenda for information or discussion, as necessary.

Procedure under article 83(1) of regulation (EC) 726/2004 (compassionate use) *(section 9)*

Compassionate use is a way of making available to patients with an unmet medical need a promising medicine which has not yet been authorised (licensed) for their condition. Upon request, the CHMP provides recommendations to all EU Member States on how to administer, distribute and use certain medicines for compassionate use.

Pre-submission issues *(section 10)*

In some cases the CHMP may discuss a medicine before a formal application for marketing authorisation is submitted. These cases generally refer to requests for an accelerated assessment for medicines that are of major interest for public health or can be considered a therapeutic innovation. In case of an accelerated assessment the assessment timetable is reduced from 210 to 150 days.

Post-authorisation issues *(section 11)*

This section lists other issues concerning authorised medicines that are not covered elsewhere in the agenda. Issues include supply shortages, quality defects, some annual reassessments or renewals or type II variations to marketing authorisations that would require specific discussion at the plenary.

Referral procedures *(section 12)*

This section lists referrals that are ongoing or due to be started at the plenary meeting. A referral is a procedure used to resolve issues such as concerns over the safety or benefit-risk balance of a medicine or a class of medicines. In a referral, the EMA is requested to conduct a scientific assessment of a particular medicine or class of medicines on behalf of the EU. Further information on such procedures can be found here.

Pharmacovigilance issues *(section 13)*

This section lists issues that have been discussed at the previous meeting of the PRAC, the EMA's committee responsible for evaluating and monitoring safety issues for medicines. Feedback is provided by the PRAC. This section also refers to the early notification system, a system used to notify the European regulatory network on proposed EMA communication on safety of medicines.

Inspections Issues *(section 14)*

This section lists inspections that are undertaken for some medicinal products. Inspections are carried out by regulatory agencies to ensure that marketing authorisation holders comply with their obligations. Inspection can relate to good manufacturing practice (GMP), good clinical practice (GCP), good laboratory practice (GLP) or good pharmacovigilance practice (GVP).

Innovation task force *(section 15)*

The Innovation Task Force (ITF) is a body set up to encourage early dialogue with applicants developing innovative medicines. Minutes from the last ITF meeting as well as any related issue that requires discussion with the CHMP are listed in this section of the agenda. Further information on the ITF can be found here.

Scientific advice working party (SAWP) *(section 16)*

This section refers to the monthly report from the CHMP's Scientific Advice Working Party (SAWP) on scientific advice given to companies during the development of medicines. Further general information on SAWP can be found here.

Satellite groups / other committees *(section 17)*

This section refers to the reports from groups and committees making decisions relating to human medicines: the Coordination Group for Mutual Recognition and Decentralised Procedures – Human (CMDh), the Committee for Orphan Medicinal Products (COMP), the Committee for Herbal Medicinal Products (HMPC), Paediatric Committee (PDCO), the Committee for Advanced Therapies (CAT) and the Pharmamacovigilance Risk Assessment Committee (PRAC).

Invented name issues *(section 18)*

This section list issues related to invented names proposed by applicants for new medicines. The CHMP has established the Name Review Group (NRG) to perform reviews of the invented names. The group's main role is to consider whether the proposed names could create a public-health concern or potential safety risk. Further information can be found here.