27 January 2015
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Procedure Management and Business Support Division

Committee for medicinal products for human use (CHMP)

Minutes of the meeting held on 15-18 December 2014

Chair: Tomas Salmonson – Vice-chair: Pierre Demolis

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).

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. 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.

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Agenda (EMA/CHMP/722409/2014 rev.3) and Annex to CHMP agenda of the CHMP plenary session to be held on 15-18 December 2014	The agenda and annex were adopted with amendments.
Timeschedule (EMA/CHMP/754937/2014 rev.3) of the CHMP plenary session to be held on 15-18 December 2014	The timeschedule was adopted.
Minutes (EMA/741182/2014) of the CHMP plenary session held on 17-20 November 2014	The Minutes of the CHMP plenary session held 17 – 20 November 2014 were adopted.
ToD / Minutes (EMA/764768/2014) of the ORGAM meeting held on 8 December 2014	The Minutes of the December 2014 CHMP ORGAM meeting held on 08 December 2014, together with all decisions taken at that meeting, were adopted.
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 on 15-18 December 2014.	See December 2014 Minutes (to be published post January 2015 CHMP meeting). The pre-meeting list was noted.
Draft Agenda of CHMP meeting to be held on 19-22 January 2015.	The draft agenda was noted.

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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)
An Oral explanation was held on Wednesday 17 December 2014 at 14.00.
See also 2.5 Products in the Decision Making Phase

(EMEA/H/C/003780), (liraglutide), (treatment of obesity)
List of Outstanding Issues adopted on 23.10.2014. List of Questions adopted on 22.05.2014.
The members were reminded of the main outstanding issues.
An Oral explanation was held on Tuesday 16 December 2014 at 11.00, mainly focusing on the wording of the indication as well as the safety profile of the drug.
After the oral explanation the members discussed whether the results from an ongoing liraglutide study could be extrapolated to the scenario.
Furthermore the Committee discussed the SmPC wording.

(EMEA/H/C/002066), (ciclosporin), (treatment of keratitis)
List of Outstanding Issues adopted on 25.09.2014. List of Questions adopted on 25.04.2014.
The CHMP noted the report from the ad-hoc expert group.
An Oral explanation was held on Wednesday 17 December 2014 at 9.00 focusing on the endpoints from the pivotal study with specific regard to the clinical effect on signs and symptoms.
The members discussed the available clinical data and the relevance of the primary endpoints and secondary endpoints.

1.2. Re-examination procedure oral explanation

No items

1.3. Post-authorisation procedure oral explanation

No items

1.4. Referral procedure oral explanation

No items

2. Initial full applications

2.1. Initial full applications; Opinions

Clopidogrel-ratiopharm (EMEA/H/C/004006), (clopidogrel), Applicant: Teva Pharma B.V., Generic of Plavix, Duplicate of Clopidogrel Teva,, (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.)
Generic application (Article 10(1) of Directive No 2001/83/EC)
List of Outstanding Issues adopted on 20.11.2014. List of Questions adopted on 25.09.2014.
The Committee confirmed that all issues previously identified in this application had been addressed.
The Committee adopted a positive Opinion recommending the granting of a marketing authorisation by consensus together with the CHMP assessment report and translation timetable.
The Icelandic and Norwegian Members were in agreement with the CHMP recommendation.
The legal status was agreed as medicinal product subject to medical prescription.
The summary of opinion was circulated for information.

Holoclar (EMEA/H/C/002450), Orphan, ATMP, (ex vivo expanded autologous corneal epithelial cells including stem cells), Applicant: Chiesi Farmaceutici S.p.A., (treatment of limbal stem cell deficiency)
New active substance (Article 8(3) of Directive No 2001/83/EC)
List of Outstanding Issues adopted on 23.10.2014. List of Questions adopted on 25.07.2013.
The CHMP was informed about the CAT opinion taken at their December 2014 Plenary meeting.
The CHMP adopted a positive opinion recommending the granting of a conditional marketing authorisation by consensus based on the CAT opinion. Furthermore the assessment report and translation timetable were adopted.
Furthermore, the CHMP considered that ex-vivo expanded autologous human corneal epithelial cells containing stem cells is a new active substance, as claimed by the applicant.
The Icelandic and Norwegian Members were in agreement with the CHMP recommendation.
The legal status was agreed as medicinal product subject to restricted medical prescription.
The summary of opinion was circulated for information.
The CHMP agreed to the wording of the EMA press release.
The Committee adopted the BWP report.

Mysimba (EMEA/H/C/003687), (naltrexone / bupropion), Applicant: Orexigen Therapeutics Ireland Limited, (indicated for the management of obesity)
Fixed combination application (Article 10b of Directive No 2001/83/EC)
List of Outstanding Issues adopted on 23.10.2014, 24.07.2014. List of Questions adopted on 20.02.2014.
The members discussed the wording of the indication as well as the safety profile of the drug and the design of an ongoing clinical trial .
The Committee confirmed that all issues previously identified in this application had been addressed.
The Committee adopted a positive opinion by majority (31 positive out of 33 votes) recommending the granting of a marketing authorisation by consensus together with the CHMP assessment report and translation timetable.
The Icelandic and Norwegian Members were in agreement with the CHMP recommendation.
The divergent position (David Lyons, Joseph Emmerich) was appended to the opinion.
The legal status was agreed as medicinal product subject to medical prescription.
The Committee noted the letter of recommendation dated 16 December 2014.
The CHMP agreed to the wording of the EMA press release.
The summary of opinion was circulated for information.

Quinsair (EMEA/H/C/002789), Orphan, (levofloxacin), Applicant: Aptalis Pharma SAS, (indicated for chronic pulmonary infections)
Known active substance (Article 8(3) of Directive No 2001/83/EC)
List of Outstanding Issues adopted on 20.11.2014, 25.09.2014. List of Questions adopted on 25.04.2014.
The Committee confirmed that all issues previously identified in this application had been addressed.
The CHMP noted the letter from the PDCO regarding the impact of adolescents' data submitted within the marketing authorisation application for Quinsair on the PIP.
The Committee adopted a positive opinion recommending the granting of a marketing authorisation by consensus together with the CHMP assessment report and translation timetable.
The Icelandic and Norwegian Members were in agreement with the CHMP recommendation.
The legal status was agreed as medicinal product subject to medical prescription.
The summary of opinion was circulated for information.

Tasermity (EMEA/H/C/003968), (sevelamer hydrochloride), Applicant: Genzyme Europe BV,, (indicated for control of hyperphosphataemia)
Informed consent application (Article 10c of Directive No 2001/83/EC)
List of Questions adopted on 25.09.2014.
The Committee confirmed that all issues previously identified in this application had been addressed.
The Committee adopted a positive opinion recommending the granting of a marketing authorisation by consensus together with the CHMP assessment report and translation timetable.
The Icelandic and Norwegian Members were in agreement with the CHMP recommendation.
The legal status was agreed as medicinal product subject to medical prescription.
The summary of opinion was circulated for information.

Xadago (EMEA/H/C/002396), (safinamide), Applicant: Zambon SpA (treatment of Parkinson's disease (PD))
New active substance (Article 8(3) of Directive No 2001/83/EC)
List of Outstanding Issues adopted on 25.09.2014. List of Questions adopted on 25.04.2014.
The Committee confirmed that all issues previously identified in this application had been addressed.
The Committee adopted a positive opinion recommending the granting of a marketing authorisation by consensus together with the CHMP assessment report and translation timetable.
Furthermore, the CHMP considered that safinamide is a new active substance, as claimed by the applicant.
The Icelandic and Norwegian Members were in agreement with the CHMP recommendation.
The legal status was agreed as medicinal product subject to medical prescription.
The Committee noted the letter of recommendation dated 17 December 2014.
The summary of opinion was circulated for information.

Xydalba (EMEA/H/C/002840), (dalbavancin), Applicant: Durata Therapeutics International B.V., (treatment of acute bacterial skin and skin structure infections (ABSSSI))
New active substance (Article 8(3) of Directive No 2001/83/EC)
List of Outstanding Issues adopted on 25.09.2014. List of Questions adopted on 25.04.2014.
The Committee confirmed that all issues previously identified in this application had been addressed.
The Committee adopted a positive opinion recommending the granting of a marketing authorisation by consensus together with the CHMP assessment report and translation timetable.
Furthermore, the CHMP considered that Dalbavancin is a new active substance, as claimed by the applicant.

The Icelandic and Norwegian Members were in agreement with the CHMP recommendation.
The legal status was agreed as medicinal product subject to medical prescription.
The summary of opinion was circulated for information.

Senshio (EMEA/H/C/002780) (Ospemifene), Applicant: Shionogi Limited, New active substance (Article 8(3) of Directive No 2001/83/EC)
Administrative changes had been made after the November 2014 CHMP meeting.
The CHMP noted the final opinion documents.

2.2. Initial full applications; Day 180 List of outstanding issues –

(EMEA/H/C/002772), Orphan, (dasiprotimut-t), Applicant: Biovest Europe Ltd, (treatment of non-Hodgkin's lymphoma (FL))
List of Questions adopted on 25.04.2014.
The Committee was reminded of the status of this application and its remaining outstanding issues,
The CHMP adopted a List of Outstanding Issues.
The Committee agreed to the request by the applicant for an extension of the clock stop.
The Committee adopted the BWP report.

(EMEA/H/C/003852), (human papillomavirus vaccine [types 6, 11, 16, 18, 31, 33, 45, 52, 58] (recombinant, adsorbed)), (treatment of HPV diseases)
List of Questions adopted on 24.07.2014.
The Committee was reminded of the status of this application and its remaining outstanding issues.
The CHMP adopted a List of Outstanding Issues with a specific timetable.
The Committee adopted the BWP report.

(EMEA/H/C/003750), Orphan, ATMP, (allogenic human heterologous liver cells), Applicant: Cytonet GmbH&Co KG, (treatment of urea cycle disorders (UCD))
List of Questions adopted on 25.04.2014.
The Committee was reminded of the status of this application and its remaining outstanding issues and the discussion at the CAT Plenary.
The CHMP agreed to the List of Outstanding Issues adopted by the CAT at their December 2014 meeting.
The Committee noted that the CAT had agreed to the request from the applicant for an extension of clock stop for the submission of responses and adopted a specific timetable.
The Committee adopted the BWP report.

(EMEA/H/C/003819), (ceritinib), (treatment of non-small cell lung cancer (NSCLC)
treatment of anaplastic lymphoma kinase (ALK)-positive locally advanced or metastatic non-small cell lung cancer (NSCLC))
List of Questions adopted on 24.07.2014.
The Committee was reminded of the status of this application and its remaining outstanding issues.
The CHMP adopted a List of Outstanding Issues with a specific timetable.

(EMEA/H/C/002807), (human fibrinogen / human thrombin), Supportive treatment where standard surgical techniques are insufficient for improvement of haemostasis)
List of Questions adopted on 20.03.2014.
Minor changes have been made to the List of Outstanding Issues after the November Plenary meeting. The CHMP noted the final List of Outstanding Issues.

2.3. Initial full applications; Day 120 List of Questions

(EMEA/H/C/004021), (aripiprazole), (treatment of schizophrenia and treatment and prevention of manic episodes in bipolar I disorder)
The Committee noted the issues identified in this application The Committee adopted the CHMP recommendation and scientific discussion together with the List of Questions

(EMEA/H/C/002734), Orphan, (isavuconazole), Applicant: Basilea Medical Ltd, (treatment of aspergillosis and mucormycosis)
The Committee discussed the issues identified in this application.
The Committee adopted the CHMP recommendation and scientific discussion together with the List of Questions.

(EMEA/H/C/004024), (pregabalin), (treatment of neuropathic pain, epilepsy and generalised anxiety disorder (GAD))
The Committee discussed the issues identified in this application.
The Committee adopted the CHMP recommendation and scientific discussion together with the List of Questions.

(EMEA/H/C/003962), (pregabalin), (treatment of neuropathic pain, epilepsy and generalised anxiety disorder (GAD))
The Committee discussed the issues identified in this application.
The Committee adopted the CHMP recommendation and scientific discussion together with the List of Questions.

(EMEA/H/C/004078), (pregabalin), (treatment of epilepsy and generalised anxiety disorder (GAD))
The Committee discussed the issues identified in this application.
The Committee adopted the CHMP recommendation and scientific discussion together with the List of Questions.

(EMEA/H/C/004010), (pregabalin), , , (treatment of neuropathic pain, epilepsy and generalised anxiety disorder)
The Committee discussed the issues identified in this application.
The Committee adopted the CHMP recommendation and scientific discussion together with the List of Questions.

(EMEA/H/C/004070), (pregabalin), (treatment of epilepsy and generalised anxiety disorder (GAD))
The Committee discussed the issues identified in this application.
The Committee adopted the CHMP recommendation and scientific discussion together with the List of Questions.

(EMEA/H/C/003900), (pregabalin), (treatment of epilepsy and Generalised Anxiety Disorder (GAD))
The Committee discussed the issues identified in this application.
The Committee adopted the CHMP recommendation and scientific discussion together with the List of Questions.

(EMEA/H/C/003772), (ceftolozane / tazobactam), (treatment of intra-abdominal urinary tract infections)
The Committee discussed the issues identified in this application.

The Committee adopted the CHMP recommendation and scientific discussion together with the List of Questions.

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

(EMEA/H/C/003769), Orphan, (mercaptamine hydrochloride), Applicant: Orphan Europe S.A.R.L., (treatment of cystinosis)
The CHMP adopted the Assessment Report on similarity.

(EMEA/H/C/003727), Orphan, (lenvatinib), Applicant: Eisai Ltd, (treatment of papillary thyroid cancer, treatment of follicular thyroid cancer)
The CHMP adopted the Assessment Report on similarity.
The QWP report was adopted.

(EMEA/H/C/003731), Orphan, (blinatumomab), Applicant: Amgen Europe B.V., (treatment of adults with Philadelphia chromosome negative relapsed or refractory B-precursor acute lymphoblastic leukaemia)
The CHMP adopted the Assessment Report on similarity.
The Committee adopted the BWP report.

(EMEA/H/C/003702), (phenylephrine hydrochloride / ketorolac trometamol), (maintenance of intraoperative mydriasis, prevention of intraoperative miosis and reduction of acute postoperative ocular pain in intraocular lens replacement (ILR) in adults)
List of Outstanding Issues adopted on 22.05.2014. List of Questions adopted on 23.01.2014.
The Committee noted the report from the ad-hoc expert group meeting held on 4 December 2014.
In addition the Committee noted the QWP report and adopted this report.

(EMEA/H/C/002739), (human alpha1-proteinase inhibitor), (treatment of lung tissue)
List of Outstanding Issues adopted on 20 November 2014. List of Questions adopted on 25.04.2014.
The Committee adopted the List of experts for the ad hoc expert group meeting.
Furthermore the CHMP agreed to the request for an extension to the clock stop to respond to the List of Outstanding Issues adopted in November 2014 with a specific timetable.

(EMEA/H/C/003800), Orphan
(ketoconazole), Applicant: Agenzia Industrie Difesa-Stabilimento Chimico Farmaceutico Militare, (treatment of Cushing's syndrome)
Day 120 List of Questions adopted in April 2014. The Committee did not agree to the request from the applicant for an extensive clock stop to respond to the Day 120 List of Questions. The CHMP agreed to limit such extension of the clock stop.
(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.
The Committee noted the report from the SAG held on 1 December 2014.

(EMEA/H/C/002616), Orphan (Pitolisant Hydrochloride), Applicant: Bioprojet Pharma, (treatment of narcolepsy)
The CHMP agreed to the request from the applicant for an extension to the clock stop to respond to Day 120 List of Questions adopted in September 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. The CHMP agreed to the request by the applicant for an extension of clock stop with a specific timetable.

(EMEA/H/C/002839), (sonidegib), (treatment of basal cell carcinoma (BCC))
The CHMP agreed to the request by the applicant for an extension to the clock stop to respond to Day 120 List of Questions adopted in September 2014 together with a specific timetable.

(EMEA/H/C/003822), Orphan, (glycerol phenylbutyrate), Applicant: Hyperion Therapeutics Limited, (treatment of patients with urea cycle disorders). The CHMP agreed to the request by the applicant for an extension to the clock stop to respond to Day 120 List of Questions adopted in October 2014 together with a specific timetable.

(EMEA/H/C/003858), (insulin human), (treatment of diabetes)
The CHMP agreed to the request by the applicant for an extension to the clock stop to respond to Day 120 List of Questions adopted in October 2014 together with a specific timetable.

(EMEA/H/C/004071), (dexamethasone acetate),(treatment of symptomatic multiple myeloma in combination with other medicinal products)
The CHMP adopted a specific timetable for the similarity assessment.

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)
The members were reminded of previous discussions.
An Oral explanation was held on Wednesday 17 December 2014 at 14.00 focused on the Applicant's derogation claim from Market Exclusivity of TOBI Podhaler.

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

Oprymea (EMEA/H/C/000941/X/0017), (pramipexole), MAH: Krka d.d. Novo mesto, Generic of Sifrol, Rapporteur: Jens Heisterberg, PRAC Rapporteur: Doris Stenver, "To add new strengths 2.62 mg and 3.15 mg prolonged-release tablets."
List of Questions adopted on 25.09.2014.
The Committee confirmed that all issues previously identified in this application had been addressed.
The Committee adopted a positive Opinion by consensus, together with the CHMP Assessment report and Translation timetable.
The Icelandic and Norwegian Members were in agreement with the CHMP recommendation.

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

Orfadin (EMEA/H/C/000555/X/0042), Orphan, (nitisinone), MAH: Swedish Orphan Biovitrum International AB, Rapporteur: Luca Pani, "To add a new strength 20 mg capsule, hard."
List of Questions adopted on 24.07.2014.
The Committee was reminded of the status of this application and its remaining outstanding issues, mainly relating to some quality aspects.
The CHMP adopted a List of Outstanding Issues with a specific timetable.

Revlimid (EMEA/H/C/000717/X/0073/G), Orphan, (lenalidomide), MAH: Celgene Europe Limited, Rapporteur: Pierre Demolis, Co-Rapporteur: Filip Josephson, PRAC Rapporteur: Arnaud Batz, "A type II variation (C.I.6) to add the following indication
Revlimid is indicated for the continuous treatment of adult patients with previously untreated multiple myeloma who are not eligible for transplant
A line extension application to add the following strength: 20 mg (21 capsules pack)"
List of Questions adopted on 24.07.2014.
The Committee was reminded of the status of this application and its remaining outstanding issues.
The members discussed the benefit/risk especially the melphalan, prednisone and lenalidomide followed by lenalidomide (MPR+R) regimen and the most appropriate patient population for this combination.
The Committee adopted a positive Opinion recommending the granting of the extension application by majority (25 positive out of 29 votes) together with the CHMP assessment report and translation timetable.
The Icelandic and Norwegian Members were in agreement with the CHMP recommendation.
The divergent position (Hubert Leufkens, Pieter de Graeff, Daniela Melchiorri, Joseph Emmerich) was appended to the opinion.
The CHMP adopted the assessment report for Revlimid on similarity by consensus. The legal status was agreed as medicinal product subject to medical prescription.
The summary of opinion was circulated for information.

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

No items

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 -

Aloxi (EMEA/H/C/000563/II/0038), (palonosetron), MAH: Helsinn Birex Pharmaceuticals Ltd., Rapporteur: Patrick Salmon, Co-Rapporteur: Arantxa Sancho-Lopez, PRAC Rapporteur: Almath Spooner, "Extension of the therapeutic indication for paediatric patients 1 month of age and older for the prevention of nausea and vomiting associated with moderately and highly emetogenic cancer chemotherapy for the IV formulation, based on the paediatric studies PALO-10-14 and PALO-10-20 and update of sections 5.1 and 5.2 of the Aloxi Oral formulation to reflect those studies. The MAH took the opportunity of this variation to update the Aloxi product information annexes in line with Version 9 of the QRD template."
Request for Supplementary Information adopted on 25.09.2014.
The Committee discussed the issues identified in this application. The main discussion focused on the wording of the indication was well as some pharmacokinetic aspects.
The Committee adopted a 2nd Request for Supplementary Information with a specific timetable.

Fycompa (EMEA/H/C/002434/II/0016), (perampanel), MAH: Eisai Europe Ltd., Rapporteur: Robert James Hemmings, Co-Rapporteur: Pierre Demolis, PRAC Rapporteur: Julie Williams, "Extension of indication as adjunctive treatment of Primary Generalised Tonic-Clonic seizures in patients with epilepsy aged 12 years and older. Sections 4.1, 4.2, 4.4, 4.5, 4.8, 5.1 and 5.2 of the SmPC and the Package Leaflet are updated accordingly. In addition, the MAH took the opportunity to propose minor changes to PL and update the contact details of the Maltese local representative."
The Committee discussed the issues identified in this application and looked at the available clinical data with specific focus on the study population and other study parameters.
The Committee adopted a Request for Supplementary Information with a specific timetable.

Perjeta (EMEA/H/C/002547/II/0010), (pertuzumab), MAH: Roche Registration Ltd, Rapporteur: Christian Schneider, Co-Rapporteur: Daniela Melchiorri, PRAC Rapporteur: Doris Stenver, "The Marketing authorisation holder (MAH) applied for an extension of the indication for Perjeta in combination with trastuzumab and docetaxel for the neoadjuvant treatment of patients with HER2-positive, locally advanced, inflammatory, or early stage breast cancer (> 2 cm in diameter) as part of the treatment for early breast cancer. Consequently changes are proposed to sections 4.1, 4.2, 4.4, 4.5, 4.8, 5.1, 5.2 of the SmPC.""
The Committee discussed the issues identified in this application, which were mainly related to the study design and analysis.
The Committee adopted a Request for Supplementary Information .
The CHMP agreed to the request by the applicant for an extension to the clock stop with a specific timetable.
The CHMP agreed to consult the SAG Oncology. The list of questions to the SAG Oncology was adopted.

Pyramax (EMEA/H/W/002319/II/0002), (pyronaridine / pyronaridine phosphate / artesunate), MAH: Shin Poong Pharmaceutical Co., Ltd., Rapporteur: Joseph Emmerich, Co-Rapporteur: Johann Lodewijk Hillege, PRAC Rapporteur: Arnaud Batz, "To amend SmPC section 4.1 (Therapeutic Indications) to remove restrictions on repeated course of treatment in any individual and use only in

areas of low transmission with evidence of artesmisinin resistance, based on further clinical experience. Consequent changes in SmPC sections 4.2 (Posology), 4.4 (Special warnings and precautions), 4.8 (Undesirable effects) and the PL are also included. A recommended change is made to SmPC Section 4.2 (Posology) in relation to dosing in mild to moderate renal impairment. A minor editorial adjustment is proposed to SmPC section 5.1 (Pharmacodynamic properties)."
Request for Supplementary Information adopted on 26.06.2014.
The Committee discussed the issues identified in this application, which were related to the systematic liver testing and monitoring, also the conditions under which the drug could be safely re-administered were discussed.
The Committee agreed to consult the SAG Anti-infectives involving additional experts and adopted the list of questions to the SAG.
The Committee adopted a 2nd Request for Supplementary Information with a specific timetable.

Resolor (EMEA/H/C/001012/II/0034), (prucalopride), MAH: Shire Pharmaceuticals Ireland Ltd., Rapporteur: Greg Markey, Co-Rapporteur: Kristina Dunder, PRAC Rapporteur: Rafe Suvarna, "This type II variation (category C.I.6) is provided on the basis of the completion of the clinical study SPD555-302. Following a meeting with the Resolor Rapporteur on 11 March 2014 it was agreed that the results of SPD555-302 would be provided to the EMA for assessment no later than 30 September 2014.
The conduct and conclusion of study SPD555-302 was completed as a Post Approval Commitment (FUM 004 (MEA 004.2)) - To perform an efficacy study in males.
Based on the results from study SPD555-302, the MAH has submitted this type II variation to extend the indication for Resolor into the male population. The results of study SPD555-302 will be reflected in the Annex (a clean and track change version of the Resolor SmPC and PIL are provided for review).
Throughout section 5.1 (Pharmacodynamic properties) of the SmPC the MAH has replaced the name of the active "prucalopride" with "Resolor" for consistency and accuracy in this section.
In support of the proposed update to the Resolor SmPC a Phase 1 Study (SPD555-104) to Investigate the Absorption, Metabolism, and Excretion of [14C] Prucalopride Succinate Following a Single Oral Dose in Healthy Male Subjects has also been provided with this variation.
In support of the proposed amendment to section 4.1 (Therapeutic indications) of the SmPC the MAH also intends to apply for extended data/market exclusivity Under Article 14 (11) of Regulation (EC) No 726/2004 or Article 10 (1) fourth subparagraph of Directive 2001/83/EC. The Marketing Authorisation Holder (MAH) hereby applies for an additional 1 year of marketing protection and supporting documentation is thus provided in module 1.5.3. An updated Risk Management Plan (RMP) (version 12) is also provided with this submission. The updated RMP is provided following the finalisation of study SPD555-302 for the male population and also following a request from the Rapporteur during the assessment of the PSUR 007 (EMEA/H/C/1012/ PSU 021) to include the important potential risk "Increase in prolactin levels". Separate to the update of the Resolor Annex with an update to the therapeutic indication, the MAH also proposes changes to section 6 of the Patient Information Leaflet with the revision of the contact details (address and/or telephone numbers for the local representatives in Belgium and Italy."
Request for 1 year of market protection for a new indication (Article 14(11) of Regulation (EC) 726/2004).
The Committee discussed the issues identified in this application. The discussion mainly focused on the data submitted to support the additional year of market protection.
The Committee adopted a Request for Supplementary Information .
The CHMP agreed to the request by the applicant for an extension to the clock stop to respond to RSI with a specific timetable.

Tresiba (EMEA/H/C/002498/II/0011), (insulin degludec), MAH: Novo Nordisk A/S, Rapporteur: Kristina Dunder, PRAC Rapporteur: Qun-Ying Yue, "Extension of the indication in children aged from 1 to 18 years. Update to sections 4.1, 4.2, 4.8 and 5.1 of the SmPC. The PL is updated accordingly. In addition, update of the Section 2 of the PL in line with the existing information in Section 4.2 of the SmPC."
Request for Supplementary Information adopted on 25.09.2014.
The Committee discussed the issues identified in this application. The Committee confirmed that all issues previously identified in this application had been resolved.
The Committee adopted a positive Opinion by consensus together with the CHMP Assessment Report and translation timetable.
The Icelandic and Norwegian CHMP members were in agreement with the CHMP recommendations.
The summary of opinion was circulated for information.

Velcade (EMEA/H/C/000539/II/0072), (bortezomib), MAH: Janssen-Cilag International N.V., Rapporteur: Daniela Melchiorri, Co-Rapporteur: Outi Mäki-Ikola, PRAC Rapporteur: Carmela Macchiarulo, "Extension of indication to include the use of Velcade in combination with rituximab, cyclophosphamide, doxorubicin and prednisone for the treatment of adult patients with previously untreated mantle cell lymphoma who are unsuitable for haematopoietic stem cell transplantation.
As a consequence, 4.1, 4.2, 4.4, 4.8 and 5.1 of the SmPC have been updated. The Package Leaflet is updated in accordance."
Request for Supplementary Information adopted on 25.09.2014.
The Committee discussed the issues identified in this application. The Committee discussed the wording of the indication and the route of administration. The Committee confirmed that all issues previously identified in this application had been resolved.
The Committee adopted a positive Opinion by consensus together with the CHMP Assessment Report and translation timetable.
The Icelandic and Norwegian CHMP members were in agreement with the CHMP recommendations.
The summary of opinion was circulated for information.
The CHMP adopted the Assessment Report on similarity.

Voncento (EMEA/H/C/002493/II/0008/G), (human coagulation factor viii / human von willebrand factor), MAH: CSL Behring GmbH, Rapporteur: Pieter de Graeff, Co-Rapporteur: Greg Markey, PRAC Rapporteur: Sabine Straus, "Extension of indication to include prophylactic treatment of patients with VWD. In addition the MAH is providing data to support treatment of paediatric patients with VWD."
The Committee discussed the issues identified in this application. The main outstanding issues related to the population pharmacokinetic model, the posology for the paediatric population as well as a clarification on an adverse event case.
The Committee adopted a Request for Supplementary Information with a specific timetable.

Xiapex (EMEA/H/C/002048/II/0044), (collagenase clostridium histolyticum), MAH: Swedish Orphan Biovitrum AB (publ), Rapporteur: Martina Weise, Co-Rapporteur: Pierre Demolis, PRAC Rapporteur: Martin Huber, "Update of the SmPC with a new indication in the treatment of adult men with Peyronie's disease with a palpable plaque and curvature deformity. The PL is updated accordingly."
Request for Supplementary Information adopted on 25.09.2014.

Request for 1 year of market protection for a new indication (Article 14(11) of Regulation (EC) 726/2004)
The Committee discussed the issues identified in this application. The Committee confirmed that all issues previously identified in this application had been resolved.
The Committee adopted a positive Opinion by consensus together with the CHMP Assessment Report and translation timetable.
The Icelandic and Norwegian CHMP members were in agreement with the CHMP recommendations.
The summary of opinion was circulated for information.
The Committee agreed by consensus on the 1 year of market protection for the new indication.

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

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."
The Committee agreed to the request by the applicant for an extension of the clock stop together with a revised timetable.

5. Ancillary medicinal substances in medical devices

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

(EMEA/H/D/003740), (human serum albumin), , (the storage, manipulation, in-vitro culture and transfer of human gametes)
List of Outstanding Issues adopted on 25.09.2014. List of Questions adopted on 23.01.2014.
The Committee was reminded of the status of this application and its remaining outstanding issues, .
The Committee adopted the 2nd List of Outstanding Issues with a specific timetable.
The Committee adopted the BWP report.

(EMEA/H/D/002831), (insulin-like growth factor-i (igf-i) segment), (hard-to-heal wounds, primarily venous leg ulcers)
List of Questions adopted on 23.01.2014. List of Outstanding Issues adopted on 25.09.2014.
The Committee adopted a revised timetable.

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

No items

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.*

(H0003790), Orphan (carfilzomib) Applicant: Amgen Europe B.V., (indicated for the treatment of adult patients with relapsed and refractory multiple myeloma who have received at least two prior therapies that included bortezomib (a proteasome inhibitor) and an immunomodulatory agent, or for whom such treatments are not appropriate),
The CHMP agreed to the request for accelerated assessment and adopted the briefing note and Rapporteurs' recommendation on the Request for Accelerated Assessment.

11. Post-authorisation issues

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

Tyverb (EMEA/H/C/000795/II/0037), (lapatinib), MAH: Glaxo Group Ltd, Rapporteur: Filip Josephson, PRAC Rapporteur: Ulla Wändel Liminga, "Update of section 5.1 of the SmPC with information on lapatinib effect on CNS metastasis further to comparative data on the incidence of CNS metastases from studies EGF108919 (COMPLETE), EGF105485 (TEACH) and EGF106708 (ALTTO) (SOB 002.4). Annex II is also updated further to the fulfilment of the specific obligation."
The Committee discussed the issues identified in this application. The members noted that with the data submission in this variation the last remaining specific obligation from the conditional marketing authorisation has been fulfilled. The Committee agreed that now comprehensive data on efficacy and safety of lapatinib are available and that a full marketing authorisation could be issued.
The CHMP adopted an opinion by consensus recommending switching from a conditional to full marketing authorisation, together with the assessment report.
The Icelandic and Norwegian Members were in agreement with the CHMP recommendation.

Tyverb (EMEA/H/C/000795/II/0038), (lapatinib), MAH: Glaxo Group Ltd, Rapporteur: Filip Josephson, "Update of sections 4.1, 4.4 and 5.1 of the SmPC further to disease-free survival (DFS)

results from hormone receptor-positive patients in the adjuvant EGF106708 (ALTTO) study (REC 037)."
The Committee discussed the issues identified in this application, mainly focusing on the wording of the indication. The CHMP adopted an opinion by consensus recommending the variation to the terms of the Marketing Authorisation together with the assessment report.
The Icelandic and Norwegian Members were in agreement with the CHMP recommendation.

INOmax (EMEA/H/C/000337) (Nitric Oxide), MAH: Linde Healthcare AB, Rapporteur: Robert James Hemmings, Co-Rapporteur: Daniel Brasseur, (treatment of hypoxic respiratory failure in newborns), Complete application (stand-alone) - Council Directive 81/851/EEC
The Committee adopted the DHPC and the communication plan.
The Committee adopted the Assessment Report.

Eperzan (EMEA/H/C/002735/II/0009), (albiglutide), MAH: GlaxoSmithKline Trading Services, Rapporteur: Kristina Dunder, "Update of section 4.8 with information on appendicitis / pancreatitis. The PL is updated accordingly. In addition, the MAH took the opportunity to correct some information in Sections 5.1 and 6.6 of the SmPC, the Annex III.A and the Package Leaflet."
Request for Supplementary Information adopted on 23.10.2014.
The members discussed the available scientific information
The CHMP agreed to consult the PRAC
The Committee adopted a Request for Supplementary Information with a specific timetable.

Xarelto (EMEA/H/C/000944/II/0033), (rivaroxaban), MAH: Bayer Pharma AG, Rapporteur: Kristina Dunder, , "Section 4.2 and 5.1: After approval of the Xarelto in 2008, a prospective, randomized, open-label, parallel-group, active
controlled phase IIIb study (X-VERT) was performed to compare Xarelto® with vitamin K antagonists (VKA) for the prevention of cardiovascular events in patients with non-valvular atrial fibrillation undergoing cardioversion. Based on the results of this X-VERT study, the applicant therefore proposes to update the information in the indication SPAF in the section 4.2 and to describe the results of this X-VERT study in section 5.1 of the SmPC."
Request for Supplementary Information adopted on 25.09.2014.
The Committee discussed the issues identified in this application, with specific focus on different sections of the SmPC in comparison with other medicines' wording in the same therapeutic group.
The Committee confirmed that all issues previously identified in this application had been resolved.
The Committee adopted a positive Opinion on the amendment of the SmPC by consensus together with the CHMP assessment report and translation timetable.
The Icelandic and Norwegian Members were in agreement with the CHMP recommendation.

Docetaxel Accord (EMEA/H/C/002539) (Docetaxel), MAH: Accord Healthcare Ltd, Rapporteur: Filip Josephson, (Article 10(1) of Directive No 2001/83/EC).
The CHMP noted the update.

Vyndaqel (EMEA/H/C/002294/II/0021), Orphan, (tafamidis), MAH: Pfizer Limited, Rapporteur: Joseph Emmerich, , "Update of section 5.1 of the SmPC in order to include information from a recently completed thorough QT study."
The Committee discussed the issues identified in this application, mainly related to the shown effect of tafamidis on QTc and how to amend section 5.1 of the SmPC.
The Committee adopted a Request for Supplementary Information with a specific timetable.

Zelboraf (EMEA/H/C/002409/II/0018), (vemurafenib), MAH: Roche Registration Ltd, Rapporteur: Filip Josephson, "Update of section 4.8 of the SmPC in order to add the adverse reaction pancreatitis reported with a frequency "uncommon" further to a cumulative review conducted by the MAH. The Package Leaflet is updated accordingly."
The Committee discussed the issues identified in this application. The Committee looked at the incidence of pancreatitis for the product class and discussed the appropriate wording of section 4.4. of the SmPC.
The Committee adopted a Request for Supplementary Information with a specific timetable.

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 20.11.2014, 26.06.2014.
The Committee discussed the issues identified in this application, mainly focusing on potential off label use of lenalidomide in CLL patients in the EU.
The Committee adopted a 3rd Request for Supplementary Information with a specific timetable.

12. Referral procedures

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

No items

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

Polymyxin-based products (EMEA/H/A-5(3)/1384) (colistin, colistimethate), Rapporteur: Robert James Hemmings, Co-Rapporteur: Martina Weise,
Review of the module 3 (quality) and European Pharmacopeia monograph. Triggered by the EMA Executive Director.

The CHMP agreed to consult the QWP and adopted a revised timetable.
Joint assessment report: 22.01.2015; Comments from CHMP: 29.01.2015; Consultation of the QWP: February 2015 QWP; Updated joint assessment report: 16.02.2015; CHMP discussion/CHMP adoption of opinion: February 2015 CHMP.

Medicinal products under development for the treatment of Ebola (EMEA/H/A-5(3)/1410)
The CHMP was updated about the procedure.

Post meeting note: a revised timetable was adopted via written procedure on 19 December 2014

Submission of responses : 16.12.2014; Updated Co-Rapporteurs' assessment reports and updated Lead Rapporteur's overall assessment report circulated to CHMP: 09.01.2015; CHMP members' comments: 14.01.2015; Updated Rapporteur' and co-Rapporteur's assessment reports circulated to CHMP, as applicable: 16.01.2015; CHMP discussion: January 2015 CHMP

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

No items

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

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
The CHMP noted the update.

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 Oral explanation held in October 2014.
The Committee discussed the outcome of GVK Working Group meeting held on 8-9 December 2014. submissions were reviewed and divided between 3 categories.
Category 1: no other evidence submitted or ongoing BE studies
Category 2: some evidence submitted (e.g. biowaiver)
Category 3: evidence submitted, e.g. other bioequivalence data.
The CHMP appointed lead member states to assess the INNs in categories 2, 3 and on-going studies , as applicable, with a specific timetable.
TC with Co-Rapporteurs/WG: 18.12.2014; TC with Co-Rapporteurs (purpose: outcomes, heads up for rapporteurs): 08.01.2015; Co-Rapporteurs ARs circulated to CHMP: 12.01.2015; Rapporteur overall AR circulated to CHMP (along with Annexes): 14.01.2015; Comments from CHMP: 16.01.2015; CHMP opinion/OE: January 2015 CHMP

Adrenaline auto injectors (EMEA/H/A-31/1398) Rapporteur: Alar Irs, Co-Rapporteur: Robert Hemmings,
Article 31 triggered by the MHRA due to the lack of robust evidence that the devices deliver the adrenaline intramuscularly in all patients.
The CHMP adopted the list of experts for the ad-hoc expert meeting together with a revised timetable.

Ad hoc expert group meeting: 23.01.2015; Re-start of the procedure: 27.01.2015; Joint assessment report circulated to CHMP: 09.02.2015; Comments: 16.02.2015; List of outstanding issues or CHMP opinion: February 2015 CHMP

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 1-4 December 2014: **For information**	The Committee noted the report. The members noted the Summary of recommendations and advices of the PRAC meeting.
List of Union Reference Dates and frequency of submission of Periodic Safety Update Reports (EURD list) for December 2014: **For adoption**	The EURD list was adopted.
Early Notification System: December 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**	See individual items

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.*
GCP Inspection Report: **For information**	
Extension of the target date of the IIR: **For adoption**	

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.*
Risk-based programme for routine pharmacovigilance inspections of MAHs connected with human centrally authorised products: **For adoption**	The CHMP adopted Confidential H PhV inspection programme 2014-2017 (2nd revision for 2014).

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.

ITF Briefing Meeting	The CHMP agreed to the ITF meeting.
ITF Briefing Meeting	The CHMP agreed to the ITF meeting.

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 1-4 December 2014. Table of conclusions: **For information**	The CHMP noted the report.
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 15-17 December 2014: **For information**	The CHMP noted the report.
CMDh question to CHMP (PKWP) regarding potential risk of longer half-life of acitretin, • Feedback from the teleconference between the Chairs of CHMP, PKWP and SWP and the DK assessor held on 2 December 2014: **For discussion** • Questions from CMDh to PKWP and SWP on a possible extension of the post-	The CHMP noted the Questions from CMDh to PKWP and SWP were agreed. The Committee discussed the feedback from the teleconference.

treatment pregnancy prevention period after use of acitretin: **For adoption**	
Reply letter from PDCO about the request from CMDh to CHMP and PDCO regarding Guidance on the development of parenteral nutrition in the paediatric population: **For adoption**	The CHMP agreed to the response letter from PDCO and adopted the final letter to CMDh.

18. Other Committees

18.1. Committee for Orphan Medicinal Products (COMP)

Press release of the COMP meeting held on 9-10 December 2014: **For information**	To be sent in the Post-mail.

18.2. Committee for Herbal Medicinal Products (HMPC)

Report from the HMPC meeting held on 24-25 November 2014: **For information**	To be sent in the Post-mail.

18.3. Paediatric Committee (PDCO)

PIPs reaching D30 at December 2014 PDCO: **For information**	To be sent in the Post-mail.
Report from the PDCO meeting held on12-14 December 2014: **For information**	The CHMP noted the report.

18.4. Committee for Advanced Therapies (CAT)

Table of Decisions of CAT meeting held on 13-14 December 2014: **For information**	The CHMP noted the Table of Decision.

19. Invented name issues

Table of Decisions of the NRG meeting held on 26 November 2014: **For adoption**	The CHMP adopted the Table of Decision.

20. Any other business

Initiative to enhance early dialogue with applicants to foster better development of products	The objectives of the initiative are to raise awareness of regulatory requirements earlier in development to enable sponsors to optimise and accelerate medicines development; improve data quality; reinforce regulatory predictability to stimulate R&D investment; foster and facilitate interaction with the Agency & the network; improve planning and optimise resource allocation across the network.
Reminder to committee members to submit e-DoI version 2 by 30 Jan 2015	The Committee noted the new e-DoI timelines.
Benefit-risk communication to medicines users. How can regulators best meet the information needs of patients and healthcare professionals? Workshop (held 2013-2014) report (EMA/581546/2014): **For information**	Series of workshops were held involving representatives of patients and healthcare professionals together with academics, members of EMA scientific committees, EMA staff and other interested stakeholders. The objective of this workshop were to review current practice in communicating benefit-risk, to examine recent initiatives into how research can help inform best practice, to discuss the role of communications in risk minimisation and to explore how they can aid patients and healthcare professionals in making decisions throughout the therapeutic journey. The members noted the link to the workshop documents, presentations and video. As a next step a survey will be prepared on how the safety communication is perceived by different stakeholders.
Benefit-Risk Effects Table Guidance: **For discussion**	The CHMP was informed on the latest updates to the Effects Table Guidance and the training which will be provided on this as part of the Benefit/Risk Workshop in January 2015. The Committee endorsed the proposed implementation plan for the Effects Table in the CHMP AR. It was agreed that the Effects Table will be constructed only by the Rapporteur for all new initial applications and significant extensions of indications which will be submitted from 1st of February 2015 onwards. The Guidance on the Effects Table was endorsed in principle but will be updated as experience with these Tables is gathered and evaluated.

Update Multinational Teams Scheme – Extended to CHMP/CAT Rapporteurships (not only Co-Rapporteurships): **For information**	Follow up from December ORGAM. Extension of Multinational Teams Scheme to rapporteurships, starts from January 2015 on and applies only for the initial MAA assessment.
PSUR repository implementation plan: **For discussion**	The members were updated on the PSUR repository implementation plan. The CHMP had no objections against the proposed starting date of 12.03.2015.
Outcome of presidency CHMP meeting in Rome • Action items: **For agreement**	The CHMP agreed on the action items
Presidency CHMP meeting in the first half of year 2015 under the Latvian EU presidency: **For discussion**	The CHMP was updated on the next presidency meeting under the Latvian EU presidency.
FDA final rule on changes to pregnancy and lactation labeling information for prescription drug and biological products: **For discussion**	FDA published the final rule that sets standards for how information about using medicines during pregnancy and breastfeeding is presented in the labelling of prescription drugs and biological products. The final rule replaces the current product letter categories – A, B, C, D and X – used to classify the risks of using prescription drugs during pregnancy with three detailed subsections: Pregnancy; Lactation; Females and Males of Reproductive Potential.
Guideline on similar biological medicinal products containing biotechnology-derived proteins as active substance: non-clinical and clinical issues: **For adoption and release for implementation**	The guideline was presented to the Committee . The CHMP adopted the guideline.
Update on Risk Management Process: **For discussion**	The schedule for the implementation of the revised RMP assessment process was presented. The members highlighted the need for further clarification on the roles of the CHMP and PRAC Rapporteurs. Furthermore a question on the applicability for generics was raised. This will be clarified at a later time. For the new MAAs this will be implemented starting from February 2015.
Follow-up on general aspects regarding SmPC labelling (indication wording): **For discussion**	CHMP agreed to have a small working group, who would work on principles about the indications' wording. Some CHMP members expressed interest:
Daklinza indication wording and general issues regarding indication wording: **For discussion**	The Committee was updated on issues related to Daklinza indication wording. The importance of indication wording for HTA bodies was highlighted.

Data gathering initiative, Pilot Exercise: **For information**	In March 2014, the project was initiated in order to gather evidence needed by European Commission in drafting future legislative proposal on fees. The objective is to assemble evidence about time spent on procedures at EMA + NCA's. The Pilot Exercise will be conducted to validate the time collection methodology and to extend the refined collection process to formal exercise addressing the major activities areas of the network.
Process for agreement of Working Parties' Work Programmes 2015: **For discussion**	Existing WPs/DGs will be asked to prepare a work programme 2015 based on the 2014-agreed paper, highlighting in track-change any updates and deadlines. Draft 2015 Work Programmes will be presented to the CHMP plenary in either January or February for adoption.
Answers to the requests for scientific advice on the impact on public health and animal health of the use of antibiotics in animals (EMA/381884/2014): **For adoption** • Overview of comments received on 'Answers to the request for scientific advice on the impact on public health and animal health of the use of antibiotics in animals' (EMA/381884/2014): **For information**	The CHMP adopted the Answers to the requests for scientific advice on the impact on public health and animal health of the use of antibiotics in animals. The requests were received from European Commission and the answers were prepared by the Antimicrobial Advice ad hoc Expert Group. The CHMP noted the overview of comments.
Draft Guidance with applicants on responses to questions received from EMA Scientific Committees during the evaluation within the centralised procedure: **For adoption**	Postponed
Update on renewal to-be process: **For discussion**	The CHMP noted the new process on renewals.
Appointment of CHMP representatives to the CAT	The current members (Sol Ruiz, Romaldas Maciulaitis, Bruno Sepodes, John Borg, Jean-Louis Robert) confirmed their willingness to continue the cross Committee membership to the CAT. The CHMP renominated Sol Ruiz, Romaldas Maciulaitis, Bruno Sepodes, John Borg, Jean-Louis Robert as CHMP representative to the CAT for another 3-year mandate. The Co-opted members Sol Ruiz and Jean-Louis Robert need to propose their CAT alternates.
Q4-14 update of 2015 expected MAAs with appointed Rapporteur for CHMP: **For information**	The CHMP noted the report.

21. List of participants: including any restrictions with respect to involvement of members / alternates / experts following evaluation of declared interests for the 15-18 December 2014 meeting.

CHMP Chair	Country	Outcome restriction following evaluation of e-DoI for the meeting	Topics on the current Committee Agenda for which restrictions applies Product/substance
Tomas Salmonson	Sweden	Full involvement	

CHMP Member	Country	Outcome restriction following evaluation of e-DoI for the meeting	Topics on the current Committee Agenda for which restrictions applies Product/substance
Andrea Laslop	Austria	Full involvement	
Daniel Brasseur	Belgium	Full involvement	
Mila Vlaskovska	Bulgaria	Full involvement	
Ivana Mikacic	Croatia	No participation in final deliberations and voting on:	(EMEA/H/C/003772), (ceftolozane / tazobactam)
Panayiotis Triantafyllis	Cyprus	Full involvement	
Ondřej Slanař	Czech Republic	No participation in final deliberations and voting on:	Tasermity (EMEA/H/C/003968), (sevelamer)
			(EMEA/H/C/003900), (pregabalin)
			(EMEA/H/C/004021), (aripiprazole)
			Tresiba (EMEA/H/C/002498/II/0011), (insulin degludec)
		No participation in discussions, final deliberations and voting on:	12.6. Community Interests - Referral under Article 31 of Directive 2001/83/EC: GVK Biosciences (EMEA/H/A-31/1408)
Jens Heisterberg	Denmark	No participation in final deliberations and voting on:	(EMEA/H/C/004021), (aripiprazole)
			12.6. Community Interests - Referral under Article 31 of Directive 2001/83/EC: GVK Biosciences (EMEA/H/A-31/1408)
Alar Irs	Estonia	Full involvement	
Outi Mäki-Ikola	Finland	No participation in final deliberations and voting on:	Tasermity (EMEA/H/C/003968), (sevelamer)
			(H0003790), Orphan (carfilzomib)
			(EMEA/H/C/003731),

CHMP Member	Country	Outcome restriction following evaluation of e-DoI for the meeting	Topics on the current Committee Agenda for which restrictions applies Product/substance
			Orphan, (blinatumomab) 12.6. Community Interests - Referral under Article 31 of Directive 2001/83/EC: GVK Biosciences (EMEA/H/A-31/1408)
Pierre Demolis	France	Full involvement Connected via TC	
Harald Enzmann	Germany	Full involvement	
Dimitrios Kouvelas	Greece	Full involvement	
Agnes Gyurasics	Hungary	Full involvement	
Kolbeinn Guðmundsson	Iceland	Full involvement Connected via TC	
David Lyons	Ireland	Full involvement	
Daniela Melchiorri	Italy	Full involvement	
Romaldas Mačiulaitis	Lithuania	Full involvement	
Jacqueline Genoux-Hames	Luxembourg	Full involvement	
John Joseph Borg	Malta	Full involvement	
Pieter de Graeff	Netherlands	Full involvement	
Karsten Bruins Slot	Norway	Full involvement	
Piotr Fiedor	Poland	Full involvement	
Bruno Sepodes	Portugal	Full involvement	
Nela Vilceanu	Romania	Full involvement	
Jan Mazag	Slovakia	Full involvement	
Stanislav Primožič	Slovenia	Full involvement	
Concepcion Prieto Yerro	Spain	Full involvement	
Kristina Dunder	Sweden	Full involvement	
Greg Markey	United Kingdom	Full involvement	
Jan Mueller-Berghaus	Co-Opted	Full involvement	
Jean-Louis Robert	Co-opted	Full involvement	
Hubert Leufkens	Co-opted	Full involvement	
Sol Ruiz	Co-opted	Full involvement	
Robert James Hemmings	Co-opted	Full involvement	

CHMP Alternate	Country	Outcome restriction following evaluation of e-DoI for the meeting	Topics on the current Committee Agenda for which restrictions applies Product/substance
Milena Stain	Austria	Full involvement	
Bart Van der Schueren	Belgium	Full involvement	
Radka Montoniová	Czech Republic	Full involvement	
Christian Schneider	Denmark	Full involvement Connected via TC	
Joseph Emmerich	France	No participation in final deliberations and voting on:	12.6. Community Interests - Referral under Article 31 of Directive 2001/83/EC: GVK Biosciences (EMEA/H/A-31/1408)
			Xarelto (EMEA/H/C/000944/II/0033), (rivaroxaban)
			Clopidogrel-ratiopharm (EMEA/H/C/004006), (clopidogrel)
Martina Weise	Germany	Full involvement	
George Aislaitner	Greece	Full involvement	
Melinda Sobor	Hungary	Full involvement	
Hrefna Gudmundsdottir	Iceland	Full involvement	
Patrick Salmon	Ireland	Full involvement	
Natalja Karpova	Latvia	Full involvement	
Johann Lodewijk Hillege	Netherlands	Full involvement	
Ingunn Hagen Westgaard	Norway	Full involvement	
Dinah Duarte	Portugal	Full involvement	
Arantxa Sancho-Lopez	Spain	Full involvement	
Filip Josephson	Sweden	Full involvement	
Tomas Salmonson	Sweden	Full involvement	
Rafe Suvarna	United Kingdom	Full involvement	

European Commission	Country	Outcome restriction following evaluation of e-DoI for the meeting	Topics on the current Committee Agenda for which restriction applies Product/ substance
	European Commission	Full involvement	
	European	Full involvement	

European Commission	Country	Outcome restriction following evaluation of e-DoI for the meeting	Topics on the current Committee Agenda for which restriction applies Product / substance
	Commission		
	European Commission	Full involvement	

CHMP Expert	Country	Outcome restriction following evaluation of e-DoI for the meeting	Topics on the current Committee Agenda for which the expert is invited Product / substance
* Experts were only evaluated against the product they have been invited to talk about.			
Christophe Focke	Belgium	Full involvement	
Anne Hasle Buur	Denmark	Full involvement	
Sabine Mayrhofer	Germany	Full involvement	
Eva Blahutova	Slovakia	Full involvement	
Ivana Pankuchova	Slovakia	Full involvement	
Jorge Camarero Jiménez	Spain	Full involvement	
Sara Camilleri	Malta	Full involvement	
Cheryl Aquilina	Malta	Full involvement	
Miriam Bartolo	Malta	Full involvement	
Judit Fehér	Malta	Full involvement	
Nathalie Morgensztejn	France	Full involvement	
Ingrid Chau	France	Full involvement	
Patrick Vrijlandt	Netherlands	Full involvement	
Paula Salmikangas	Finland	Full involvement	
Sandra Krall-Choma	Germany	Full involvement	

CHMP Expert by phone	Country	Outcome restriction following evaluation of e-DoI for the meeting	Topics on the current Committee Agenda for which the expert is invited Product / substance
* Experts were only evaluated against the product they have been invited to talk about.			
Jens Ersbøll	Denmark	Full involvement	
Sinan B. Sarac	Denmark	Full involvement	
Bronwyn Grimshaw	United Kingdom	Full involvement	

CHMP Expert by phone	Country	Outcome restriction following evaluation of e-DoI for the meeting	Topics on the current Committee Agenda for which the expert is invited Product/substance
Enrike Potthast	Germany	Full involvement	
Pekka Kurki	Finland	Full involvement	
Bertil Jonsson	Sweden	Full involvement	
Maarten Simoons	EMA	Full involvement	

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)*

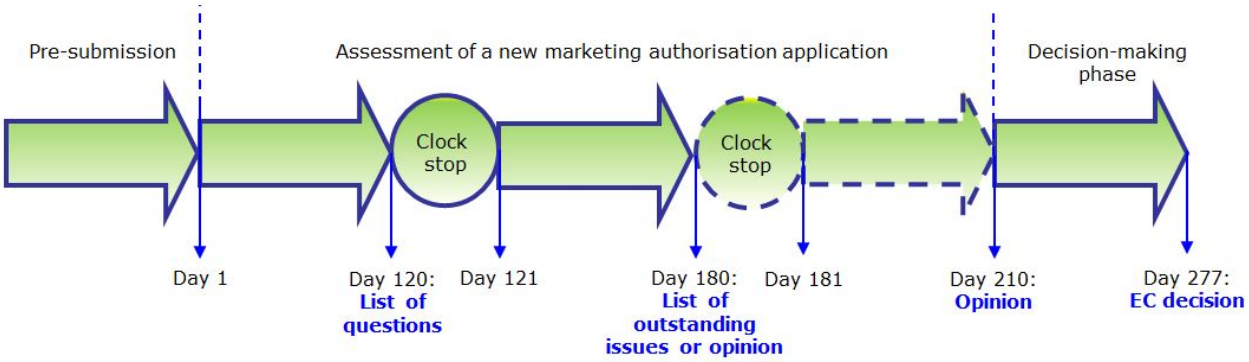
The items listed in this section are those for which marketing authorisation holders (MAHs) or applicants have been invited to the CHMP plenary meeting to address questions raised by the Committee. Oral explanations normally relate to on-going applications (section 2 and 3) or referral procedures (section 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.