27 May 2016
EMA/CHMP/363014/2016 Rev. 1
Procedure Management and Committees Support Division

Committee for medicinal products for human use (CHMP)

Minutes for the meeting on 25-28 April 2016

Chair: Tomas Salmonson – Vice-Chair: Pierre Demolis

Health and safety information

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

Disclaimers

Some of the information contained in these minutes is considered commercially confidential or sensitive and therefore not disclosed. With regard to intended therapeutic indications or procedure scopes listed against products, it must be noted that these may not reflect the full wording proposed by applicants and may also vary during the course of the review. 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.

Of note, these minutes are a working document primarily designed for CHMP members and the work the Committee undertakes.

Note on access to documents

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

30 Churchill Place ● Canary Wharf ● London E14 5EU ● United Kingdom
Telephone +44 (0)20 3660 6000 **Facsimile** +44 (0)20 3660 5520
Send a question via our website www.ema.europa.eu/contact

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1. Introduction

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

In accordance with the Agency's policy on handling of declarations of interests of scientific committees' members and experts, based on the declarations of interest submitted by the Committee members, alternates and experts and based on the topics in the agenda of the current meeting, the Committee Secretariat announced that no restriction in the involvement of meeting participants in upcoming discussions was identified as included in the pre-meeting list of participants and restrictions. See (current) April 2016 CHMP minutes for the 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 held 25-28 April 2016.

Participants in this meeting were asked to declare any changes, omissions or errors to their declared interests and/or additional restrictions concerning the matters for discussion. No new or additional interests or restrictions were declared.

Discussions, deliberations and voting took place in full respect of the restricted involvement of Committee members and experts in line with the relevant provisions of the Rules of Procedure and as included in the list of participants. All decisions taken at this meeting were made in the presence of a quorum of members (i.e. 22 or more members were present in the room). All decisions, recommendations and advice were agreed by consensus, unless otherwise specified.

1.2. Adoption of agenda

CHMP agenda for 25-28 April 2016.

The CHMP adopted the agenda.

1.3. Adoption of the minutes

CHMP minutes for 29 March - 1 April 2016.

The CHMP adopted the minutes.

2. Oral Explanations

2.1. Pre-authorisation procedure oral explanations

2.1.1. - sirolimus - Orphan - EMEA/H/C/003978

Santen Oy; treatment of chronic non-infectious uveitis

Scope: Oral explanation, Report from Ad-hoc expert group meeting held on 5 April 2016.

Action: Oral explanation to be held on Tuesday 26 April 2016 at 14.00. Report from ad-hoc expert meeting held on 5 April 2016.

List of Outstanding Issues adopted on 17.12.2015. List of Questions adopted on 25.06.2015.

The CHMP noted the minutes from the ad-hoc expert group meeting held on 5 April 2016.

Based on the available data the expert were not able to clearly identify a patient sub-population that might benefit most from intravitreal sirolimus. There was general agreement amongst the experts that Opsiria could in principle be a promising treatment option based on the local route of administration and lack of corticosteroid side effects such as increased intraocular pressure and cataract. Given the chronic nature of uveitis, the potential to avoid the complication of long-term corticosteroid treatment would be of value. However, the results from the pivotal trial supporting the application, while showing some effect based on a limited improvement in vitreous haze, were considered disappointing.

An oral explanation was held on Tuesday 26 April 2016 at 14.00. The applicant presented data to support a clinically relevant beneficial effect and made proposals for possible restricted target populations with a more favourable benefit-risk profile. The bell-shaped dose response curve was also addressed.

2.1.2. - saxagliptin / dapagliflozin - EMEA/H/C/004057

treatment of type 2 diabetes

Scope: Oral explanation

Action: Oral explanation to be held on Tuesday 26 April 2016 at 16.00.

List of Outstanding Issues adopted on 25.02.2016. List of Questions adopted on 24.09.2015.

An oral explanation was held on Tuesday 26 April 2016 at 16.00. The presentation focused on the data supporting the fixed dose combination of saxagliptin and dapagliflozin.

See also 3.2.10

2.2. Re-examination procedure oral explanations

No items

2.3. Post-authorisation procedure oral explanations

2.3.1. Adcetris - brentuximab vedotin - Orphan - EMEA/H/C/002455/II/0025

Takeda Pharma A/S

Rapporteur: Pieter de Graeff, Co-Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Sabine Straus

Scope: Oral explanation, report by Steen Werner Hansen from SAG Oncology meeting held on 14th April 2016

Action: Oral explanation to be held on Tuesday 26 April 2016 at 11.00.

“Extension of Indication to include new indication for Adcetris (Adcetris is indicated for the treatment of adult patients at increased risk of relapse or progression following ASCT”). as a consequence, sections 4.1, 4.2, 4.4, 4.8 and 5.1 of the SmPC are updated. The Package Leaflet is updated in accordance.” Request for 1 year of market protection for a new indication (Article 14(11) of Regulation (EC) 726/2004).

Action: For adoption

Request for Supplementary Information adopted on 01.04.2016, 22.10.2015, 25.06.2015.

An oral explanation was held on Tuesday 26 April 2016 at 11.00. The presentation focused on

the clinical data on progression free survival and overall survival as well as safety data and analysis performed. There was further discussion on the target population and on the significant clinical benefit over current standard of care.

See also 5.1.1

2.4. Referral procedure oral explanations

2.4.1. Otipax 1% (11 mg/ml) ear drops, solution – lidocaine hydrochloride - EMEA/H/A-29/1426

Biocodex Benelux SA/NV

Rapporteur: Daniel Brasseur, Co-Rapporteur: Martina Weise,

RMS: BE, CMS: AT, CY, DE, DK, EL, ES, FI, IT, NO, PL, PT, SE, Mutual recognition procedure number: BE/H/0213/001/MR

Scope: Oral explanation and Opinion

Disagreement regarding efficacy and the evidence of well-established use.

Action: Oral explanation to be held on Tuesday 26 April 2016 at 9.00.

List of Questions adopted on 22 October 2015. List of outstanding issues adopted on 25.02.2016.

Oral explanation was held on Tuesday 26 April 2016 at 9.00. The Company's presentation focused on publications on the local use of lidocaine, the formulation of Otipax and its safety.

See also 10.4.1

3. Initial applications

3.1. Initial applications; Opinions

3.1.1. EndolucinBeta - lutetium (177 lu) chloride - EMEA/H/C/003999

ITG Isotope Technologies Garching GmbH; radiolabelling of carrier molecules, which have been specifically developed for radiolabelling with this radionuclide

Scope: Opinion

Action: For adoption

Known active substance (Article 8(3) of Directive No 2001/83/EC)

List of Outstanding Issues adopted on 25.02.2016. List of Questions adopted on 24.09.2015.

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 lutetium (177 Lu) chloride is not 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.

3.1.2. Enzepi - pancreas powder - EMEA/H/C/002070

Aptalis Pharma SAS; treatment in exocrine pancreatic insufficiency

Scope: Opinion

Action: For adoption

Known active substance (Article 8(3) of Directive No 2001/83/EC)

List of Outstanding Issues adopted on 25.02.2016. List of Questions adopted on 25.06.2015.

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.

The Committee adopted the BWP report.

3.1.3. Odefsey - emtricitabine / rilpivirine / tenofovir alafenamide - EMEA/H/C/004156

Gilead Sciences International Ltd; treatment of HIV-1

Scope: Opinion

Action: For adoption

Known active substance (Article 8(3) of Directive No 2001/83/EC)

List of Outstanding Issues adopted on 01.04.2016. List of Questions adopted on 17.12.2015.

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.

3.1.4. Ongentys - opicapone - EMEA/H/C/002790

Bial - Portela & Cª, S.A.; Parkinson's disease and motor fluctuations

Scope: Opinion

Action: For adoption

New active substance (Article 8(3) of Directive No 2001/83/EC)

List of Outstanding Issues adopted on 25.02.2016, 19.11.2015, 24.09.2015. List of Questions adopted on 23.04.2015.

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 opicapone 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 CHMP noted the letter of recommendation dated 25.04.2016.

The summary of opinion was circulated for information.

3.1.5. Sialanar - glycopyrronium bromide - PUMA - EMEA/H/C/003883

Proveca Limited; treatment of sialorrhoea

Scope: Opinion

Action: For adoption

Well-established use application (Article 10a of Directive No 2001/83/EC)

An oral explanation was held in March 2016.

List of Outstanding Issues adopted on 17.12.2015. List of Questions adopted on 25.06.2015.

The CHMP noted that there was a lack of adequate data on the medicine's risks as well as insufficient toxicology data from non-human studies. As a consequence, the data provided were not detailed enough to show that glycopyrronium bromide has been used with an acceptable level of safety to treat persistent drooling in patients with neurological conditions. Finally, the CHMP noted that Sialanar has not been shown to improve quality of life. The Committee therefore concluded that the benefits of Sialanar did not outweigh its risks and recommended that the marketing authorisation be refused.

The CHMP adopted a negative opinion by consensus recommending the refusal of the granting of paediatric use marketing authorisation together with the Assessment Report.

The Icelandic and Norwegian Members were in agreement with the CHMP recommendation.

The refusal question and answers document was circulated for information.

3.1.6. Umbipro - chlorhexidine - Article 58 - EMEA/H/W/003799

Accelerated assessment

GlaxoSmithKline Trading Services; prophylaxis of omphalitis

Scope: Opinion

Action: For adoption

Known active substance (Article 8(3) of Directive No 2001/83/EC)

List of Questions adopted on 25.02.2016.

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.

3.1.7. Zavicefta - ceftazidime / avibactam - EMEA/H/C/004027

AstraZeneca AB; treatment of complicated Intra-Abdominal Infection (cIAI), complicated Urinary Tract Infection (cUTI), including pyelonephritis, nosocomial pneumonia, including ventilator associated pneumonia (VAP), infections due to aerobic Gram-negative organisms in patients with limited treatment options

Scope: Opinion

Action: For adoption

New active substance (Article 8(3) of Directive No 2001/83/EC)

List of Outstanding Issues adopted on 25.02.2016. List of Questions adopted on 24.09.2015.

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 avibactam 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.

3.1.8. Zinbryta - daclizumab - EMEA/H/C/003862

Biogen Idec Ltd; treatment of multiple sclerosis (RMS)

Scope: Opinion

Action: For adoption

Known active substance (Article 8(3) of Directive No 2001/83/EC)

List of Outstanding Issues adopted on 17.12.2015. List of Questions adopted on 23.07.2015.

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 daclizumab is not 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 CHMP noted the letter of recommendation dated 26.04.2016.

The summary of opinion was circulated for information.

3.2. Initial applications; Day 180 list of outstanding issues

3.2.1. - atazanavir - EMEA/H/C/004048

treatment of HIV-1

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Outstanding Issues adopted on 28.01.2016, 22.10.2015. List of Questions adopted on 21.05.2015.

The Committee was reminded of the status of this application and its remaining outstanding issues.

The Committee adopted a 3rd list of outstanding issues with a specific timetable.

3.2.2. - bortezomib - EMEA/H/C/004207

treatment of multiple myeloma

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 28.01.2016.

The Committee was reminded of the status of this application and its remaining outstanding issues.

The Committee adopted a list of outstanding issues with a specific timetable.

The CHMP adopted the CHMP assessment report on similarity

3.2.3. - chenodeoxycholic acid - Orphan - EMEA/H/C/004061

Sigma-tau Arzneimittel GmbH; treatment of inborn errors of primary bile acid synthesis

Scope: Day 150 List of Outstanding Issues

Action: For adoption

List of Questions adopted on 25.02.2016.

The Committee was reminded of the status of this application and its remaining outstanding issues.

The Committee adopted a list of outstanding issues with a specific timetable.

3.2.4. - parathyroid hormone - Orphan - EMEA/H/C/003861

NPS Pharma Holdings Limited; treatment of hypoparathyroidism

Scope: 2nd Day 180 list of outstanding issue, Report from ad hoc expert group meeting held on 20 January 2016.

Action: For adoption

List of Outstanding Issues adopted on 24.09.2015. List of Questions adopted on 26.03.2015.

The members noted the report from the ad-hoc expert meeting held on 20.01.2016.

The Committee was reminded of the status of this application and its remaining outstanding issues.

The Committee adopted a list of outstanding issues .

The CHMP agreed to the request by the applicant for an extension to the clock stop to respond to the 2nd LoOI with a specific timetable.

The Committee adopted the BWP report.

3.2.5. - cediranib - Orphan - EMEA/H/C/004003

AstraZeneca AB; treatment of platinum sensitive relapsed (PSR) ovarian cancer
relapsed (PSR) ovarian cancer

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 19.11.2015.

The Committee was reminded of the status of this application and its remaining outstanding issues.

The Committee adopted a list of outstanding issues with a specific timetable.

3.2.6. - methotrexate - EMEA/H/C/003983

treatment of active rheumatoid arthritis; severe, active juvenile idiopathic arthritis; severe recalcitrant disabling psoriasis

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 17.12.2015.

The Committee was reminded of the status of this application and its remaining outstanding issues.

The Committee adopted a list of outstanding issues with a specific timetable.

3.2.7. - reslizumab - EMEA/H/C/003912

treatment of asthma and elevated blood eosinophils who are inadequately controlled on inhaled corticosteroids

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 19.11.2015.

The Committee was reminded of the status of this application and its remaining outstanding issues.

The Committee adopted a list of outstanding issues with a specific timetable.

The Committee adopted the BWP report.

3.2.8. - enoxaparin sodium - EMEA/H/C/004264

prophylaxis of thromboembolic disorders of venous origin

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 23.07.2015.

The Committee was reminded of the status of this application and its remaining outstanding issues.

The Committee adopted a list of outstanding issues .

The CHMP agreed to the request by the applicant for an extension to the clock stop to respond to the list of outstanding issues with a specific timetable.

The Committee adopted the BWP report.

3.2.9. - enoxaparin sodium - EMEA/H/C/003795

prophylaxis of thromboembolic disorders of venous origin

Scope: Day 180 list of outstanding issue

Action: For adoption

List of Questions adopted on 23.07.2015.

The Committee was reminded of the status of this application and its remaining outstanding issues,

The Committee adopted a list of outstanding issues .

The CHMP agreed to the request by the applicant for an extension to the clock stop to respond to the list of outstanding issues with a specific timetable.

The Committee adopted the BWP report.

3.2.10. - saxagliptin / dapagliflozin - EMEA/H/C/004057

treatment of type 2 diabetes

Scope: Oral explanation

Action: Oral explanation to be held on Tuesday 26 April 2016 at 16.00.

List of Outstanding Issues adopted on 25.02.2016. List of Questions adopted on 24.09.2015.

An oral explanation was held on Tuesday 26 April 2016 at 16.00. The presentation focused on the data supporting the fixed dose combination of saxagliptin and dapagliflozin.

The Committee adopted a 2nd list of outstanding issues with a specific timetable.

See also 2.1.2

3.3. Initial applications; Day 120 list of questions

3.3.1. - lonoctocog alfa - EMEA/H/C/004075

treatment of haemophilia A

Scope: Day 120 list of questions

Action: For adoption

The Committee discussed the issues identified in this application.

The Committee adopted the CHMP recommendation and scientific discussion together with the list of questions.

The Committee adopted the BWP report.

3.3.2. - adalimumab - EMEA/H/C/004212

treatment of rheumatoid arthritis, juvenile idiopathic arthritis, axial spondyloarthritis, psoriatic arthritis, psoriasis, paediatric plaque psoriasis, hidradenitis suppurativa, Crohn's disease, paediatric Crohn's disease and Ulcerative colitis

Scope: Day 120 list of questions

Action: For adoption

The Committee discussed the issues identified in this application.

It was agreed to consult SmPC Advisory Group. Furthermore a list of questions to the BMWP was adopted.

The Committee adopted the CHMP recommendation and scientific discussion together with the list of questions.

The Committee adopted the BWP report.

3.3.3. - adalimumab - EMEA/H/C/004373

treatment of rheumatoid arthritis, juvenile idiopathic arthritis, axial spondyloarthritis, psoriatic arthritis, psoriasis, paediatric plaque psoriasis, hidradenitis suppurativa, Crohn's disease, paediatric Crohn's disease and Ulcerative colitis

Scope: Day 120 list of questions

Action: For adoption

The Committee discussed the issues identified in this application.

It was agreed to consult SmPC Advisory Group.

The Committee adopted the CHMP recommendation and scientific discussion together with the list of questions.

The Committee adopted the BWP report.

3.3.4. - fluciclovine (18f) - EMEA/H/C/004197

Diagnostic agent for PET of adult men with suspected recurrence of prostate cancer

Scope: Day 120 list of questions

Action: For adoption

The Committee discussed the issues identified in this application.

The Committee adopted the CHMP recommendation and scientific discussion together with the list of questions.

3.3.5. - trientine tetrahydrochloride - Orphan - EMEA/H/C/004005

GMP-Orphan SA; Wilson’s disease

Scope: Day 120 list of questions

Action: For adoption

The Committee discussed the issues identified in this application.

The Committee adopted the CHMP recommendation and scientific discussion together with the list of questions. The Committee agreed to consult an ad-hoc expert group.

3.3.6. - emtricitabine / tenofovir disoproxil - EMEA/H/C/004215

treatment of HIV-1 infection

Scope: Day 120 list of questions,

Action: For adoption

The Committee discussed the issues identified in this application.

The Committee adopted the CHMP recommendation and scientific discussion together with the list of questions.

The Committee adopted a list of questions to the PKWP who will also liaise with the QWP core group.

3.3.7. - iloperidone - EMEA/H/C/004149

treatment of schizophrenia

Scope: Day 120 list of questions

Action: For adoption

The Committee discussed the issues identified in this application,

The Committee adopted the CHMP recommendation and scientific discussion together with the list of questions . The Committee agreed to consult the SAG on central nervous system and cardiovascular issues.

3.3.8. - insulin aspart - EMEA/H/C/004046

treatment of diabetes mellitus in adults

Scope: Day 120 list of questions

Action: For adoption

The Committee discussed the issues identified in this application

The Committee adopted the CHMP recommendation and scientific discussion together with the list of questions.

The Committee adopted the BWP report.

3.3.9. - insulin glargine - EMEA/H/C/004101

treatment of diabetes mellitus

Scope: Day 120 list of questions

Action: For adoption

The Committee discussed the issues identified in this application.

The Committee adopted the CHMP recommendation and scientific discussion together with the list of questions.

The Committee adopted the BWP report.

3.3.10. - sodium zirconium cyclosilicate - EMEA/H/C/004029

for the treatment of hyperkalaemia

Scope: Day 120 list of questions

Action: For adoption

The Committee discussed the issues identified in this application.

The Committee adopted the CHMP recommendation and scientific discussion together with the list of questions.

3.3.11. - teriparatide - EMEA/H/C/004368

treatment of osteoporosis

Scope: Day 120 list of questions

Action: For adoption

The Committee discussed the issues identified in this application.

The Committee adopted the CHMP recommendation and scientific discussion together with the list of questions.

3.3.12. - vosaroxin - Orphan - EMEA/H/C/004118

Sunesis Europe Ltd; treatment acute myeloid leukaemia

Scope: Day 120 list of questions

Action: For adoption

The Committee discussed the issues identified in this application.

The Committee adopted the CHMP recommendation and scientific discussion together with the list of questions . Furthermore the CHMP adopted the updated similarity assessment report.

3.3.13. - dimethyl fumarate - EMEA/H/C/002157

treatment of moderate to severe plaque psoriasis in adults in need of systemic drug therapy, treatment of plaque psoriasis

Scope: Day 120 list of questions

Action: For adoption

The Committee discussed the issues identified in this application.

The Committee adopted the CHMP recommendation and scientific discussion together with the list of questions.

3.3.14. - teriparatide - EMEA/H/C/003916

treatment of osteoporosis

Scope: Day 120 list of questions

Action: For adoption

The Committee discussed the issues identified in this application.

The Committee adopted the CHMP recommendation and scientific discussion together with the list of questions.

The Committee adopted the BWP report.

3.4. Update on on-going initial applications for Centralised procedure

3.4.1. lenvatinib - EMEA/H/C/004224

in combination with everolimus for the treatment of unresectable advanced or metastatic renal cell carcinoma (RCC)

Scope: Assessment Report on similarity

Action: For adoption

The CHMP adopted the similarity assessment report

3.4.2. - alendronic acid / colecalciferol - EMEA/H/C/004172

treatment of postmenopausal osteoporosis

Scope: Letter from the applicant dated 11 April 2016 requesting an extension of clock stop to submit responses to Day 120 List of Questions adopted in February 2016.

Action: For information

List of Questions adopted on 25.02.2016.

The CHMP agreed to the request by the applicant for an extension ot the clock stop to respond to the list of questions adopted in February 2016.

3.4.3. - grazoprevir / elbasvir - EMEA/H/C/004126

treatment of chronic hepatitis C (CHC) in adults

Action: For discussion

List of Outstanding Issues adopted on 01.04.2016, 25.02.2016. List of Questions adopted on 19.11.2015.

The members were updated on the status of the product

3.4.4. - irinotecan - Orphan - EMEA/H/C/004125

Baxter Innovations GmbH; treatment of pancreatic cancer

Rapporteur: Filip Josephson, Co-Rapporteur: Daniela Melchiorri, PRAC Rapporteur: Ingebjørg Buajordet

Scope: Letter from the applicant dated 15 April 2016 requesting an extension of clock stop to respond to the Day 180 list of outstanding issues adopted on 25.02.2016

Action: For adoption by written procedure

List of Outstanding Issues adopted on 25.02.2016. List of Questions adopted on 24.09.2015.

The CHMP agreed to the request by the applicant for an extension to the clock stop to respond to the list of outstanding issues via written procedure on 22.04.2016.

3.4.5. - miglustat - EMEA/H/C/004016

treatment of Gaucher disease

Scope: Response from PKWP on CHMP question

Letter from the applicant received 26 April 2016 requesting an extension of clock stop to submit responses to List of Outstanding Issues adopted on 01.04.2016.

Action: For adoption

List of outstanding issues adopted on 01.04.2016. List of Questions adopted on 23.07.2015.

The CHMP adopted the response from the PKWP and agreed to the request by the applicant for an extension to the clock stop to respond to the list of outstanding issues adopted during the March 2016 CHMP. The Committee discussed the PKWP response.

3.5. Re-examination of initial application procedures under Article 9(2) of Regulation no 726/2004

No items

3.6. Initial applications in the decision-making phase

No items

3.7. Withdrawals of initial marketing authorisation application

No items

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

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

4.1.1. Reyataz - atazanavir / atazanavir sulfate - EMEA/H/C/000494/X/0094/G

Bristol-Myers Squibb Pharma EEIG

Rapporteur: Joseph Emmerich, Co-Rapporteur: Nithyanandan Nagercoil, PRAC Rapporteur: Arnaud Batz

Scope: “An extension application covering a new pharmaceutical form (oral powder), a new strength for the oral powder presentation (50mg), and a new paediatric indication (patients from 3 months of age and weighing at least 5kg); a type II variation (C.1.6) to updated Reyataz capsules in light of new paediatric data; a type IB (C.I.11) variation to make minor revisions to the RMP with regards to nephrolithiasis, folllowing PRAC's assesment of RMP version 7.3.”

Action: For adoption

List of Outstanding Issues adopted on 28.01.2016. List of Questions adopted on 23.07.2015.

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 CHMP members were in agreement with the CHMP recommendations.

The CHMP noted the letter of recommendation dated 27.04.2016.

The summary of opinion was circulated for information.

4.1.2. Daklinza - daclatasvir - EMEA/H/C/003768/X/0013

Bristol-Myers Squibb Pharma EEIG

Rapporteur: Filip Josephson

Scope: "To add a new strength of 90 mg for Daklinza with the same pharmaceutical form (film-coated tablets) and route of administration (oral administration) as the currently approved Daklinza 30 mg and 60 mg film-coated tablets."

Action: For adoption

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 CHMP members were in agreement with the CHMP recommendations.

The summary of opinion was circulated for information.

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

No items

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

4.3.1. Stelara - ustekinumab - EMEA/H/C/000958/X/0049/G

Janssen-Cilag International N.V.

Rapporteur: Greg Markey, Co-Rapporteur: David Lyons, PRAC Rapporteur: Julie Williams

Scope: "An extension application covering a new pharmaceutical form (concentrate for solution for infusion), a new strength (130mg) and a new route of administration (intravenous use); a type II variation (C.1.6.a) to add a new indication (Crohn`s Disease)." Request for 1 year of market protection for a new indication (Article 14(11) of Regulation (EC) 726/2004)

Action: For adoption

The Committee discussed the issues identified in this application and agreed that further explanations are needed on some issues.

The Committee adopted the CHMP recommendation and scientific discussion together with the list of questions.

The CHMP agreed by consensus on the one additional year of market protection for a new indication.

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

No items

4.5. Re-examination procedure of extension of marketing authorisation according to Annex I of Commission Regulation (EC) No 1234/2008

No items

5. Type II variations - variation of therapeutic indication procedure according to Annex I of Commission Regulation (EC) No 1234/2008

5.1. Type II variations - variation of therapeutic indication procedure according to Commission Regulation (EC) No 1234/2008; Opinions or Requests for supplementary information

5.1.1. Adcetris - brentuximab vedotin - Orphan - EMEA/H/C/002455/II/0025

Takeda Pharma A/S

Rapporteur: Pieter de Graeff, Co-Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Sabine Straus

Scope: “Extension of Indication to include new indication for Adcetris (Adcetris is indicated for the treatment of adult patients at increased risk of relapse or progression following ASCT”). as a consequence, sections 4.1, 4.2, 4.4, 4.8 and 5.1 of the SmPC are updated. The Package Leaflet is updated in accordance.” Request for 1 year of market protection for a new indication (Article 14(11) of Regulation (EC) 726/2004)

Report by Steen Werner Hansen from SAG Oncology meeting held on 14th April 2016

Action: For adoption

Request for Supplementary Information adopted on 01.04.2016, 22.10.2015, 25.06.2015.

An oral explanation was held on Tuesday 26 April 2016 at 11.00.

See also 2.3.1

The CHMP noted the report from SAG Oncology meeting held on 14th April 2016. The SAG report concluded that the investigator-based PFS benefit observed with brentuximab vedotin in the study population with $\geqslant$2 risk factors was clinically relevant.

An oral explanation was held on Tuesday 26 April 2016 at 11.00. The presentation focused on the clinical data on progression free survival and overall survival as well as safety data and analysis performed. There was further discussion on the target population and on the significant clinical benefit over current standard of care.

The Committee discussed the issues identified in this application. The members discussed the clinical data and possible reasons why the effect seen on progression free survival does not show on the overall survival. The view was expressed that although no effect on the OS could be seen the totality of data could justify a positive benefit/risk. Furthermore the members discussed the appropriate target population and considerd that further discussion on this would be required.

The Committee adopted a 4th request for supplementary information with a specific timetable.

5.1.2. Afinitor - everolimus - EMEA/H/C/001038/II/0048

Novartis Europharm Ltd

Rapporteur: Harald Enzmann, Co-Rapporteur: Filip Josephson, PRAC Rapporteur: Martin Huber

Scope: "Extension of Indication to include a new indication for the treatment of unresectable or metastatic, well-differentiated non-functional neuroendocrine tumours of gastrointestinal or lung origin in adults with progressive disease; as a consequence, sections 4.1, 4.4, 4.8 and 5.1 of the SmPC are updated. The Package Leaflet is updated in accordance. Furthermore, the PI is brought in line with the latest QRD template version 9.1."

Action: For adoption

Request for Supplementary Information adopted on 25.02.2016, 19.11.2015.

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 CHMP members were in agreement with the CHMP recommendations.

The summary of opinion was circulated for information.

5.1.3. Arzerra - ofatumumab - Orphan - EMEA/H/C/001131/II/0041

Novartis Europharm Ltd

Rapporteur: Sinan B. Sarac, Co-Rapporteur: Bjorg Bolstad, PRAC Rapporteur: Doris Stenver

Scope: Opinion / Request for Supplementary Information, Report by Steen Werner Hansen from SAG Oncology meeting held on 14 April 2016

Scope: "Extension of Indication to include maintenance therapy in Chronic Lymphocytic Leukemia (CLL).
As a consequence, sections 4.1, 4.2, 4.8, 5.1, 5.2 of the SmPC are updated. The Package Leaflet is updated in accordance.
In accordance with the new QRD template version 9.1, the MAH is also taking the opportunity of this procedure to update the Annex II and combine the 2 SmPCs for the 100mg an 1,000mg vials."

Action: For adoption

Request for Supplementary Information adopted on 25.02.2016, 22.10.2015.

The CHMP noted the report from the SAG Oncology meeting held on 14 April 2016. The SAG report concluded that there are concerns about the rationale and potential added value of ofatumumab as maintenance in patients with complete repsonse/partial response. The clinical relevance of the PFS gain was considered doubtful. According to report, there was no effect on OS and QoL, and therefore the maintenance regimen was considered not justifiable. However, maintenance therapy may be considered justified in patients at high risk of symptomatic progression or death or for a treatment with a very good safety profile.

The Committee discussed the issues identified in this application related to trial population and found that there are still issues, that should be addressed by the applicant - the prognosis of the studied population, including molecular characterisation at study entry, response to previous therapy and response duration. Furthermore, the applicant should explain the efficacy and safety of ofatumumab in high-risk patients.

The Committee adopted a 3rd request for supplementary information with a specific timetable.

5.1.4. Avastin - bevacizumab - EMEA/H/C/000582/II/0086

Roche Registration Limited

Rapporteur: Sinan B. Sarac, Co-Rapporteur: Bjorg Bolstad, PRAC Rapporteur: Doris Stenver

Scope: "Extension of indication to extend the use of Avastin in combination with erlotinib for the first line treatment of patients with unresectable advanced, metastatic or recurrent non-squamous NSCLC with EGFR activating mutations. As a consequence sections 4.1, 4.2, 4.5, 4.8 and 5.1 of the SmPC are proposed to be updated. The Package Leaflet and RMP are updated in accordance."

Action: For adoption

Request for Supplementary Information adopted on 01.04.2016, 22.10.2015.

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 CHMP members were in agreement with the CHMP recommendations.

The summary of opinion was circulated for information.

5.1.5. Ferriprox - deferiprone - EMEA/H/C/000236/II/0103

Apotex Europe BV

Rapporteur: Pierre Demolis, Co-Rapporteur: Concepcion Prieto Yerro

Scope: "Extension of Indication to include a new indication for Ferriprox in combination with another chelator.
As a consequence, sections 4.1, 4.2, 4.4, 4.5, 4.8 and 5.1 of the SmPC are updated. The Package Leaflet is updated in accordance.
In addition, the MAH took the opportunity of this procedure to update the Product Information

in compliance with the QRD template version 9.1 and combine the SmPC for the 500mg and 1000mg tablets. The contact details of France and Portugal have been updated in the PL."

Action: For adoption

Request for Supplementary Information adopted on 25.02.2016, 17.12.2015.

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 CHMP members were in agreement with the CHMP recommendations.

The summary of opinion was circulated for information.

5.1.6. Gazyvaro - obinutuzumab - Orphan - EMEA/H/C/002799/II/0007

Roche Registration Limited

Rapporteur: Sinan B. Sarac, Co-Rapporteur: Pierre Demolis, PRAC Rapporteur: Julie Williams

Scope: "Extension of indication to add the treatment of patients with follicular lymphoma based on the results of the pivotal study GAO4753g. As a consequence, sections 4.1, 4.2, 4.5, 4.8, 5.1 and 5.2 of the SmPC have been updated. The Package Leaflet has been updated accordingly. Furthermore, the MAH took the opportunity to make editorial changes to sections 4.4, 4.6, 5.3 and 6.6 of the SmPC."

Action: For adoption

Request for Supplementary Information adopted on 17.12.2015.

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 CHMP members were in agreement with the CHMP recommendations.

The summary of opinion was circulated for information.

5.1.7. Humira - adalimumab - EMEA/H/C/000481/II/0146

AbbVie Ltd.

Rapporteur: Kristina Dunder, Co-Rapporteur: Daniela Melchiorri, PRAC Rapporteur: Ulla Wändel Liminga

Scope: Request for Supplementary Information / Report from ad-hoc expert group meeting held on 5 April 2016.

"Extension of Indication to include treatment of non-infectious intermediate, posterior and panuveitis in adult patients for Humira.
As a consequence, sections 4.1, 4.2, 4.4, 4.8, 5.1 and 5.2 of the SmPC are updated. The

Package Leaflet is updated in accordance."

Action: For adoption

Request for Supplementary Information adopted on 17.12.2015.

The CHMP noted the report from an ad-hoc expert group meeting held on 5 April 2016. The experts were of the view that based on the totality of the available data, a clinically relevant beneficial effect of Humira had been shown in the treatment of non-infectious uveitis. The expert group considered the use of Humira appropriate in uveitis patients who have had an inadequate response to corticosteroids, in need of corticosteroid-sparing, or in whom corticosteroid treatment is inappropriate. Also treatment duration was discussed and the experts agreed with the proposal of regular check-ups of the patients including re-evaluation of treatment continuation on a yearly basis.

The Committee discussed the issues identified in this application related to proposed indication wording. Further clarifications are needed from applicant related to PK and PK/PD modelling and efficacy.

The Committee adopted a 2nd request for supplementary information with a specific timetable.

5.1.8. HyQvia - human normal immunoglobulin - EMEA/H/C/002491/II/0021

Baxalta Innovations GmbH

Rapporteur: Jan Mueller-Berghaus, Co-Rapporteur: Andrea Laslop,

Scope: "Extension of indication to include paediatric population for all authorised indications: as a consequence, sections 4.1, 4.2, 4.4, 4.5, 4.8 and 5.1 of the SmPC are updated. The Package Leaflet is updated in accordance."

Action: For adoption

Request for Supplementary Information adopted on 28.01.2016.

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 CHMP members were in agreement with the CHMP recommendations.

The summary of opinion was circulated for information.

5.1.9. Imbruvica - ibrutinib - Orphan - EMEA/H/C/003791/II/0016

Janssen-Cilag International NV

Rapporteur: Filip Josephson, PRAC Rapporteur: Julie Williams

Scope: "Extension of indication to add the treatment of adult patients with previously untreated chronic lymphocytic leukaemia (CLL) based on the results from the final CSR of study PCYC-1115-CA (MEA 021). As a consequence, sections 4.1, 4.4, 4.6, 4.8, 5.1 and 5.3 of the SmPC have been updated. The Package Leaflet has been updated accordingly. In addition,

the Marketing authorisation holder (MAH) took the opportunity to make minor editorial changes to the SmPC. A revised version of the RMP (version 5.0.3) has been approved as part of this application.

Action: For adoption

Request for Supplementary Information adopted on 01.04.2016, 28.01.2016.

The Committee discussed the indication and suitable patient groups for first line treatment based on study data and whether indication should be restricted to patients, for whom full-dose fludarabine-based therapy is not appropriate.

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 CHMP members were in agreement with the CHMP recommendations.

The summary of opinion was circulated for information.

5.1.10. Keytruda - pembrolizumab - EMEA/H/C/003820/II/0007

Merck Sharp & Dohme Limited

Rapporteur: Daniela Melchiorri, Co-Rapporteur: Jan Mueller-Berghaus, PRAC Rapporteur: Sabine Straus

Scope: “Extension of Indication to include a new indication for Keytruda in second line Non-Small Cell Lung Cancer (NSCLC). As a consequence, sections 4.1, 4.2, 4.8, 5.1 and 5.2 of the SmPC are updated. The Package Leaflet is updated in accordance.”

Action: For adoption

The Committee discussed the issues identified in this application. The main discussion focused on the overall benefit /risk as well as the posology. Furthermore the members looked at the diagnostic assay used in the clinical trial and the one proposed as diagnostic companion.

The Committee adopted a request for supplementary information with a specific timetable.

5.1.11. Nimenrix - meningococcal group a, c, w135 and y conjugate vaccine - EMEA/H/C/002226/II/0049

Pfizer Limited

Rapporteur: Greg Markey, Co-Rapporteur: Karsten Bruins Slot, PRAC Rapporteur: Rafe Suvarna

Scope: “Extension of Indication to include a wider paediatric population starting from 6 weeks of age for Nimenrix. As a consequence, sections 4.1, 4.2, 4.5, 4.8 and 5.1 of the SmPC are updated. The Package Leaflet and the RMP are updated in accordance.”

Action: For adoption

The Committee discussed the issues identified in this application and noted that further information is needed from the applicant.

The Committee adopted Request for Supplementary Information with a specific timetable.

5.1.12. Ryzodeg - insulin degludec / insulin aspart - EMEA/H/C/002499/II/0017

Novo Nordisk A/S

Rapporteur: Kristina Dunder, PRAC Rapporteur: Qun-Ying Yue

Scope: "Extension of Indication to include paediatric population from 1 to 18 year of age for Ryzodeg. As a consequence, sections 4.1, 4.2, 4.4, 4.8, 5.1 and 5.2 of the SmPC are updated. The Package Leaflet is updated in accordance. Furthermore, the PI is brought in line with the latest QRD template version 9.1."

Action: For adoption

Request for Supplementary Information adopted on 28.01.2016.

The members discussed the outstanding issues. The main discussion focused on the use in children below the age of 6 years. The members discussed whether a list of questions was required to request clarification on the adjustment of the dosing in small children, or whether the proposed wording in the SmPC was considered sufficient and an opinion could be taken at this meeting. Although a higher rate of hypoglycaemic events was noted in the young age group, it was also considered that the drug will be prescribed for these children by specialists who would be able to make a judgement on the correct use of the drug.

The Committee adopted Request for Supplementary Information with a specific timetable.

5.1.13. Truvada - emtricitabine / tenofovir disoproxil - EMEA/H/C/000594/II/0126

Gilead Sciences International Ltd

Rapporteur: Greg Markey, Co-Rapporteur: Pierre Demolis, PRAC Rapporteur: Julie Williams

Scope: "Extension of the indication to add Pre-exposure prophylaxis (PrEP) in combination with safer sex practices to reduce the risk of sexually acquired HIV-1 in adults at high risk. As a consequence, sections 4.1, 4.2, 4.3, 4.4, 4.8, 4.9, 5.1, 5.2 and 5.3 of the SmPC are updated. The Package Leaflet is updated in accordance."

Action: For adoption

The Committee discussed the issues identified in this application. The Committee mainly discussed the posology and possible food effects. Furthermore the CHMP supported the involvement of patient representatives in this procedure.

The Committee adopted a request for supplementary information with a specific timetable.

5.1.14. Victoza - liraglutide - EMEA/H/C/001026/II/0038

Novo Nordisk A/S

Rapporteur: Pieter de Graeff, PRAC Rapporteur: Menno van der Elst

Scope: "Extension of indication to include monotherapy with liraglutide when diet and exercise

alone do not provide adequate glycaemic control in patients for whom use of metformin is considered inappropriate; additionally, the MAH updated information related to the hepatic and renal impairment. As a consequence, sections 4.1, 4.2, 5.1 and 5.2 of the SmPC have been updated with new efficacy and safety information. The Package Leaflet and RMP (v. 25.1) are updated in accordance. Furthermore, the Marketing authorisation holder (MAH) took the opportunity to align the PI with the QRD template version 9.1."

Action: For adoption

Request for Supplementary Information adopted on 01.04.2016.

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 CHMP members were in agreement with the CHMP recommendations.

The summary of opinion was circulated for information.

5.1.15. Vimpat - lacosamide - EMEA/H/C/000863/II/0060/G

UCB Pharma S.A.

Rapporteur: Filip Josephson, Co-Rapporteur: Luca Pani, PRAC Rapporteur: Qun-Ying Yue

Scope: "C.I.6.a - Extension of Indication to add a new indication as monotherapy in the treatment of partial-onset seizures . As a consequence, sections 4.1, 4.2, 4.8 and 5.1 of the SmPC are updated. The Package Leaflet is updated accordingly.

In addition, the applicant took the opportunity to update the PI in line with the latest QRD template."

Action: For adoption

The Committee discussed the issues identified in this application, which were related to the B/R of the maximum recommended maintenance dose and the proposed starting dose for the monotherapy indication.

The Committee adopted Request for Supplementary Information with a specific timetable.

5.1.16. Zinforo - ceftaroline fosamil - EMEA/H/C/002252/II/0022

AstraZeneca AB

Rapporteur: Greg Markey

Scope: "Extension of indication for Zinforo to include a new population, children from the age of 2 months; as a consequence, sections 4.1, 4.2, 4.8, 5.1, 5.2, 5.3 and 6.6 of the SmPC have been updated with new information on dosing, PK and safety. The Package Leaflet is updated in accordance. In addition, the Marketing Authorisation Holder (MAH) took the opportunity to update the list of local representatives in the Package Leaflet and to align the PI with the latest QRD template 10.0."

Action: For adoption

Request for Supplementary Information adopted on 17.12.2015, 24.09.2015.

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 CHMP members were in agreement with the CHMP recommendations.

The summary of opinion was circulated for information.

5.2. Update on on-going Type II variation; variation of therapeutic indication procedure according to Commission Regulation (EC) No 1234/2008

5.2.1. Caprelsa - vandetanib - EMEA/H/C/002315/II/0016

AstraZeneca AB

Rapporteur: Pierre Demolis,

Scope: "Extension of Indication to include paediatric indication population for Caprelsa. As a consequence, sections 4.1, 4.2, 4.6, 4.8, 5.1 and 5.2 of the SmPC are updated in update the safety information. The Package Leaflet is updated in accordance."

Scope: Letter from the MAH dated 11 April 2016 requesting an extension of timeframe to respond to the Request for Supplementary Information adopted on 01.04.2016 .

Action: For adoption

Request for Supplementary Information adopted on 01.04.2016, 19.11.2015.

The CHMP agreed to the request by the applicant for an extension to the clock stop to respond to the request for supplementary information adopted in March 2016.

5.3. Re-examination of Type II variation; variation of therapeutic indication procedure according to Commission Regulation (EC) No 1234/2008

No items

6. Ancillary medicinal substances in medical devices

6.1. Ancillary medicinal substances in medical devices; Opinions/ Day 180 list of outstanding issues / Day 120 list of questions

No items

6.2. Update of Ancillary medicinal substances in medical devices

No items

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

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

No items

8. Pre-submission issues

8.1. Pre-submission issue

8.1.1. - 177Lu-DOTA0-Tyr3-Octreotate - Orphan - H0004123

Advanced Accelerator Applications - Saint Genis Pouilly; treatment of metastatic or unresectable, well differentiated, midgut (jejunum, ileum, appendix and ascending colon) neuroendocrine tumours, which overexpresssomatostatin receptors

Scope: Letter from the company dated 12 April 2016 requesting an accelerated assessment

Action: For adoption

The CHMP agreed to the request for accelerated assessment and adopted the briefing note and Rapporteurs' recommendation on the Request for Accelerated Assessment.

8.1.2. - Recombinant Human TriPeptidyl Peptidase 1 (rhTPP1) - Cerliponase alfa - Orphan - H0004065

BioMarin International Limited; Treatment of Neuronal Ceroid Lipofuscinosis Type 2 (CLN2)

Scope: Letter from the company dated 30 March 2016 requesting an accelerated assessment

Action: For adoption

The CHMP agreed to the request for accelerated assessment and adopted the briefing note and Rapporteurs' recommendation on the Request for Accelerated Assessment.

8.1.3. - Neisseria meningitidis serogroup B recombinant lipoprotein (rLP2086; subfamily A; Escherichia coli),Neisseria meningitidis serogroup B recombinant lipoprotein (rLP2086; subfamily B; Escherichia coli) - H0004051

prevention of invasive meningococcal disease caused by Neisseria meningitidis (IMD) serogroup B in individuals 10 years and older.

Scope: Letter from the company dated 12 April 2016 requesting an accelerated assessment

Action: For adoption

The CHMP did not agree to the request for accelerated assessment and adopted the briefing note and Rapporteurs' recommendation on the Request for Accelerated Assessment.

8.2. Priority Medicines (PRIME)

Disclosure of information related to priority medicines cannot be released at present time as these contain commercially confidential information

Scope: List of applications received

Action: For information

The CHMP noted the list of applications received.

9. Post-authorisation issues

9.1. Post-authorisation issues

9.1.1. Translarna - ataluren – Orphan - EMEA/H/C/002720/R/0022

MAH: PTC Therapeutics International Limited, treatment of Duchenne muscular dystrophy

Rapporteur: Johann Lodewijk Hillege, Co-Rapporteur: Concepcion Prieto Yerro, PRAC Rapporteur: Sabine Straus

Scope: Request for supplementary information / Opinion

Renewal of Conditional Marketing Authorisations

Action: For discussion

The CHMP was updated on the procedure and noted the timetable for the on-going type II variation II/12.

The CHMP adopted a Request for Supplementary information with a specific timetable.

The CHMP rediscussed the timing of the planned SAG meeting.

9.1.2. Taxotere – docetaxel – EMEA/H/C/000073

MAH: Aventis Pharma S.A., Rapporteur: Pierre Demolis, Co-Rapporteur: Harald Enzmann, PRAC Rapporteur: Martin Huber

Scope: DHPC and communication plan

Action: For adoption

The CHMP agreed to the wording of the DHPC letter and the communication plan.

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

10.1.1. Invokana - canagliflozin, Vokanamet - canagliflozin / metformin - (EMEA/H/A-20/1442)

Janssen-Cilag International N.V.; treatment of type 2 diabetes mellitus

Rapporteurs for the Article 20 procedure: Rapporteur: Valerie Strassmann, Co-Rapporteur: Menno van der Elst

CHMP Rapporteurs for Ivokana and Vokanamet: Rapporteur: Martina Weise, Co-Rapporteur: Kristina Dunder

Scope: Article 20 procedure to PRAC was triggered by European Commission on 15 April 2016, Signal of potential increased risk of lower limb amputations

Action: For information

DHPC and communication plan recommended by the PRAC for the signal adopted by written procedure on 15 April 2016.

The CHMP noted the documents adopted via written procedure on 15.04.2016.

10.1.2. Zydelig - idelalisib - (EMEA/H/A-20/1439)

Gilead Sciences International Ltd; treatment of chronic lymphocytic leukaemia (CLL) and follicular lymphoma (FL).

Rapporteurs for the Article 20 procedure: Rapporteur: Rafe Suvarna, Co-Rapporteur: Ulla Waendel Liminga

CHMP Rapporteurs for Zydelig: Rapporteur: Filip Josephson, Co-Rapporteur: Pieter de Graeff

Scope: PRAC list of questions to the IC SAG Oncology , draft list of experts

Action: For information

The CHMP noted the list of questions to the IC SAG Oncology as adopted by the PRAC and the draft list of experts.

10.1.3. Direct-acting antivirals (DAAV) indicated for the treatment of hepatitis C (interferon free - EMEA/H/A-20/1438

Daklinza - Daclatasvir Dihydrochloride - EMEA/H/C/003768

MAH: Bristol-Myers Squibb Pharma EEIG, Rapporteur: Filip Josephson, Co-Rapporteur: Robert James Hemmings,

Exviera - Dasabuvir Sodium - EMEA/H/C/003837

MAH: AbbVie Ltd., Rapporteur: Filip Josephson, Co-Rapporteur: Johann Lodewijk Hillege,

Viekirax - Ombitasvir,Paritaprevir,Ritonavir - EMEA/H/C/003839

MAH: AbbVie Ltd., Rapporteur: Filip Josephson, Co-Rapporteur: Johann Lodewijk Hillege,

OLYSIO - Simeprevir - EMEA/H/C/002777

MAH: Janssen-Cilag International N.V., Rapporteur: Arantxa Sancho-Lopez, Co-Rapporteur: Daniela Melchiorri,

Sovaldi - Sofosbuvir - EMEA/H/C/002798

MAH: Gilead Sciences International Ltd, Rapporteur: Filip Josephson, Co-Rapporteur: Alar Irs,

Harvoni - Ledipasvir,Sofosbuvir - (EMEA/H/C/003850)

MAH: Gilead Sciences International Ltd, Rapporteur: Filip Josephson, Co-Rapporteur: Joseph Emmerich,

PRAC Rapporteur: Margarida Guimarães; PRAC Co-rapporteur: Dolores Montero Corominas

Scope: Article 20 procedure to PRAC was triggered by European Commission on 17 March 2016, Extension of the scope for the review of the benefit-risk balance of DAAV following notification by the European Commission of a referral under Article 20 of Regulation (EC) No 726/2004, based on pharmacovigilance data

Action: For information

The CHMP noted the status of the PRAC referral.

10.2. Requests for CHMP Opinion under Article 5(3) of Regulation (EC) No 726/2004

No items

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

No items

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

10.4.1. Otipax 1% (11 mg/ml) ear drops, solution – lidocaine hydrochloride - EMEA/H/A-29/1426

Biocodex Benelux SA/NV

Rapporteur: Daniel Brasseur, Co-Rapporteur: Martina Weise,

RMS: BE, CMS: AT, CY, DE, DK, EL, ES, FI, IT, NO, PL, PT, SE, Mutual recognition procedure number: BE/H/0213/001/MR

Scope: Oral explanation and Opinion

Disagreement regarding efficacy and the evidence of well-established use

Action: For adoption

List of Questions adopted on 22 October 2015. List of outstanding issues adopted on 25.02.2016.

Oral explanation was held on Tuesday 26 April 2016 at 9.00. The company's presentation focused on publications on the local use of lidocaine, the formulation of Otipax and its safety.

The Committee discussed the information available from studies and recommendations of clinical guidelines. The Committee had different views on B/R of the product with regard to the indication of short-term treatment of pain of the external ear canal, and some members were of the opinion that the published evidence was limited to support this indication.

The CHMP noted that the MAH withdrew the marketing authorisation in the RMS Belgium and all the marketing authorisation applications in the involved CMS.

See also 2.4.1

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

10.5.1. Durogesic transdermal patches – fentanyl - EMEA/H/A-30/1413

Janssen-Cilag group of companies and associated companies

Rapporteur: Johann Lodewijk Hillege, Co-Rapporteur: Martina Weise,

Scope: List of outstanding Issues

Action: For adoption

List of outstanding Issues adopted 28 January 2016.

The Committee discussed issues related section 4.2 posology: adjustments in equianalgesic potency conversion table and section on opioid-naïve patients.

The CHMP agreed to consult the SmPC Advisory Group on SmpC section 4.2 related to expression of equianalgesic potency conversion.

The CHMP adopted a list of outstanding Issues with a specific timetable.

Submission of responses: 23.05.2016

Re-start of the procedure: 26.05.2016

Joint assessment report circulated to CHMP: 08.06.2016

Comments: 13.06.2016

Updated joint assessment report circulated to CHMP: 16.06.2016

CHMP opinion: June 2016 CHMP

10.5.2. Haldol and associated names (EMEA/H/A-30/1393) (haloperidol)

Janssen-Cilag Group of companies and associated companies

Rapporteur: Martina Weise, Co-Rapporteur: Katarina Vučić,

Scope: List of Questions to SAG, Letter from the MAH dated 14 April 2016 requesting an extension of time frame to respond to the List of outstanding issues adopted during the March 2016 CHMP.

Action: For adoption

The CHMP adopted a revised timetable for the submission of the responses to the list of outstanding issues. The CHMP also adopted List of Questions to SAG Psychiatry.

Submission of responses: 28.07.2016

Re-start of the procedure: 15.09.2016

Joint assessment report circulated to CHMP: 28.09.2016

Comments: 03.10.2016

Updated joint assessment report(s) circulated to CHMP: 06.10.2016

CHMP list of outstanding issues or CHMP opinion: October 2016 CHMP

10.5.3. Haldol decanoate and associated names (EMEA/H/A-30/1405) (haloperidol) Janssen-Cilag Group of companies and associated companies

Rapporteur: Martina Weise, Co-Rapporteur: Katarina Vučić,

Scope: Letter from the MAH dated 14 April 2016 requesting an extension of time frame to respond to the List of outstanding issues adopted during the March 2016 CHMP.

Action: For adoption

The CHMP adopted a revised timetable for the submission of the responses to the list of outstanding issues.

Submission of responses: 28.07.2016

Re-start of the procedure: 15.09.2016

Joint assessment report circulated to CHMP: 28.09.2016

Comments: 03.10.2016

Updated joint assessment report(s) circulated to CHMP: 06.10.2016

CHMP list of outstanding issues or CHMP opinion: October 2016 CHMP

10.5.4. Lovenox and associated names – enoxaparin - EMEA/H/A-30/1429

Sanofi Aventis group of companies and associated companies

Rapporteur: Joseph Emmerich, Co-Rapporteur: Pieter de Graeff,

Scope: List of outstanding Issues

Harmonisation exercise for Lovenox and associated names. The review was triggered by France, due to the need of harmonisation of the Summary of Product Characteristics across Member States.

List of Questions adopted on 19.11.2015.

Action: For adoption

The members discussed several aspects of the SmPC, mainly concerning the expression of strength, the risk minimisation of medication errors, the use in patients with severe renal impairment, and the use in patients with a history of heparin-induced thrombocytopenia.

The CHMP adopted a list of outstanding issues with a specific timetable.

Submission of responses: 09.06.2016

Re-start of the procedure: 23.06.2016

Rapporteur / co-rapporteur joint assessment report circulated to CHMP: 06.07.2016

Comments: 11.07.2016

Updated rapporteur / co-rapporteur joint assessment report circulated to CHMP: 14.07.2016

Adoption of list of outstanding issues / CHMP opinion: July 2016 CHMP

10.5.5. Novantrone and associated names - mitoxantrone - EMEA/H/A-30/1399

MEDA group of companies and associated companies

Rapporteur: Pieter de Graeff, Co-Rapporteur: Robert Hemmings,

Scope: Opinion

Harmonisation exercise for Novantrone and associated names. The review was triggered by the European Commission, due to the need of harmonisation of the Summary of Product Characteristics across Member States.

Action: For adoption

Scientific Advisory Group meeting held on 06.11.2015. List of outstanding issues adopted on 28.01.2016, 19.11.2015, 23.07.2015, 26.03.2015.

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.

The Icelandic and Norwegian CHMP members were in agreement with the CHMP recommendations.

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

10.6.1. Inhaled corticosteroids (ICS)-containing medicinal products indicated in the treatment of chronic obstructive pulmonary disease: beclomethasone (NAP); beclomethasone, formoterol (NAP); budesonide (NAP); budesonide, formoterol – BIRESP SPIROMAX (CAP); BUDESONIDE FORMOTEROL TEVA (CAP); DUORESP SPIROMAX (CAP); VYALER SPIROMAX (CAP); flunisolide, salbutamol (NAP); fluticasone (NAP); fluticasone, salmeterol (NAP); fluticasone, vilanterol – RELVAR ELLIPTA (CAP); REVINTY ELLIPTA (CAP) – EMEA/H/A-31/1415

Applicant: Glaxo Group Ltd, Teva Pharma B.V., Teva Pharmaceuticals Europe, various

PRAC Rapporteur: Rafe Suvarna; PRAC Co-rapporteur: Jan Neuhauser

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

A referral procedure under Article 31 of Directive 2001/83/EC for inhaled corticosteroids (ICS)-containing products (beclomethasone; budesonide; flunisolide; fluticasone proprionate; fluticasone furoate) reviewing the risk of pneumonia in patients with chronic obstructive pulmonary disease (COPD) is to be concluded.

Scope: Opinion

Action: For adoption

The Committee confirmed that all issues previously identified in this referral had been addressed. The CHMP noted the PRAC recommendation.

The CHMP adopted an opinion by consensus recommending changes to the SmPCs, labelling

and package leaflets. The assessment report was adopted.

The Icelandic and Norwegian Members were in agreement with the CHMP recommendation.

10.6.2. Alkem Laboratories Limited, India - EMEA/H/A-31/1436

Rapporteur: Harald Enzmann, Co-Rapporteur: Greg Markey,

Scope: Letter from Krka d.d. dated 22.04.2016 requesting extension of timeframe to submit responses to the list of questions adopted 1 April 2016

Action: For adoption

The CHMP did not agree to the request by the applicant for an additional extension to the clock stop to respond to the list of questions adopted during the March 2016 CHMP as the Committee considered that the originally adopted timelines should remain.

10.6.3. Semler Research Centre Private Ltd - EMEA/H/A-31/1443

Rapporteur: Pieter de Graeff, Co-Rapporteur: Concepcion Prieto-Yerro,

Scope: Appointment of (Co)Rapporteur, List of Questions and timetable

Article 31 triggered by the the UK, Germany, Spain, Denmark and the Netherlands in relation to findings of non-compliance with GCP at the Semler bioanalytical and clinical facilities in Bangalore, India.

Action: For adoption

The CHMP noted the letters from the MHRA, BfArM, AEMPS and the MEB dated 27 April 2016 and the letter from the Danish Medicines Agency dated 28 April 2016 notifying of a official referral under Article 31 and its grounds.

The CHMP appointed Pieter de Graeff as Rapporteur (interest level 1) and Concepcion Prieto-Yerro as Co-Rapporteur (interest level 1).

Ther CHMP adopted a list of questions with a specific timetable.

Submission of responses: 09.06.2016

Re-start of the procedure: 23.06.2016

Rapporteur/co-rapporteur assessment report(s) circulated to CHMP: 06.07.2016

Comments: 11.07.2016

Updated Rapporteur/co-rapporteur assessment reports circulated to CHMP: 14.07.2016

CHMP LoOI/ opinion: July 2016 CHMP

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

No items

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

No items

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

No items

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

No items

10.11. 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

11. Pharmacovigilance issue

11.1. Early Notification System

April 2016 Early Notification System on envisaged CHMP/CMDh outcome accompanied by communication to the general public

Action: For information

The CHMP noted the information.

12. Inspections

12.1. GMP inspections

Disclosure of information related to GMP inspections will not be published as it undermines the purpose of such inspections

12.2. GCP inspections

Disclosure of information related to GCP inspections will not be published as it undermines the purpose of such inspections

12.3. Pharmacovigilance inspections

Disclosure of information related to Pharmacovigilance inspections will not be published as it undermines the purpose of such inspections

12.4. GLP inspections

Disclosure of information related to GLP inspections will not be published as it undermines the purpose of such inspections

13. Innovation Task Force

13.1. Minutes of Innovation Task Force

Action: For information

The CHMP noted the minutes.

13.2. Innovation Task Force briefing meetings

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

Scope: ITF Briefing Meeting

Meeting date: 13th May 2016

Action: For adoption

The CHMP agreed to the meeting.

Scope: ITF Briefing Meeting

Meeting date: 10th May 2016

Action: For adoption

The CHMP agreed to the meeting.

Scope: ITF Briefing Meeting

Action: For adoption

The CHMP agreed to the meeting.

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

No items

13.4. Nanomedicines activities

No items

14. Organisational, regulatory and methodological matters

14.1. Mandate and organisation of the CHMP

14.1.1. Discussion on area of expertise of Co-opted Member

Scope : Area of expertise of Co-opted Member

Action: For discussion

Mandate of Sol Ruiz and Jean-Louis Robert expires in July 2016. Sol Ruiz's area of expertise has been in Quality (biotech and biological), with expertise in advanced therapies (gene, cell and tissue therapies) and Jean-Louis Robert's in Quality (non-biologicals, synthetic chemicals).

The CHMP was informed about the upcoming expiry of mandates of the two CHMP co-opted members Sol Ruiz and Jean-Louis Robert. The Committee was asked to consider the current expertise of the CHMP for discussion and agreement on the required areas of expertise for the co-opted members during the May 2016 CHMP Plenary.

14.1.2. Joint CHMP and COMP strategic review and learning meeting to be held in Utrecht on 31 May-1 June 2016

CHMP: Pieter De Graeff

Scope: Agenda

Action: For information

The CHMP noted the draft agenda for meeting in Utrecht on 31 May-1 June 2016.

14.1.3. Best Practice Guide for CHMP plenaries

Action: For adoption

The CHMP noted the Best Practice Guide. Further comments are awaited from members and their assessment teams and comments should be sent by 16 May 2016.

14.1.4. New procedure for assessing imposed PASS final study results for CAPs and NAPs under Art 107q of Directive 2010/84/EU

Scope: Appointment of CHMP sponsor

Action: For discussion

The CHMP noted that a new procedure for assessing non-interventional imposed PASS final study results (CAPs and NAPs) which fall under Article 107q (Directive 2010/84/EU) is to be developed. For the implementation of the new procedure for assessing imposed PASS Results 107q, CHMP sponsor was invited to be involved.

14.2. Coordination with EMA Scientific Committees

14.2.1. Pharmacovigilance Risk Assessment Committee (PRAC)

Summary of recommendations and advice of PRAC meeting held on **11-14 April 2016**

Action: For information

The CHMP noted the Summary of recommendations and advice.

List of Union Reference Dates and frequency of submission of Periodic Safety Update Reports (EURD list) for April 2016

Action: For adoption

The CHMP adopted the list.

14.2.2. Committee for Advanced Therapies (CAT)

CAT draft minutes of meeting held on 21-22 April 2016
Action: For information

The CHMP noted the minutes.

14.2.3. Committee for Herbal Medicinal Products (HMPC)

Report from the HMPC meeting held on 4-7 April 2016

Action: For information

The CHMP noted the report.

14.2.4. Paediatric Committee (PDCO)

PIPs reaching D30 at April 2016 PDCO

Action: For information

The CHMP noted the information.

Report from the PDCO meeting held on 27-29 April 2016

Action: For information

The CHMP noted the report.

14.2.5. Committee for Orphan Medicinal Products (COMP)

Report from the COMP meeting held on 19-21 April 2016
Action: For information

The CHMP noted the report.

14.2.6. CMDh

Report from the Coordination Group for Mutual Recognition and Decentralised Procedures – Human (CMDh) on the meeting held on 25-27 April 2016

Action: For information

The CHMP noted the report.

Scope: Letter from CHMh dated 21 April 2016 regarding regarding administration of crushed/disintegrated tablets, List of questions to PKWP

Action: For adoption

The CHMP adopted the List of questions to PKWP.

Letter from the CMDh dated 22 April 2016 regarding low dose acetylsalicylic acid gastro-resistant formulations in fixed dose combinations with substitution indication, List of questions to the PKWP

Action: For adoption

The CHMP adopted the List of questions to PKWP.

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

14.3.1. Scientific Advice Working Party (SAWP)

Report from the SAWP meeting held on 11-14 April 2016. Table of conclusions

Action: For information

Scientific advice letters: The CHMP noted the report.

Disclosure of information related to scientific advice letters cannot be released at present time as these contain commercially confidential information.

14.3.2. Vaccines Working Party (VWP)

Vice-Chair: Daniel Brasseur (Acting as Chair)

Scope: Nomination of new core members:

- Current membership list

Action: For adoption

The CHMP appointed 4 experts as new core members to VWP: Isabelle Bekeredjian-Ding (PEI(DE)), Svein Rune Andersen (NO), Agustín Portela Moreira (ES) and Nele Berthels (BE). Furthermore additional expert to VWP was appointed - Kaatje Smith (BE).

The CHMP appointed Claudia Gramiccioni (IT) as observer to VWP.

14.3.3. Pharmacogenomics Working Party (PGWP)

Chair: Krishna Prasad

Scope: Guideline on good pharmacogenomic practice (EMA/CHMP/268544/2016)

Action: For adoption for 5-months public consultation

The CHMP adopted the guideline for 5-months public consultation.

14.3.4. Name Review Group (NRG)

Table of Decisions of the NRG plenary meeting held on 6 April 2016

Action: For adoption

The CHMP adopted the Table of Decisions.

14.3.5. Safety Working Party (SWP)

Scope: Concept paper on the revision of the guideline on the environmental risk assessment

Action: For adoption for 6-month public consultation

The CHMP adopted the concept paper for 6-month public consultation.

14.4. Cooperation within the EU regulatory network

No items

14.5. Cooperation with International Regulators

No items

14.6. Contacts of the CHMP with external parties and interaction with the Interested Parties to the Committee

No items

14.7. CHMP work plan

No items

14.8. Planning and reporting

No items

14.9. Others

No items

15. Any other business

15.1.1. Update on Ebola, Zika virus and other issues

Action: For information

The CHMP noted the update.

16. List of participants

List of participants including any restrictions with respect to involvement of members/alternates/ experts following evaluation of declared interests for the 25 – 28 April 2016 meeting.

Name	Role	Member State or affiliation	Outcome restriction following evaluation of e-DoI	Topics on agenda for which restrictions apply
Tomas Salmonson	Chair	Sweden	No interests declared	
Andrea Laslop	Member	Austria	No interests declared	
Milena Stain	Alternate	Austria	No interests declared	
Daniel Brasseur	Member	Belgium	No interests declared	
Bart Van der Schueren	Alternate	Belgium	No interests declared	
Mila Vlaskovska	Member	Bulgaria	No interests declared	
Ines Baotic	Member	Croatia	No restrictions applicable to this meeting	
Katarina Vučić	Alternate	Croatia	No interests declared	
Ondřej Slanař	Member	Czech Republic	No interests declared	
Radka Montoniová	Alternate	Czech Republic	No interests declared	
Sinan B. Sarac	Alternate	Denmark	No interests declared	
Alar Irs	Member	Estonia	No restrictions applicable to this meeting	
Outi Mäki-Ikola	Member	Finland	No restrictions applicable to this meeting	
Tuomo Lapveteläinen	Alternate	Finland	No interests declared	

Name	Role	Member State or affiliation	Outcome restriction following evaluation of e-DoI	Topics on agenda for which restrictions apply
Pierre Demolis	Member (Vice-Chair)	France	No interests declared	
Joseph Emmerich	Alternate	France	No interests declared	
Harald Enzmann	Member	Germany	No interests declared	
Martina Weise	Alternate	Germany	No restrictions applicable to this meeting	
Dimitrios Kouvelas	Member	Greece	No interests declared	
George Aislaitner	Alternate	Greece	No interests declared	
Agnes Gyurasics	Member	Hungary	No interests declared	
Kolbeinn Gudmundsson	Member	Iceland	No interests declared	
David Lyons	Member	Ireland	No restrictions applicable to this meeting	
Patrick Salmon	Alternate	Ireland	No interests declared	
Daniela Melchiorri	Member	Italy	No interests declared	
Natalja Karpova	Alternate	Latvia	No interests declared	
Romaldas Mačiulaitis	Member	Lithuania	No restrictions applicable to this meeting	
Jacqueline Genoux-Hames	Member	Luxembourg	No interests declared	
John Joseph Borg	Member	Malta	No interests declared	
Pieter de Graeff	Member	Netherlands	No interests declared	

Name	Role	Member State or affiliation	Outcome restriction following evaluation of e-DoI	Topics on agenda for which restrictions apply
Johann Lodewijk Hillege	Alternate	Netherlands	No interests declared	
Karsten Bruins Slot	Member	Norway	No interests declared	
Bjorg Bolstad	Alternate	Norway	No restrictions applicable to this meeting	
Piotr Fiedor	Member	Poland	No interests declared	
Aldona Paluchowska	Alternate	Poland	No interests declared	
Bruno Sepodes	Member	Portugal	No interests declared	
Nela Vilceanu	Member	Romania	No interests declared	
Jan Mazag	Member	Slovakia	No interests declared	
Jana Schweigertova	Alternate	Slovakia	No restrictions applicable to this meeting	
Nevenka Tršinar	Alternate	Slovenia	No interests declared	
Concepcion Prieto Yerro	Member	Spain	No interests declared	
Arantxa Sancho-Lopez	Alternate	Spain	No restrictions applicable to this meeting	
Kristina Dunder	Member	Sweden	No interests declared	
Filip Josephson	Alternate	Sweden	No interests declared	
Greg Markey	Member	United Kingdom	No interests declared	
Nithyanandan Nagercoil	Alternate	United Kingdom	No restrictions applicable to this	

Name	Role	Member State or affiliation	Outcome restriction following evaluation of e-DoI	Topics on agenda for which restrictions apply
			meeting	
Robert James Hemmings	Co-opted member	United Kingdom	No restrictions applicable to this meeting	
Koenraad Norga	Co-opted member	Belgium	No restrictions applicable to this meeting	
Jan Mueller-Berghaus	Co-opted member	Germany	No interests declared	
Jean-Louis Robert	Co-opted member	Luxembourg	No interests declared	
Sol Ruiz	Co-opted member	Spain	No interests declared	
Ana Alonso Gutierrez	Expert - in person*	Spain	No interests declared	
Christophe Focke	Expert - in person*	Belgium	No interests declared	
Kristina Bech Jensen	Expert - in person*	Denmark	No interests declared	
Maarit Gustafsson	Expert - in person*	Finland	No interests declared	
Taru Kuittinen	Expert - in person*	Finland	No participation in discussions, final deliberations and voting on:	5.1.4. Avastin - bevacizumab, EMEA/H/C/000582/II/0086 5.1.6. Gazyvaro - obinutuzumab - Orphan, EMEA/H/C/002799/II/0007
Hanne Lomholt Larsen	Expert - via telephone*	Denmark	No interests declared	
Johannes Pohly	Expert - via telephone*	Germany	No interests declared	
Maarten Lagendijk	Expert - via telephone*	Netherlands	No interests declared	

Name	Role	Member State or affiliation	Outcome restriction following evaluation of e-DoI	Topics on agenda for which restrictions apply
Karoline Buhre	Expert - via telephone*	Germany	No interests declared	
Paula Boudewina van Hennik	Expert - in person*	Netherlands	No interests declared	
Sabine Mayrhofer	Expert - in person*	Germany	No interests declared	
Janet Schriever	Expert - via telephone*	Germany	No interests declared	
Rafe Suvarna	Expert - via telephone*	United Kingdom	No interests declared	
Jo Lyn Chooi	Expert - via telephone*	United Kingdom	No restrictions applicable to this meeting	
Else Carrière	Expert - via telephone*	Netherlands	No interests declared	
Steen Werner Hansen	Expert - via telephone*	Denmark	No interests declared	
Nathalie Morgensztejn	Expert - via telephone*	France	No interests declared	
Carolien Versantvoort	Expert - via telephone*	Netherlands	No interests declared	
Representative from the European Commission attended the meeting				
Meeting run with support from relevant EMA staff				

* Experts were only evaluated against the product(s) they have been invited to talk about.

17. 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 2)*

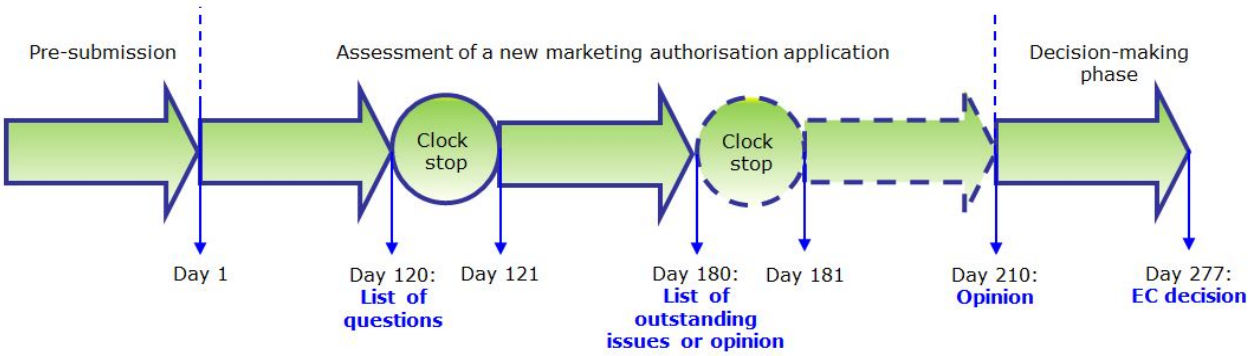
The items listed in this section are those for which marketing authorisation holders (MAHs) or applicants have been invited to the CHMP plenary meeting to address questions raised by the Committee. Oral explanations normally relate to on-going applications (section 3, 4 and 5) or referral procedures (section 10) but can relate to any other issue for which the CHMP would like to discuss with company representatives in person.

Initial applications *(section 3)*

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

Section 3.1 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 3.2 (**Day 180 List of outstanding issues**) and 3.3 (**Day 120 list of questions**).

CHMP discussions may also occur at any other stage of the evaluation, and these are listed under section 3.4, **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 3.6, **products in the decision making phase**.

Extension of marketing authorisations according to Annex I of Reg. 1234/2008 *(section 4)*

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 5)*

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 4. 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 6)*

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 3.5)*

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 *(section5.3)*

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 3.7)*

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 7)*

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 8)*

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.

PRIME

PRIME is a scheme launched by the European Medicines Agency (EMA) to enhance support for the development of medicines that target an unmet medical need. This voluntary scheme is based on enhanced interaction and early dialogue with developers of promising medicines, to optimise development plans and speed up evaluation so these medicines can reach patients earlier

Post-authorisation issues *(section 9)*

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 10)*

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 11)*

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 12)*

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 13)*

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 14.3.1)*

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 14.2)*

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 14.3)*

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.

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