05 September 2022
EMA/CHMP/670236/2022
Human Medicines Division

Committee for medicinal products for human use (CHMP)

PROM[1] agenda for the meeting on 5 September 2022

Chair: Harald Enzmann – Vice-Chair: Bruno Sepodes

5 September 2022, 09:00–16:00, virtual meeting/room 08-A

Disclaimers

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Of note, agendas and minutes are working documents primarily designed for CHMP members and the work the Committee undertakes.

Note on access to documents

Some documents mentioned in the agenda/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).

[1] The CHMP Preparatory and Organisational matters (PROM) is a meeting to discuss CHMP organisational matters and other topics in preparation for the CHMP Plenary meeting. It is a virtual meeting, which usually takes place on Monday before the CHMP Plenary meeting. CHMP members, working party chairs and national experts together with EMA staff are participating in this forum. Depending on the nature of the issue and availability of documents and experts some PROM topics can be discussed at the CHMP Plenary.

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1. Agenda and Minutes

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

1.2. Adoption of agenda

CHMP PROM agenda for 5 September 2022 meeting.

1.3. Adoption of the minutes

CHMP PROM Minutes of 5 September 2022 meeting will be adopted at the September CHMP plenary.

2. Quality Domain

2.1. Biologics Working Party (BWP)

Chairs: Sol Ruiz, Sean Barry

2.1.1. Agenda and minutes

- Draft agenda for BWP meeting to be held virtually on 5-7 September 2022
- Final minutes for BWP meeting held virtually on 13-15 June 2022

Action: For information

2.1.2. EDQM letters to BWP and CMDh requesting input on the limit for residual protein in monographs of unfractionated heparin (Heparin sodium (0333) and Heparin calcium (0332)

Proposal to continue scientific discussion at BWP and prepare a joint response from CMDh and CHMP (BWP) to EDQM, for requested deadline of 30 November 2022.

Action: For information

2.1.3. PMF dossier requirements. Questions and Answers for PMF Holders

This Q&A supplements the data requirements in the published guidelines on Plasma Master File (PMF) Scientific requirements (EMEA/CHMP/BWP/3794/03) and dossier requirements on PMF epidemiological data (EMA/CHMP/BWP/548524/2008) on topics related to inspection approval status, information on audits and epidemiology alert limits.

Action: For adoption

2.2. Quality Working Party (QWP)

Chair: Blanka Hirschlerova

2.2.1. Agenda and minutes

- Final agenda for QWP-CT meeting held virtually on 13 July 2022
- Final minutes for QWP-CT meeting held virtually on 13 July 2022

Action: For information

2.2.2. Guideline on the development and manufacture of synthetic peptides: concept paper for public consultation

The 3-year workplan of QWP foresees the preparation of a guideline (scope includes human and veterinary medicinal products) on the development and manufacture of synthetic peptides, and a guideline on the development and manufacture of synthetic oligonucleotides. In view of this, two concept papers were prepared by the drafting group, discussed in QWP and CVMP and are now proposed for adoption. A 3-month public consultation for the concept papers is proposed.

CHMP: Blanka Hirschlerova

Action: For adoption

2.2.3. Guideline on the development and manufacture of synthetic oligonucleotides: concept paper for public consultation

See 2.2.2

A 3-month public consultation for the concept papers is proposed.

CHMP: Blanka Hirschlerova

Action: For adoption

2.2.4. Nomination of new alternate to the QWP

Nomination of new QWP alternate to replace Ewa Wyzsecka-Kaszuba representing Poland.

Action: For endorsement

2.2.5. Nomination of new member to the QWP

Nomination of new QWP member to replace Kim Notenboom representing the Netherlands.

Action: For endorsement

2.3. Biosimilar Medicinal Product Working Party (BMWP)

No topics

2.4. Quality Innovation Group (QIG)

No topics

2.5. Formulation Expert Group (FEG)

No topics

3. Non-Clinical Domain

3.1. Non-Clinical Working Party (NCWP)

Chairs: Susanne Brendler-Schwaab, Karen van Malderen

3.1.1. Agenda and minutes

- Draft minutes for NcWP meeting held virtually on 12-13 July 2022
- Draft agenda for NcWP meeting to be held virtually on 1 and 7 September 2022

Action: For information

3.1.2. CMDh questions to NcWP on new nitrosamines

CMDh requests that the NcWP determines the acceptable intake for the following nitrosamines based on lifetime daily exposure including information on the points of departure and methodology used.

- N-nitroso-moxifloxacin
- N-nitroso-fluoxetine
- N- nitroso-nebivolol
- N-nitroso-bisoprolol
- N-nitroso-biotin
- N-nitroso-diclofenac
- N-Nitroso-betahistine
- N-nitroso-Bupropion

Action: For adoption

3.1.3. Third party request for clarifications on the SWP response to CMDh on genotoxicity and contraception

In February 2020, based on a request from CMDh, the Safety working party (SWP) published advice on the duration of contraception in male and female patients after cessation of treatment with a genotoxic drug in the context both of clinical trial applications as well as marketing authorisation applications (EMA/CHMP/SWP/74077/2020). In July 2022, the CMDh and EMA have received questions from a third party for which NcWP view is sought.

Action: For adoption

3.1.4. Creation of an excipients drafting group

The Non-clinical domain governance adopted the creation of an ad hoc excipients drafting group for finalising the excipients safety information documents which have been on hold due to the Business Contingency Plan. The following excipients will be covered:

- Lactose
- Dextrans
- Polysorbate
- Proline

A kick-off meeting will take place in September

Action: For information

3.1.5. New nomination in the ERA drafting group

Following the stepping away from Caroline Moermond, nomination of a member of the ERA drafting group to finalise the revision of the Environmental Risk Assessment guideline.

Action: For endorsement

3.1.6. Non-clinical software

EMA has acquired non-clinical on behalf of NCAs for an initial 5-year period. These tools are widely used for non-clinical and quality applications and include in silico prediction software for toxicity, metabolic fate and chemical degradation of chemicals, chemical database software for chemical toxicity information management and software for the calculation of purge factors. In order to install the software in the NCAs and get credentials, every NCA should confirm to NcWP secretariat their interest in using the software. A training will be organised and made available via the EU NTC to all assessors.

Action: for information

3.2. Joint 3Rs Replacement, Reduction and Refinement Working Party (3Rs)

Chair(s): Vacant

3.2.1. Nomination of the J3RsWP Chair and Vice-chair

Upon constitution of the J3RsWP following the implementation of the new Working Parties Operating Model (WOM), a first meeting was held on 18 July 2022 to elect the Chair and Vice-Chair. During the meeting, it was agreed to select a Chair from the area of human medicinal products, while the Vice-Chair would represent the area of veterinary products. The following members were elected:

- Dr Sonja Beken (Belgian NCA) as Chair
- and Dr Sarah Adler-Flindt (German NCA) as Vice-chair.

Action: For endorsement

4. Methodology Domain

4.1. Methodology Working Party (MWP)

Chairs: Kit Roes, Kristin Karlsson

4.1.1. Agenda and minutes

- Final agenda and minutes for MWP meetings held virtually on 14 and 28 July 2022

Action: For information

4.2. Biostatistics Operational Expert Group (BOEG)

No topics

4.3. Modelling and Simulation Operational Expert Group (MSOEG)

No topics

4.4. Real World Data Operational Expert Group (RWDOEG)

No topics

4.5. Pharmacokinetics Working Party (PKWP)

No topics

5. Clinical Domain

5.1. Central Nervous System Working Party (CNSWP)

Chair: Andre Elferink

5.1.1. Launch call for nomination for new member for CNSWP

The current CNS WP has only 7 members. It is proposed to add an additional member to bring the Working Party to full capacity. It is proposed to launch a call for nomination for a new member for CNSWP.

Nominations should be sent to the Agency by **9 September 2022**. Candidates are kindly asked to submit a brief CV in support of their candidature together with a cover letter highlighting their expertise.

Action: For information

5.2. Cardiovascular Working Party (CVSWP)

Chairs: Alar Irs, Patrick Vrijlandt

5.2.1. CVSWP response to the PKWP LoQs regarding considerations for digoxin as NTI drug

The CVSWP input was requested by the PKWP (July 2022) on the qualification of digoxin as a drug of narrow therapeutic index (NTI).

CHMP: Alar Irs

Action: For adoption

5.3. Oncology Working Party (ONCWP)

Chair: Pierre Demolis

5.3.1. Nomination of Oncology ESEC experts

- Nomination by ONCWP of the experts to enter the Oncology European Specialised Expert Community (ESEC)

Action: For endorsement

5.3.2. Update on the Oncology ESEC

Update on the start of projects and training/presentations.

Action: For information

5.4. Rheumatology and Immunology Working Party (RIWP)

No topics

5.5. Infectious Disease Working Party (IDWP)

No topics

5.6. Vaccines Working Party (VWP)

No topics

5.7. Haematology Working Party (HaemWP)

Chair: Daniela Philadelphy

5.7.1. Agenda

- Draft agenda of the HAEMWP to be held virtually on 9 September 2022

Action: For information

5.7.2. Minutes Blood cluster TC

- Final minutes of the Blood cluster TC held virtually on 24 June 2022

Action: For information

5.7.3. Minutes annual meeting with PPTA and IPFA

- Final minutes of the annual meeting with PPTA and IPFA held on 7 June 2022

Action: For information

5.8. Scientific Advisory Groups (SAGs) and Ad-hoc Expert Groups (AHEG)

No topics

6. Patients, Healthcare Professionals and Consumers

6.1. Patients and Consumers Working Party (PCWP) Healthcare Professionals Working Party (HCPWP)

No topics

7. Harmonisation and consistency groups

7.1. International Council on Harmonisation (ICH)

7.1.1. ICH E19 – A Selective Approach to Safety Data Collection in Specific Late-Stage Pre-approval or Post-Approval Clinical Trials

Following the public consultation that closed in Q4 2019, the ICH E19 working group has been revising the draft guideline. The final version has now been agreed and is brought to CHMP for adoption. After this, implementation is foreseen (6 months from date of publication).

Action: For adoption

7.1.2. ICH S1B(R1) – Revision of the guideline on testing for carcinogenicity of pharmaceuticals

Following the Non-Clinical WP endorsement of the final guideline version and the conclusion of the ICH process, this guideline is to be adopted by CHMP. Implementation will follow 6 months from publication.

Action: For adoption

7.2. Guideline Consistency Group (GCG)

No topics

7.3. Summary of product characteristics Advisory Group

No topics

8. Joint groups and collaboration with other Scientific committees

8.1. Joint CHMP/CVMP/CMDh/CMDv Working Group on Active Substance Master File Procedures (ASMF WG)

No topics

8.2. Collaboration with other Scientific committees

8.2.1. PRAC report to CHMP

Chair: Sabine Straus

Summary of recommendations and advice of PRAC meeting held on 29 August-01 September 2022.

Action: For information

9. Regulatory/Organisational matters

9.1. Regulatory Issues/new legislation

No topics

9.2. CHMP organisation/templates

9.2.1. CHMP learnings

Collection, discussion and recording of CHMP learnings.

CHMP: Outi Mäki-Ikola

Action: For discussion

9.2.2. CHMP Rules of Procedure amendments

Following the entry into force of both EU Regulations on medical devices, i.e. Regulation (EU) 2017/745 on medical devices (MDR) and Regulation (EU) 2017/746 on in vitro diagnostic medical devices (IVDR), as well as the Regulation (EU) 2022/123 on a reinforced role for the European Medicines Agency in crisis preparedness and management for medicinal products and medical devices, an update in the CHMP's Rules of Procedure (RoP) is required.

Action: For adoption

10. Product development support

10.1. Scientific Advice Working Party (SAWP)

Chair: Paolo Foggi

10.1.1. Agenda and Table of Decisions

- Agenda from 29 August until 01 September 2022 SAWP meeting held face-to-face
- Draft Table of Decisions from 29 August until 01 September 2022 SAWP meeting held face-to-face

Action: For information

10.1.2. Appointment of CHMP peer review for SA

Action: For information

10.1.3. Nomination of new members and alternates to the Scientific Advice Working Party

Nomination of new SAWP members and alternates.

Required areas of expertise: Haematology/ onco-haematology, cardiology, biosimilars.

Action: For endorsement

10.2. Innovation Task Force

10.2.1. ITF meeting

Meeting date: 7 September 2022

Action: For adoption

10.2.2. ITF meeting

Meeting date: 15 September 2022

Action: For adoption

10.2.3. ITF meetingMeeting date: 29 September 2022

Action: For adoption

10.2.4. ITF meeting

Meeting date: 23 September 2022 or 28 October 2022 tbc

Action: For adoption

10.2.5. ITF meeting

Meeting date: 10 October 2022

Action: For adoption

11. Product related topics

11.1. Preview CHMP Plenary

CHMP: Harald Enzmann

Action: For information

11.2. COVID-19 ongoing and upcoming procedures

List of currently ongoing and upcoming (imminently, i.e. expected within the next 2 months) applications for COVID-19 vaccines and therapeutics.

Action: For information

11.3. gefapixant - EMEA/H/C/005476 and EMEA/H/C/005884

Treatment of refractory or unexplained chronic cough

Scope: CHMP list of questions for consultation to the MWP

Action: For adoption

12. Any Other Business

12.1. Rapporteurships

Action: For information

12.2. PRIME implementation of 5-year review recommendations

Presentation of the proposals for implementation of the recommendations arising from the first 5 years' experience with the scheme (see also PRIME 5-year report), as discussed and agreed by the PRIME oversight group.

Action: For adoption

12.3. Update on the upcoming proof-of-concept raw data pilot

Update on the proof-of-concept raw data pilot. The pilot, which is expected to start in Q3 2022, will analyse individual patient data in electronic structured format (raw data) from selected marketing authorisation applications. The pilot aims at generating relevant learnings about accessing and analysing raw data during the assessment process. The proof-of-concept raw data pilot is part of EMA's Lifecycle Regulatory Submissions Raw Data project, which is focusing on utilising raw data to support regulatory decision-making and was listed as an action in CHMP's work plan for 2022.

Action: For information

12.4. Real World Evidence pilot and DARWIN EU®

Update on the establishment of DARWIN EU® containing an update on the list of data partners selected for Year 1 and an update on the progress of studies. Additionally, update

on the RWE pilot with CHMP, with a reminder to submit all research questions to the EMA RWE team.

Action: For discussion

12.5. Call to CHMP members to be part of the Immunisation and Vaccine Monitoring Board (IVMAB) of the ECDC/EMA Vaccine Monitoring Platform (VMP)

The vaccine monitoring platform (VMP), under the EMA and ECDC's extended mandates, has been established for the joint coordination of vaccine safety and effectiveness studies. The VMP will feature, as part of its governance, an EU Immunisation and Vaccine Monitoring Board (IVMAB) that will have a consultative and facilitative (non-binding) role, to help inform regulatory or public health action. Request to three CHMP members to join the IVMAB.

Interested members should express their interest by **8 September 2022**.

Action: For information