13 March 2024
EMA/CAT/81614/2024
Human Medicines Division

Committee for Advanced Therapies (CAT)

Draft agenda for the meeting on 13-15 March 2024

Chair: Ilona Reischl; Vice-Chair: Kieran Breen

13 March 2024, 14:00 – 18:30

14 March 2024, 09:00 – 18:30

15 March 2024, 09:00 – 13:00

Disclaimers

Some of the information contained in this agenda is considered commercially confidential or sensitive and therefore not disclosed. With regard to intended therapeutic indications or procedure scopes listed against products, it must be noted that these may not reflect the full wording proposed by applicants and may also vary during the course of the review. Additional details on some of these procedures will be published in the CAT meeting reports once the procedures are finalised.

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

Note on access to documents

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

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1. Introduction

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

Pre-meeting list of participants and restrictions in relation to declarations of interests applicable to the items of the agenda for the CAT plenary session to be held 13-15 March 2024. See 13-15 March 2024 CAT minutes (to be published post 17-19 April 2024 CAT meeting).

1.2. Adoption of agenda

CAT agenda for 13-15 March 2024 meeting

1.3. Adoption of the minutes

CAT minutes for 14-16 February 2024 meeting

2. Evaluation of ATMPs

2.1. Opinions

No items

2.2. Oral explanations

No items

2.3. Day 180 list of outstanding issues

2.3.1. Fidanacogene elaparvovec - PRIME - EMEA/H/C/004774

Indicated for the treatment of severe and moderately severe haemophilia B

Scope: Day 180 list of outstanding issues

Action: for adoption

List of questions adopted on 08.09.2023.

2.4. Day 120 list of questions

2.4.1. Beremagene geperpavec - PRIME - Orphan - EMEA/H/C/006330

Krystal Biotech Netherlands B.V.; Treatment of patients from birth with dystrophic epidermolysis bullosa (DEB) with mutation(s) in the collagen type VII alpha 1 chain (COL7A1) gene

Scope: Day 120 list of questions

Action: for adoption

2.5. Day 80 assessment reports

No items

2.6. Update on ongoing initial applications

No items

2.7. New applications

No items

2.8. Withdrawal of initial marketing authorisation application

No items

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

No items

2.10. Companion diagnostics

2.10.1. Initial consultation

No items

2.10.2. Follow-up consultation

No items

2.11. Type II variations and variations of therapeutic indication procedure according to Commission Regulation (EC) No 1234/2008

2.11.1. Breyanzi - Lisocabtagene maraleucel / Lisocabtagene maraleucel - EMEA/H/C/004731/II/0037/G

Bristol-Myers Squibb Pharma EEIG

Rapporteur: Concetta Quintarelli

Scope: Quality, request for supplementary information

Action: for adoption

2.12. Extension applications

No items

2.13. Other Post-Authorisation Activities

2.13.1. Breyanzi - Lisocabtagene maraleucel / Lisocabtagene maraleucel - EMEA/H/C/004731/REC/015.1

Bristol-Myers Squibb Pharma EEIG

Rapporteur: Concetta Quintarelli, CHMP Coordinator: Paolo Gasparini

Scope: Quality, opinion

Action: for adoption

2.13.2. Breyanzi - Lisocabtagene maraleucel / Lisocabtagene maraleucel - EMEA/H/C/004731/REC/018

Bristol-Myers Squibb Pharma EEIG

Rapporteur: Concetta Quintarelli, CHMP Coordinator: Paolo Gasparini

Scope: Quality, request for supplementary information

Action: for adoption

2.13.3. ROCTAVIAN - Valoctocogene roxaparvovec - Orphan - EMEA/H/C/005830/ANX/002.2

BioMarin International Limited

Rapporteur: Violaine Closson Carella, CHMP Coordinator: Jean-Michel Race

Scope: Clinical, opinion

Annual Status Report - Study Protocol 270-601 [A Non-Interventional, Multi- National, Longitudinal Study of Patients Treated with Roctavian (valoctocogene roxaparvovec)].

Action: for adoption

2.13.4. ROCTAVIAN - Valoctocogene roxaparvovec - Orphan - EMEA/H/C/005830/ANX/004.2

BioMarin International Limited

Rapporteur: Violaine Closson Carella, CHMP Coordinator: Jean-Michel Race

Scope: Clinical, opinion

Annual Registry - Study 270-801 [A Retrospective Cohort Study of Patients Treated with Roctavian (valoctocogene roxaparvovec): An Analysis of Patient Registries].

Action: for adoption

2.13.5. Zolgensma - Onasemnogene abeparvovec - Orphan - EMEA/H/C/004750/P46/023

Novartis Europharm Limited

Rapporteur: Emmely de Vries, CHMP Coordinator: Peter Mol

Scope: Clinical, request for supplementary information

Paediatric studies submitted in accordance with Article 46 of Regulation (EC) No1901/2006, as amended.
Clinical study report of Study No. COAV101A1IC01 (OFELIA) [Phase IV Open-label, single-arm, single-dose, multicentre study to evaluate the safety, tolerability and efficacy of gene replacement therapy with intravenous OAV101(AVXS101) in paediatric patients from Latin America with spinal muscular atrophy (SMA)].

Action: for adoption

2.14. GMP and GCP inspections requests

No items

3. Certification of ATMPs

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

3.1. Opinion

No items

3.2. Day 60 Evaluation Reports

No items

3.3. New Applications

4. Scientific Recommendation on Classification of ATMPs

Timetable:

-Start of the procedure:	25.03.2024
-EMA Coordinator's draft report:	02.04.2024
-CAT Coordinator's comments:	10.04.2024
-Revised scientific recommendation:	12.04.2024
-CAT's discussion of scientific recommendation:	19.04.2024

4.1. New requests – Appointment of CAT Coordinator

4.1.1. mRNA encoding ARCUS nuclease

For treatment of chronic hepatitis B (CHB) virus infection

Scope: Appointment of CAT Coordinator and adoption of timetable

Action: for adoption

4.1.2. Allogeneic human corneal endothelial cells (neltependocel) and a low molecular weight Rho kinase inhibitor (Y-27632)

For treatment of corneal oedema due to corneal endothelial dysfunction

Scope: Appointment of CAT Coordinator and adoption of timetable

Action: for adoption

4.1.3. Lymphocyte concentrate

For improvement of the pregnancy outcomes among women with unexplained repeated pregnancy loss and HLA sharing among partners

Scope: Appointment of CAT Coordinator and adoption of timetable

Action: for adoption

4.2. Day 30 ATMP scientific recommendation

4.2.1. Allogeneic human induced pluripotent stem cells-derived corneal limbal stem cells

For treatment of limbal stem cell deficiency

Scope: ATMP scientific recommendation

Action: for adoption

4.2.2. Olfactory glial cells isolated from autologous human olfactory bulb, expanded in culture

For treatment of complete spinal cord injuries

Scope: ATMP scientific recommendation

Action: for adoption

4.2.3. Circular RNA capable to bind to mutated regions of the messenger RNA from the DMPK gene

For treatment of myotonic dystrophy type 1

Scope: ATMP scientific recommendation

Action: for adoption

4.3. Day 60 revised scientific recommendation (following list of questions)

No items

4.4. Finalisation of procedure

4.4.1. Modified measles vaccine virus

For the treatment of solid cancer tumours

Scope: European Commission raised no comments. ATMP scientific recommendation

Action: for adoption

4.5. Follow-up and guidance

No items

5. Scientific Advice

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

5.1. New requests - appointment of CAT Rapporteurs

5.1.1. Ongoing scientific advice procedures - Appointment of CAT Peer Reviewers

Timetable:

- Start of procedure at SAWP: 04-07.03.2024
- Appointment of CAT Peer Reviewers: 13-15.03.2024

- SAWP first reports:	02.04.2024
- CAT Peer Reviewer comments (NC/C):	05.04.2024
- CAT Peer Reviewer comments (Q):	10.04.2024
- Discussion at SAWP:	08-11.04.2024
- Discussion at CAT and feedback to SAWP:	17-19.04.2024

5.1.2. Scientific advice procedures starting at the next SAWP meeting

Timetable:

- Start of procedure at SAWP:	08-11.04.2024
- Appointment of CAT Peer Reviewers:	17-19.04.2024
- SAWP first reports:	06.05.2024
- CAT Peer Reviewer comments (NC/C):	10.05.2024
- CAT Peer Reviewer comments (Q):	15.05.2024
- Discussion at SAWP:	13-16.05.2024
- Discussion at CAT and feedback to SAWP:	22-24.05.2024

No items

5.2. Procedures discussed at SAWP – 1st reports, D40 JRs, LoIs

5.3. Finalisation of D70 procedures – feedback from the discussion meeting

5.4. Final Advice Letters for procedures finalised the previous month

6. Pre-Authorisation Activities

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

6.1. Paediatric investigation plans

6.2. ITF briefing meetings in the field of ATMPs

6.3. Priority Medicines (PRIME) – Eligibility requests

6.3.1. Month 0 - Start of the procedure

Timetable for assessment:

Procedure start:	04-07.03.2024
SAWP recommendation:	11.04.2024
CAT recommendation:	19.04.2024
CHMP adoption of report and final recommendation:	25.04.2024

6.3.2. Month 1 – Discussion of eligibility

6.3.3. Month 2 – Recommendation of eligibility

6.3.4. Ongoing support

No items

7. Organisational, regulatory and methodological matters

7.1. Mandate and organisation of the CAT

7.1.1. CAT membership

Action: for information

7.1.2. Vote by proxy

Action: for information

7.1.3. CAT Strategic Review & Learning meeting (SRLM) under the Belgian presidency – 15-17 May 2024

CAT: Claire Beuneu

Scope: Draft agenda of the upcoming SRLM

Action: for discussion

7.2. Coordination with EMA Scientific Committees

7.2.1. EU NTC training webinar on the regulatory/HTA interface under the HTA Regulation

Scope: Presentation on the upcoming EU NTC training webinar on the regulatory/HTA interface to take place on 02.05.2024

Action: for information

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

7.3.1. Workshop - Challenges in drug development, regulation and clinical practice in hemoglobinopathies

Scope: Presentation on the draft agenda of the workshop to take place on 01.07.2024

Action: for discussion

7.3.2. Revision of the Variations Framework

Scope: Presentation on the main points of the proposal

Action: for discussion

7.4. Cooperation with the EU regulatory network

No items

7.5. Cooperation with international regulators

7.5.1. ATMP cluster teleconference with US-FDA, Health Canada and PMDA (Japan)

CAT: Ilona Reischl

Scope: Feedback from the teleconference of 22.02.2024

Action: for information

7.5.2. International Pharmaceutical Regulatory Programme (IPRP) Gene and Cell therapy working group

CAT: Pille Säälik

Scope: Feedback from the teleconference of 27.02.2024

Action: for information

7.6. CAT work plan

7.6.1. Guideline on quality, non-clinical and clinical requirements for investigational ATMPs in clinical trials

CAT: Ilona Reischl

Scope: Adopted guideline, for release of a second public consultation

Action: for information

7.7. Planning and reporting

7.7.1. Business Pipeline Report

Scope: Q1/2024 Update of the Business Pipeline report for the human scientific committees

Action: for information

7.8. Others

No items

8. Any other business

No items

Date of next CAT meeting:

17-19 April 2024

9. Explanatory notes

The Notes give a brief explanation of relevant agenda items and should be read in conjunction with the agenda.

For a list of acronyms and abbreviations, see:

List of abbreviations used in EMA human medicines scientific committees and CMDh documents, and in relation to EMA's regulatory activities

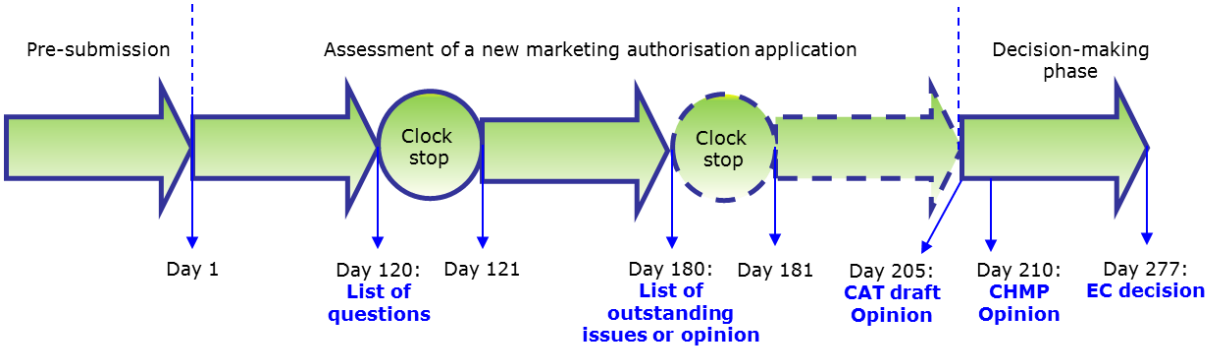
Evaluation of ATMPs (section 2)

This section lists applications for marketing authorisations of new Advanced Therapy Medicinal Products (ATMPs) that are to be discussed by the Committee. It also lists Post-authorisation activities (section 2.11-2.13) and any ATMP related inspection requests (section 2.14).

New applications (sections 2.1. to 2.9.)

Section 2.1 is for ATMPs nearing the end of the evaluation and for which the CAT is expected to adopt a draft **opinion** at this meeting on whether marketing authorisation should be granted. Once adopted, the CAT opinion is transmitted to the CHMP for final adoption. The CHMP opinion will be forwarded to the European Commission for a final legally binding decision valid throughout the EU. More information on the evaluation of ATMPs can be found here.

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 CAT. The clock stop happens after day 120 and may also happen after day 180, when the CAT has adopted respectively a **Day 120 list of questions** (section 2.4) or a List of outstanding issues to be addressed by the company, which is listed in the agenda under sections 2.7 (**Ongoing evaluation procedures** (section 2.3). Section 2.6 also includes the CAT discussions at any other timepoint of the evaluation procedure of new applications.

Oral explanation (section 2.2.)

Prior to adoption of the CAT opinion, marketing authorisation applicants are normally invited to the CAT plenary meeting to address questions raised by the Committee.

Oral explanations normally relate to ongoing applications, but they can also relate to any other issue for which the CAT would like to discuss with company representatives in person.

New applications (section 2.7.)

In this section, information is included on upcoming marketing authorisation applications for ATMPs, as well as information on appointment of Rapporteurs for new ATMP applications.

Withdrawal of applications (section 2.8.)

This section includes information on marketing authorisation applications that are withdrawn by the applicant. Applicants may decide to withdraw applications at any stage during the assessment and a CAT opinion will therefore not be issued. Withdrawals are included in the agenda for information or discussion, as necessary.

Re-examination procedures (new applications) under article 9(2) of regulation no 726/2004 (section 2.9.)

This section lists applications for new marketing authorisation for ATMPs for which the applicant has requested a re-examination of the opinion previously issued by the CHMP. Similar to the initial evaluation of a marketing authorisation of an ATMP, CAT will adopt a draft re-examination opinion, which is transmitted to the CHMP for final adoption.

Companion diagnostics (section 2.10)

This section lists applications for initial and follow-on consultation of companion diagnostics.

Post-authorisation activities (section 2.11-2.13.)

Section 2.11 lists type II variations, including extension of indication applications and re-examination procedures for type II variations for which the applicant has requested re-examination of the opinion previously issued by the CHMP. Section 2.12 list extension application according to Annex I of Reg. 1234/2008 and section 2.13 includes all other post-authorisation activities concerning authorised ATMPs that are not covered elsewhere in the agenda such as post-authorisation measures, annual reassessments, 5-year renewals, supply shortages, qualify defects. 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, will also be included here.

GMP and GCP Inspections Issues (section 2.14.)

This section lists inspections that are undertaken for ATMPs. 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).

Certification of ATMPs (section 3)

This section includes the scientific evaluation by the CAT of quality and non-clinical data that small and medium-sized enterprises have generated at any stage of the ATMP development process. More information on the ATMP certification procedure can be found here.

Scientific Recommendation on Classification of ATMPs (Section 4)

This section includes the scientific recommendation by the CAT on whether medicines based on genes, cells or tissues meet the scientific criteria that define ATMPs. More information on the ATMP classification procedure, including the outcomes of finalised classifications, can be found here.

Scientific Advice (section 5)

This section includes all scientific advice given to companies during the development of an ATMP. Information related to the number of ATMP related scientific advices discussed by CAT can be found in the CAT Monthly reports. Further information on SAWP can be found here.

Pre-Authorisation (section 6)

Paediatric Investigation Plan (PIP)

This section includes the discussion of an ATMP before a formal application for marketing authorisation

is submitted. These cases refer for example 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.

CAT contributes to the evaluation of a Paediatric Investigation Plan (PIPs) for ATMPs by the Paediatric Committee. These PIPs are included in this section of the Agenda.

ITF Briefing meeting in the field of ATMPs

This section refers to briefing meetings of the Innovation Task Force and International co-operations activities of the CAT

The Innovation Task Force (ITF) is a body set up to encourage early dialogue with applicants developing innovative medicines. Minutes of meetings with applicants developing ATMPs and of other ITF meetings of interest to the CAT are included in this section of the agenda. Further information on the ITF can be found here.

Priority Medicines (PRIME)

This section includes the new requests for eligibility to PRIME for ATMPs under development, the discussions in CAT of these eligibility requests and the final recommendations for eligibility of ATMPs adopted by CHMP.

CAT will appoint one of its members as the CAT sponsor for each new ATMP eligibility request who will lead the CAT discussion based on the recommendation from the SAWP.

Organisational, regulatory and methodological matters (section 7)

This section includes topics related to regulatory and procedural guidance, CAT workplan, CAT meeting organisation (including CAT membership), planning and reporting, co-ordination with other committees, working parties and scientific advisory groups.

Furthermore, this section refers to the activities of the CAT drafting groups developing scientific guidelines for gene therapy medicinal products and for cell-based medicinal products, cooperation within the EU regulatory network and international regulators as well as direct interaction with interested parties. It also includes topics of scientific interest for the Committee that are not directly related to the work of the CAT drafting groups or CAT associated working parties.

Any other business (section 8)

This section is populated with miscellaneous topics not suitable under the previous headings.

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