17 January 2018
EMA/CAT/34202/2018
Inspections, Human Medicines Pharmacovigilance and Committees Division

Committee for Advanced Therapies (CAT)

Agenda for the meeting on 17-19 January 2018

Chair: Martina Schüßler-Lenz; Vice-Chair: Ilona Reischl

17 January 2018, 14:00 – 17:30, room 03-E

18 January 2018, 09:00 – 18:00, room 03-E

19 January 2018, 09:00 – 12:00, room 03-E

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 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).

30 Churchill Place ● Canary Wharf ● London E14 5EU ● United Kingdom
Telephone +44 (0)20 3660 6000 **Facsimile** +44 (0)20 3660 5555
Send a question via our website www.ema.europa.eu/contact

Table of contents

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 18-19 January 2018. See January 2018 CAT minutes (to be published post-February 2018 CAT meeting).

1.2. Adoption of agenda

CAT agenda for 17-19 January 2018 meeting

1.3. Adoption of the minutes

CAT minutes for the 06-08 December 2017 meeting

1.4. Technical information

2. Evaluation of ATMPs

2.1. Opinions

No items

2.2. Oral explanations

No items

2.3. Day 180 list of outstanding issues

No items

2.4. Day 120 list of questions

No items

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 marking 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. GMP and GCP inspections requests

No items

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

2.11.1. Zalmoxis - allogeneic T cells genetically modified with a retroviral vector encoding for a truncated form of the human low affinity nerve growth factor receptor (ΔLNGFR) and the herpes simplex I virus thymidine kinase (HSV-TK Mut2) - Orphan - EMEA/H/C/002801/II/0005/G

MolMed SpA

Rapporteur: Hans Ovelgönne, CHMP Coordinator: Paula Boudewina Van Hennik

Scope: quality

Action: for adoption

Request for Supplementary Information adopted on 06.10.2017

2.12. Other Post-Authorisation Activities

2.12.1. MACI - matrix applied characterised autologous cultured chondrocytes - EMEA/H/C/002522/R/0017

Vericel Denmark ApS

Rapporteur: Christiane Niederlaender, Co-Rapporteur: Johannes Hendrikus Ovelgönne, CHMP Coordinator: Greg Markey, PRAC Rapporteur: Julie Williams

Scope: Five-year renewal of Marketing Authorisation

Action: for adoption

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. Ongoing Applications

3.4. New Applications

No items

4. Scientific Recommendation on Classification of ATMPs

4.1. New requests – Appointment of CAT Coordinator

4.1.1. Expanded autologous auricular chondrocytes - H0004979

Intended for the surgical implantation for the repair of microtia

Scope: appointment of CAT Coordinator and adoption of timetable

Action: for adoption

4.1.2. Elastin recombinamer (ELR)-encapsulated allogeneic pancreatic islets - H0004980

Intended for treatment of severe forms of type 1 diabetes

Scope: appointment of CAT Coordinator and adoption of timetable

Action: for adoption

4.1.3. Autologous CD31+ Cells - H0004981

Intended as adjunct therapy during primary care of proximal humeral fracture to decrease incidence of non-union and secondary displacement

Scope: appointment of CAT Coordinator and adoption of timetable

Action: for adoption

4.2. Day 30 ATMP scientific recommendation

4.2.1. Autologous dendritic cells pulsed with allogeneic tumour cell lysate - H0004949

Intended for the treatment of malignant mesothelioma

Scope: scientific recommendation

Action: for adoption

4.2.2. Allogeneic mesenchymal stem cells suspended in cell supernatant - H0004952

Intended for the treatment of osteoarthritis

Scope: 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. Autologous CD34+ cells derived from bone marrow - H0004941/0001

Intended for the improvement of neurologic function in patients with non-lacunar acute ischemic stroke infarctions Scope: no comments received by the European Commission. Final ATMP scientific recommendation

Action: for information

4.4.2. Stromal vascular fraction (SVF) – H0004926

Intended to diminish cancer-related lymphedema in breast cancer patients

Scope: no comments received by the European Commission. Final ATMP scientific recommendation

Action: for information

4.5. Follow-up and guidance

4.5.1. Regulation of non-viable tissues in the EU

CAT: Ilona Reischl

Scope: message from the European Commission (DG Santé B4) to the National Competent Authorities for tissues and cells (September 2017)

Action: for information

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

Timetable:

-Final Briefing Package:	31.01.2018
-Start of the procedure at SAWP:	05-08.02.2018
-CAT report due by:	09.02.2018
-CAT recommendation:	16.02.2018

5.2. CAT reports

5.3. List of Issues

5.4. Finalisation of SA procedures

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 of Medicines (PRIME) – Eligibility requests

6.3.1. Month 0 - Start of the procedure

Timetable for assessment:

Start of the procedure:	11.01.2018
SAWP recommendation:	08.02.2018
CAT recommendation:	16.02.2018
CHMP adoption of report and final recommendation:	22.02.2018

6.3.2. Month 1 – Discussion of eligibility

6.3.3. Month 2 – Recommendation for eligibility

6.3.4. Month 3 – Nomination of Rapporteurs

6.3.5. Ongoing support

7. Organisational, regulatory and methodological matters

7.1. Mandate and organisation of the CAT

7.1.1. CAT membership

Scope: membership changes

Italy: Luca Sangiorgi – alternate membership ended on 08 January 2017

Slovakia: Ján Kyselovič – nominated as the new member from 22 January 2018

Slovakia: Mikuláš Hrubiško – membership ended on 22 January 2018

Sweden: Lisbeth Barkholt has been nominated as the new member from 01 January 2018

Action: for information

7.1.2. Strategic Review & Learning meeting – Joint CHMP/PDCO/CAT, Oslo, Norway, 07-09 May 2019

CAT resources: Helga Olsen, Rune Kjeken

Scope: Strategic Review & Learning meeting in partnership with the CHMP and PDCO to be hosted by Norway in Oslo on 07-09 May 2018 under the auspices of the Bulgarian Presidency of the Council of the European Union.

Action: for information

7.1.3. Strategic Review & Learning meeting – Tallinn, Estonia, 15-17 November 2017

Scope: CAT Strategic Review & Learning meeting (SRLM)

CAT: Martina Schüßler-Lenz, Toivo Maimets

Scope: minutes of the meeting

Action: for adoption

7.2. Coordination with EMA Scientific Committees

7.2.1. Committee for Medicinal Products for Human Use (CHMP)

Scope: Summary of Outcomes (SoO) for the December 2017 meeting

Action: for information

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

7.3.1. BWP mandate and workload

Scope: presentation on the BWP on role, responsibilities, product exposure and 2018 workplan

CAT: Sol Ruiz

Action: for information

7.3.2. Guideline on quality, non-clinical and clinical aspects of medicinal products containing genetically modified cells

CAT Rapporteur: Marcos Timón

Scope: feedback on the drafting group meeting that took place on 17 January 2018

Action: for information

Drafting group: Marcos Timón, Ilona Reischl, Christiane Niederlaender, Belaïd Sekkali, Tiina Palomäki, Guido Pantè, Matthias Renner, Marcel Hoefnagel, Brigitte Anliker, Olli Tenhunen, Paolo Gasparini, Martina Schüßler-Lenz

7.3.3. ATMP guideline on safety and efficacy follow-up and risk management

Drafting group: Simona Badoi, Tomas Boráň, Violaine Closson-Carella, Romaldas Mačiulaitis, Maura O'Donovan, Sol Ruíz

Action: for adoption for publication for external consultation

Note: the guideline has been adopted by PRAC and reviewed by the Guideline Consistency Group.

7.3.4. PRIME for ATMPs

CAT: Martina Schüßler-Lenz

Scope: feedback from the PRIME oversight group; discussion of general issues ; revision of PRIME related documents – provision of ATMP specific comments.

Action: for discussion

7.4. Cooperation within the EU regulatory network

7.4.1. ATMP training curriculum

CAT: Ilona Reishl

Scope: presentation of the updated draft curriculum following discussions at the Strategic Review & Learning meeting (Tallinn, 15-17 November 2017),

Action: for discussion

7.5. Cooperation with international regulators

7.5.1. ATMP cluster teleconference with FDA, Health Canada and PMDA

The teleconference will take place

CAT: Martina Schüßler-Lenz

Scope: draft agenda

Action: for adoption

7.5.2. International Pharmaceutical Regulators Forum (IPRF)

CAT: Martina Schüßler-Lenz

Scope: presentation of role and activities of IPRF

Action: for nomination / re-nomination of CAT members for the gene therapy and cell therapy group.

7.5.3. International Pharmaceutical Regulators Forum Gene Therapy Working Group (IPRF-GTWG)

CAT experts: Björn Carlsson, Tiina Palomäki

Scope: IPRF Reflection paper on biodistribution (BD) for gene therapy products.

Action: for comments by 2 February 2018

Background: This reflection paper describing the expectations for BD for GTMP has been developed by a drafting group from the IPRF-GTWG. Drafting took place virtually and during the in-person IPRF-GTWG meeting that was organised at EMA on 2-3 May 2017 (see minutes of May 2017 CAT meeting, agenda point 7.5.1). Björn Carlsson and Tiina Palomäki were part of this drafting group. CAT members are asked to review the reflection paper and provide comments not later than 2 February 2018.

7.6. CAT work plan

7.6.1. Registry requirements for chimeric antigen receptor T (CAR-T) cells

CAT: Martina Schüßler-Lenz

Scope: feedback on the activitiesorganisation of the workshop with all stakeholders (9 February 2018)

Action: for information

Note: this will be a cross-committee activity, involving members/experts from CAT, SAWP, PRAC, CHMP and PDCO.

7.6.2. Expert meeting on adeno-associated viral vectors, 06 September 2017, EMA, London

CAT: Martina Schüßler-Lenz

Scope: report of the meeting that took place on 6 September 2017

Action: for adoption

7.6.3. Environmental assessment of gene therapy medicinal products

CAT: Rocío Salvador-Roldán – European Commission

Scope: feedback from discussions with GMO authorities regarding the assessment of human cells genetically modified.

Action: for discussion

7.7. Planning and reporting

7.7.1. Planning estimates of forthcoming ATMP MAAs

Scope: Q4/2017 update of the business pipeline report for the human scientific committees

Action: for information

7.8. Others

7.8.1. European Biopharmaceutical Enterprise (EBE)-Escher-ATMP project results

Escher group: Andre Broekmans and Renske ten Ham; EBE delegation: Barbara Freischem and Veronique Debaut

Scope: presentation by the Esther ATMP Project analysis findings

Action: for discussion

Note: the Escher-ATMP research project consists of two work packages. WP1 is to provide an overview of product and developer characteristics and identify barriers in regulation and market access in Europe and WP2 is to examine factors associated with successful ATMP development and commercialisation in Europe.

8. Any other business

8.1. CAT's Journal Club

Scope: placeholder in the agenda each month to list relevant publications on ATMPs, regulation, science

Action: for discussion

Note: the idea of a CAT's Journal Club was first introduced in July 2010

Date of next CAT meeting:
15-16.02.2018

Explanatory notes

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

Abbreviations / Acronyms

AR: Assessment Report

ATMP: Advanced Therapy Medicinal Product

BWP: Biologics Working Party

CAT: Committee for Advanced Therapies

CHMP: Committee for Medicinal Product for Human Use

COMP: Committee for Orphan Medicinal Products

CTFG: Clinical Trial Facilitation Group

DG: Drafting Group

EC: European Commission

ERA: Environmental Risk Assessment

FDA: Food and Drug Administration

FL: Final Letter

GCP: Good Clinical Practice

GLP: Good Laboratory Practice

GMO: Genetically-modified organism

GMP: Good Manufacturing Practice

HTA: Health Technology Assessment Bodies

HSPC: Hematopoietic Stem and Progenitor Cells

ITF: Innovative Task Force

JR: Joint Report

LoOI: List of outstanding issues

LoQ: List of questions

MA: Marketing Authorisation

MAA: Marketing Authorisation Applicant

MAH: Marketing Authorisation Holder

MSC: Mesenchymal stem cells

PDCO: Paediatric Committee

PMDA: Pharmaceuticals and Medical Devices Agency (Japan)

PIP: Paediatric Investigation Plan

PL: Package leaflet

PRAC: Pharmacovigilance and Risk Assessment Committee #

PRIME: Priority Medicines

RMP: Risk Management Plan

RP: Reflection paper

RSI: Request for supplementary information

SAs: Scientific Advices

SAG-O: Scientific Advisory Group Oncology

SAWP: Scientific Advice Working Party

SR: Summary Report

SWP: Scientific Working Party

SME: Small and medium size enterprises

SmPC: Summary of Products Characteristics

TT: Timetable

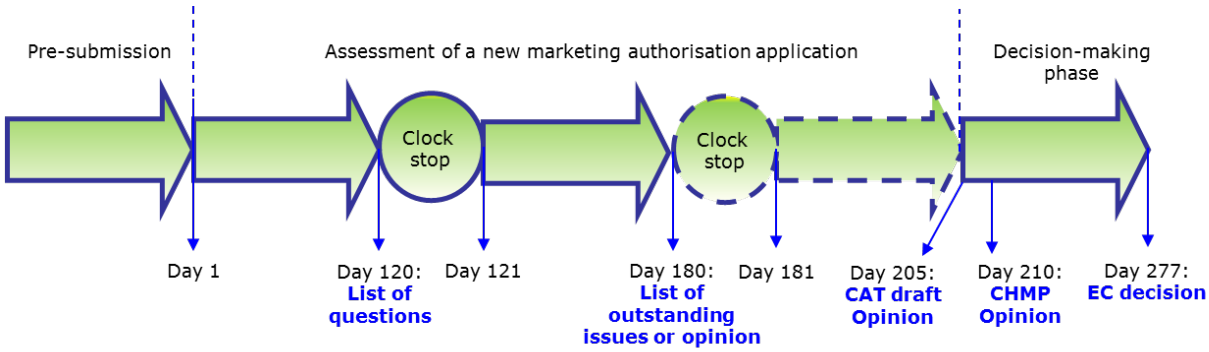
Evaluation of ATMPs (section 2)

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 any ATMP related inspection requests (section 2.9) and Post-authorisation activities (section 2.10).

New applications (sections 2.1. to 2.12.)

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 2.3) 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 2.7 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.

Re-examination procedures (new applications) under article 9(2) of regulation no 726/2004 (section 2.6.)

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.

Withdrawal of applications (section 2.7.)

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.

New applications (section 2.9.)

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.

GMP and GCP Inspections Issues (section 2.10.)

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).

Post-authorisation activities (section 2.12.)

This section lists type II variations, extension application according to Annex I of Reg. 1234/2008, re-examination procedures 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 and other issues concerning authorised medicines that are not covered elsewhere in the agenda such as 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.

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/