16 June 2017
EMA/CAT/305411/2017
Inspections, Human Medicines Pharmacovigilance and Committees Division

Committee for Advanced Therapies (CAT)

Minutes for the meeting on 10-12 May 2017

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

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 CAT meeting reports once the procedures are finalised.

Of note, the minutes are a working document primarily designed for CAT 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).

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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.

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. Marc Turner declared a past involvement in a data safety monitoring board for a clinical trial with a CAR-T product: as this product is not on the agenda, no additional restrictions were applied.

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

The CAT agenda for 10-12 May 2017 meeting was adopted.

1.3. Adoption of the minutes

The CAT minutes for 10-12 April 2017 meeting were adopted.

2. Evaluation of ATMPs

2.1. Opinions

2.1.1. Spheroids of human autologous matrix-associated chondrocytes - EMEA/H/C/002736

treatment of repair of symptomatic articular cartilage defects of the femoral condyle and the patella of the knee (International Cartilage Repair Society [ICRS] grade III or IV) with defect sizes up to 10 cm^2 in adults

Scope: Opinion

Action: for adoption

List of Outstanding Issues adopted on 17.02.2017 and 12.04.2017. List of Questions adopted on 19.04.2013.

Further to the discussion in the April 2017 CAT meeting, CAT discussed the conditions for

approval (Annex II conditions, Risk Management Plan (RMP) and Recommendations).

The final vote was taken on 11.05.2017. The outcome of the vote was: 17 members agree, 12 members disagree, Norway voted in favour.

CAT adopted by majority the CAT assessment report and the positive CAT draft opinion , for transmission to the CHMP. The divergent opinion signed by the following CAT members was appended to the CAT draft opinion: P. Salmikangas, A. Tsifsoglou, T. Tiido, I. Haunerova, H. Ovelgönne, U. Riekstina, M. Menezes Ferreira, P. Gasparini, R. Mačiulaitis, S. Badoi, B Gänsbacher, M. Turner.

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

2.6.1. Expanded human allogeneic mesenchymal adult stem cells extracted from adipose tissue – Orphan - EMEA/H/C/0004258

TiGenix S.A.U.; treatment of complex perianal fistula(s)

Scope: applicant's request to extend the clock-stop to respond to the D180 LoOI

The CAT noted this information and adopted the revised timetable.

2.7. New applications

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

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

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

No items

2.12. Other Post-Authorisation Activities

2.12.1. Imlygic - talimogene laherparepvec - EMA/H/C/002771/MEA/006

Amgen Europe B.V.; indicated for the treatment of adults with unresectable melanoma that is regionally or distantly metastatic (stage IIIB, stage IIIC and stage IV with distant skin, subcutaneous or nodal metastates (stage IV M1a)) with no bone, brain, lung or other visceral disease

Rapporteur: Olli Tenhunen, CHMP Coordinator: Tuomo Lapveteläinen

Scope: extension of the submission time (from February 2021 to October 2021) of study 20120324 (category 3): a phase 2, multicentre, single-arm trial to evaluate the biodistribution and shedding of talimogene laherparepvec in subjects with unresected, stage IIIB to IVM1c melanoma.

Action: for adoption

Note: Extension agreed by Rapporteur in Variation IB/0007

The extension was adopted.

2.12.2. Imlygic – talimogene laherparepvec - EMEA/H/C/002771/PSUSA/10459/201610

Amgen Europe B.V.; indicated for the treatment of adults with unresectable melanoma that is regionally or distantly metastatic (stage IIIB, IIIC and IVM1a) with no bone, brain, lung or other visceral disease.

Rapporteur: Olli Tenhunen, CHMP Coordinator: Tuomo Lapveteläinen; PRAC Rapporteur: Brigitte Keller-Stanislawski

Scope: PRAC assessment of PSUR (period 17.04.2016 – 26.10.2016), of the following adverse event:

Action: for information

CAT noted the outcome of the PRAC assessment. No action is proposed by PRAC in the approved indication (melanoma). Linked to the fatal case of arterial haemorrhage/carotid blow out syndrome following administration in a clinical trial of Imlygic in combination with pembrolizumab for the treatment of subjects with recurrent or metastatic squamous cell carcinoma of the head and neck, the company will provide a follow-up on this adverse event in the next PSUR.

2.12.3. Glybera - Alipogene tiparvovec – Orphan - EMEA/H/C/002145/R/0062

UniQure; indicated for the long term correction of lipoprotein lipase deficiency, to control or abolish symptoms and prevent complications in adult patients clinically diagnosed with lipoprotein lipase deficiency (LPLD)

Rapporteur: Christiane Niederlaender; Co-Rapporteur: Egbert Flory; CHMP Coordinators: Greg Markey, Jan Mueller-Berghaus

Scope: Formal notification letter by the MAH informing the EMA of the withdrawal of the 5-year renewal application in view of allowing the marketing authorisation to expire on 25 October 2017 based on economic reasons

Action: for information

The CAT noted the notification letter.

2.12.4. Zalmoxis – Allogeneic T cells genetically modified with a retroviral vector encoding for a truncated form of the human low affinity nerve growth factor receptor (ðlngfr) and the herpes simplex I virus thymidine kinase (hsv-tk mut2) – Orphan - EMEA/H/C/002801/R/0003

MolMed SpA; indicated for adjunctive treatment in haploidentical haematopoietic stem cell transplantation of adult patients with high-risk haematological malignancies

Rapporteur: Hans Ovelgönne; CHMP Coordinator: Paula Boudewina van Hennik

Scope: 1st annual reassessment for renewal of conditional MA. Opinion

Action: for adoption

Note: CAT adopted an RSI at is 10-12 April 2017 meeting

The renewal opinion was adopted.

3. Certification of ATMPs

Disclosure of 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

3.3. New Applications

No items

4. Scientific Recommendation on Classification of ATMPs

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

4.1. New requests – Appointment of CAT Coordinator

4.1.1. Stromal vascular fraction cells for autologous use - EMA/H0004838

Intended for the relief of symptoms of osteoarthritis

Scope: appointment of CAT coordinator and adoption of timetable

Action: for adoption

Nominations were received. The CAT member was appointed as CAT coordinator for this procedure.

4.1.2. Autologous human keratinocytes - EMA/H0004841

Intended for the treatment of burns and chronic, severe wounds

Scope: appointment of CAT coordinator and adoption of timetable

Action: for adoption

Nominations were received. The CAT member was appointed as CAT coordinator for this procedure.

4.1.3. Autologous human chondrocytes - EMA/H0004840

Intended for the repair of single symptomatic cartilage defect of the knee or ankle

Scope: appointment of CAT coordinator and adoption of timetable

Action: for adoption

Nominations were received. The CAT member was appointed as CAT coordinator for this procedure.

4.1.4. Allogeneic human umbilical cord blood-derived mesenchymal stem cells (hUCB-MSC) - EMA/H0004839

Intended for the treatment of atopic dermatitis

Scope: appointment of CAT coordinator and adoption of timetable

Action: for adoption

Nominations were received. The CAT member was appointed as CAT coordinator for this procedure.

4.2. Day 30 ATMP scientific recommendation

4.2.1. Autologous adipose derived mesenchymal stem cells - EMA/H0004813

Intended for the treatment of chronic wound

Scope: scientific recommendation

Action: postponed

The start of this procedure was postponed, pending additional information from the applicant.

4.2.2. Allogenic human mesenchymal stem cells - Mesenchymal stem cells isolated from umbilical cord - EMA/H0004815

Intended for the treatment of chronic obstructive pulmonary disease

Scope: scientific recommendation

Action: for adoption

CAT discussed the ATMP classification report. CAT adopted by consensus the ATMP classification report. CAT secretariat to send the draft scientific recommendation to the European Commission for comments by 29 May 2017.

The CAT recommendation will be considered final if no comments are received from the European Commission. The final report will be sent thereafter to the applicant.

4.2.3. Bilayer, engineered, collagen hydrogel-based skin graft composed of autologous keratinocytes and fibroblasts - EMA/H0004817

Intended for the treatment of partial deep dermal and full thickness burn wounds

Scope: scientific recommendation

Action: for adoption

CAT discussed the ATMP classification report. CAT adopted by consensus the ATMP classification report. CAT secretariat to send the draft scientific recommendation to the European Commission for comments by 29 May 2017.

The CAT recommendation will be considered final if no comments are received from the European Commission. The final report will be sent thereafter to the applicant.

4.2.4. Replication incompetent adenoviral serotype 5 vector encoding the human interleukin-12 p70 (hIL-12) transgene under the control of the activator ligand, veledimex - EMA/H0004805

Intended for the treatment of recurrent or progressive glioblastoma

Scope: scientific recommendation

Action: for adoption

CAT discussed the ATMP classification report. CAT adopted by consensus the ATMP classification report. CAT secretariat to send the draft scientific recommendation to the European Commission for comments by 29 May 2017.

The CAT recommendation will be considered final if no comments are received from the European Commission. The final report will be sent thereafter to the applicant.

4.2.5. Adenovirus-associated viral vector serotype 5 containing CRISPR Cas9 and guide ribonucleic acids (RNAs) targeting intron 26 of the centrosomal protein 290 gene (AAV5-GRK1-SauCas9-CEP290gRNA 323/64) - EMA/H0004818

Intended for the treatment of patients aged 3 years and older with Leber congenital amaurosis type 10 (LCA10) caused by a homozygous or compound heterozygous intron 26 mutation, c.2991+1655 A>G, in the CEP290 gene

Scope: scientific recommendation

Action: for adoption

CAT discussed the ATMP classification report. CAT adopted by consensus the ATMP classification report. CAT secretariat to send the draft scientific recommendation to the European Commission for comments by 29 May 2017.

The CAT recommendation will be considered final if no comments are received from the European Commission. The final report will be sent thereafter to the applicant.

4.2.6. Autologous human adipose perivascular stromal cells genetically modified to secrete soluble tumour necrosis factor-related apoptosis-inducing ligand (sTRAIL) - EMA/H0004820

Intended for the treatment of TRAIL-sensitive cancers such as Ewing sarcoma and pancreatic ductal adenocarcinoma

Scope: scientific recommendation

Action: for adoption

CAT discussed the ATMP classification report. CAT adopted by consensus the ATMP classification report. . CAT secretariat to send the draft scientific recommendation to the European Commission for comments by 29 May 2017.

The CAT recommendation will be considered final if no comments are received from the European Commission. The final report will be sent thereafter to the applicant.

4.2.7. Resorbable, viscoelastic matrix for use with autologous stromal vascular fraction (SVF) - EMA/H0004819

A resorbable matrix to be used for the delivery of autologous SVF adipose derived cells for the treatment of Human immunodeficiency virus (HIV)-related facial lipoatrophy

Scope: scientific recommendation

Action: for adoption

CAT discussed the ATMP classification report. CAT adopted by consensus the ATMP classification report. CAT secretariat to send the draft scientific recommendation to the European Commission for comments by 29 May 2017.

The CAT recommendation will be considered final if no comments are received from the European Commission. The final report will be sent thereafter to the applicant.

Even though not requested by the applicant in this classification application, CAT discussed the status of the combination of this matrix with stromal vascular cells (which is the intended function of the matrix). This information will be included in the cover letter to the applicant.

4.2.8. Allogeneic unexpanded amniotic fluid derived cells suspended with dried and cryofractured amniotic tissue - EMA/H0004816

Intended for the treatment of chronic wound care

Scope: scientific recommendation

Action: for adoption

CAT discussed the ATMP classification report. CAT adopted by consensus the ATMP classification report. . CAT secretariat to send the draft scientific recommendation to the European Commission for comments by 29 May 2017.

The CAT recommendation will be considered final if no comments are received from the European Commission. The final report will be sent thereafter to the applicant.

4.2.9. Human autologous stromal vascular fraction (SVF) - EMA/H0004822

Intended for the treatment of articular cartilage and bone defects

Scope: scientific recommendation

Action: for adoption

CAT discussed the ATMP classification report. CAT adopted by consensus the ATMP classification report. . CAT secretariat to send the draft scientific recommendation to the European Commission for comments by 29 May 2017.

The CAT recommendation will be considered final if no comments are received from the European Commission. The final report will be sent thereafter to the applicant.

4.2.10. Human autologous adipose-derived stromal/stem cells (ADSCs) - EMA/H0004823

Intended for the treatment of articular cartilage and bone defects

Scope: scientific recommendation

Action: for adoption

CAT discussed the ATMP classification report. CAT adopted by consensus the ATMP classification report. . CAT secretariat to send the draft scientific recommendation to the European Commission for comments by 29 May 2017.

The CAT recommendation will be considered final if no comments are received from the European Commission. The final report will be sent thereafter to the applicant.

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

4.3.1. Cultured autologous Wharton jelly derived mesenchymal stem cells - EMA/H0004795

Intended for the treatment of amyotrophic lateral sclerosis (ALS)

Scope: revised scientific recommendation

Action: for adoption

CAT discussed the ATMP classification report. CAT adopted by consensus the ATMP classification report. CAT secretariat to send the draft scientific recommendation to the European Commission for comments by 29 May 2017.

The CAT recommendation will be considered final if no comments are received from the European Commission. The final report will be sent thereafter to the applicant.

4.3.2. Cultured allogeneic Wharton jelly derived mesenchymal stem cells - EMA/H0004796

Intended for the treatment of amyotrophic lateral sclerosis (ALS)

Scope: revised scientific recommendation

Action: for adoption

CAT discussed the ATMP classification report. CAT adopted by consensus the ATMP classification report. CAT secretariat to send the draft scientific recommendation to the European Commission for comments by 29 May 2017.

The CAT recommendation will be considered final if no comments are received from the European Commission. The final report will be sent thereafter to the applicant.

4.3.3. Cultured autologous adipose derived regenerative mesenchymal stem cells - EMA/H0004797

Intended for the treatment of autoimmune drug resistant epilepsy

Scope: revised scientific recommendation

Action: for adoption

CAT discussed the ATMP classification report. CAT adopted by consensus the ATMP classification report. CAT secretariat to send the draft scientific recommendation to the European Commission for comments by 29 May 2017.

The CAT recommendation will be considered final if no comments are received from the European Commission. The final report will be sent thereafter to the applicant.

4.3.4. Autologous adipose derived mesenchymal stem cells - EMA/H0004798

Intended for the treatment of autoimmune drug resistant epilepsy

Scope: revised scientific recommendation

Action: for adoption

CAT discussed the ATMP classification report. CAT adopted by consensus the ATMP classification report. . CAT secretariat to send the draft scientific recommendation to the European Commission for comments by 29 May 2017.

The CAT recommendation will be considered final if no comments are received from the European Commission. The final report will be sent thereafter to the applicant.

4.3.5. Cultured autologous adipose derived mesenchymal stem cells - EMA/H0004799

Intended for the treatment of autoimmune drug resistant epilepsy

Scope: revised scientific recommendation

Action: for adoption

CAT discussed the ATMP classification report. CAT adopted by consensus the ATMP classification report. CAT secretariat to send the draft scientific recommendation to the European Commission for comments by 29 May 2017.

The CAT recommendation will be considered final if no comments are received from the European Commission. The final report will be sent thereafter to the applicant.

4.4. Finalisation of procedure

4.4.1. Stimulated resistant cells suspension cancer vaccine - H0004763

Intended for the treatment of colorectal cancer

Scope: revised scientific recommendation following comments by the European Commission

Action: for adoption

The classification report was amended in response to the comments raised by the European Commission. The revised report was adopted by consensus and will be sent to the applicant.

4.4.2. Human induced pluripotent stem cell derived natural killer cells expressing high-affinity non-cleavable CD16 Fc - EMA/H0004784

Intended for the treatment of advanced solid tumour malignancies

Scope: No comments raised by the European Commission

Action: for information

The CAT noted this information.

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 Coordinators

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

No items

6.2. ITF briefing meetings in the field of ATMPs

6.3. Priority Medicines (PRIME) – Eligibility requests

6.3.1. Month 1 – Discussion of eligibility

No items

6.3.2. Month 2 – Recommendation of eligibility

6.3.3. Month 3 – Nomination of Rapporteurs

6.3.4. Ongoing support

7. Organisational, regulatory and methodological matters

7.1. Mandate and organisation of the CAT

7.1.1. Strategic Review & Learning meeting – Malta, June 2017

CAT Strategic Review & Learning meeting (SRLM) will take place in Gozo, Malta on 1-2 June 2017 under the auspices of the Maltese Presidency of the Council of the European Union

CAT: John-Joseph Borg

Scope: draft programme

Action: for information

CAT discussed the draft programme and proposed amendments to take into account the availability of CAT members. There was a short discussion on the possibility to develop guidance already on gene editing technologies (see also 7.3.2). It was agreed that a discussion on this topic could take place during the upcoming SRLM meeting.

7.1.2. Good manufacturing practice (GMP) for advanced therapy medicinal products

CAT: Ilona Reischl

Scope: feedback from drafting group meeting

Action: for information

Feedback was provided on the discussion that took place during the drafting group meeting with CAT experts and GMP inspectors. The aim was to discuss and make final changes to the GMP for ATMP guideline. As a next step the updated document will be sent to the GMP-Inspectors Working Group and the CAT for final discussion and endorsement.

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 18-21 April 2017 meeting

Action: for information

The CAT noted this information.

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

7.3.1. Guideline on quality, non-clinical and clinical aspects of gene therapy medicinal products

CAT: drafting group (DG) members:
-Quality: M. Menezes-Ferreira, C. Niederlaender, S. Ruiz and P. Salmikangas
-Non-clinical: K. Breen, B. Sarkadi, M. Renner, Tiina Palomäki
-Clinical: P. Gasparini, B. Klug, M. Hystad, O. Tenhunen, B. Gänsbacher

Scope: feedback on the revision of the guideline by the DG in their meeting of 10 May 2017

Action: for discussion

A short feedback was provided from the outcome of the drafting group meeting. The document is now almost ready (pending editorial changes and a discussion of the comment on the pharmacovigilance section). The updated guideline will be presented to the CAT in June 2017.

7.3.2. Guideline on genetically modified cells

CAT: Marcos Timón, Paolo Gasparini, Olli Tenhunen

Scope: review of draft concept paper; call for additional clinical expertise to join the drafting group

Action: for discussion

The concept paper for the revision of this guideline was presented. CAT members were asked to provide comments by 5 June 2017.

CAT appointed Paolo Gasparini and Olli Tenhunen to join the drafting group (clinical aspects).

7.4. Cooperation within the EU regulatory network

7.4.1. Orphan similarity for ATMPs

CAT drafting group: Simona Badoi, Violaine Closson-Carella, Michele Lipucci, Margarida Menezes-Ferreira, Christiane Niederlaender, Ilona Reischl, Paula Salmikangas

Scope: Reflection from the perspective of ATMPs on the concept of 'similar active substance' as referred to in Art 3(3)c of Reg (EC) No 847/2000 of April 2000 laying down the provisions for implementation of the criteria for designation of a medicinal product as an orphan medicinal product and definitions of the concept 'similar medicinal product' and 'clinical superiority'. Review of comments received from the public consultation

Action: for discussion

Consultation document published by the European Commission
http://ec.europa.eu/health/sites/health/files/files/orphanmp/2016_07_pc_orphan/2016_07_consultation_paper.pdf

The comments received on the European Commission's consultation document were discussed by the CAT drafting group in the margins of the April CAT meeting.

The CAT input in the orphan similarity proposal (ATMP part) and the overview of comments was presented. Due to time constraints, it was agreed that the documents will be sent to the CAT members for agreement via a written procedure.

Post-meeting note: Following the written procedure, it was decided to put this item on the June 2017 CAT meeting agenda for discussion.

7.5. Cooperation with international regulators

7.5.1. International Pharmaceutical Regulators Forum (IPRF) gene therapy working group (GTWG)

Scope: feedback from the meeting of 2-3 May 2017 on the topic of IPRF biodistribution reflection paper studies

Action: for information

Agenda item postponed to the June 2017 CAT meeting.

7.6. CAT work plan

7.6.1. Expert meeting on adeno-associated viral vectors

Scope: feedback from the organising committee

Action: for information

Note: expert meeting to take place on 06 September 2017

Feedback was provided from the discussions in the Organising Committee (meetings on 8 and 10 May). Experts have been identified and a draft agenda is being prepared: this will be further discussed in the CAT SRLM on 1 June 2017 (see 7.1.1).

7.7. Planning and reporting

None

7.8. Others

7.8.1. EMA framework of collaboration with academia

Scope: presentation on the scope, objectives and methodology

Action: for information

Note: The EMA's management board adopted the framework on 16 March 2017

EMA presented the framework of collaboration with academia, as agreed by the EMA Management Board. It was noted that the EMA framework includes the collaboration with the innovation network (Innovation offices) in the member states. CAT members noted that for academia, the national authorities should be the first point of call for advice. Another comment made was the need to develop procedures for academia (or simplify existing procedures) to facilitate the interactions with the EMA.

7.8.2. EMA's ATMP matrix team

Scope: presentation of the new EMA – ATMP team

Action: for discussion

The EMA ATMP team was presented to CAT.

7.8.3. Impact of Brexit on committees

Scope: presentation and Q&A on the impact of Brexit on the work of the CAT

Action: for information

8. Any other business

No items

Date of next CAT meeting:
15-16 June 2017. The meeting be held via Adobe Connect (virtual)

9. 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

DNA: Deoxyribonucleic acid

DG: Drafting Group

EC: European Commission

EDQM: European Directorate for the Quality of Medicines

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

GTMP: Gene therapy medicinal product

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

MNAT: Multinational Assessment Team

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

RNA: Ribonucleic acid

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

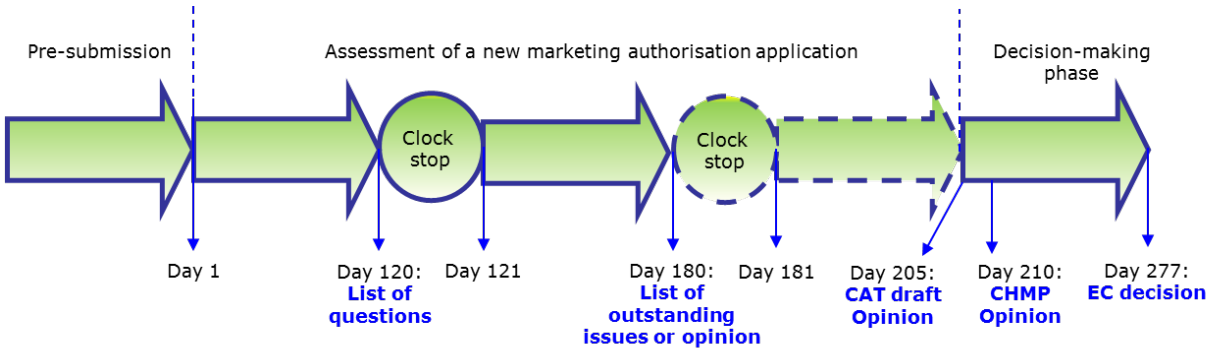
Evaluation of ATMPs (section 2)

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.

PRIME

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/

10. List of participants

including any restrictions with respect to involvement of members / alternates / experts following evaluation of declared interests for the 10-12 May 2017 meeting.

Name	Role	Member state or affiliation	Outcome restriction following evaluation of e-DoI	Topics on agenda for which restrictions apply
Martina Schüssler-Lenz	Chair	Germany	No interests declared	
Ilona Reischl	Member	Austria	No interests declared	
Claire Beuneu	Member	Belgium	No interests declared	
Belaïd Sekkali	Alternate	Belgium	No interests declared	
Rozalina Kulaksazova	Member	Bulgaria	No interests declared	
Ivica Malnar	Alternate	Croatia	No interests declared	
Marina Ieridi	Member	Cyprus	No interests declared	
Ivana Haunerova	Alternate. Replacing member.	Czech Republic	No interests declared	
Anne Pastoft	Alternate. Replacing member.	Denmark	No interests declared	
Tarmo Tiido	Alternate. Replacing member.	Estonia	No interests declared	
Paula Salmikangas	Member	Finland	No interests declared	
Violaine Closson	Member	France	No interests declared	
Jan Mueller-Berghaus	Member	Germany	No interests declared	
Egbert Flory	Alternate	Germany	No interests declared	
Asterios Tsiftsoglou	Member	Greece	No interests declared	
Angeliki Roboti	Alternate	Greece	No interests declared	
Krisztian Fodor	Member	Hungary	No interests declared	
Niamh Curran	Alternate. Replacing member.	Ireland	No interests declared	
Paolo Gasparini	Member	Italy	No interests declared	
Una Riekstina	Member	Latvia	No interests declared	
Romaldas Mačiulaitis	Member (CHMP member)	Lithuania	No restrictions applicable to this meeting	
Johannes Hendrikus Ovelgönne	Member	Netherlands	No interests declared	
Rune Kjeken	Alternate. Replacing member.	Norway	No restrictions applicable to this meeting	
Dariusz Śladowski	Member	Poland	No restrictions applicable to this meeting	
Margarida Menezes-Ferreira	Alternate (to CHMP representative). Replacing member.	Portugal	No interests declared	
Simona Badoi	Member	Romania	No interests declared	
Ján Kyselovič	Alternate. Replacing	Slovakia	No interests declared	

Name	Role	Member state or affiliation	Outcome restriction following evaluation of e-DoI	Topics on agenda for which restrictions apply
	member.			
Metoda Lipnik-Stangelj	Member	Slovenia	No interests declared	
Marcos Timón	Alternate (to CHMP representative)	Spain	No interests declared	
Lennart Åkerblom	Member	Sweden	No interests declared	
Christiane Niederlaender	Member	United Kingdom	No interests declared	
James McBlane	Alternate	United Kingdom	No interests declared	
Marc Turner	Member	Healthcare Professionals' Representative	No restrictions applicable to this meeting	
Bernd Gänsbacher	Member	Healthcare Professionals' Representative	No interests declared	
Kieran Breen	Member	Patients' Representative	No restrictions applicable to this meeting	
Mariëtte Driessens	Member	Patients' Representative	No restrictions applicable to this meeting	
Erik Briers	Alternate	Patients' Representative	No restrictions applicable to this meeting	
Barbara Bonamassa	Expert - in person*	Italy	No interests declared	
Tiina Palomäki	Expert - in person*	Finland	No interests declared	
Christos Sotirelis	Expert - in person*	Patients' Representative	No interests declared	
Tony Maderson	Expert - in person*	Visitor from TGA - Australia	No interests declared	
Tuomo Lapveteläinen	Expert - via telephone*	Finland	No interests declared	
Markus Funk	Expert - via telephone*	Finland	No interests declared	
Monique Wakelkamp	Expert - via telephone*	Sweden	No restrictions applicable to this meeting	
Kristina Dunder	Expert - via telephone*	Sweden	No interests declared	
A representative from the European Commission attended the meeting.				
Meeting run with support from relevant EMA staff				

* Experts were only evaluated against the agenda topics or activities they participated in.