4 November 2020
EMA/CAT/647967/2020
Human Medicines Division

Committee for Advanced Therapies (CAT)

Minutes the meeting on 07-09 October 2020

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

Disclaimers

Some of the information contained in these minutes are considered commercially confidential or sensitive and therefore not disclosed. Regarding 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, these 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

The Chairperson opened the 7-9 October 2020 meeting by welcoming all participants. In light of the current crisis (COVID-19 outbreak), the EMA Business Continuity Plan (BCP) and exceptional measures taken to protect the staff members and all delegates, experts and members of the Committee are maintained. This entails that the participation and the voting from remote are allowed, based on the current exceptional circumstances.

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. No new or additional interests or restrictions were declared.

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

The CAT chair welcomed Tomas Boráň as new alternate member for the Czech Republic.

1.2. Adoption of agenda

The CAT agenda for 07-09 October 2020 meeting was adopted.

1.3. Adoption of the minutes

The CAT minutes of the 09-11 September 2020 meeting were adopted with an addition of agenda point 2.3.2 and a correction in agenda point 5.1.3.

2. Evaluation of ATMPs

2.1. Opinions

2.1.1. Autologous peripheral blood T cells CD4 and CD8 selected and CD3 and CD28 activated transduced with retroviral vector expressing anti-CD19 CD28/CD3-zeta chimeric antigen receptor and cultured - Orphan - EMEA/H/C/005102

Kite Pharma EU B.V.; treatment of adult patients with relapsed or refractory Mantle cell

lymphoma (MCL)

Scope: Opinion

Action: for adoption

List of Outstanding Issues adopted on 11.09.2020. List of Questions adopted on 20.05.2020.

The Rapporteurs presented their assessment of the responses to the list of outstanding issues.

The CAT draft opinion and assessment report were adopted by consensus.

2.2. Oral explanations

2.2.1. Autologous CD34+ cell enriched population that contains hematopoietic stem and progenitor cells transduced ex vivo using a lentiviral vector encoding the human arylsulfatase A gene - Orphan - EMEA/H/C/005321

Orchard Therapeutics (Netherlands) BV; treatment of metachromatic leukodystrophy (MLD)

Scope: oral explanation

Action: oral explanation

List of Outstanding Issues adopted on 11.09.2020. List of Questions adopted on 20.03.2020.

The Rapporteurs presented their assessment of the responses to the list of outstanding issues.

The CAT draft opinion and assessment report were adopted by consensus.

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

2.5.1. Lisocabtagene maraleucel - Orphan - EMEA/H/C/004731

Accelerated assessment

Celgene Europe BV; treatment of large B-cell lymphoma, diffuse large B-cell lymphoma (DLBCL), primary mediastinal large B-cell lymphoma (PMBCL) and follicular lymphoma grade 3B (FL3B)

Scope: Day 80 assessment report

Action: for information

CAT members were asked to provide comments on the Day 80 assessment report.

2.6. Update on ongoing initial applications

No items

2.7. New applications

2.7.1. Autologous glioma tumour cells, inactivated / autologous glioma tumour cell lysates, inactivated / allogeneic glioma tumour cells, inactivated / allogeneic glioma tumour cell lysates, inactivated - Orphan - EMEA/H/C/003693

Epitopoietic Research Corporation-Belgium (E.R.C.); treatment of glioma

Scope: Timetable for assessment

Action: for adoption

The assessment timetable was adopted.

2.7.2. Elivaldogene autotemcel - Orphan - EMEA/H/C/003690

Accelerated assessment

bluebird bio (Netherlands) B.V; treatment of patients less than 18 years of age with an *ABCD1* genetic mutation and early cerebral adrenoleukodystrophy

Scope: Timetable for assessment

Action: for adoption

The assessment timetable was adopted.

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. GMP and GCP inspections requests

2.10.1. Good manufacturing practice (GMP) - forthcoming inspections for ATMPs

Scope: forthcoming inspections

Action: for information

Note: these requests were adopted by CHMP at its September 2020 meeting

The information was noted.

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

2.11.1. Kymriah - tisagenlecleucel - Orphan - EMEA/H/C/004090/II/0026/G

Novartis Europharm Limited

Rapporteur: Rune Kjeken

Scope: Quality. Request for supplementary information (RSI)

Action: for adoption

The RSI was adopted.

2.11.2. Kymriah - tisagenlecleucel - Orphan - EMEA/H/C/004090/II/0027

Novartis Europharm Limited

Rapporteur: Rune Kjeken

Scope: Quality. Request for supplementary information (RSI)

Action: for adoption

The company will be reminded that, under the terms of the EU marketing authorisation, a dose can constitute one or more bags. A change to the question 3 of the RSI was proposed . The revised RSI was adopted.

2.11.3. Yescarta - axicabtagene ciloleucel - Orphan - EMEA/H/C/004480/II/0028

Kite Pharma EU B.V.

Rapporteur: Jan Müller-Berghaus, PRAC Rapporteur: Anette Kristine Stark

Scope: Clinical. To update SmPC sections; 4.4 on cytokine release syndrome (CRS) grading and neurologic adverse reactions; 4.8 on safety profile summary; 5.1 on follow up analysis; to update the safety information based on updates from study KTE-C19-101, entitled "A Phase 1/2 Multicentre Study Evaluating the Safety and Efficacy of KTE-C19 in Subjects with refractory aggressive non-Hodgkin lymphoma (ZUMA-1)", the pivotal study for Yescarta. The updates include the Phase 2 safety management ZUMA-1 cohort 4, which was intended to assess the impact of earlier interventions (tocilizumab and/or corticosteroids, in addition to prophylactic levetiracetam) on the rate and severity of CRS and neurologic events; and data from a 36-month analysis from ZUMA-1 Cohorts 1 and 2. The updated RMP version 3.1 has also been submitted. Request for supplementary information (RSI).

Action: for adoption

The Rapporteurs presented the outcome of the assessment. The RSI was adopted.

2.11.4. Zolgensma - onasemnogene abeparvovec - Orphan - EMEA/H/C/004750/II/0003/G

AveXis EU Limited

Rapporteur: Hans Ovelgönne

Scope: Quality. Opinion
Action: for adoption

Request for Supplementary Information adopted on 11.09.2020.

The variation was adopted.

2.11.5. Zolgensma - onasemnogene abeparvovec - Orphan - EMEA/H/C/004750/II/0006

AveXis EU Limited

Rapporteur: Hans Ovelgönne

Scope: Quality. Request for supplementary information (RSI)
Action: for adoption

The RSI was adopted.

2.12. Extension applications

No items

2.13. Other Post-Authorisation Activities

2.13.1. Holoclar - ex vivo expanded autologous human corneal epithelial cells containing stem cells - Orphan - EMEA/H/C/002450/R/0032

Holostem Terapie Avanzate s.r.l.

Rapporteur: Egbert Flory, Co-Rapporteur: Paolo Gasparini, PRAC Rapporteur: Rhea Fitzgerald

Scope: 1-year Renewal of Marketing Authorisation

Action: for adoption

CAT noted the request for a 1-year delay to finalise to agreed conditions, due to the Covid-19 pandemic.

The renewal was adopted, as well as the request of extension of the due date for the obligation.

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

No items

4. Scientific Recommendation on Classification of ATMPs

Timetable:

-Start of the procedure:	09.10.2020
-Draft CAT co-ordinator's report:	23.10.2020
-ITF peer-review comments:	28.10.2020
-Revised scientific recommendation:	30.10.2020
-Adoption of scientific recommendation by CAT:	06.10.2020

4.1. New requests – Appointment of CAT Coordinator

4.1.1. Autologous CD34+ cells transduced with a lentiviral vector encoding human cystinosin

Intended for the treatment of cystinosis

Scope: appointment of CAT Coordinator and adoption of timetable

Action: for adoption

The CAT coordinator was appointed.

4.1.2. Autologous tumour-infiltrating lymphocytes

Intended for the treatment of advance melanoma

Scope: appointment of CAT Coordinator and adoption of timetable

Action: for adoption

The CAT coordinator was appointed.

4.1.3. Delolimogene mupadenorepvec (oncolytic adenovirus expressing two immunostimulatory transgenes (TMZ-CD40L and 4-1BBL))

Intended for the treatment of cancer

Scope: appointment of CAT Coordinator and adoption of timetable

Action: for adoption

The CAT coordinator was appointed.

4.1.4. Allogeneic cord tissue-derived mesenchymal stromal cells

Intended for the treatment of inflammatory and immunological diseases (acute graft-versus-host disease, systemic lupus erythematosus, systemic sclerosis, acute respiratory distress syndrome)

Scope: appointment of CAT Coordinator and adoption of timetable

Action: for adoption

The CAT coordinator was appointed.

4.2. Day 30 ATMP scientific recommendation

4.2.1. 3D bio-printed bionic pancreas composed of islets of Langerhans and non-viable printable porcine-derived matrix plus porcine–derived decellularised blood vessel – H0005801

Intended for the treatment of late-chronic pancreatitis

Scope: ATMP scientific recommendation

Action: for adoption

CAT discussed the ATMP classification report. Additional information and clarifications are needed before concluding on this classification. The applicant is asked to address the questions

agreed by CATThe list of issues was adopted by CAT and the procedure is stopped awaiting responses from the applicant.

4.2.2. 3D bio-printed bionic pancreas composed of insulin- and glucagon-releasing cells and non-viable printable porcine-derived matrix plus porcine–derived decellularised blood vessel – H0005802

Intended for the treatment of brittle diabetes mellitus type I

Scope: ATMP scientific recommendation

Action: for adoption

CAT discussed the ATMP classification report. Additional information and clarifications are needed before concluding on this classification. The applicant is asked to address the questions agreed by CAT. The list of issues was adopted by CAT and the procedure is stopped awaiting responses from the applicant.

4.2.3. Recombinant serotype 9 adeno-associated virus (rAAV9) encoding a codon-optimised human neuronal ceroid lipofuscinosis-5 (CLN5) transgene – H0005800

Intended for the treatment of neuronal ceroid lipofuscinosis type 5

Scope: ATMP scientific recommendation

Action: for adoption

CAT discussed the ATMP classification report. CAT secretariat to send the draft scientific recommendation to the European Commission for comments by 30 October 2020.

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. Autologous adipose-derived mesenchymal stem cell , diabetic foot syndrome - H0005699

Intended for the treatment of diabetic foot syndrome

Scope: Responses from the applicant to the LoQs. Revised ATMP scientific recommendation

Action: for adoption

CAT discussed the revised ATMP classification report. CAT secretariat to send the draft scientific recommendation to the European Commission for comments by 30 October 2020.

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. Allogeneic CRISPR/Cas9-mediated genetically modified chimeric antigen receptor (CAR) T-cells targeting CD70 - H0005771

Intended for the treatment or renal cell carcinoma and haematological malignancies

Scope: the European Commission raised no comments. ATMP scientific recommendation

Action: for information

The information was noted.

4.4.2. Umbilical cord derived CD34+ cells expanded and umbilical cord derived non-expanded CD34- cells - H0005772

Intended for the treatment in haematopoietic stem cell transplantation

Scope: the European Commission raised no comments. ATMP scientific recommendation

Action: for information

The information was noted.

4.4.3. Autologous human endometrial stem cells - H0005773

Intended for the treatment of stem cell therapy for ovarian insufficiency includes diminished ovarian reserve (DOR), premature ovarian failure (POF), primary ovarian insufficiency (POI) and poor ovarian response (POR)

Scope: the European Commission raised minor comments. ATMP scientific recommendation

Action: for information

The information was noted.

4.4.4. Irradiated allogeneic induced-pluripotent stem cells expressing pluripotent genes and cancer-specific embryonic neo-antigens – H0005108/0002

Intended for the treatment malignant solid tumours including all epithelial cancers in sub-group type harbouring a stemness mesenchymal-like signature and haematopoietic malignancies

Scope: the European Commission raised no comments. ATMP scientific recommendation

Action: for information

The information was noted.

4.4.5. Autologous human T cells genetically modified ex vivo with a lentiviral vector encoding a chimeric antigen receptor (CAR) directed against G protein-coupled receptor family C group 5 member D (GPRC5D) - H0005774

Intended for the treatment of patients with relapsed or refractory multiple myeloma

Scope: the European Commission raised no comments. ATMP scientific recommendation

Action: for information

The information was noted.

4.4.6. Recombinant serotype 9 adeno-associated virus (rAAV9) encoding a codon-optimised human neuronal ceroid lipofuscinosis-7 (CLN7) transgene - H0005770

Intended for the treatment of neuronal ceroid lipofuscinosis type 7

Scope: the European Commission raised no comments. ATMP scientific recommendation

Action: for information

The information was noted.

4.4.7. Autologous regulatory T lymphocytes (Treg), with the marker profile of CD3+, CD4+, CD25high, CD127-, FoxP3+ - H0004575/0002

Intended for the treatment and prevention of progression of, multiple sclerosis (MS) [relapsing remitting (RRMS), primary progressing (PPMS), secondary progressing (SPMS)]

Scope: the European Commission raised no comments. ATMP scientific recommendation

Action: for information

The information was noted.

4.4.8. Adeno-associated viral vector serotype 9 encoding human ATP7B - H0005775

Intended for the treatment of Wilson disease

Scope: the European Commission raised no comments. ATMP scientific recommendation

Action: for information

The information was noted.

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

Timetable:

-Final Briefing Package:	23.10.2020
-Start of the procedure at SAWP:	29.10.2020
-CAT report due by:	29.10.2020
-CAT recommendation:	06.11.2020

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 0 - Start of the procedure

Timetable for assessment:

Procedure start:	01.10.2020
SAWP recommendation:	29.10.2020
CAT recommendation:	06.11.2020
CHMP adoption of report and final recommendation:	12.11.2020

6.3.2. Month 1 – Discussion of eligibility

6.3.3. Month 2 – Recommendation of eligibility

6.3.4. Ongoing support

7. Mandate and organisation of the CAT

7.1.1. CAT membership

Czech Republic: Tomáš Boráň – membership mandate started on 01 October 2020

Action: for information

The information is noted. The chair welcomed the new alternate from the Czech Republic.

7.1.2. Strategic Review & Learning (virtual) meeting (SRLM) under the German presidency, 22 October 2020

CAT: Martina Schüßler-Lenz, Egbert Flory

Scope: final agenda

Action: for discussion

Note: to further develop the agenda for the joint meeting, a teleconference (Adobe Connect) with all interested CAT members took place on 28 September 2020.

The final agenda of the joint CAT-PRAC-PDCO session was presented.

7.1.3. CAT meeting dates for the period 2022-2024

Action: for adoption

The meeting dates for 2022-2024 were adopted.

7.2. Coordination with EMA Scientific Committees

7.2.1. CAT-PDCO interaction

CAT: Martina Schüßler-Lenz, Ilona Reischl, Maja Sommerfelt, Claire Beuneu
PDCO: Koen Norga (PDCO Chair), Sabine Scherer (PDCO Vice Chair), Karen van Melderen (Non-clinical expert), Sylvie Benchetrit (Clinical expert)

Scope: feedback from the brainstorming meeting that took place on 17 September 2020

Action: for information

Topics and proposal discussed at the brainstorming meeting on CAT-PDCO interactions were presented. CAT members made some additional proposals .

This topic might be discussed further during the joint SRLM (see 7.1.2).

7.2.2. New Readers Guidance template

Scope: introduction to the new template

Action: for information

The new template of the rolling Readers Guidance was presented. The aim of this document was explained (prepared for the use of Committee members, to identify the list of topics for discussion, but not replacing the Overview). The rolling document will include the previous reader guidance for products and therefore allows the committee members to easily retrieve previous issues identified and discussed during the evaluation procedure.

CAT members can find the template in .

7.2.3. Scientific Coordination Board (SciCoBo) – meeting of 21 September 2020

CAT: Martina Schüßler-Lenz

Scope: feedback on the outcome of the SciCoBo meeting that took place on 21 September 2020

Action: for information

A short feedback was provided from the last SciCoBo meeting.

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

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

CAT Rapporteurs: Martina Schüßler-Lenz, Marcos Timón, Tiina Palomäki

Scope: Guideline

Action: for adoption

Note: comments from the public consultations have been incorporated.

Further to presentation of the draft guideline during the July 2020 CAT meeting, comments from CAT and BWP members were received and incorporate where considered appropriate (in consultation with the drafting group members). Thereafter, the guideline was sent to the Guideline Consistency Group (GCG) for final agreement. The Biostatistics Working Party was consulted on request of the GCG to discuss one paragraph in the clinical efficacy section: the revised paragraph was presented.

BWP during its September meeting already agreed the content of the quality part of the guideline.

CAT subsequently adopted the revision of the Guideline on quality, non-clinical and clinical aspects of medicinal products containing genetically modified cells. The revised guideline will be published after formal adoption by CHMP.

The Rapporteur (Marcos Timon) thanked everyone who was involved in the revision of this guideline.

7.3.2. Biostatistics Working Party (BSWP) – reflection paper on the importance of randomisation for confirmatory evidence

CAT: Martina Schüssler-Lenz

Scope: CAT representative to take part in the development of the 'reflection paper on the importance of randomisation for confirmatory evidence'

Action: for appointment

The BSWP is developing the above-mentioned reflection paper, which will include guidance on single arm trials (SAT). Because many ATMPs include SAT, involvement of CAT in the development of the guidance is welcomed. A call for interest for volunteer in the drafting was presented to CHMP (ORGAM) and to CAT, for nomination by 16 October 2020.

Post meeting note: Carla Herberts was nominated as CAT representative, Jan Mueller Berghaus will join as CHMP representative.

7.3.3. Draft BWP/QWP/IWG toolbox guidance on scientific elements and regulatory tools to support quality data packages for PRIME marketing authorisation applications

Scope: scientific and regulatory approaches for consideration by Applicants to facilitate the development and preparation of robust quality data packages for PRIME products.

Action: for information

Background:
Experience to date has shown that applicants face challenges to complete quality and manufacturing development and data requirements during development of medicines for early access (PRIME). In order to address this, a workshop[1] with industry took place to discuss quality and GMP deficiencies in PRIME products and possible scientific and regulatory approaches which could be used to facilitate development and preparation of robust quality data packages, to enable timely access to medicines for patients whilst providing assurance that patient safety, efficacy and product quality are not compromised.

As a follow-up action, the toolbox guidance was prepared. The toolbox considers which flexibility could be accepted to defer data to the post-approval phase considering the available data and the benefit/risk of the product.

A detailed feedback on the background and content of the toolbox was provided. The European Commission's representative expressed concerns that the possibility to rely on previous knowledge could be misinterpreted for ATMPs and reference was made to the change in Notice to Applicants regarding limitations of literature data in case of ATMPs where there has been substantial manipulation. Additional comments were made on process validation (need to align to GMP for ATMPs), comparability (clarify that, for ATMPs, applicants should follow Q&A on

[1] See meeting report:
https://www.ema.europa.eu/en/documents/report/report-workshop-stakeholders-support-quality-development-early-access-approaches-ie-prime_en.pdf

comparability for ATMPs) and risk-based approach (always applicable to ATMPs even if no PRIME status).
CAT members are asked to provide feedback by end of October 2020 .

7.3.4. Workshop on guideline on registry-based studies

Scope: announcement of a workshop scheduled on Monday, 19 October 2020 from 12:30-17:00 (CET) where the guideline will be presented and where stakeholders will have the opportunity to ask questions for clarification

Action: nomination of CAT member to join the workshop

Note:
-the draft Guideline on registry-based studies has been published for consultation until 31 December 2020: LINK
-an e-mail was sent out on 25.09.20 to all committees inviting participation in the workshop

The agenda of the upcoming workshop was presented. Following CAT members will participate to the workshop: Ilona Reischl, Liesbeth Barkholt, Carla Herberts and Maura O'Donovan.

7.4. Cooperation within the EU regulatory network

7.4.1. Provision on non-conforming batches of ATMPs

Scope: feedback from the IWG discussion on provision on non-conforming ATMPs batches

Action: for discussion

Feedback was provided from the discussion in the IWG on this topic.

7.5. Cooperation with international regulators

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

CAT: Martina Schüssler-Lenz, Ilona Reischl, Violaine Closson-Carella, Rocío Salvador-Roldán

Scope: feedback from the teleconference that took place on 17 September 2020

Action: for information

A short feedback was provided from the discussion at the last ATMP cluster teleconference.

7.6. CAT work plan

None

7.7. Planning and reporting

None

7.8. Others

None

8. Any other business

8.1. Clinical presentation of aromatic l-amino acid decarboxylase (AADC) deficiency

CAT: Martina Schüßler-Lenz

Scope: presentation by Prof. Thomas Opladen, Centre for Paediatric and Adolescent Medicine, Heidelberg University Hospital (Germany)

Action: for information

The presentation from Prof Opladen, an expert for the treatment of children with AADC deficiency, was well received by CAT. The CAT chair, on behalf of the CAT members, thanked Prof Opladen for the interesting presentation and the open discussion with CAT members.

Date of next CAT meeting:

04-06/11/2020

9. Explanatory notes

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

Abbreviations / Acronyms

AAV: Adeno-Associated Virus

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

EU NTC: European Union Network Training Centre

ERA: Environmental Risk Assessment

FDA: Food and Drug Administration

FL: Final Letter

GCG: Guideline Consistency Group

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

IWG: Inspectors Working Group

JR: Joint Report

LoOI: List of outstanding issues

LoQ: List of questions

MA: Marketing Authorisation

MAA: Marketing Authorisation Application

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

QRD: Quality Review of Documents

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

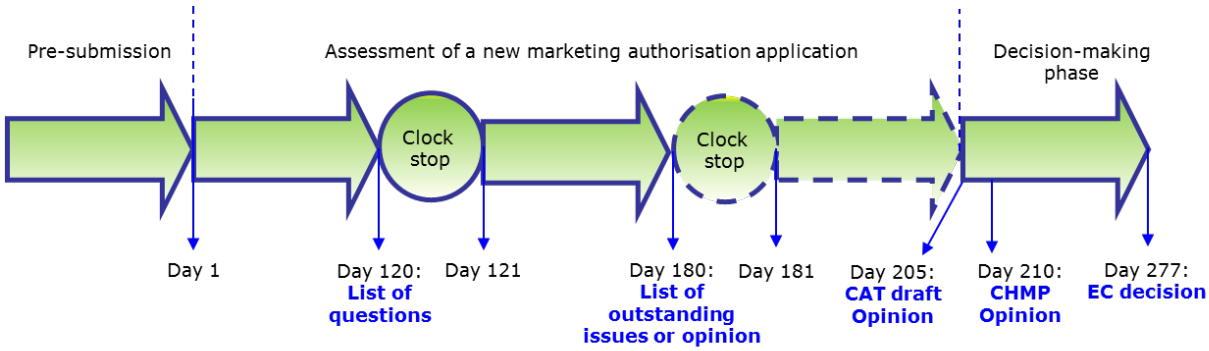
Evaluation of ATMPs (section 2)

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/

10. List of participants

including any restrictions with respect to involvement of members / alternates / experts following evaluation of declared interests for the 07-09 October 2020 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 (Vice-Chair)	Austria	No interests declared	
Silke Dorner	Alternate	Austria	No interests declared	
Claire Beuneu	Member	Belgium	No interests declared	
Rozalina Kulaksazova	Member	Bulgaria	No interests declared	
Petra Sokol	Alternate	Croatia	No interests declared	
Rafaella Pontou	Member	Cyprus	No interests declared	
Ivana Haunerova	Member	Czech Republic	No interests declared	
Tomas Boran	Alternate	Czech Republic	No interests declared	
Anne Pastoft	Member	Denmark	No interests declared	
Toivo Maimets	Member	Estonia	No interests declared	
Pille Saalik	Alternate	Estonia	No interests declared	
Heli Suila	Member	Finland	No interests declared	
Olli Tenhunen	Alternate	Finland	No interests declared	
Violaine Closson	Member	France	No interests declared	
Jan Mueller-Berghaus	Member (CHMP co-opted member)	Germany	No interests declared	
Egbert Flory	Alternate (to CHMP representative)	Germany	No interests declared	
Angeliki Rompoti	Alternate	Greece	No interests declared	
Katalin Lengyel	Member	Hungary	No interests declared	
Maura O'Donovan	Member	Ireland	No interests declared	
Niamh Curran	Alternate	Ireland	No restrictions applicable to this meeting	

Name	Role	Member state or affiliation	Outcome restriction following evaluation of e-DoI	Topics on agenda for which restrictions apply
Paolo Gasparini	Member	Italy	No interests declared	
Giulio Pompilio	Alternate	Italy	No restrictions applicable to this meeting	
Una Riekstina	Member	Latvia	No interests declared	
Romaldas Mačiulaitis	Member (CHMP member)	Lithuania	No interests declared	
Guy Berchem	Member	Luxembourg	No restrictions applicable to this meeting	
John J. Borg	Member (CHMP member)		No interests declared	
Anthony Samuel	Alternate (to CHMP representative)	Malta	No interests declared	
Carla Herberts	Member	Netherlands	No interests declared	
Johannes Hendrikus Ovelgonne	Alternate	Netherlands	No interests declared	
Rune Kjeken	Member	Norway	No restrictions applicable to this meeting	
Maja Sommerfelt Grønvold	Alternate	Norway	No restrictions applicable to this meeting	
Dariusz Śladowski	Member	Poland	No restrictions applicable to this meeting	
Maria Isabel Borba Vieira	Alternate (to CHMP representative)	Portugal	No interests declared	
Felicia Ciulu-Costinescu	Member	Romania	No interests declared	
Lukas Slovak	Member	Slovakia	No interests declared	
Alexandra Padova	Alternate	Slovakia	No interests declared	
Metoda Lipnik-Stangelj	Member	Slovenia	No interests declared	
Sol Ruiz	Member (CHMP co-opted member)	Spain	No interests declared	

Name	Role	Member state or affiliation	Outcome restriction following evaluation of e-DoI	Topics on agenda for which restrictions apply
Marcos Timón	Alternate (to CHMP representative)	Spain	No interests declared	
Lisbeth Barkholt	Member	Sweden	No interests declared	
Maria Luttgen	Alternate	Sweden	No restrictions applicable to this meeting	
Bernd Gänsbacher	Member	Healthcare Professionals' Representative	No interests declared	
Alessandro Aiuti	Member	Healthcare Professionals' Representative	No restrictions applicable to this meeting	No participation in discussions, final deliberations and voting on: 2.2.1 and 5.2.7
Kerstin Sollerbrant	Member	Patients' Representative	No interests declared	
Lydie Meheus	Alternate	Patients' Representative	No interests declared	
Kieran Breen	Member	Patients' Representative	No restrictions applicable to this meeting	
Roland Pochet	Alternate	Patients' Representative	No interests declared	
Giuseppa Pistritto	Expert - virtual	AIFA-IT	No interests declared	
Marja Bovenkamp	Expert - virtual	CBG-MEB-NL	No interests declared	
Marcel Hoefnagel	Expert - virtual	CBG-MEB-NL	No interests declared	
Rou-Afza Gunput	Expert - virtual	CBG-MEB-NL	No interests declared	
Loes den Otter	Expert - virtual	CBG-MEB-NL	No interests declared	
Jolien de Groot	Expert - virtual	CBG-MEB-NL	No interests declared	
Leon Bongers	Expert - virtual	CBG-MEB-NL	No interests declared	
Armando Genazzani	CHMP member	AIFA-IT	No interests declared	
Svetlana Lorenzano	Expert - virtual	AIFA-IT	No restrictions applicable to this meeting	
Antonella Isgrò	Expert - virtual	AIFA-IT	No interests declared	
Miguel Angel	Expert - virtual	AEMPS-ES	No restrictions	

Name	Role	Member state or affiliation	Outcome restriction following evaluation of e-DoI	Topics on agenda for which restrictions apply
Ortiz-Rosales			applicable to this meeting	
Marcel Kwa	Expert - virtual	CBG-MEB-NL	No interests declared	
Ingrid Wang	CHMP member	NOMA-NO	No interests declared	
Kristine Moltu	Expert - virtual	NOMA-NO	No interests declared	
Louise Lauritsen	Expert - virtual	DKMA-DK	No interests declared	
Karin Nylen	Expert - virtual	MPA-SE	No interests declared	
Nuala Kelly	Expert - virtual	HPRA-IE	No interests declared	
Peter Kiely	Expert - virtual	HPRA-IE	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.