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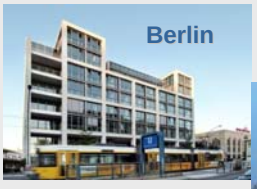
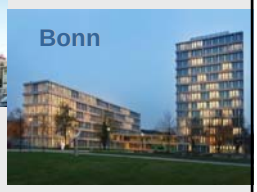
Federal Ministry of Health





European Legislation:

The “New” Regulation on Clinical Trials on Medicinal Products for Human Use

EU Portal and Database

Bundesministerium für Gesundheit
Federal Ministry of Health

1. EU CT Regulation
   - Scope, History, Entry into Force
2. EU Portal & Database (General)
   - Legal Basis, Setting up and Maintenance
   - Working Groups: MS, EC, EMA (Stakeholder)
3. EU Portal & Database (Details)
   - Functional Specifications to be Audited
   - EU Portal and other Systems
   - User Registration and User Rights
4. Summary EU Portal & Database

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EU CT Regulation Scope

REGULATION (EU) No 536/2014 OF THE EUROPEAN PARLIAMENT AND OF THE COUNCIL of 16 April 2014 on clinical trials on medicinal products for human use, and repealing Directive 2001/20/EC

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EU CT Regulation History

17.07.2012	Regulation Proposal made by EC
24.07.2012	Start of Negotiations in the Council in Brussels
09 / 2012	Hearing in Germany in the Ministry of Health
01 / 2013	Decision of German Bundestag (BT Drs 17/12183)
29.05.2013	Voting in ENVI Committee of EP >900 Amendments
20.12.2013	Agreement of Council, EP and Commission on a joint position to the Regulation Proposal (informal trialogue)
02.04.2014	Voting in Plenum of the EP, Approval of ENVI on 22.01.14
14.04.2014	Formal Adoption of the Council
27.05.2014	Publication in the Official Journal of the EU
16.06.2014	Entry into Force

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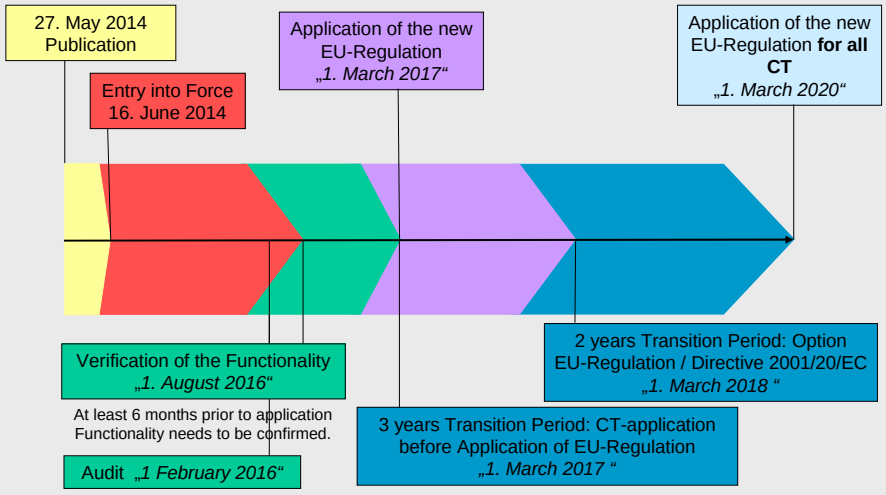
EU CT Regulation
Enty into Force, Transition Periods (Examples using fictious Dates)



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EU Portal & Database
Legal Basis

The EMA shall, in collaboration with the MS and the EC set up and maintain an EU portal & database.
(Regulation (EU) No 536/2014, Articles 80, 81)

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EU Portal & Database
Transparency

The EU database shall be publicly accessible unless, for all or part of the data and information contained therein, **confidentiality is justified** on any of the following grounds (Regulation (EU) No 536/2014, Articles 80, 81)**:**

- Protecting **personal data**
- Protecting **commercially confidential information**
- Protection **confidential communication between MS**
- Ensuring **effective supervision**

CT Application and Approval Process

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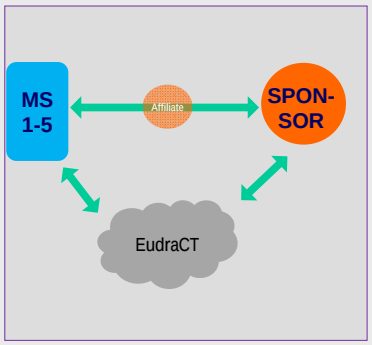
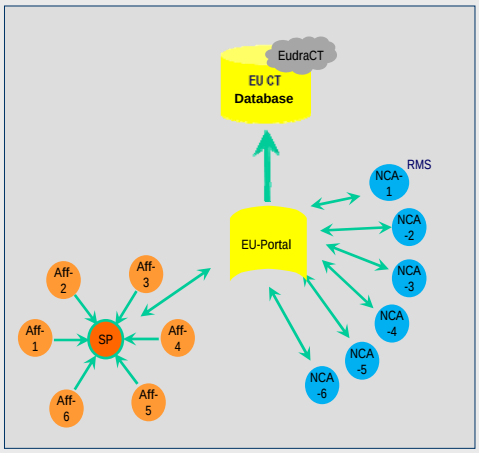
Future Situation

Current Situation





EU Portal & Database Working Groups

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- Expert Group
  to work on a concept "functional specifications to be audited":
  MS (and Ethic Committee) (BE, CZ, DE, IR, NL, ES, SE, UK, EC, EMA)
- Subgroups to work on details
  - A Data fields, forms
  - B Structure and content of the application dossier (Sponsor driven activities)
  - C Process flow (application, validation, evaluation, authorisation) (MS driven activities)
  - D Metadata search and reporting
  - E User roles and access rights (User management and other requirements)
  - F System and reporting
  - G Inspections (Inspections)
  - H Safety reporting
- Group "All MS":
  Information/adoption of proposals drafted by the Expert Group
- EU Portal and Database Stakeholder Meetings: organisations invited by EMA

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   - User Registration, Administration and User Rights
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EU Portal & Database
Functional Specifications

EUROPEAN MEDICINES AGENCY
SCIENCE MEDICINES HEALTH

19 December 2014
EMA/42176/2014
Compliance and Inspections

Functional specifications for the EU portal and EU database to be audited

EU Portal & Database

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Functional specifications – high level business requirements

- ➔ The functional specifications are to describe the high level requirements for the EU portal and database.
- ➔ An Addendum to the functional specifications for defining transparency rules was added.
- ➔ These will be the basis for the audit of the EU portal and database.
- ➔ In parallel, the detailed business requirements that will enable the preparation of the use cases will be collected.
- ➔ Open issues: details transparency rules, project "safety reporting."

EU Portal & Database
Scope of Functional Specifications and Audit

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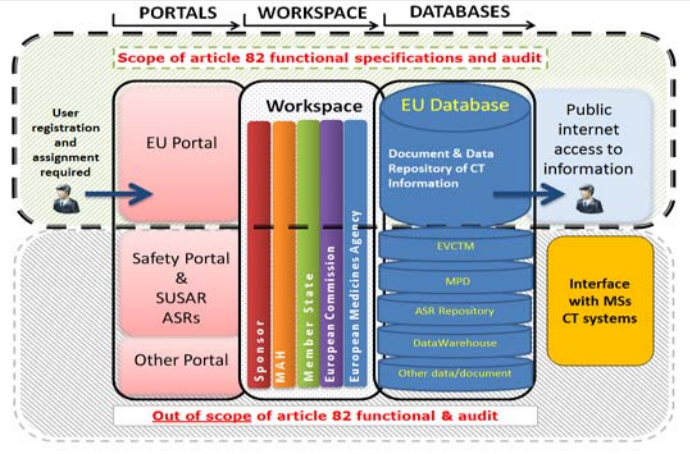

Source EMA

EU Portal & Database
Additional Requirements to be made available in the Workspace (not to be audited)

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1. Communication features for MS general discussion
2. Exchange of documents and data between MS systems and EU system
3. Links to CT systems allowing a full range of statistics
4. Delegation of the RMS role to another MS
5. Initiation of an ad hoc-workflow/case management for assessment of CT related issues (e.g. safety)
6. Cooperation in assessment and surveillance of safety
7. Link to EudraVigilance database
8. Uploading of CT dossiers prepared outside of the workspace

EU Portal & Database
Project Safety Reporting

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➔Upgrade of the Eudra Vigilance CT Modules

➔Creation of an e-web based form for the reporting of SUSARs

➔Development of an Annual Safety Report (ASR) Repository

➔Add a functionality to allow the forwarding of SUSARs and ASRs to the MS

Source EMA

EU Portal & Database: Functional Specifications
Annex 2 (Submission through the EU Portal)

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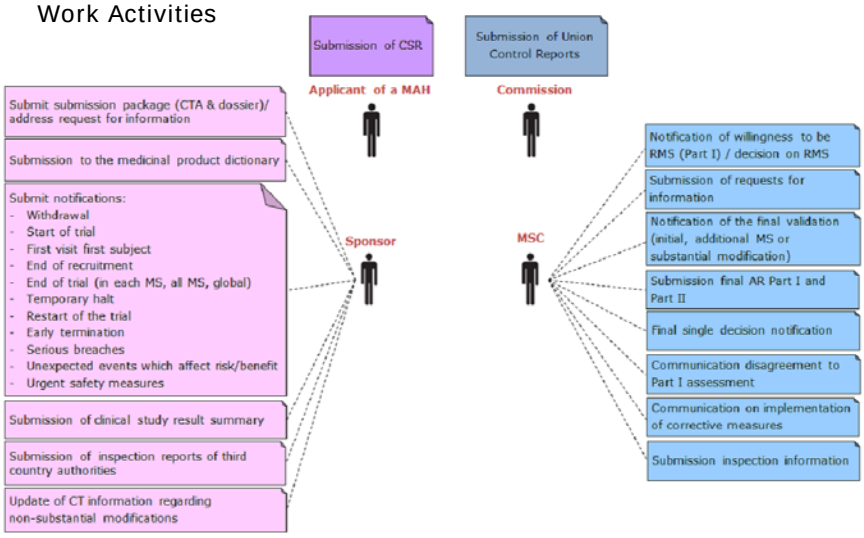
Work Activities



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EU Portal & Database Access

➔Single log in to access workspace and other systems (DWH, EV CTM, ASR repository) and in order to submit to the database through the EU Portal

➔All users will need to be registered in the EMA systems before they can access the EU Portal & Database.

➔A user name and password will be provided to access the system.

➔Access to the Portal functions will be provided according to user's role. The system will display the appropriate data to users and make available the appropriate activities to be executed according to the user's role in the system.

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EU Portal & Database
Involved Parties in Germany

➔Bundesoberbehoerden: BfArM, PEI

➔Ethic Committees

➔“Laender”: Inspectorates

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EU Portal & Database
Users I

<u>Marketing Authorisation Holder</u>

➔To upload and submit the CT reports.

➔To manage “MAH Users”.

<u>European Commission (EC)</u>

➔To upload and submit the Union control reports.

➔To manage “EC Users”.

EU Portal & Database
Users II

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Sponsor

➔To prepare and submit applications, notifications and responses to requests for additional information.

➔To manage “Sponsor Users”.

➔To monitor the work flow of the CT.

MS

➔To prepare and submit the assessment feedback (validation, assessment, decision, answer to questions of MS).

➔To monitor the work flow of the CT.

➔To collaborate between MS (for multi state application).

➔To communicate with sponsor and between MS.

➔To manage “MS Users”.

➔To supervise the CT through out it’s life cycle.

User Structure
Proposed Relationship between Users

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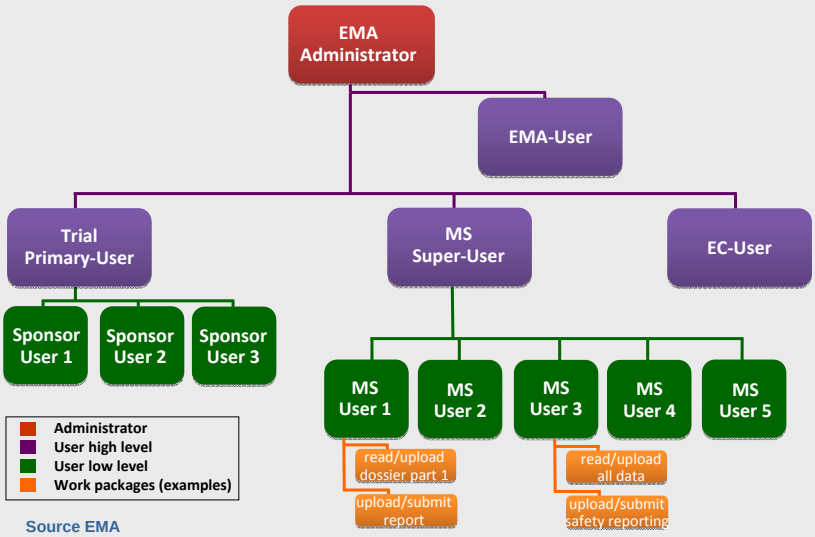

Source EMA

EU Portal & Database
Super User I

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Super User (authority)

➔MS define their super user

➔Maintain “full rights” for all activities from EMA Admin; rights are transferable to another user (Back-Up) -consolidation of work packages for MS users-

➔Access to all documents related to part I, for all CMS regardless whether they are RMS or CMS

Super User (Sponsor/MAH)

➔System allocated and registered upper user automatically during the submission procedure of CT

➔Maintain “full rights” for submitted CT; rights are transferable to other users of this CT -consolidation of work packages-

EU Portal & Database
Super User II (Work Packages)

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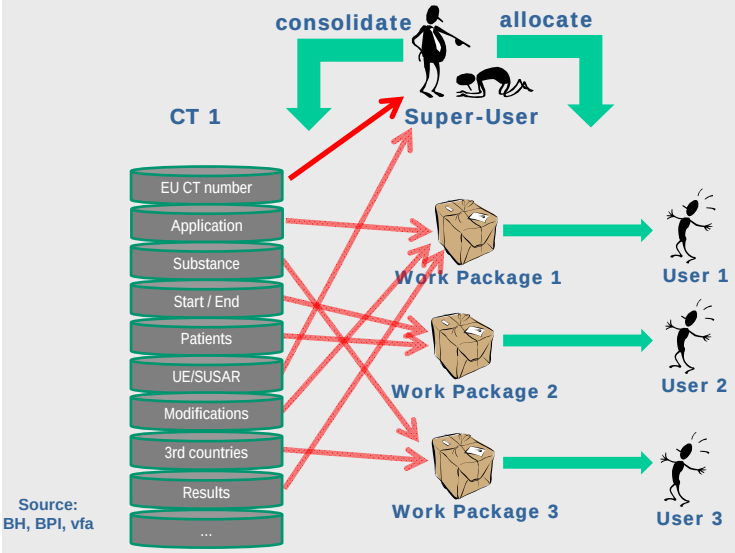

Source:
BH, BPI, vfa

EU Portal & Database
Initial Registration for a new User

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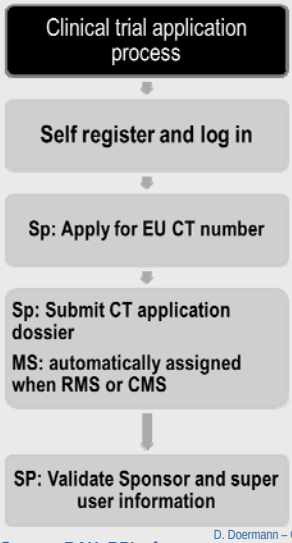

User has no link to sponsor or authority.

- **User has to register himself in the system.**
- **User has no link to sponsor or authority.**
- **User will be assigned automatically to sponsor / super user with all rights. Super user can delegate activities for one or for all general tasks for all CT.**
- **SP User submits the CT dossier.**
- **Pre defined MS super user can assign registered users of the authority for CT conducted in his MS**
- **MS can validated the identity of the sponsor (users)**

Source: BAH, BPI, vfa

EU Portal & Database
Summary I

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- **Set up and maintained in collaboration with EMA, EC and MS**
- **Publicly accessible but protecting personal data, commercially confidential information, confidential communication between MS and ensuring effective supervision**
- **Requirements to be audited are described in document “Functional Specifications” with addendum describing transparency rules**
- **Additional requirements (not to be audited) will be made available in the workspace**

EU Portal & Database Summary II

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- **Single log in to access workspace, EU database and other systems; access to portal functions will be provided according to user's role**
- **Application procedure and user registration are separate procedures and independently**
- **For registration each user must be assigned to an organization, which is "known" by the portal**
- **User management is using the principle of "super user"**
- **Planned verification of functionality in August 2016**

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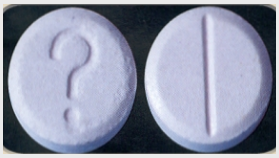
Information related to Clinical Trials



- **http://ec.europa.eu**
  = European Commission
- **http://www.ema.europa.eu**
  = European Medicines Agency
- **http://www.hma.eu**
  = (National) Medicines Authorities in the European Union
- **http://www.bmg.de**
  = Federal Ministry of Health