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AI-Augmented RBQM Solutions

QbD in Action:
ICH E6(R3) Structure Overview

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ICH E6(R3)

Quality by Design in Action

ICH E6(R3)

Structure and Hierarchy

Structure & Hierarchy

Main Guidance Document (ICH E6(R3)):

- **Scope:** Applies to all interventional clinical trials of investigational products intended for regulatory submission.
- **Core Sections:**
  - Principles of Good Clinical Practice (GCP)
  - Institutional roles (e.g., Sponsor, Investigator, IRB/IEC)
  - Data governance and management.
- **Emphasis:** Foundational principles for trial design, conduct, oversight, and data handling, with references to Quality by Design (QbD) and proportional risk-based approaches.

Annex 1:

- Expands on the principles provided in the main guideline.
- Offers detailed guidance on practical applications across different trial settings.
- Aimed at making the principles operationally feasible.

Annex 2:

- Focuses on **emerging designs** and **real-world data (RWD)** considerations.
- Highlights decentralized trial elements, pragmatic approaches, and integration of digital health technologies.
- Not comprehensive but provides targeted GCP considerations for trials incorporating innovative methods.

ICH E6(R3)

From Application to Innovation

Key Differences Between Main Document & Annexes

Aspect	Main Document (ICH E6(R3))	Annex 1: APPLICATION	Annex 2: INNOVATION
Scope	Broad application across all trials.	Guidance on applying principles in standard trials.	Specific considerations for innovative designs.
Focus	Foundational GCP principles and regulatory compliance.	Operational implementation of GCP.	Advanced trial designs (e.g., decentralized).
Target Audience	All stakeholders (Sponsors, IRBs, Investigators).	Practitioners implementing standard trials.	Innovators adopting RWD and decentralized models.
Content Specificity	General principles and requirements.	Practical examples and applications.	Case studies and emerging challenges.

ICH E6(R3)

When?

Timeline & Development

Document	Development Stage	Status	Date
ICH E6	Initial guideline	Adopted	May 1996
ICH E6(R1)	Editorial updates	Post-Step 4 corrections	June 1996
ICH E6(R2)	Integrated Addendum	Adopted	November 2016
ICH E6(R3) (Main)	*Updated guideline*	*Public consultation*	*May 2023*
ICH E6(R3) Annex 2	*Specific GCP for new designs*	*Public consultation*	*November 2024*

ICH E6(R3)

Quality by Design in Action

Document Relationship

- **Main Document** serves as the comprehensive set of principles.
- **Annex 1** ensures the principles are practically applicable in conventional trials.
- **Annex 2** extends the guidelines to address the evolving trial landscape, focusing on digital innovation and RWD.

LINK TO THE DOCUMENTS:

- Main Document, incl. Annex 1
- Annex 2

Quality by Design in Action

Office Europe
Altenhöferallee 3
60438 Frankfurt (M), Germany
o. +49 (0)6192-470-113-50

Office USA
220 Juana Avenue
San Leandro, CA 94577
o. +1 888.812.RBQM

Contact Us
post@cyntegrity.com