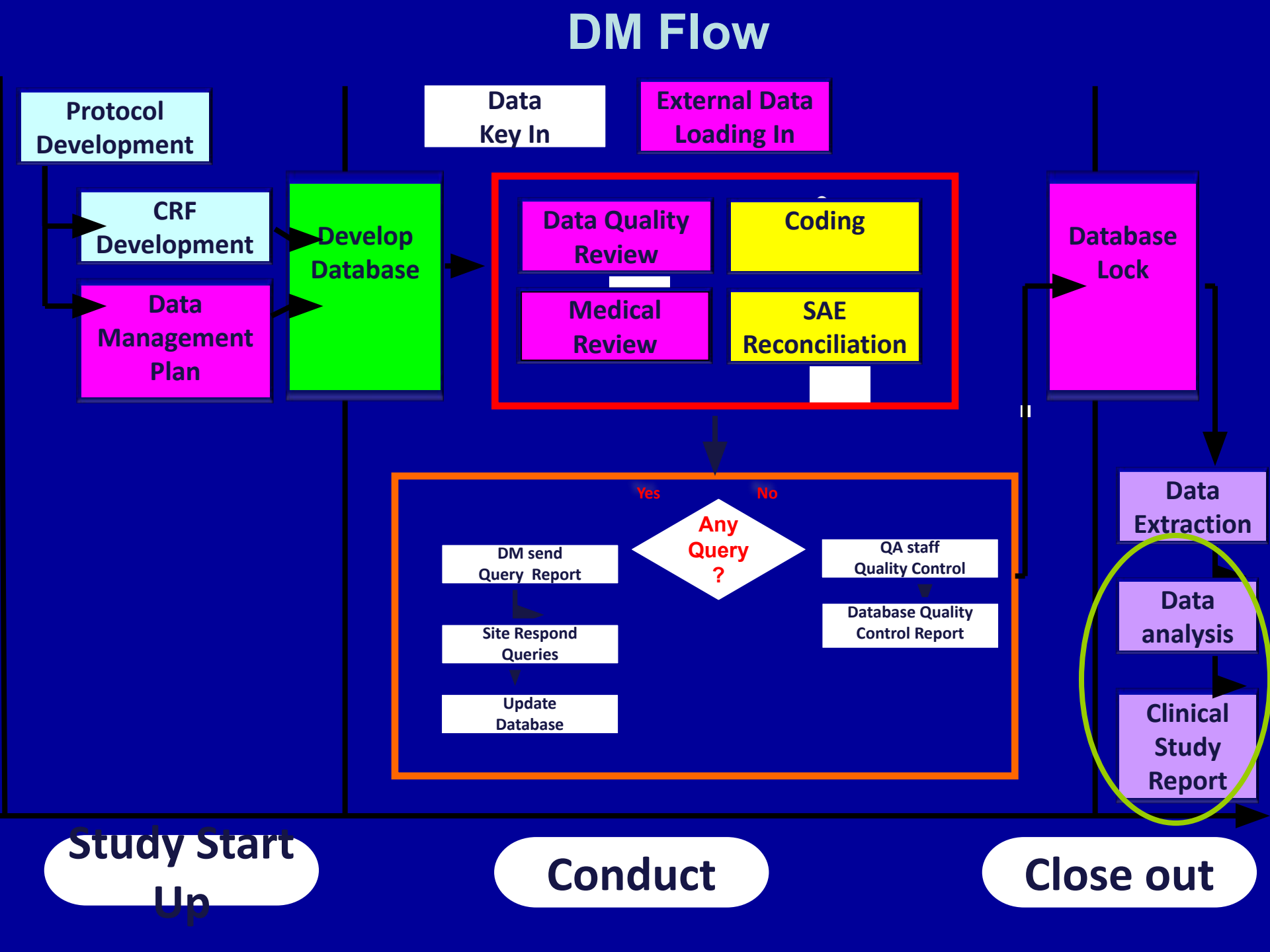
Before Presentation…

- This slide deck is based on Jain Chung's presentation for the 1st CDM training course in 2008.


DM Flow
Protocol Development
CRF Development
Data Management Plan
Develop Database
Data Key In
External Data Loading In
Data Quality Review
Coding
Medical Review
SAE Reconciliation
Database Lock
Yes
No
Any Query ?
DM send Query Report
Site Respond Queries
Update Database
QA staff Quality Control
Database Quality Control Report
Data Extraction
Data analysis
Clinical Study Report
Study Start Up
Conduct
Close out

ICH E9 Statistical Principles

INTERNATIONAL CONFERENCE ON HARMONISATION OF TECHNICAL REQUIREMENTS FOR REGISTRATION OF PHARMACEUTICALS FOR HUMAN USE

ICH HARMONISED TRIPARTITE GUIDELINE

STATISTICAL PRINCIPLES FOR CLINICAL TRIALS

E9

Current *Step 4* version

dated 5 February 1998

ICH E3 Clinical Study Reports

INTERNATIONAL CONFERENCE ON HARMONISATION OF TECHNICAL REQUIREMENTS FOR REGISTRATION OF PHARMACEUTICALS FOR HUMAN USE

ICH HARMONISED TRIPARTITE GUIDELINE

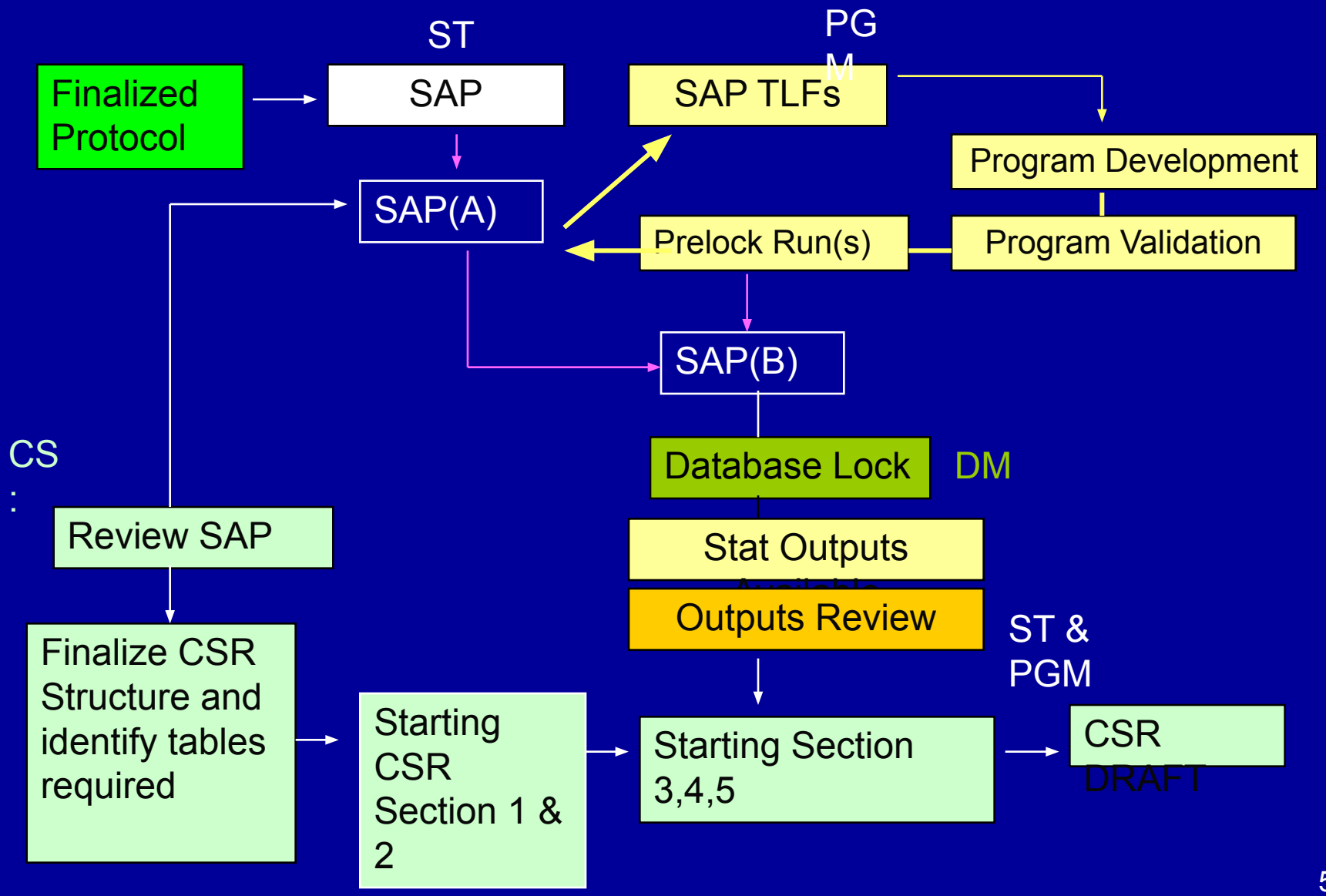
STRUCTURE AND CONTENT OF CLINICAL STUDY REPORTS

E3

Current *Step 4* version

dated 30 November 1995

Process for Development Clinical Study Report



Sample of CSR Report Body

In the format of the Journal-Style scientific paper

1. Background, Rationale and Objectives
2. Materials And Methods
3. Results
   - 3.1 Study Population
   - 3.2 Efficacy Results
   - 3.3 Pharmacodynamic, Pharmacokinetic and PK/PD Modeling
   - 3.4 Safety Analysis
4. Discussion
5. Conclusion
6. References

Appendices

Sample of CSR Report Body

In the format of ICH E3 "Structure and Content of Clinical Study Reports"

1. Title page
2. Synopsis
3. Table of contents
4. List of abbreviations
5. Ethics
6. Investigators and study administrative structure
7. Introduction
8. Study objectives
9. Investigational plan
10. Study patients
11. Efficacy evaluation
12. Safety evaluation
13. Discussion and overall conclusions
14. Tables, figures and graphs referred to but not included in the text
15. Reference list
16. Appendices

CSR Section 3 - Results

3.1 Study Population

- 3.1.1 Disposition of Patients
- 3.1.2 Patients Withdrawn Prematurely from treatment
- 3.1.3 Overall of Analysis Populations
- 3.1.4 Protocol Violations
- 3.15 Demographic Data and Baseline Characteristics
- 3.1.6 Previous Concomitant Medications and Diseases

CSR Section 3 - Results

3.2 Efficacy Results

3.2.1 Primary Efficacy Parameter

3.2.2 Secondary Efficacy Parameter (s)

3.1.3 Subgroup and Exploratory Analyses

3.3 Pharmacodynamic, Pharmacokinetic and PK/PD Modeling

CSR Section 3- Results

3.4 Safety Analysis

3.4.1 Extent of Exposure to Trial Medication
3.4.2 Overview of Safety
3.4.3 Adverse Events

3.4.3.1 Overview Adverse Events
3.4.3.2 Deaths
3.4.3.3 Serious Adverse Events
3.4.3.4 Adverse Events and Laboratory abnormalities Leading to Withdrawal from treatment
3.4.3.5 Dose Modifications for Safety Reasons

CSR Section 3 - Results

3.4.4 Laboratory Parameters

3.4.4.1 Mean (or Median) Change from Baseline

3.4.4.2 Shift from Baseline

3.4.5 Vital Signs

3.4.6 ECGs

Other CSR Sections: 4, 5, and 6

4. Discussion
5. Conclusion
6. References

   Appendices

Review CSR, final TLFs

- Validation
- Consistency
- Interpretations
- Discussions

CSR Section 1: Background, Rationale and Objectives

1.1 Background
1.2 Rationale
1.3 Objective

CSR Section 2 - Materials and Methods

2.1 Overall Study Design

2.1.1 Protocol Amendments

2.2 Study Population

2.2.1 Overview

2.2.2 Inclusion Criteria

2.2.3 Exclusion Criteria

2.2.4 Criteria for Withdrawal from Treatment or Study and Replacement Policy

2.2.5 Concomitant Medication, Treatments and Procedures

2.3 Compliance with Good Clinical Practice

2.3.1 Ethics

2.3.2 Audits

2.3.3 Data Quality Assurance

2.4 Trial Medication

2.4.1 Rationale for Dosage Selection

2.4.2 Formulation and Packaging

2.4.3 Assignment to Treatment Group/Sequence

2.4.4 Blinding

2.4.5 Drug Administration

2.4.6 Dose Modification

2.4.7 Dose Accountability and Compliance

ICH E3 Structure and Content of Clinical Study Reports

1. Title page
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15. Reference list
16. Appendices

** Details for Sections 9 – 12 on next slides*

ICH E3 Structure and Content of Clinical Study Reports (cont.)

9. Investigational plan

9.1 Overall study design and plan description

9.2 Discussion of study design, including the choice of control groups

9.3 Selection of study population

- 9.3.1 Inclusion Criteria
- 9.3.2 Exclusion Criteria
- 9.3.3 Removal of Patients from Therapy or Assessment

9.4 Treatments

- 9.4.1 Treatments Administered
- 9.4.2 Identity of Investigational Product(s)
- 9.4.3 Method of Assigning Patients to Treatment Groups
- 9.4.4 Selection of Doses in the Study

9.4 Treatments (cont.)

- 9.4.5 Selection and Timing of Dose for each Patient
- 9.4.6 Blinding
- 9.4.7 Prior and Concomitant Therapy
- 9.4.8 Treatment Compliance

9.5 Efficacy and safety variables

- 9.5.1 Efficacy and Safety Measurements Assessed and Flow Chart
- 9.5.2 Appropriateness of Measurements
- 9.5.3 Primary Efficacy Variable(s)
- 9.5.4 Drug Concentration Measurements

9.6 Data quality assurance

9.7 Statistical methods planned in the protocol & determination of sample size

9.8 Changes in the conduct of the study or planned analyses

ICH E3 Structure and Content of Clinical Study Reports (cont.)

10 Study patients

10.1 Disposition of patients

10.2 Protocol deviations

11. Efficacy evaluation

11.1 Data sets analyzed

11.2 Demographic and other baseline characteristics

11.3 Measurements of treatment compliance

11.4 Efficacy results and tabulations of individual patient data

12. Safety evaluation

12.1 Extent of exposure

12.2 Adverse events (AEs)

12.3 Deaths, other SAEs, and other significant adverse events

12.4 Clinical laboratory evaluation

12.5 Vital signs, physical findings and other observations related to safety

12.6 Safety conclusions

References

- ICH Guidelines www.ich.org
  - E9 Statistical Principles for Clinical Trials
  - E3 Structure and Content of Clinical Study Reports