

FDA

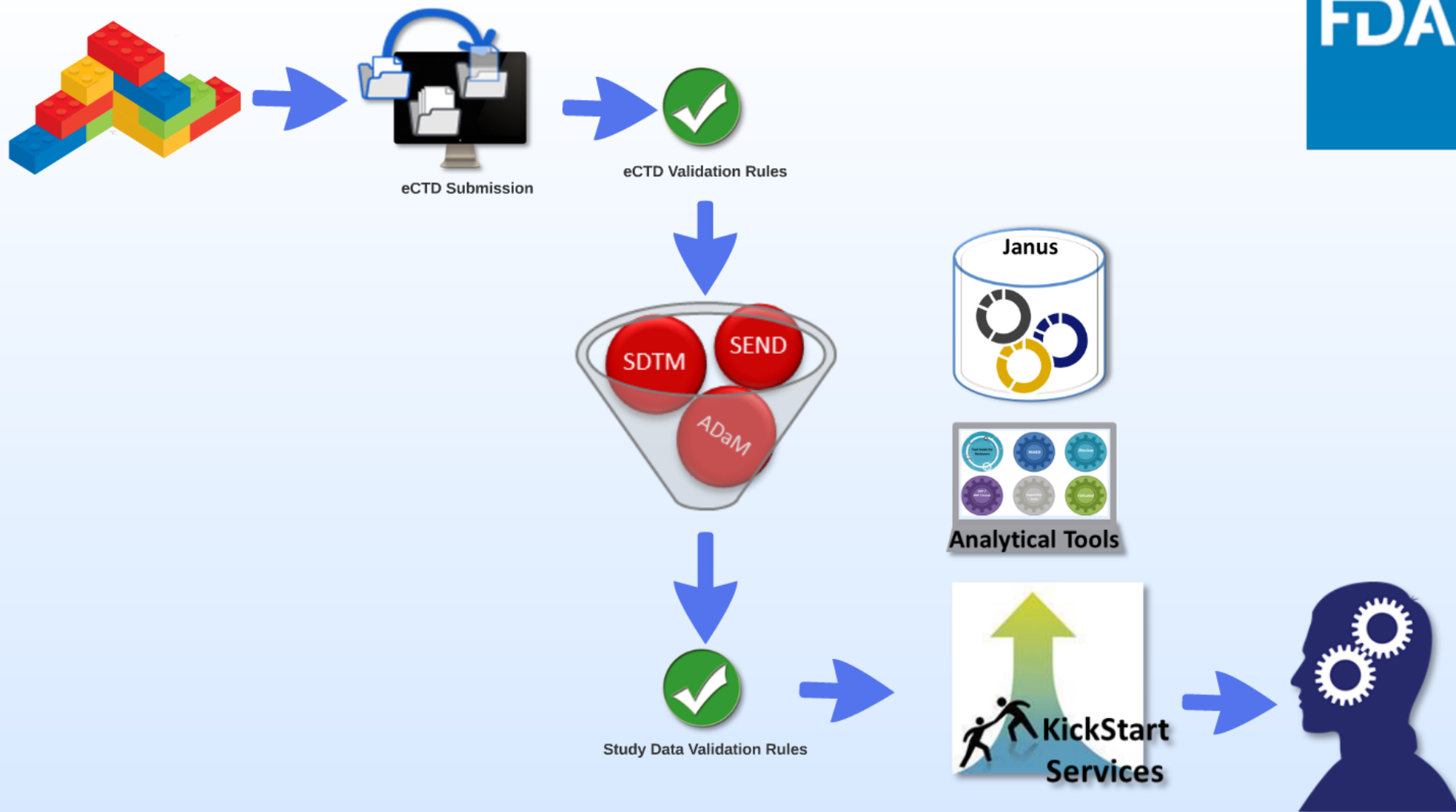
**U.S. FOOD & DRUG
ADMINISTRATION**

OFFICE OF COMPUTATIONAL SCIENCE

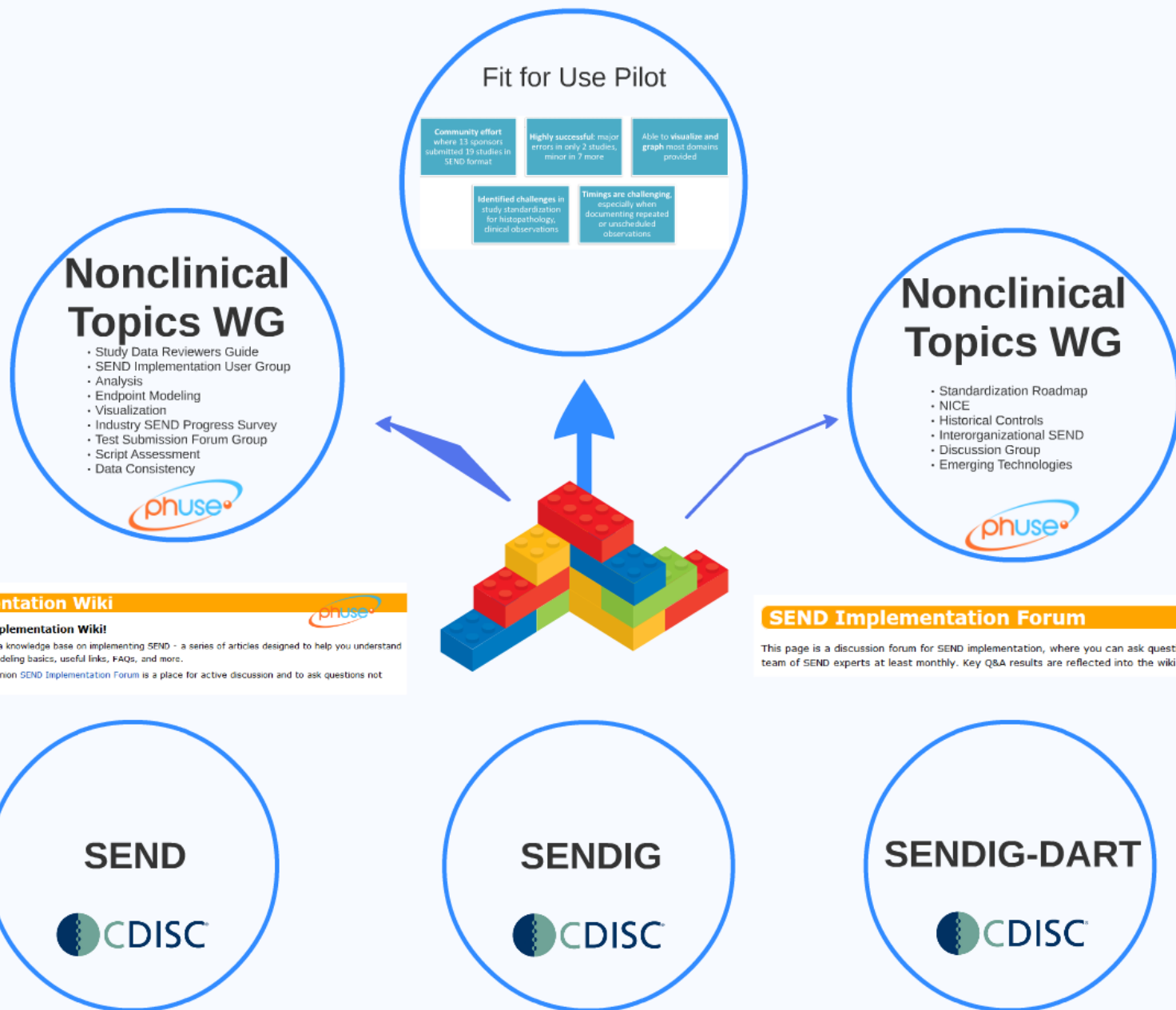
Nonclinical: Pulling it All Together

Lilliam Rosario, Ph.D.
Director, Office of Computational Science

Better Data | Better Tools | Better Decisions



Better Data | Better Tools | Better Decisions



**Final Published
Dec 17, 2014**

FDASIA Guidance

**Final Published
May 5, 2015**

24 months after
final guidance,
sponsors must use
standards identified
by FDA (NDAs,
ANDAs, BLAs)

eStudy Guidance

Binding Guidance – Requires that studies are compliant with the standards outlined in the FDA Data Standards Catalog.



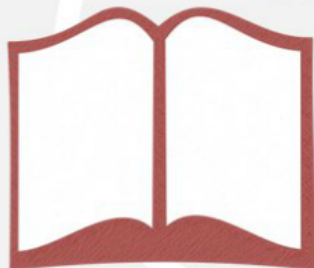
eCTD Guidance

Binding Guidance – Requires that content be submitted to the agency electronically in the format specified in the guidance.



Data Standards Catalog

Lists supported and/or required standards.



Technical Conformance Guide

Provides specifications, recommendations, and general considerations on how to submit standardized study data.



Data Standards Catalog

Study Data...SDTM, ADaM, SEND, Define.XML

Technical Conformance Guide

FDA Data Standards Catalog v4.5.1 (08-31-2016) - Supported and Required Standards

This table contains a listing of the data exchange, file formats and terminology standards supported at FDA. These standards have gone through all the steps necessary to make this part of the regulatory review process, including posting of regulatory guidance documents and associated implementation guidelines and technical specifications. The submission of standardized data using any standard not listed, or to an FDA Center not listed, should be discussed with the Agency in advance. This catalog is incorporated by reference in the guidance to industry, *Providing Regulatory Submissions in Electronic Format-Standardized Study Data* (<http://www.fda.gov/downloads/Drugs/Guidances/UCM292334.pdf>).

Use	Data Exchange Standard	Exchange Format	Standards Development Organization (SDO)	Supported Version	Implementation Guide Version	FDA Center(s)	Date Support Begins (MM/DD/YYYY)	Date Support Ends (MM/DD/YYYY)	Date Requirement Begins (MM/DD/YYYY)	Date Requirement Ends	Regulatory Reference and Information Sources
Regulatory Applications (IND, NDA, ANDA, BLA, master files)	Electronic Common Technical Document (eCTD)	Extensible Markup Language (XML)	International Council for Harmonization (ICH)	3.2.2	M2 eCTD: Electronic Common Technical Document Specifications	CDER, CBER	05/01/2008		05/05/2017 [5] 05/05/2018 [6]		Electronic Submissions: Electronic Common Technical Document (eCTD)
Product Labeling Submissions	Structured Product Labeling (SPL)	XML	Health Level 7 (HL7)	Release 5		CDER, CBER	Ongoing		04/01/2006 [3] 12/11/2003 [4]		Structured Product Labeling (SPL) Implementation Guide with Validation Procedures
Postmarketing Safety Reporting - Adverse Events for Medical Devices	Individual Case Safety Report										Electronic Medical Device Reporting (eMDR) - Device Reporting and Guidance
Case 1				N/A		CDRH	Ongoing				Veterinary Adverse Event Reporting for Manufacturers
Case 2				N/A		CVM	Ongoing				
Case 2				ICH E2B		CDER, CBER	Ongoing				FDA Adverse Events Reporting System (FAERS) Electronic Submissions
Case 2				N/A		CDER, CBER	Ongoing				FDA Adverse Events Reporting System (FAERS) Electronic Submissions

Data Standards Catalog

Lists supported and/or required standards.



www.fda.gov

STUDY DATA TECHNICAL CONFORMANCE GUIDE

Technical Specifications Document

This Document is incorporated by reference into the following Guidance Document(s):

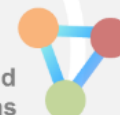
Guidance for Industry *Providing Regulatory Submissions in Electronic Format - Standardized Study Data*

For questions regarding this technical specifications document, contact CDER at cdcr-stda@fda.hhs.gov or CBER at cbcr-disc@fda.hhs.gov

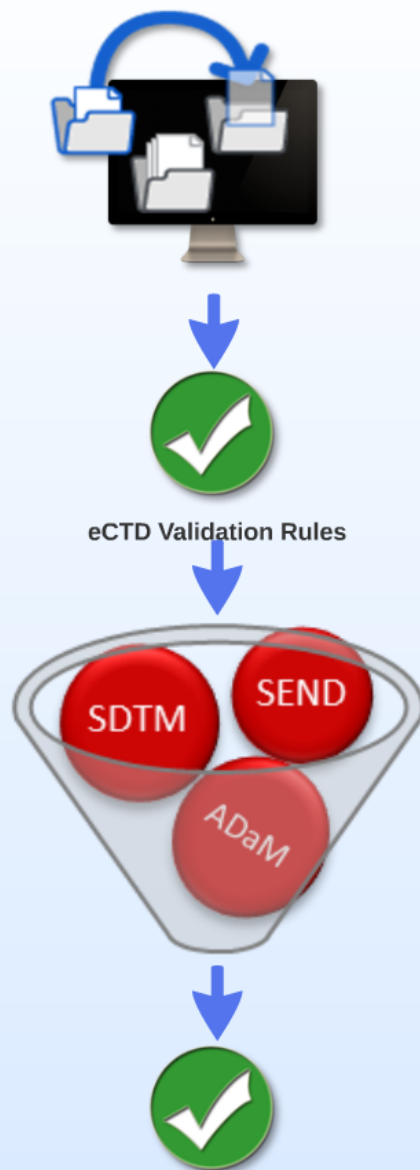
Health and Human Services
Administration
Center for Drug Evaluation and Research (CDER)
Center for Biologics Evaluation and Research (CBER)

Technical Conformance Guide

Provides specifications, recommendations, and general considerations on how to submit standardized study data.



2016



FDA

eCTD Study Data Validation Rules and Severity

High

Demographic dataset (DM) and the define.xml must be submitted in Module 4 and the DM dataset Subject level analysis dataset (ADSL) and define.xml must be submitted in Module 5

High

Trial Summary (TS) dataset must be presented for each study in Module 4 or 5

Medium

Correct STF file-tags must be used for all standardized datasets

- Data-tabulations-dataset-sdtm
- Analysis-dataset-adam
- Data-tabulations-dataset-send

Medium

For each study, no more than one dataset of the same type should be submitted as new

FDA Business Rule ID

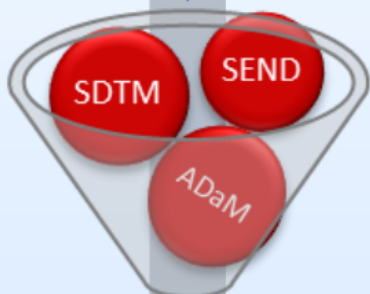
FDA Business Rule

Nonclinical Only

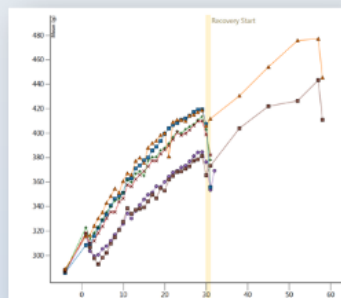
FDAB046	All Trial Set data should be submitted for nonclinical data.
FDAB047	Required timing variables for identification of the day on which group summaries (Group Means and Incidences) are calculated should be populated for nonclinical data.
FDAB048	Result modifiers for the MA and MI domains should be included in SUPPMA and SUPPMI unless all findings are normal for nonclinical data.
FDAB049	Set Code should match Trial Sets for nonclinical data.
FDAB050	Trial Set Parameter Code should be unique for each study set for nonclinical data.
FDAB051	Variable intended only for human studies should not be included for nonclinical data.
FDAB052	Variables should conform to SEND IG for nonclinical data.
FDAB053	When range is used for Age, then it should be provided in a 'number-number' format for nonclinical data.
FDAB054	Pool ID should match Pool Definition for nonclinical data.



eCTD Validation Rules



Study Data Validation Rules



Graphs

Tables



Visualizations



KickStart



**Janus
NonClinical**

Data

**Available
accurate
reviewable**

Review



Training

Best Practices

**Knowledge
Management**

**OCS Service
Desk**

Standard



SEND 3.0/3.1



**DART
Provisional**



**Genetox, Dermal
Ocular, CNS**

Tools & Services



KickStart



**Janus
NonClinical**



Other Tools

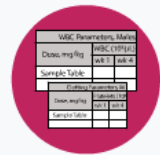
Template



IND Template



NDA Review



**Additional
Review**

Decision



**Safety &
Dose**



Labeling



**Exploratory
Research**

Standard



SEND 3.0/3.1



**DART
Provisional**



**Genetox, Dermal
Ocular, CNS**

Tools & Services



KickStart



**Janus
NonClinical**



Other Tools

Template



IND Template



NDA Review



**Additional
Review**

Decision



**Safety &
Dose**



Labeling



**Exploratory
Research**

Standard

SEND 3.0/3.1

**DART
Provisional**

**Genetox, Dermal
Ocular, CNS**

Tools & Services

KickStart

**Janus
NonClinical**

Other Tools

Template

IND Template

NDA Review

Additional
Review

Decision

Safety &
Dose

Labeling

Exploratory
Research

Standard

SEND 3.0/3.1

DART
Provisional

**Safety
Pharmacology**

Tools & Services

KickStart

Janus
NonClinical

Other Tools

Template

IND Template

NDA Review

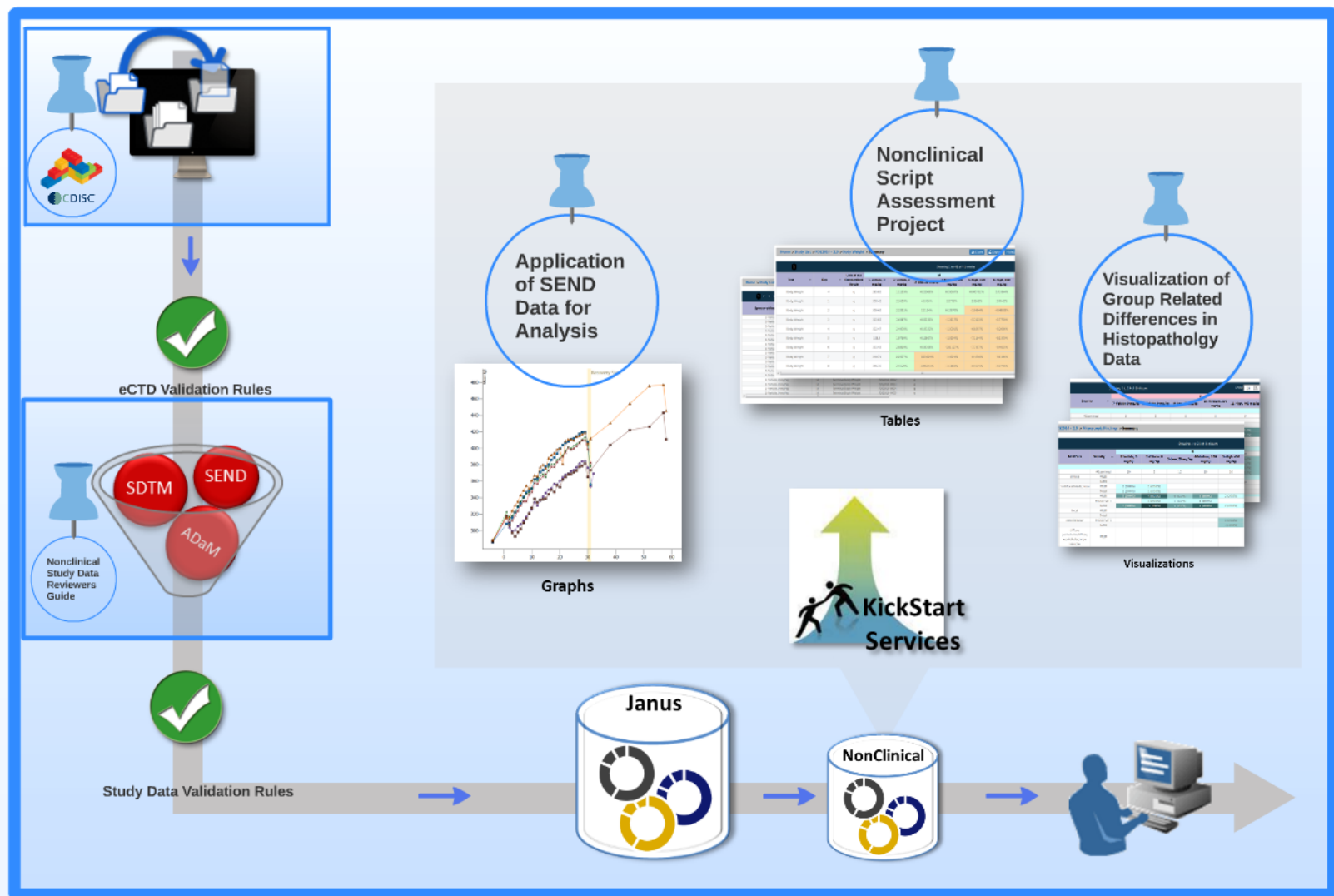
**Additional
Review**

Decision

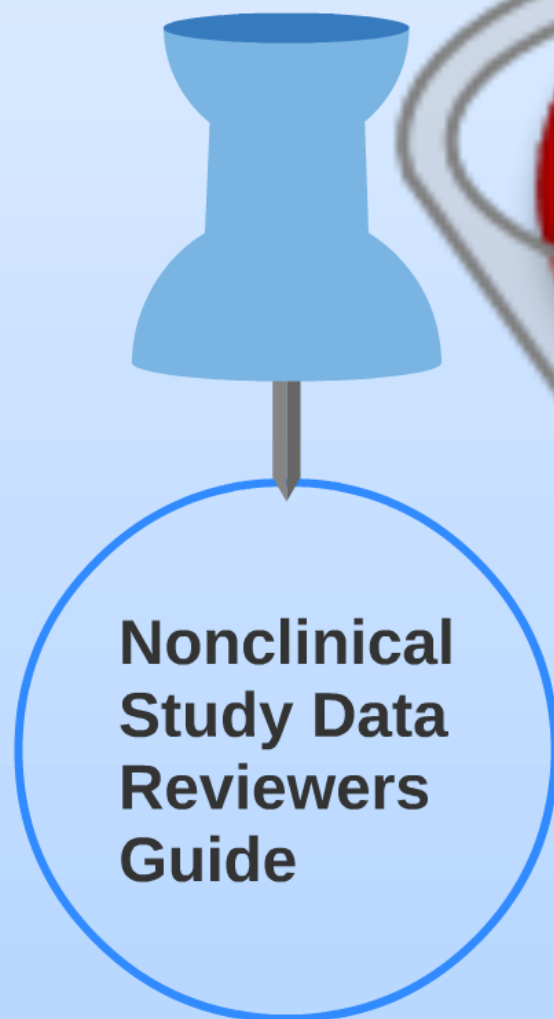
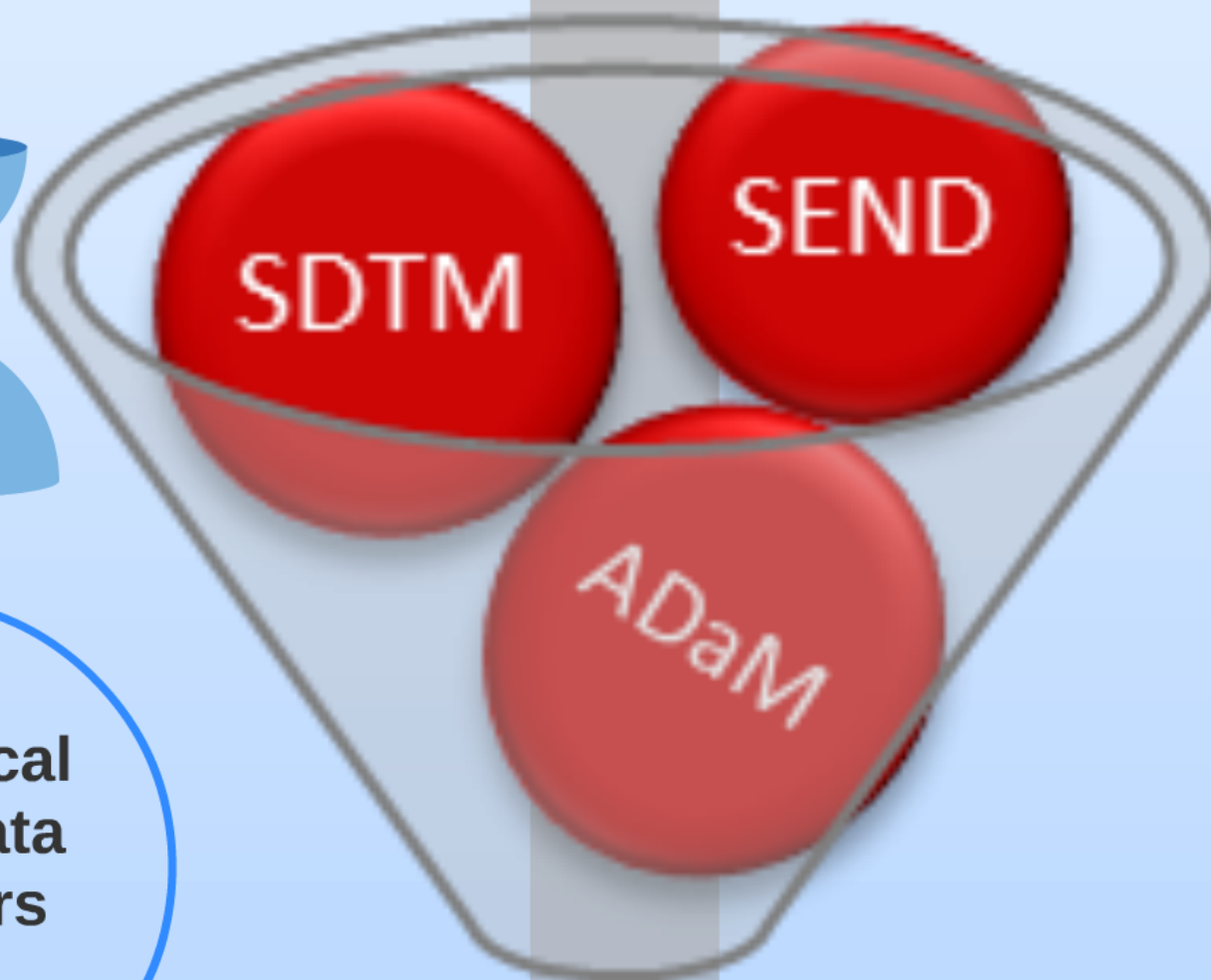
Safety &
Dose

Labeling

**Exploratory
Research**

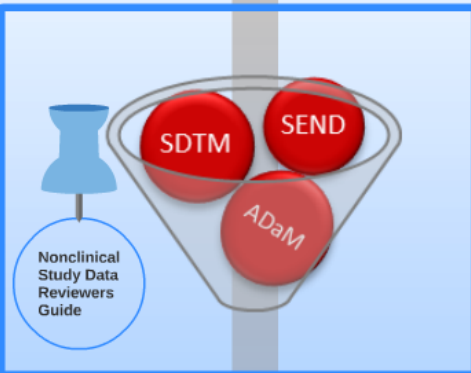




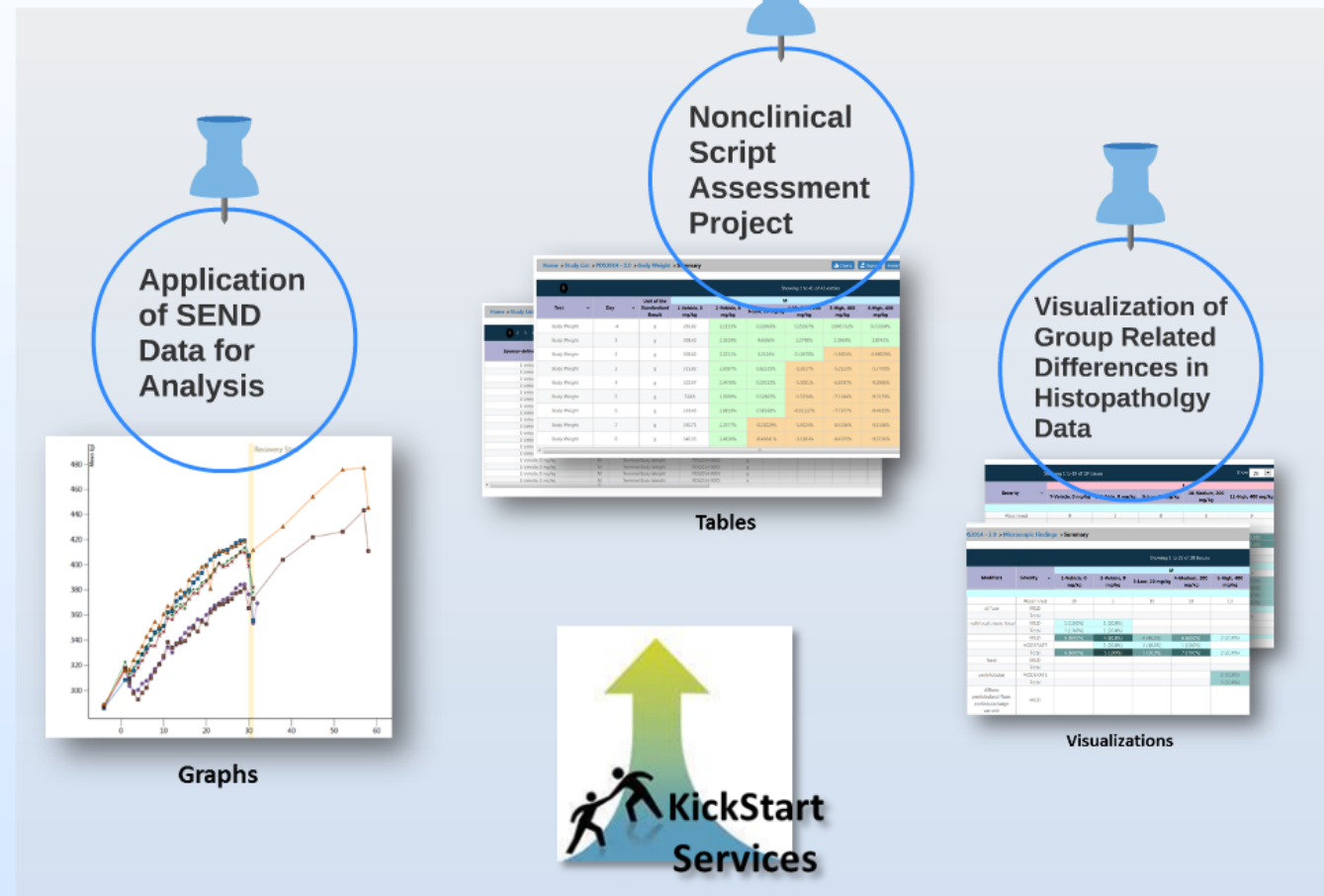




eCTD Validation Rules



Study Data Validation Rules





**SEND
Implementation
User Group**



**Industry SEND
Progress
Survey**



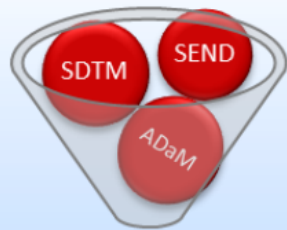
**Test
Submission
Forum Group**



**Data
Consistency:
SEND Datasets
and the Study
Report**



**Investigating
Endpoint
Modeling**



Smart Template

DEPARTMENT OF HEALTH AND HUMAN SERVICES
PUBLIC HEALTH SERVICE
FOOD AND DRUG ADMINISTRATION
CENTER FOR DRUG EVALUATION AND RESEARCH

PHARMACOLOGY TOXICOLOGY AND REVIEW AND INSPECTION

Application number: [\[Link\]](#)
 Review number: [\[Link\]](#)
 Reporting number: [\[Link\]](#)
 PERM review number: [\[Link\]](#)
 Other number: [\[Link\]](#)
 Product: [\[Link\]](#)
 Indication: [\[Link\]](#)
 Therapeutic use: [\[Link\]](#)
 Sponsor: [\[Link\]](#)
 Review Division: [\[Link\]](#)
 Reviewer: [\[Link\]](#)
 Supervisor/Team Leader: [\[Link\]](#)
 Division Director: [\[Link\]](#)
 Project Manager: [\[Link\]](#)

Template version: August 28, 2015

