CDISC Italian User Network 2019

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Regulatory News, Updates and Final Q&A

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CDISC EU Committee / Italian UN Representative

Overview of Agencies Accepting CDISC Data Submission

Mandatory for all studies **started** after December 16, 2016

- Mandatory for submission to CDER and CBER
- Recommended for submission to CDRH

Mandatory for all submissions after 31-Mar-2020

Similar process i.e. Study Data Technical Conformance Guide, Concept of Catalog of Standards

Minor Differences i.e. naming convention for reviewer guide csdrg.pdf vs study-data-reviewers-guide.pdf (for SDTM)

ARM still nice to have for FDA, PMDA would like to see it

Other Small Differences i.e. FDA to have all warnings/errors commented vs PMDA requiring only Errors

FDA Last 18 Months Update

- FDA issues every March and every October a new version of the **Study Data Technical Conformance Guide (SDTCG)** → latest update Version 4.2 October 2018 (minor update February 2019 v4.2.1)
  - → Webinar are usually run by the FDA to present changes from previous version
- **Biomedical Monitoring Technical Conformance Guide** – February 2018
  - Part I: Clinical Study Level Information
  - Part II: Subject-Level Data Line Listings by Clinical Site
  - Part III: Summary-Level Clinical Site Dataset (CLINSITE)

  These tasks can be standardized.
- **Study Data Standardization Plan** is now a key document when meeting with FDA

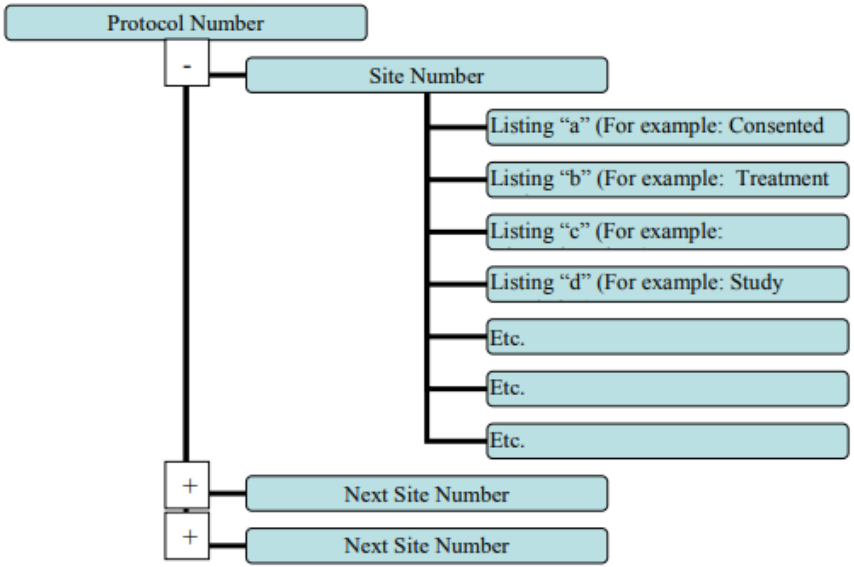
FDA Last 18 Months Update - SDTCG

- Clarified requiremt for Software Programs to be submitted
  - ADaM and Tables/Figures supporting primary and secondary efficacy analyses
  - *The main purpose of requesting the submission of these programs is to **understand the process** by which the variables for the respective analyses were created and **to confirm the analysis algorithms and results***
- Special Characters Usage
  - Printable Values below ASCII 128
  - UTF-8 not recommended
  - LBSTRESC and LBTEST do not contain ASCII values between 160-191
- Trial Summary (TS) Parameters for Submission Recommended Parameters
- Reccomendations on how to handle screen failures / multiple enrolment in SDTM
- Include DV (Protocol Deviations) in SDTM → to facilitate BIMO clinical investigator site selection process

FDA Last 18 Months Update - BIMO

- Part II: Subject-Level Data Line Listings by Clinical Site

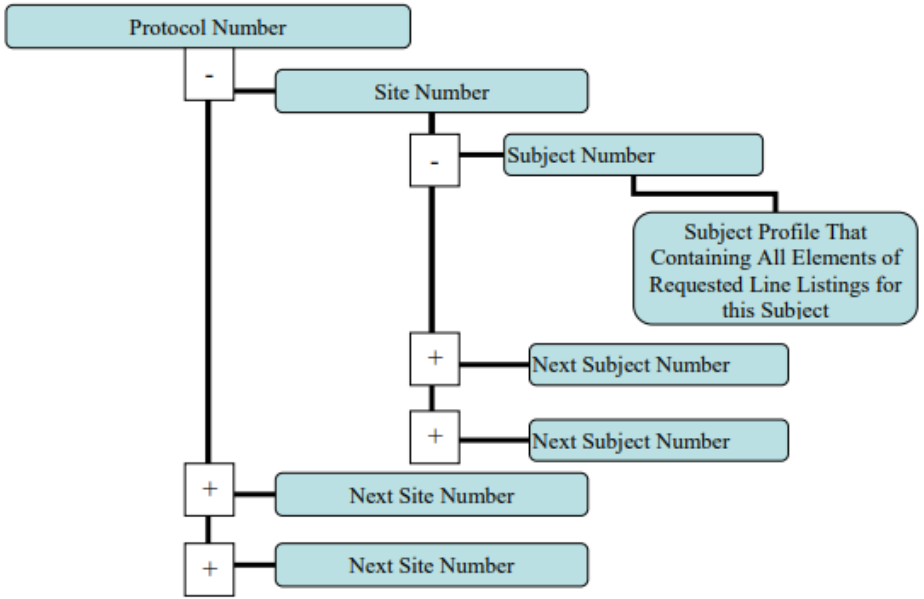
By Site, By Listing Option A:



FDA Last 18 Months Update - BIMO

- Part II: Subject-Level Data Line Listings by Clinical Site

By Site, By Subject Profile Option B:



FDA Last 18 Months Update - BIMO

- Part III: Summary-Level Clinical Site Dataset (CLINSITE)

  List of the critical variables associated with efficacy to be submitted for each study and investigator site.

STUDYID	STUDYTL	SITEID	ARM	SAFPOP	SCREEN	DISCSTUD
ABC-123	Double blind...	001	Active	26	61	3
ABC-123	Double blind...	001	Placebo	25	61	4
ABC-123	Double blind...	002	Active	23	54	2
ABC-123	Double blind...	002	Placebo	25	54	4
ABC-123	Double blind...	003	Active	27	62	3
ABC-123	Double blind...	003	Placebo	26	62	5

TRTEFFR Treatment Efficacy Result
SITEEFFE Site-Specific Treatment Effect and their Standard Deviations (TRTEFFS and SITEEFFS)

DISCTRT	ENDPOINT	ENDTYPE	TRTEFFR	TRTEFFS	SITEEFFE	SITEEFFS
2	Percent Responders	Binary	0.48	0.0980	0.34	0.1405
1	Percent Responders	Binary	0.14	0.0694	0.34	0.1405
1	Percent Responders	Binary	0.48	0.1042	0.33	0.1427
3	Percent Responders	Binary	0.14	0.0694	0.33	0.1427
0	Percent Responders	Binary	0.54	0.0959	0.35	0.1448
3	Percent Responders	Binary	0.19	0.0769	0.35	0.1448

FDA Last 18 Months Update - SDSP

- Through the PhUSE and FDA collaboration, the SDSP template was streamlined to support both CDER and CBER. In March 2018, PhUSE released the **SDSP (v1) template*** that incorporated a CBER Appendix.
- SDSP
  - It's a plan that outlines the **clinical and non-clinical studies** in the compound and the use data standards (CDISC, MSSO, CDRH);
  - First published in the FDA Study Data Technical Conformance Guide (June 2015);
  - Promotes **early interactions** between sponsors and FDA review divisions;
  - **Living document** that should be updated as the development program expands and additional studies are planned;
  - Implementation assist in identifying **potential data standardization issues** early in the development program.

* PhUSE Study Data Standardization Plan

Template (v1) https://www.phuse.eu/documents//sop/wp/phuse-tp001-study-data-standardization-plan-v1-8409.docx

Completion Guideline (v1) https://www.phuse.eu/documents//sop/wp/phuse-wp001-sdsp-completion-guideline-v1-8407.pdf

Recent Webinars

CDISC **Library Launch** - 21 Feb 2019

CDISC Library (formerly known as CDISC SHARE), CDISC's metadata repository (MDR), provides users with machine-readable versions of CDISC standards. CDISC Library includes a number of significant improvements over 1.0.

CDISC Members Only Webinar: **Visual Define-XML Editor** - 5 Feb 2019

CDISC Public Webinar: Package 36 **Controlled Terminology** Publication Update/Package 37 Public Review - 9 Jan 2019

CDISC Public Webinar: **Controlled Terminology Mapping & Alignment** Across Codelists - 8 Jan 2019

Public Courses

At Europe Interchange in **Amsterdam**, Netherlands 6 - 10 May 2019

ADaM Primer, ADaM Theory and Application, CDASH Implementation , CDISC for Newcomers, Controlled Terminology, Define-XML, ODM Implementation, SDTM Theory and Application , SDTM Theory and Application for Medical Devices, SEND Implementation

In **Frankfurt** (Sulzbach), Germany, 3 - 7 Jun 2019

ADaM Primer, ADaM Theory and Application, CDASH Implementation , Define-XML, SDTM Theory and Application

In **Brussels**, Belgium, 9 - 13 Sep 2019

ADaM Primer, ADaM Theory and Application, CDASH Implementation , Define-XML, SDTM Theory and Application

In **Paris**, France, 7 - 11 Oct 2019

ADaM Primer, ADaM Theory and Application, CDASH Implementation , Define-XML, SDTM Theory and Application

European CDISC Interchange

https://www.cdisc.org/2019-cdisc-europe-interchange

Early bird fee expires today.

Program **key sessions**:

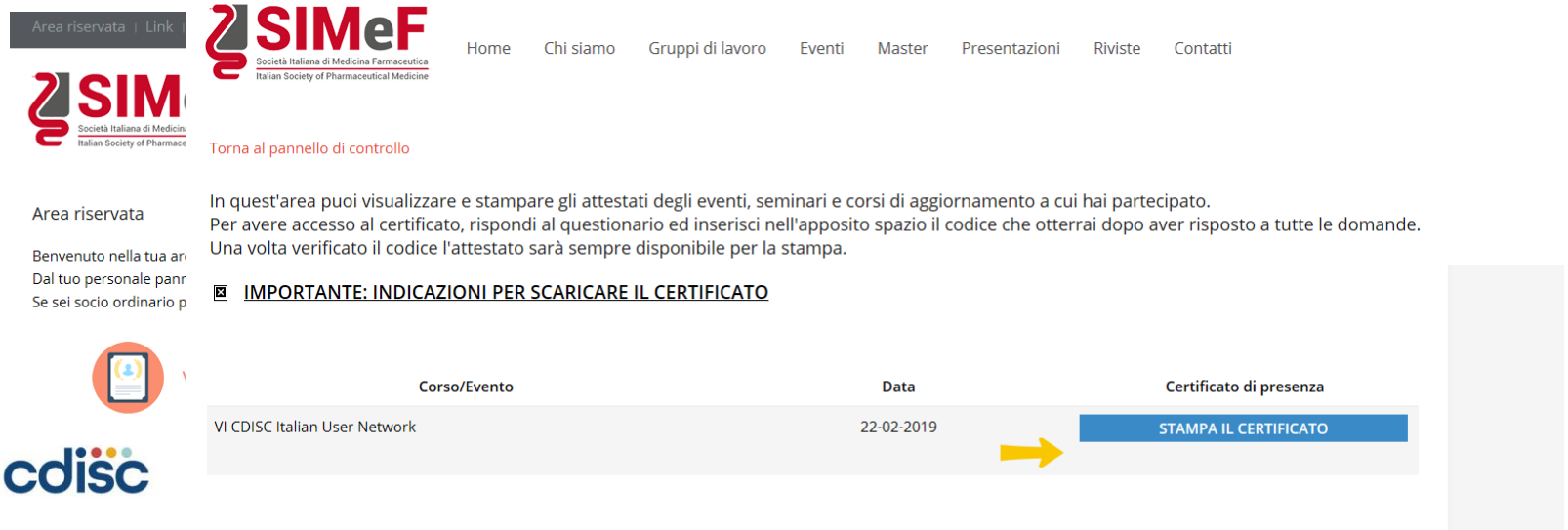
- Data governance
- Use cases
- Submissions
- End-to-end
- Foundational
- Rules
- RWE/Observational
- Regulatory

5 discounted tickets for Italian CDISC UN: 3/4 of them assigned to today speakers, 1/2 assigned by Italian CDISC UN representatives.

Attestato di partecipazione

Sul sito SIMeF www.simef.it

- effettuare il Login (cliccando su Area Riservata in alto a sinistra),
- Visualizza i tuoi certificati,
- Rispondere al questionario, al termine verrà fornito il codice del questionario da copiare e incollare nel box e quindi sarà possibile stampare il certificato.



Final Q&A

Topics covered today

- CDISC 360 Project Introduction
- CDASH 2.0
- SDTM Ig 3.3
- Clinical Trial – AI Value Cases
- ADaM 1.2 e altro da ADaM Team
- Define-XML v2.1 Guideline and PhUSE WG
- Standards Management & Study Metadata Management
- Use Case: Real Case in Vaccine sull'implementazione dei dati da e-diary in CDISC format
- TAUGs Update
- Novità Mondo Regolatorio

Any question?

Thank You!

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