



2023

CHINA

INTERCHANGE

BEIJING | 25-26 AUGUST



## The Big Changes for Annotated CRF Submission to Health Authorities

Presented by 彭兰岚 Lanlan Peng, Associate Manager, Statistical Programming,  
Biostatistics and Programming Department , Sanofi



## Meet the Speaker

彭兰岚 Lanlan Peng

**Title:** Associate Manager, Statistical Programming

**Organization:** Sanofi

More than 5 years' experience in Biostatistics and Programming department of Sanofi and be Associate Manager from early 2023, specializes in the clinical data submission workflow optimization, and clinical submission tool development.

Graduated from Claude Bernard University Lyon 1 with Master degree in Economics and Statistics.



# Disclaimer and Disclosures

- *The views and opinions expressed in this presentation are those of the author(s) and do not necessarily reflect the official policy or position of CDISC.*



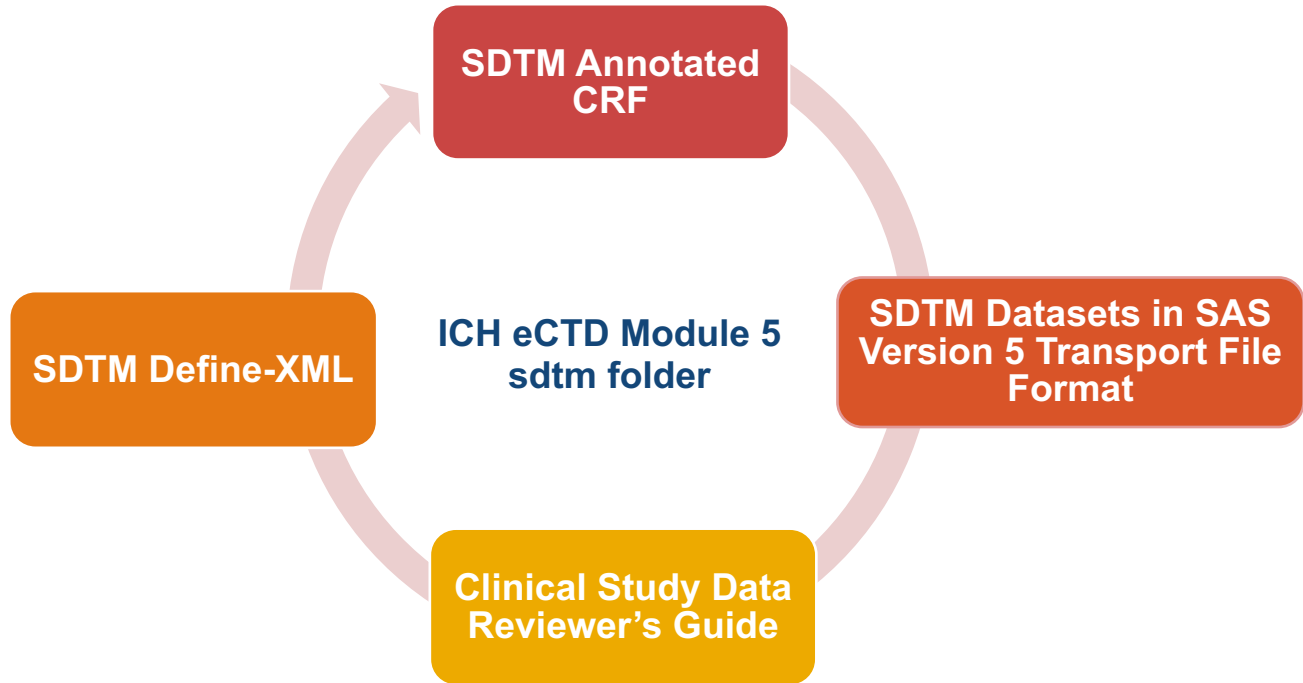
## Agenda

1. WHY SDTM-MSG is Important for Annotated CRF Generation?
2. HOW to Prepare Submitted Annotated CRF?
3. DEMO

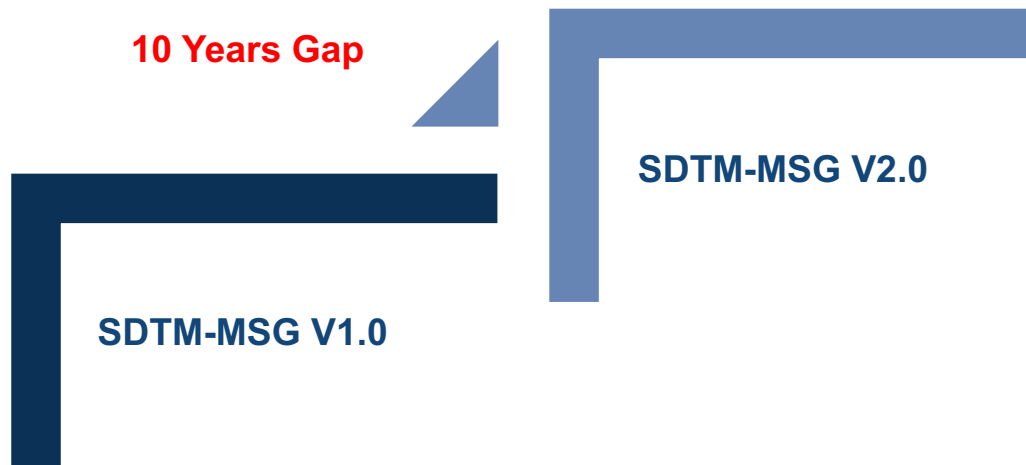


## WHY SDTM-MSG is Important for Annotated CRF Generation?

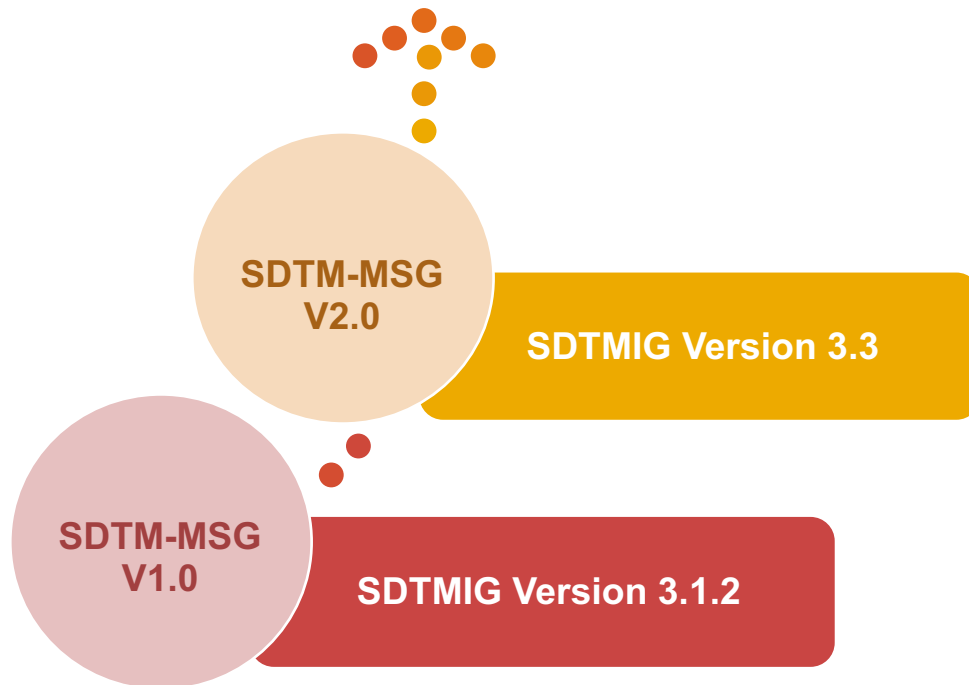
# Background



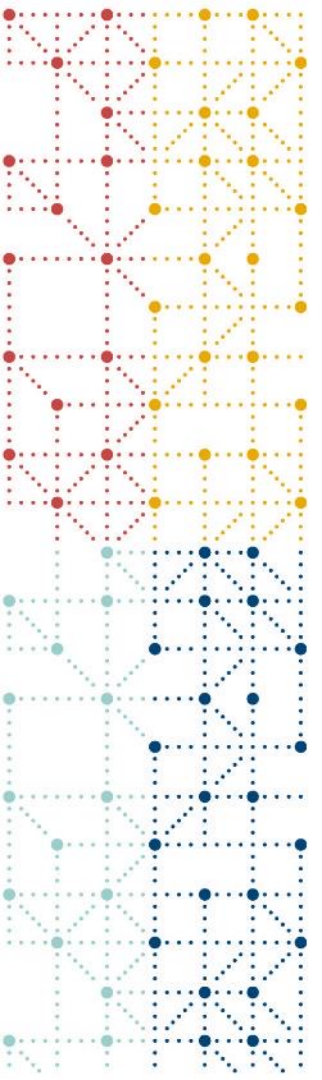
# Background



# Scope







## HOW to Prepare Submitted Annotated CRF?



# Annotating Unique CRF Pages

- The SDTM-MSG V2.0 recommended that sponsors include and annotate **unique form** only.
- **Unique Form:**

If a CRF page has a data collection point that is added, removed, or otherwise differs from other instances of the CRF page, then the page is considered to be unique.

# Annotating Unique CRF Pages

## Case 1

Data collection point that is added, removed.

**VS (Vital Signs)**

**VITAL SIGNS** [NOT SUBMITTED]

Vital Signs Collected?  Yes  No VSSTAT = NOT DONE when VSTESTCD = VSALL

Visit  VISIT

Date    VSDTC

VSTESTCD = WEIGHT Weight  VSORRES pounds  VSORRESU

VSTESTCD = TEMP Temperature  VSORRES F  VSORRESU

VSPPOS

Pulse and Blood Pressure (Supine)

VSTESTCD = PULSE Pulse  VSORRES bpm  VSORRESU

VSTESTCD = SYSBP Systolic  VSORRES mmHg  VSORRESU

VSTESTCD = DIABP Diastolic  VSORRES mmHg  VSORRESU

**VS (Vital Signs)**

**VITAL SIGNS** [NOT SUBMITTED]

Vital Signs Collected?  Yes  No VSSTAT = NOT DONE when VSTESTCD = VSALL

Visit  VISIT

Date    VSDTC

VSTESTCD = WEIGHT Weight  VSORRES pounds  VSORRESU

VSTESTCD = HEIGHT Height  VSORRES inches  VSORRESU

VSTESTCD = TEMP Temperature  VSORRES F  VSORRESU

VSPPOS

Pulse and Blood Pressure (Supine)

VSTESTCD = PULSE Pulse  VSORRES bpm  VSORRESU

VSTESTCD = SYSBP Systolic  VSORRES mmHg  VSORRESU

VSTESTCD = DIABP Diastolic  VSORRES mmHg  VSORRESU

# Annotating Unique CRF Pages

## Case 2

Allowable values are changed.

Subcategory of Reaction	Administration Site	<input type="checkbox"/>
	Associated Signs and Symptoms	<input type="checkbox"/>
	Systemic	<input checked="" type="checkbox"/>
	Trigger Factors	<input type="checkbox"/>

Subcategory of Reaction	Administration Site	<input checked="" type="checkbox"/>
	Associated Signs and Symptoms	<input type="checkbox"/>
	Systemic	<input type="checkbox"/>
	Trigger Factors	<input type="checkbox"/>



# Bookmarking

The aCRF is required to have **dual bookmarking**:

## 1. bookmarks by **chronology**

- It should be ordered chronologically based on study schedule of activities (SOA), with each timepoint bookmark, form bookmark should appear orderly that they in the aCRF.

## 2. bookmarks by CRF **topics or forms**

- It should be ordered alphabetically, within each topic bookmark, timepoint bookmark should appear chronologically according to SOA.

# Bookmarking

## SDTM-MSG V1.0

- CDISC01
  - Printable Table of Contents
  - Visits
    - Screening
  - Baseline
    - Randomization
    - Vital Signs
    - Mini-Mental State Examination
    - Cornell Scale For Depression in Dementia
  - Week 2
  - Week 24
  - End of Study / Termination
  - Running Records
  - Domains
    - 12-Lead ECG
      - Screening
      - Week 24

## SDTM-MSG V2.0

- Printable Table Of Contents
- Visit
  - Screening 1
    - Informed Consent
    - Demographics
    - Eligibility
    - Medical History
    - Vital Signs (Weight, Height And Temperature)
      - Ophthalmic Examination
  - Screening 2
  - Baseline
  - Week 2
  - Week 4
  - Week 6
  - Week 8
  - Week 12
  - Week 16
  - Week 20
  - Week 24
  - Week 26
  - Early Discontinuation Retrieval
  - Running Records
  - Forms
  - Adverse Events

# Table of Contents

In the aCRF, the table of contents (TOC) is used to facilitate review which may be included at the beginning of the aCRF.

## Bookmarking by Visits

## Bookmarking by Forms

Visits	Forms
Screening 1	Adverse Events
Informed Consent	Running Records
Demographics	Auditory Verbal Learning Test (AVL02)
Eligibility	Screening 2
Medical History	Week 4
Vital Signs (Weight, Height and Temperature)	Week 8
Version 2	Week 12
Version 1	Week 16
Ophthalmic Examination	Week 24
Screening 2	Early Discontinuation Retrieval
Vital Signs (Temperature)	Concomitant Medications
Version 2	Running Records
Version 1	Death
Auditory Verbal Learning Test	Running Records
Ophthalmic Examination	Demographics
Baseline	Screening 1
Vital Signs (Weight and Temperature)	Electroencephalogram
Version 2	Running Records
Version 1	Eligibility
Patient Health Questionnaire-9 (PHQ-9)	Screening 1
Satisfaction With Life Survey (SWLS)	End of Treatment
Hamilton Depression Rating Scale - 17 (HAMD 17)	Week 26
Ophthalmic Examination	Exposure
Week 2	Running Record
Vital Signs Vital Signs (Weight and Temperature)	Hamilton Depression Rating Scale - 17 (HAMD 17)
Version 2	Baseline
Version 1	Early Discontinuation Retrieval
Auditory Verbal Learning Test (AVL02)	Informed Consent
Ophthalmic Examination	Screening 1

# Appearance of Annotation

## Basic Information

Annotation Font	Arial
Annotation Font Size	12 Points
Annotation Background Color	Coblis Color Blindness Simulator

BLUE	<b>DM (Demographics)</b>	191, 255, 255
YELLOW	<b>DS (Disposition)</b>	255, 255, 150
GREEN	<b>SC (Subject Characteristics)</b>	150, 255, 150
ORANGE	<b>VS (Vital Signs)</b>	255, 190, 155



# Appearance of Annotation

## Domain Annotations

### SDTM-MSG V1.0

<b>DM=Demographics</b>	
CDISC Study CDISC01	
<b>Enrollment Form</b>	
Enroll the subject by entering the 3-digit Site # and the 5-digit Subject ID#	

### SDTM-MSG V2.0

<b>DS (Disposition)</b>	DSCAT = PROTOCOL MILESTONE
<b>INFORMED CONSENT</b>	DSTERM / DSDECOD = INFORMED CONSENT OBTAINED
Informed Consent Date	<input type="text"/> <input type="text"/> <input type="text"/>
<b>DM (Demographics)</b>	DSSTDTC
<b>DEMOGRAPHICS</b>	RFICDTC

1. Black text with bold formatting
2. Change DM = Demographics to DM (Demographics)

# Appearance of Annotation

## Variable Annotations

### SDTM-MSG V1.0

Date of Birth: \_\_\_\_\_ / \_\_\_\_\_ / **BRTHDTC**

**SEX**

Gender:  Male  Female

**ETHNIC**

Ethnicity:  Hispanic or Latino  Not Hispanic or Latino

### SDTM-MSG V2.0

**EXPOSURE**

Did the subject take Zanomaline?  Yes  No **ECOCCUR**

If No, Reason Not Administered  **ECREASOC in SUPPEC**

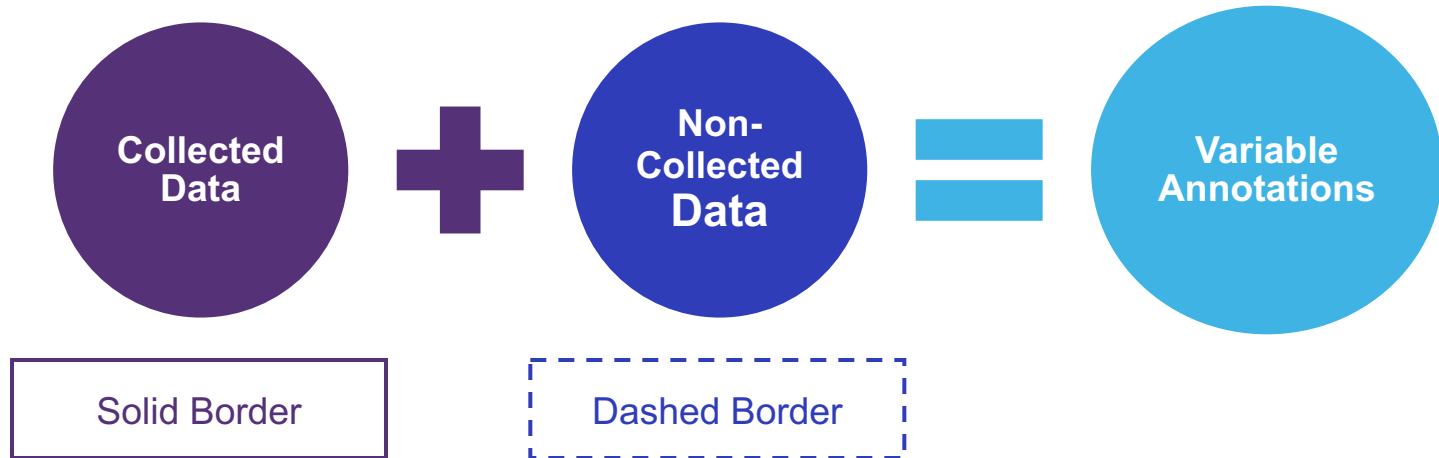
Date    **ECSTDTC**  
**ECENDTC**

Lot Number  **ECLLOT**

1. Black text without bold formatting
2. Use slash “/” to separate the variables

# Appearance of Annotation

## Variable Annotations



# Appearance of Annotation

## Variable Annotations → Collected Data

### Case 1

**EXPOSURE**

Did the subject take Zanomaline?  Yes  No **ECOCCUR**

If No, Reason Not Administered  **ECREASOC in SUPPEC**

Date    **ECSTDTC**  
**ECENDTC**

Lot Number  **ECLOT**

What was the subject's treatment status?

COMPLETED **DSTERM / DSDECOD**

ADVERSE EVENT

DEATH

LACK OF EFFICACY

# Appearance of Annotation

## Variable Annotations → Collected Data

### Case 2

Single Annotation: --ORRES when -TESTCD = <value>

Location	<input type="text"/>	OELOC
Abnormality	<input type="text"/>	OEORRES when OETESTCD = ABDETAIL

Separate Annotation

VSTESTCD = WEIGHT	Weight	VSORRES
VSTESTCD = HEIGHT	Height	VSORRES
VSTESTCD = TEMP	Temperature	VSORRES



# Appearance of Annotation

## Variable Annotations → Non-Collected Data

### Case 1

Default Value

Dose      ECDOSE 5 mL ECDOSU

Explicit Value Pertaining to Variable Annotation

QS (Questionnaires)      QSCAT = PHQ-9

**PATIENT HEALTH QUESTIONNAIRE-9  
(PHQ-9)**

# Appearance of Annotation

## Variable Annotations → Non-Collected Data

### Case 2

Not Submitted

Met Criteria  Yes  No [NOT SUBMITTED]

Not Submitted: Finding Forms

**VITAL SIGNS** [NOT SUBMITTED]

Vital Signs Collected?  Yes  No [VSSTAT = NOT DONE when VSTESTCD = VSALL]

# Appearance of Annotation

## Other Annotations → Non-Collected Data

### Case 1

Note: Explain a Situation and Not Direct Variable Annotations

Race (Check all that apply)

- White
- Black or African American

When multiple values are selected then RACE = MULTIPLE and individual responses are RACE1, RACE2, RACE3, etc. in SUPPDM



# Appearance of Annotation

## Other Annotations → Non-Collected Data

### Case 2

RELREC: There are Relationships Between Data Collection on CRF Form

**DS=Disposition**

CDISC  
Study CDISC01      Assessment Date: \_\_\_/\_\_\_/\_\_\_ **[NOT SUBMITTED]**

**TERMINATION**

Did patient complete the study?  Yes  No  
**DSDECOD / DSTERM = COMPLETED when Yes**

If patient did not complete the study, indicate the date of termination and check one primary reason to indicate why:

Date of Termination: \_\_\_/\_\_\_/\_\_\_ **DSSDTC**

**DSDECOD** Patient did not meet Inclusion/Exclusion Criteria at Screening or baseline (specify): **DSTERM**

Discontinued due to lack of Therapeutic Response **DSDECOD**

Discontinued due to Adverse Event  
Adverse Event No. \_\_\_\_\_ (Enter the number from the ADVERSE EVENTS Form)  
**Linked to related AE record via RELREC**

Protocol Violation (specify): \_\_\_\_\_ **DSTERM**

**SDTM-MSG V1.0**

If the subject's status is Adverse Event, Death or Other  
SPECIFY

---

If the subject's status is Adverse Event or Death  
PRIMARY AE #

**RELREC when  
DSLKNID = AELNKID**

**SDTM-MSG V2.0**

# Appearance of Annotation

## Little Tips

- It represents the data that would have been submitted had they been collected. For the case no data were collected, it is not necessary to re-annotate, it should indicate in the Define-XML document using the “HasNoData” attribute for datasets and variables.

Race (Check all that apply)

White

Black or African American

When multiple values are selected then RACE = MULTIPLE and individual responses are RACE1, RACE2, RACE3, etc. in SUPPDM

<p>► <b>QNAM</b> = "RACE4" [No Data]</p>	Race 4	text	Result Qualifier	41	<p><b>Race</b></p> <ul style="list-style-type: none"> <li>"AMERICAN INDIAN OR ALASKA NATIVE" = "American Indian Or Alaska Native"</li> <li>"ASIAN" = "Asian"</li> <li>"BLACK OR AFRICAN AMERICAN" = "Black Or African American"</li> <li>"NATIVE HAWAIIAN OR OTHER PACIFIC ISLANDER" = "Native Hawaiian Or Other Pacific Islander"</li> <li>"WHITE" = "White"</li> </ul>	<p>Collected (Source: Investigator) Annotated CRF [ <a href="#">L</a> ]</p> <p>Since no subjects had more than 3 Races, RACE4 was not used.</p>
--	--------	------	------------------	----	--	---

# Appearance of Annotation

## Little Tips

- To distinguish the domain level annotations from the variable annotations a larger font can be used for the domain annotations.

Domain Annotation Font Size	Variable Annotation Font Size
14 Points	12 Points

- Instructional text and comments should be case sensitive.

Race (Check all that apply)

White

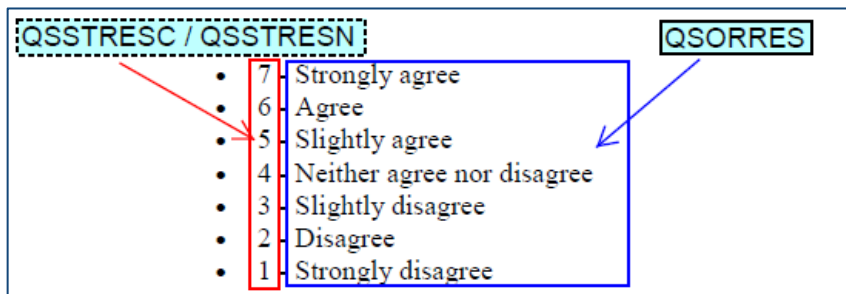
Black or African American

When multiple values are selected then RACE = MULTIPLE and individual responses are RACE1, RACE2, RACE3, etc. in SUPPDM

# Appearance of Annotation

## Little Tips

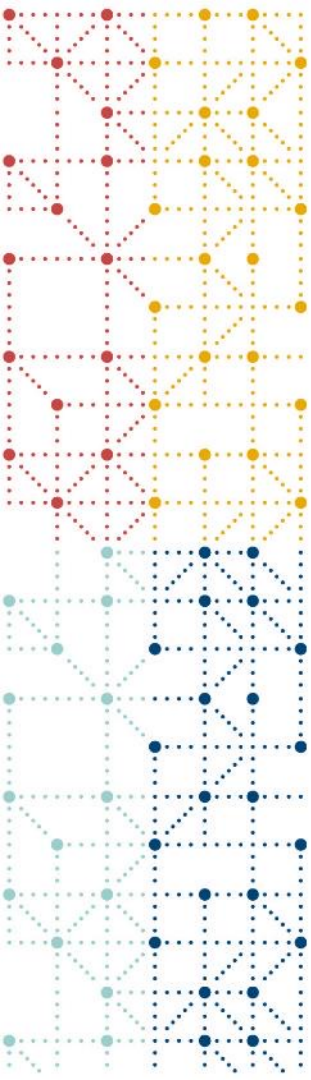
- Supplemental references—via boxes, arrows, and lines—can be used to further clarify annotations.



- To annotate Similar CRF form, the SDTM-MSG V2.0 recommends annotating. To annotate “See Page <n> for Annotations” on similar CRF pages that makes reviewability more difficult.



DEMO



**Thank You!**

**cdisc**