

CDISC Public Webinar – Standards Updates and Additions

13 OCT 2016



Strength *through Collaboration*

Agenda

- Ebola TA Public Review

- Rhonda Facile, MS, Vice President, Standards Development
- Maura Kush, CDISC Specialist and Data Standards Consultant, Pharmastat
- Shannon Labout CCDM, Vice President of Education, CDISC
- Bess LeRoy, Metadata Engineer, CDISC
- Jon Neville, PSM, Program Director, Data Standards and Management
- Diane Wold, Senior Director, Standards Development and Modeling, CDISC

- Malaria TA Public Review

- Lesley Workman, Scientific Coordinator, Division of Clinical Pharmacology, University of Cape Town
- Bess LeRoy, Metadata Engineer, CDISC
- Alana St. Clair, Associate Project Manager, CDISC
- Jon Neville, PSM, Program Director, Data Standards and Management
- Diane Wold, Senior Director, Standards Development and Modeling, CDISC
- Rhonda Facile, MS, Vice President, Standards Development

Question & Answer

- 'Panelist': Question
- OR
- 'Presentation': Question

Examples:

Maura: Where can I find the Ebola TA Guide?

Malaria Team: Are there known issues with the Malaria TA analysis section?

EVD Therapeutic Area User Guide

Maura Kush
EVD Project Co-Lead

Panelists: Bess LeRoy, Diane Wold, Jon Neville, Shannon
Labout



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Agenda

- Background on project and participants
- Scope and gap analysis
- Additional requirements
- Known issues and focus of public review
- Public review dates

Background

- Sense of urgency during the Ebola outbreak
- Collaboration with the Oxford University Clinical Research Unit (OUCRU) in Vietnam
- Needed to address certain challenges of EVD studies
 - Outbreak setting
 - Many different stakeholders, all with different use cases and research goals
- Led to a gap analysis and a narrowed scope with more focused development

EVD TAUG Scope

- Majority based on data elements from WHO CORE case report form (CRF),
 - Includes some epidemiological and vaccination data
 - Does not include pediatric or survivor data, or vaccine trials as a whole
 - Does not include ADaM
- Describes how to represent the most common data for Ebola studies

EVD TAUG Documentation

- SDTM v1.4
- SDTMIG v3.2
- SDTMIG – AP
- Controlled Terminology
- CDASH v1.1
- Influenza v2.0, Tuberculosis v2.0, Malaria v1.0

Ebola Table of Contents

- 1 INTRODUCTION
 - 1.1 How to Read this Document
 - 1.2 Organization of this Document
 - 1.3 Known Issues
- 2 SUBJECT AND DISEASE CHARACTERISTICS
 - 2.1 Subject Characteristics
 - 2.2 EVD Exposure and Risk Factors
 - 2.2.1 Environmental Risk Factors
 - 2.2.2 Source Case Investigation
 - 2.2.3 Contact Case Investigation
- 3 DISEASE ASSESSMENTS
 - 3.1 Daily Assessment of Signs and Symptoms
 - 3.2 Diagnosis and Confirmation of Infection
 - 3.3 Immune Response
 - 3.4 Reactogenicity
 - 3.5 Questionnaires, Ratings, and Scales
- 4 ROUTINE DATA
 - 4.1 Concomitant Medications
 - 4.2 Measured/Estimated Weight
 - 4.3 Medication Redosing
 - 4.4 Reproductive Findings
 - 4.5 End of Study/Early Termination

Threshold Cycle CDASH CRF and SDTM Examples

mb.xpt

Row	STUDYID	DOMAIN	USUBJID	SPDEVID	MBSEQ	MBREFID	MBTESTCD	MBTEST	MBTSTDTL	MBORRES	MBSTRESC	MBSPEC	MBMETHOD	VISITNUM	VISIT	MBDTC	CTVAL
1	EVD123	MB	EVD01-01	ABI7500	1	SAMP0101	ZEBOV	Zaire Ebolavirus	IDENTIFICATION	NEGATIVE	NEGATIVE	BLOOD	QUANTITATIVE REVERSE TRANSCRIPTASE POLYMERASE CHAIN REACTION	1	VISIT 1	2014-08-06	42
2	EVD123	MB	EVD01-02	ABI7500	2	SAMP0102	ZEBOV	Zaire Ebolavirus	IDENTIFICATION	POSITIVE	POSITIVE	BLOOD	QUANTITATIVE REVERSE TRANSCRIPTASE POLYMERASE CHAIN REACTION	1	VISIT 1	2014-08-06	26
3	EVD123	MB	EVD01-03	ABI7500	3	SAMP0103	ZEBOV	Zaire Ebolavirus	IDENTIFICATION	NEGATIVE	NEGATIVE	BLOOD	QUANTITATIVE REVERSE TRANSCRIPTASE POLYMERASE CHAIN REACTION	1	VISIT 1	2014-08-06	41

MB NSV Metadata

Variable	Label	Type	Role	Origin
CTVAL	Threshold Cycle Value	text	Non-Standard Variable Qualifier	CRF

Date	<input type="text" value="mm / dd / yyyy"/>
MBDAT MBDTC <i>dd-MON-yyyy</i>	
Time	<input type="text" value="-- : -- --"/>
MBTIM MBDAT <i>hh:mm:ss</i>	
Method	<input type="text" value="qRT-PCR"/>
MBMETHOD <i>Pre-specified</i>	
Test Name	<input type="text" value="ZEBOV"/>
MBTEST <i>Pre-specified</i>	
Device Type	<input type="text"/>
DIVAL where DIPARM = 'Device Type'	
Specimen Type	<input type="radio"/> Blood <input type="radio"/> Serum <input type="radio"/> Other
MBSPEC	
Specimen Type Other	<input type="text"/>
SPTYPOTH MBSPEC	
Accession Number	<input type="text"/>
MBREFID	
Kit Number	<input type="text"/>
SPDEVID	
Result	<input type="radio"/> Positive <input type="radio"/> Negative <input type="radio"/> Unknown
MBORRES	
Threshold Cycle Result	<input type="text" value=""/> cycles
SUPPMB.QVAL where QNAM = 'CTVAL'	

Sponsors are responsible for understanding and implementing CDISC Controlled Terminology where applicable.

Viral Identification

- Historically viral identification shown in LB
- This has changed, and this guide has examples that show it in MB where
MBTSTDTL='IDENTIFICATION'

mb.xpt

mb.xpt

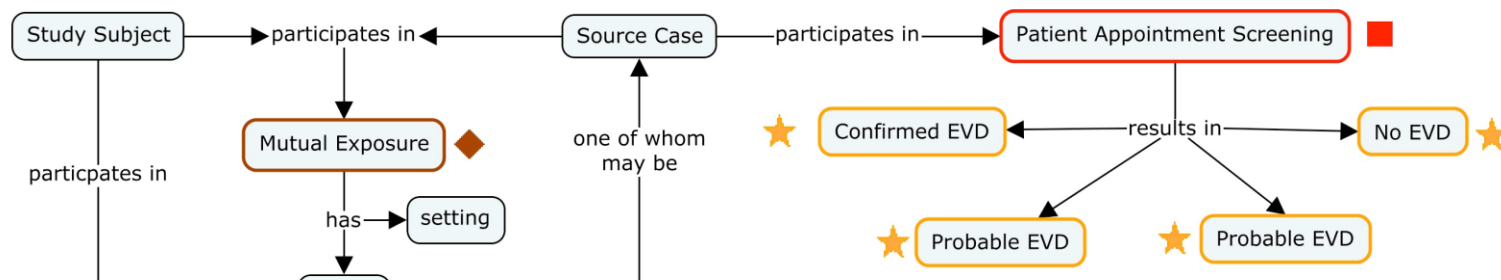
Row	STUDYID	DOMAIN	USUBJID	SPDEVID	MBSEQ	MBREFID	MBTESTCD	MBTEST	MBTSTDTL	MBORRES	MBSTRESC	MBSPEC	MBMETHOD	VISITNUM	VISIT	MBDTC	CTVAL
1	EVD123	MB	EVD01-01	ABI7500	1	SAMP0101	ZEBOV	Zaire Ebola virus	IDENTIFICATION	NEGATIVE	NEGATIVE	BLOOD	QUANTITATIVE REVERSE TRANSCRIPTASE POLYMERASE CHAIN REACTION	1	VISIT 1	2014- 08-06	42
2	EVD123	MB	EVD01-02	ABI7500	1	SAMP0102	ZEBOV	Zaire Ebola virus	IDENTIFICATION	POSITIVE	POSITIVE	BLOOD	QUANTITATIVE REVERSE TRANSCRIPTASE POLYMERASE CHAIN REACTION	1	VISIT 1	2014- 08-06	26
3	EVD123	MB	EVD01-03	ABI7500	1	SAMP0103	ZEBOV	Zaire Ebola virus	IDENTIFICATION	NEGATIVE	NEGATIVE	BLOOD	QUANTITATIVE REVERSE TRANSCRIPTASE POLYMERASE CHAIN REACTION	1	VISIT 1	2014- 08-06	41

MB NSV Metadata

Variable	Label	Type	Role	Origin
CTVAL	Threshold Cycle Value	text	Non-Standard Variable Qualifier	CRF

New Domain

- ER (Environmental Risk Factors)
 - Out for review with SDTM v3.3 Batch 3
 - Events observation class domain for data collected on potential exposures to, or diseases by way of environmental contact or through participation in activities associated with risk



er.xpt

Row	STUDYID	DOMAIN	USUBJID	ERSEQ	ERTERM	ERCAT	ERDTC	ERSTDTC	ERENDTC	CNCTYP
1	VHF-2015	ER	VHF-15-101	1	Exposure to EVD	SOURCE CASE INVESTIGATION	2015-02-05	2015-01-02	2015-01-10	Touched or shared linens, clothes, dishes or utensils of case

ER NSV Metadata

Variable	Label	Type	Role	Origin
CNCTYP	Contact Type	text	Non-Standard Record Qualifier	CRF

Known Issues

- Non Standard Variables (NSV) - proposed changes
- Signs and symptoms in Findings About (FA)
- Personally identifying information in Subject Characteristics (SC)
- Controlled Terminology is not final
- NHOID='Zaire Ebola Virus' - provides enough detail without an OI domain
- Measured/Estimated Weight

Public Review

- Public Review comments due: 4 November 2016
- Public review is a key part of the standards development process.

We look forward to getting your feedback!

DEVELOPING A DATA STANDARD FOR UNCOMPLICATED MALARIA

Clinical **D**ata **I**nterchange **S**tandards **C**onsortium (CDISC)
World**W**ide **A**ntimalarial **R**esistance **N**etwork (WWARN)

- WWARN collaboration
- 13 October, 2016

www.wwarn.org

Twitter: @WWARN



Topics

- Project Scope - Alana
- Project Background - Lesley
- New Domains and Examples - Bess
- Known issues - Bess
- Public review

Malaria Standards Development Project Scope

- Describe how to represent the most common data for uncomplicated malaria studies
 - Deliverables: CDISC Therapeutic Area User Guide, concept maps, CDASH compliant CRF, SDTM examples and controlled terminology
- Based on data elements from WWARN case report form (CRF)
 - Includes examples for parasite genotyping, combination therapy, dosing with food and post dose vomiting, among others
 - Includes an analysis considerations section

Project Mission:

To support efficient, scientifically valid generation and reporting of clinical malaria data to streamline antimalarial development, regulatory submission and post-marketing research, as well as enable data sharing, comparison and aggregation.

Aim:

Consensus-based development of a single, freely available data standard to ensure the consistent use of existing CDISC standards, and to facilitate alignment and development of new standards for the electronic acquisition, exchange, submission and archiving of clinical malaria data collection, analysis and reporting

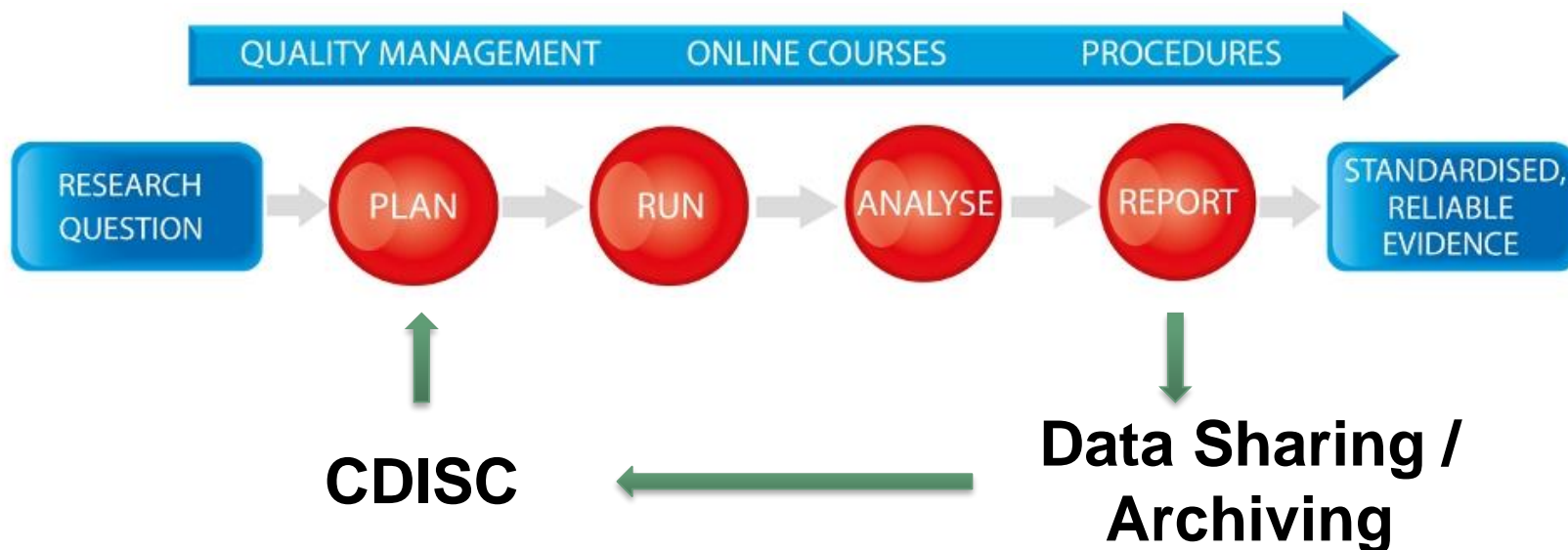
Scope (Year 1):

- Uncomplicated falciparum malaria
- Uncomplicated vivax malaria

Preventive treatment (e.g SMC, IPT, MDA), **Induced Blood Stage Malaria (Human Challenge Studies)**, severe malaria to follow, as needed

Role of WWARN

- Facilitate the development of a CDISC data standard for malaria.
- Enable data sharing
- Conduct pooled data analyses to quantify effects
- Provide the (long-term) storage infrastructure and maintaining of the antimalarial data repository / archive



The role of stakeholders

Current Stakeholders include:

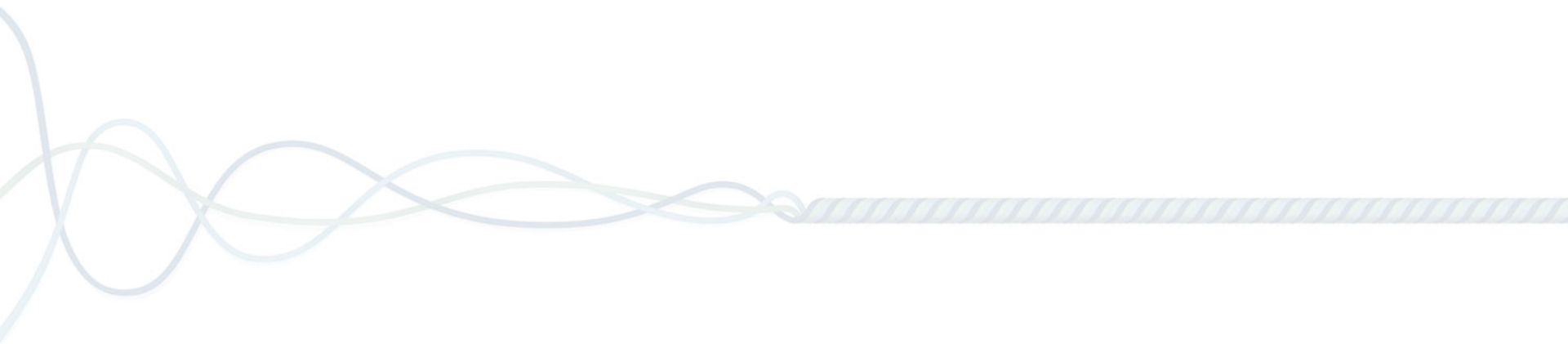
- *CDISC, C-PATH*
- *WWARN members*
- *WHO GMP / TDR*
- *BMGF*
- *GHT, LSTM,*
- *Pharma:*
 - *GlaxoSmithKline*
 - *Medicines for Malaria Venture*
 - *Merck*
 - *Novartis*
 - *Sanofi*
 - *Shin Poon*
 - *Sigma Tau*
 - *Takeda*
 - *UCB*

Review draft data standards

Share relevant experience:

- Recent CRF templates / Database Structures / Statistical Analysis Plans
- Identification of critical issues in regulatory submissions.

CDASH CRF and SDTM Examples



New Domain #1

- ER – Environmental Risk Factors

Household Characteristics (ER)	Occur EROCCUR
Has the Malaria Control Program performed Indoor Residual Insecticide Spraying (IRS) of your home in the last 12 months? EHTERM=Indoor residual insecticide spraying	<input type="radio"/> Yes <input type="radio"/> No
Did you sleep under an insecticide treated bed-net last night? ERTERM=Sleep under an insecticide treated bed-net	<input type="radio"/> Yes <input type="radio"/> No

Row 1: Shows that subject ABC-001's household had indoor residual insecticide spraying within the last 12 months (EREVLINT).

Row 2: Shows that subject ABC-001 slept under an insecticide treated bed-net last night (EREVINTX). [MAL-112](#) - Getting issue details... STATUS

er.xpt

Row	STUDYID	DOMAIN	USUBJID	ERSEQ	ERTERM	ERCAT	ERPRES	EROCCUR	ERDTC	EREVLINT	EREVINTX
1	ABC	ER	ABC-001	1	Indoor residual insecticide spraying	HOUSEHOLD CHARACTERISTICS	Y	Y	2015-05-15	-P12M	
2	ABC	ER	ABC-001	2	Sleep under an insecticide-treated bed net	HOUSEHOLD CHARACTERISTICS	Y	Y	2015-05-15		LAST NIGHT

New Domain #2

- SI – Site Summary

si.xpt

Row	STUDYID	DOMAIN	SITEID	SEQ	SIGRPID	SIPARMCD	SIPARM	SIVAL	SIVALNF	SIVALCD	SIVCDREF	SIVCDVER
1	ABC	SI	123	1		TRANINTY	Transmission Intensity	HIGH		C22222	CDISC	2016-03-25
2	ABC	SI	345	1		TRANINTY	Transmission Intensity	MEDIUM		C22222	CDISC	2016-03-25
3	ABC	SI	567	1		TRANINTY	Transmission Intensity	HIGH		C22222	CDISC	2016-03-25

Known Issues

- Representation of combination therapy data
 - We show an example using a combination of EC and EX as a possible solution. This is different from the modeling strategy used in the TAUG-TB (example using a combination of EX and FA).
- Use of --EVINTX for representation of timing around food intake and post dose vomiting
 - There is disagreement regarding whether or not this is an appropriate use of --EVINTX.

30-Day Public Review

- Public Review dates:
 - **October 6 – November 6**
- Public review is a key part of the standards development process.

We look forward to getting your feedback!

Q&A



CDISC Online Education & Event Updates

John Ezzell, CDISC



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Upcoming Webinars

Presenter	Topic	Webinar Date
John Owen, Project Manager, CDISC	Prostate Cancer TA Public Review	19 OCT 2016
Dr. Lauren Becnel, CDISC Anthony Chow, Sr. Manager, CDISC	SHARE Deep Dive	20 OCT 2016
Dr. Lauren Becnel, CDISC	BRIDG Overview	TBD

Webinar details and registration at www.cdisc.org/webinars

Standard currently out for review

- Terminology Call for Public Review Package 28
Comments Due by: October 7, 2016
- Draft Content SDTMIG v3.3 Batch 3 Now Available for Public Review
Comments Due by: October 21, 2016
- New Draft Ebola v1 Now Available for Public Review
Comments Due by: November 4, 2016

UPCOMING NORTH AMERICA PUBLIC COURSES

Location	Dates	Courses Offered:	Discount period ends:	Late fees kick(ed) in:	Host
Miami, FL	23-27 Jan 2017	SDTM, CDASH, ADaM Primer, ADaM T&A	23 Oct 2016	23 Dec 2106	 biorasi CLINICAL RESEARCH OPTIMIZED™ Powered by TALOS™
Raleigh, NC	27 Feb – 3 Mar 2017	SDTM, CDASH, ADaM Primer, ADaM T&A	28 Nov 2016	28 Jan 2017	
Audubon, PA	3-7 Apr 2017	SDTM, CDASH, ADaM Primer, ADaM T&A, CT, Define-XML	3 Jan 2017	3 Mar 2017	 BIOCLINICA SEE MORE CLEARLY
Toronto, ON	5-9 June 2017	SDTM, CDASH, ADaM Primer, ADaM T&A, Define-XML	6 Mar 2017	5 May 2017	 McDOUGALL SCIENTIFIC INSIGHTS YOU CAN TRUST

Visit cdisc.org/public-courses for information on other CDISC Public Training events.

UPCOMING EUROPE PUBLIC COURSES

Location	Dates	Courses Offered:	Discount period ends	Late fees kick(ed) in:	Host
Basel, Switzerland	7-11 Nov 2016	SDTM, ADaM Primer, ADaM T&A, Define-XML	Ended	7 Oct	
London, UK	24-28 Apr 2017	TBD	TBD	TBD	

Visit cdisc.org/public-courses for information on other CDISC Public Training events.

UPCOMING ASIA PUBLIC COURSES

Location	Dates	Courses Offered	Discount period ends:	Late fees kick(ed) in:	Host
Beijing, China	18-21 Oct 2016	SDTM, CDASH, ADaM Primer, ADaM T&A, ODM, Define-XML	Ended	None	
Shanghai, China	24-27 Oct	SDTM, CDASH, ADaM Primer, ADaM T&A, ODM, Define-XML	Ended	None	
Tokyo, Japan	5-9 Dec	SDTM, CDASH, ADaM Primer, ADaM T&A, ODM, Define-XML	Ended	5 Nov	

Visit cdisc.org/public-courses for information on other CDISC Public Training events.

CDISC Online Training Production Update

- Just Released
 - ADaM Module 4 Online Training
 - Mini-Training: Ensuring USUBJID is Unique for an Individual in an Application

Drag and Drop Exercise: Required, Conditionally Required and Permissible ADSL Variables

Instructions: Drag the Required, Conditionally Required and Permissible variables into the correct barrels. When you are complete, click the "Submit" button to check your answers.

SITEGRY: pooled group of sites used for analysis

USUBJID: unique subject identifier

AGEGRY: pooled age group

REGIONY: pooled groups of sites into geographic regions

AGE and AGEU

SITEID: unique site identifier

RACE

SUBJID: subject identifier used within study

RACEGRY: pooled race group

STUDYID: study identifier

Required or Conditionally Required variables

Permissible variables

CDISC © CDISC 2016 Submit

Online Courses in Development
TA Alzheimer's
TA QT Studies
TA Vaccines
TA Prostate Cancer
TA Rheumatoid Arthritis
TA Pain
ADaM Modules 5-8
CT Module 1 & 2
Define-XML

Any more questions?

Thank you for attending this webinar.

**CDISC's vision is to:
Inform Patient Care & Safety Through Higher Quality Medical Research**



Strength *through collaboration.*

CDISC Members Drive Global Standards

Thank you for your support!



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Authoritative. Global. Vendor neutral.