CDISC Implementation Committee

Neesha Desai

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Agenda

- NCI Requirement
- Project Updates
- LPO Status Updates
- LPO Support Questions
- LPO Impact Analysis Review
- Project Timeline

NCI Requirement

- **January 2020** **Date will be assessed based on LPO Impact Analysis*
  - All CTEP IND studies activated on/after 1/1/2020 shall be CDISC compliant.

Project Updates *(1)*

- **CDISC Mailbox**
  - ncicdiscsupport@nih.gov

- **CDISC Wiki**
  - NCI maintenance concluded
  - CDISC Webinars and slides will also be posted on the CTSU Collaboration Portal the ***last week of May***
  - CDISC LPO Impact Analysis Feedback and Support Request details will be posted to the CDISC Wiki and the CTSU Collaboration Portal the ***1st week in June***

Project Updates *(2)*

- **CDISC Harmonization Working Group**
  - NCI Project team working on the roles/responsibilities (R/R) of group members
  - Will bring R/R to the CDISC Implementation Committee for review

Project Updates *(3)*

- **LPO Support Webinars/Sessions**
  - May 22nd **"Overview of FDA Submission Process"**
    - Session 1: 10- 11:30am ET
    - Session 2: 3- 4:30pm ET
  - May 24th CDISC SME Session
    - *Rescheduling due to the holiday weekend and low attendance*
  - May 28th **"CDASH/ SDTM/ ADaM"**
    - Session 1: 10- 11:30am ET
    - Session 2: 3- 4:30pm ET
  - May 29th **"Using TA User Guides Webinar"**
    - Session 1: 10 - 11:30am ET
    - Session 2: 3- 4:30pm ET

LPO Status Updates

NCI/Group	Status	Beta Release IA: GLIB	Beta Release IA: CTSU Standard Forms
Alliance	Initial planning phase Discussing internally with IT staff, resourcing and prioritization, evaluating the next beta release	<Submitted>	<Submitted>
Theradex	Planning phase Waiting for tools; DMU is ready for CDISC implementation	<Submitted>	
NRG	Rolling out using CDISC from start to finish; Initial planning phase – Near completion of content for study build, next step to go to the global library and do a comparison on global library	May not be able to submit comments due to other activities	
SWOG	Initial planning phase Made change to standard forms to try CDASH		
ECOG-ACRIN	Started internal WG that includes study builders, programmers and stats, working on building simple forms, assessing and evaluating the standard forms ALS	Did not submit, not far enough along	
CCTG	Started an internal WG with stats, programmers. Going through GLIB, domain by domain. Started the CDISC CRFs, running into questions for custom data fields.	<Submitted>	
COG	Initial planning phase	Not far enough to go through the GLIB	

LPO Support Questions *(1)*

LPO Question	Response	LPO Feedback
I have a piece of general feedback that came out of the variables training session today. I like the pattern that Shannon described for variables and I think it would make a lot of sense to adopt that type of pattern in building our forms. The ALS that we were given does not follow this pattern, however, I think it would be easy enough for us to add the additional descriptors while creating our template study.	**CDISC Int. FG:** Pattern-based naming is being implemented for CDISC variables in the CDISC GLIB and CDISC CTSU Standard Forms ALS documents. The two-letter domain prefix (e.g., 'DM_', 'EX_') will be added to Rave Field OIDs and Variable OIDs. Study level variables are excluded from this update (e.g., site ID, patient ID, etc)	
The concern I have is more about the AE forms, which I think are supposed to be provided to us next week. On those forms, we have to use them as they are provided and cannot change the variables. Would CTEP consider using this pattern on those forms? The AE domain is a critical piece of the SDTM and it would be a shame to have a pattern that works for everything else but have to do something completely different just for that form. Could be worth a discussion to see if all the groups agree on using that pattern for variables.	**CDISC Int. FG:** Pattern-based naming is being implemented for CDISC variables in the CDISC GLIB and CDISC CTSU Standard Forms ALS documents. The two-letter domain prefix (e.g., 'DM_', 'EX_') will be added to Rave Field OIDs and Variable OIDs. Study level variables are excluded from this update (e.g., site ID, patient ID, etc)	

LPO Support Questions *(2)*

LPO Question	Response	LPO Feedback
Following the QRS presentation, I wanted to bring up a question regarding non-standard questionnaires. QRS is for standardized, named instruments. How do you handle a "questionnaire" that is used to collect information regarding a topic but is not a standardized form? For instance, I recently built some forms for a study that had several questionnaires. Some of the forms were standardized, named instruments, but others were asking a number of questions related to the disease and of interest to the investigator but were not captured on a standardized form. How would this type of data be handled?	**CDISC SME:** If the form is not standardized (not named, not version controlled, not scored, etc) it is probably not a QRS instrument. Will review internally with Theradex to review the form.	
I've been very focused on Alliance, but at Mayo Clinic we have other groups that we support for study build as well, including CPN. Is CPN included in the CDASH mandate from NCI? Assuming the mandate extends beyond NCTN trials, is there somewhere the full scope is listed?	**CDISC Int. FG:** All CTEP IND trials must be CDASH compliant. We encourage use for all other studies.	
There's a "Serious" question (AESER) that is Expected in SDTM. How would we get that from this form? Would that be derived based on the report recommendation? (SWOG)	**CDISC SME:** You could derive it from the Serious Criteria (which are strongly recommended on the AE CRF because FDA wants these values.)	Follow up on a SME call Follow up with Shanda and NCI to review process of how the data is flowing

LPO Support Questions *(3)*

LPO Question	Response	LPO Feedback
I know we've discussed the use of "Ongoing" a few times and the various ways that you can map that to timing variables in SDTM but it would be nice to see an example of how that would be done for this form.	CDISC SME will show examples from the CDASHIG and SDTMIG	
There are a lot of MedDRA fields in SDTM (AELLT, AEPTCD, AEHTL, AESOC, etc.) that look a little bit confusing to me. It would be helpful to see an example of what that data would look like.	CDISC SME will show a small example dataset	

LPO Impact Analysis: Beta Release GLIB *(1)*

#	Document	LPO Findings/ Recommendations/ Questions	Decision
1	GLIB ALS	04/18/2019: I'm assuming a new release of the Controlled Terminology (CT) will be released prior to July. I wanted to confirm the final ALS release will contain the most up-to-date information. KL With the final release in July, will the GLIB and other ALS files be updated using the newest CDASH and SDTM controlled terminology packages?	CTSU will do an impact analysis (for CT Packages released by CDISC before July 2019) to understand the potential impact to our timeline. The intent it to provide documents as current as possible to CDISC releases/ updates; this must be done in consideration of the implementation timelines.
2	GLIB Release Notes: Section 3.2 Standard Conventions – Table 6	This table states that we cannot modify the Field OID. However, on the Variable Naming NVS webinar (3/28), Shannon presented several instances where it is expected and best practice to rename field OIDs in Rave for data collection purposes. In addition, the guide provides examples in 6.3 where it is OK/expected to change the field OIDs thus conflicting with the table. Please confirm Table 6 will be updated to reflect this.	05/02/2019 CDISC Integration FG: • LPOs can not change the field OIDs on Standard Forms • LPOs can change the field OIDS on Non-Standard forms if they have a strong reason to do so. "Standard Forms" applies to the CTSU Standard Forms ALS, "Non-Standard Forms applies to variables not specified in the CTSU Standard Forms ALS.
3	GLIB Release Notes: Section 3.2 Standard Conventions – Table 7	Given the needed updates to the Field OID to be specific (see situations/comment from 2), the SAS label should be modified for Rave data collection to ensure our programmers using CDASH Rave outputs can correctly identify the fields using the OIDs and Labels. Therefore, the SAS Label should be allowed to be modified when the Form OID is changed. (When data is transformed into SDTM format, the SAS Label can also be updated)	Maintain the same SAS label as in the CDISC GLIB for consistency across LPOs.
4	GLIB ALS	In the ALS, the Field name is the Variable Label + PID. Why the Variable Label? At the Alliance, we find having the Field Name match the Field OID + PID is much easier for specification writing, debugging and general viewing. (Medidata help guides state that in most cases, the Field name is the same as the Field OID.) As far as we understand, the Field Name isn't used for anything site facing or outputs. If the global ALS will not be changed, can the Alliance modify this?	No changes to the Rave Field Names in the CDISC GLIB ALS and CDISC CTSU Standard Forms ALS; the default naming convention will remain as-is, "the caDSR Long Name of the CDE + PID details". LPOs can use "the caDSR Long Name of the CDE + PID details" or choose to use "the caDSR Short Name of the CDE + PID details" as the Rave field name as needed. If the LPO chooses to use "the caDSR Short Name of the CDE + PID details" for the Rave field name, they are responsible for this update in Rave.

LPO Impact Analysis: Beta Release GLIB *(2)*

#	Document	LPO Findings/ Recommendations/ Questions	Decision
5	GLIB Release Notes: Table 6	For the User Data String, Table 6 states we must use the NCI Preferred term. However, CDISC training has stated we can use the NCI Preferred Term or CDISC Synonym. Please clarify which is correct.	LPOs can decide to use the NCI Preferred Term or the CDISC Synonym for the User Data String in Rave. If LPOs decide to use the CDISC Synonym instead of the NCI Preferred Term, they will be responsible for related study build activities and mapping activities for SDTM submission. The 'CDISC GLIB Release Notes' document will be updated to provide details for use of the CDSIC Synonym versus the NCI Preferred Term. The CDISC GLIB ALS will remain as is, no changes will be made to the User Data String and NCI Preferred Terms.
6	GLIB ALS	Why is the Data Dictionary Name not modifiable? Some dictionaries will be used across several fields; however, different sub-sets of responses will be needed for the questions. i. Rave does not allow us to subset the dictionary choices at the field level – a separate dictionary must be created ii. Example: NY. Sometimes Unknown or Not Applicable are not valid responses for a question iii. The Alliance would be open to a standard naming convention in this situation.	LPOs should use the vetted CSC Data Elements Work Group (DEWG) naming convention of appending integer when subsetting, and appending the letter F behind any Data dictionary name that ends with a number (e.g., “_1F”, ”_2F”, “_3F”) No changes will be made to the CDISC GLIB ALS. The CDISC GLIB Release Notes will be updated to include this naming convention for subsetted dictionary names.
7	GLIB ALS	When the GLIB is updated/versioned, can we get a detailed change log with the future releases?	A change log will be provided in the release notes for each updated GLIB release.

LPO Impact Analysis: Beta Release GLIB *(3)*

#	Document	LPO Findings/ Recommendations/ Questions	Decision
8	NCI/CDISC Best Practices Document	04/18/2019: In this section, it talked about all of these documents needed to be provided. I'm curious if there is any guidance to expected/required format and/or the level that we have to have prepared for CTEP-IND trials. We do not submit data directly to the FDA; I'm curious if we have to have annotated CRFs, and all these other documents that are listed for all trials. And if the trial has intent to submit to FDA, can we negotiate with our pharma partner on these. KL Section 4.3: What does this mean for us as LPO's? We are not translating all of these statements into what actions need to be done on Alliance trials. (Maybe the training is still coming?)	All CTEP IND studies activated on/after 1/1/2020 must be CDISC compliant. CDISC Compliance is highly recommended for studies that activate prior to the 2020 date. The CDISC SME to provide additional feedback at the "Overview of FDA Submission Process" Webinar on May 22nd, 2019.
9	NCI/CDISC Best Practices Document	Could a real life example/demo of the described best practices for the Eligibility Checklist Forms be provided? We're not following the workflows, when to curate/not curate, etc.	The CBIIT caDSR team presented the curation support/ process for eligibility criteria at the December Committee Meeting and an overview of this process at the April Committee Meeting.
10	NCI/CDISC Best Practices Document	Does NCI/CTEP submit data to FDA on NCTN trials? If so, study by study or aggregated across studies?	No decision required, NCI does not submit to the FDA.
11	NCI/CDISC Best Practices Document	Throughout the entire document, please be specific what CDISC compliant means. CDASH, SDTM, ADaM, etc. a. Additionally, our understanding from previous meetings is that the requirement is to be CDASH compliant only. We understood that it is up to the LPOs discretion and negotiations with the Pharma partner on SDTM programming. Please be specific in the document about whether we are referring to CDISC-CDASH or CDISC-SDTM.	The FDA requirement is SDTM. the NCI requirement is CDASH. CDASH compliance makes SDTM compliance easier to achieve. CDISC PM to review and update the NCI CDISC Best Practices document as needed to ensure clear communication of the NCI requirement for CDASH compliance.

CDISC Implementation Timeline

Milestone	Completion Date
CDISC Rave GLIB ALS	
✓ 1) Created Beta ALS	February 15, 2019
✓ 2) Completed Internal Compliance review on Beta ALS	February 27, 2019
✓ 3) Released Beta ALS to LPOs	March 5, 2019
✓ 4) Complete Impact Analysis **LPOs**	April 2, 2019
➤ **5) Manage Impact Analysis updates**	May 17, 2019
6) Complete Internal Compliance review on Final ALS	June 14, 2019
7) Release Final CDISC Rave GLIB ALS to LPOs	July 9, 2019
CDISC-compliant CTSU Standard Forms ALS v7.0	
✓ 1) Created Beta ALS	March 7, 2019
✓ 2) Completed Internal Compliance review on Beta ALS	March 20, 2019
✓ 3) Release Beta ALS to LPOs	April 1, 2019
4) Complete Impact Analysis **LPOs**	May 13, 2019
➤ **5) Manage Impact Analysis updates**	June 13, 2019
6) Complete Internal Compliance review on Final ALS	June 28, 2019
7) Release Final CDISC-compliant CTSU Standard Forms ALS v7.0 to LPOs	July 9, 2019
Build Seed Study in collaboration with CDISC SME **LPOs**	December 31, 2019
All Groups to use CDISC Standards on <u>All CTEP IND Trials</u>	**January 1, 2020**

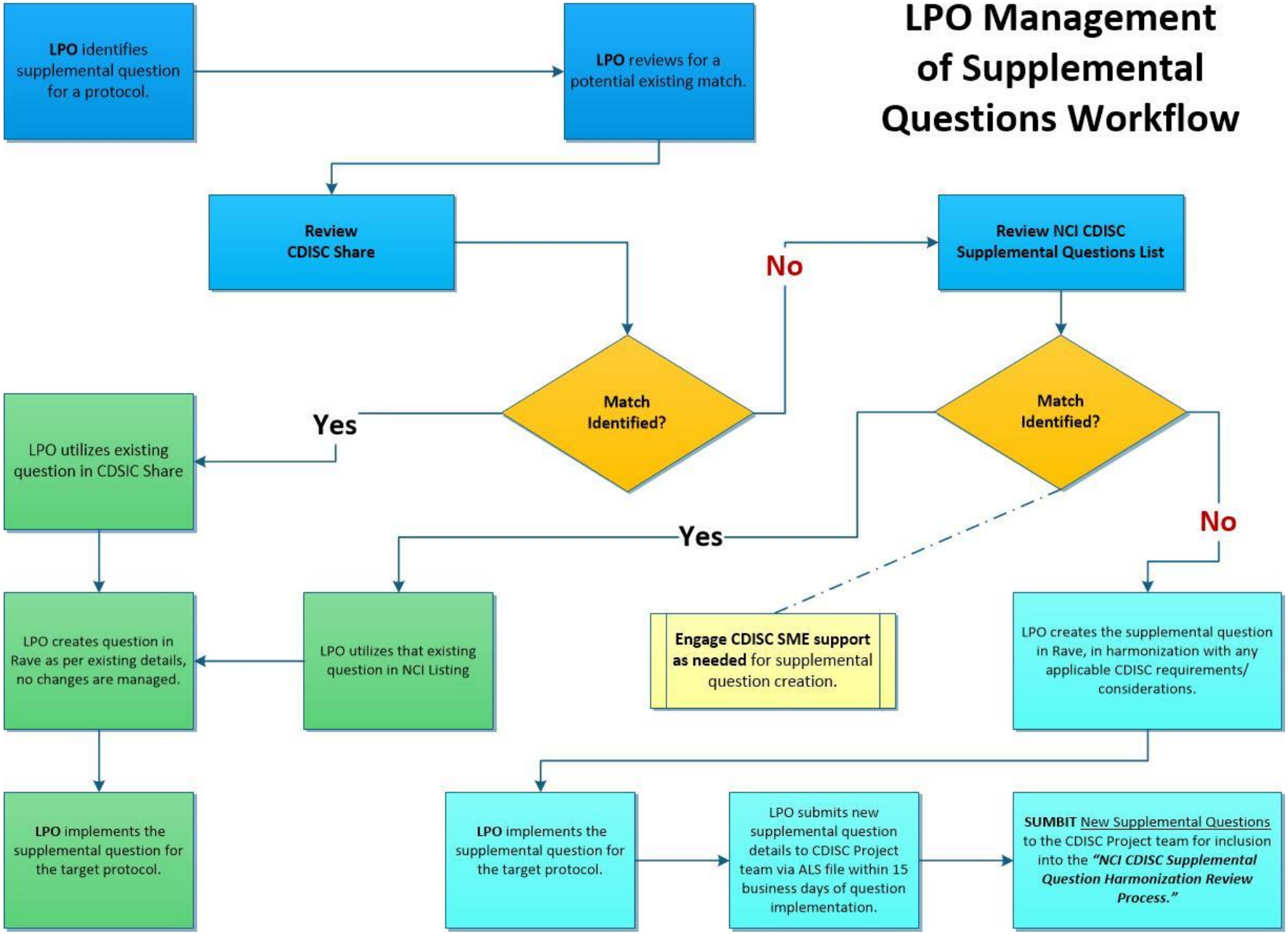
Next Steps

- Next Meeting is June 26th, 2019
- REMINDER: Send email to ncicdiscsupport@nih.gov specifying 1 primary representative and 1 secondary representative for the CDISC Harmonization WG

Appendix

CDISC Supplemental Questions Workflow
CDISC Harmonization Workflow
NCI CBIIT: CDEs for EC Forms Overview
CDISC Implementation Workflow
CTSU Change Matrix
eCRF Build Scenarios

LPO Management of Supplemental Questions Workflow



CDISC Harmonization Workflow *v01/09/2019*

1) LPO submits new supplemental question details to CDISC Project team via ALS file within 15 business days of question implementation.
2) CDISC Project team adds the new supplemental question to the **CDISC LPO Supplemental Question Listing**[1].
3) CDISC Project team prepares supplemental questions (partial or no match) for the **CDISC Harmonization Working Group (WG)** for harmonization/ standardization review.
4) **CDISC Harmonization WG** reviews, standardizes and harmonizes the supplemental question/s.
5) CDISC Project team prepares a supplemental questions review request for submission to CDSIC.
6) CDISC Project team submits and tracks review requests to CDISC.

Footnotes

1. *The **CDISC LPO Supplemental Question Listing** is a document created and maintained by the NCI CDISC Project team. This document will specify:*
   a. *Supplemental questions that have been recently created and implemented by an LPO for a specific protocol;*
   b. *Supplemental questions that have been harmonized/ standardized by the **CDISC Harmonization WG** and submitted by the CDISC Project team to CDISC for review.*

NCI CBIIT: CDEs for EC Forms Overview

1. EC CDEs – *Inclusion/Exclusion Criteria*

- CDASH does not provide a variable for individual inclusion/exclusion questions
- Continue to curate these CDEs using your current process

TI – Examples for Trial Inclusion/Exclusion Criteria Dataset

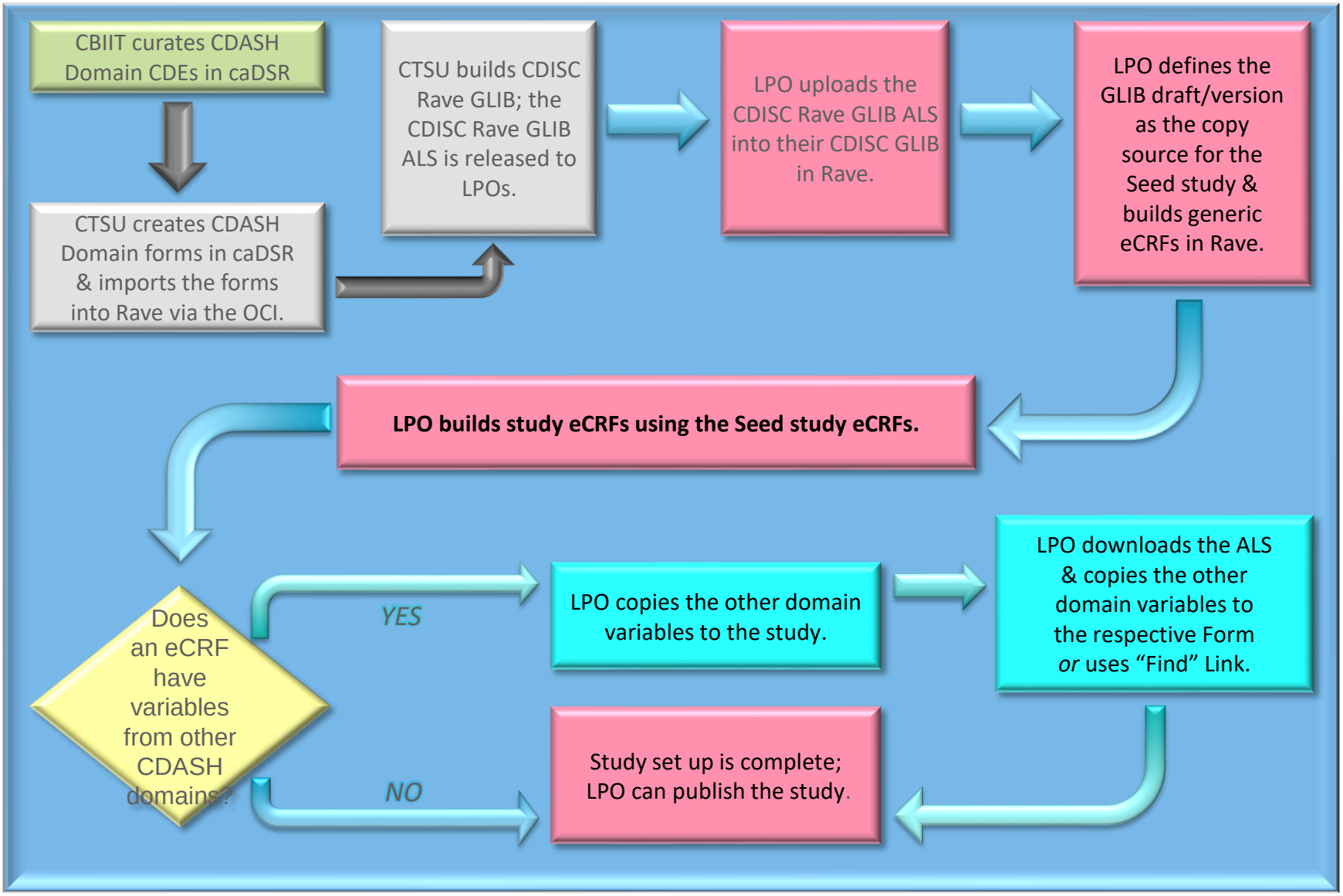
This example shows records for a trial that had two versions of inclusion/exclusion criteria.
Rows 1-3 show the two inclusion criteria and one exclusion criterion for version 1 of the protocol.
Rows 4-6 show the inclusion/exclusion criteria for version 2.2 of the protocol, which changed the minimum age entry from 21 to 18.

Row	STUDYID	DOMAIN	IETESTCD	IETEST	IECAT	TIVERS
1	XYZ	TI	INCL01	Has disease under study	INCLUSION	1
2	XYZ	TI	INCL02	Age 21 or greater	INCLUSION	1
3	XYZ	TI	EXCL01	Pregnant or lactating	EXCLUSION	1
4	XYZ	TI	INCL01	Has disease under study	INCLUSION	2.2
5	XYZ	TI	INCL02A	Age 18 or greater	INCLUSION	2.2
6	XYZ	TI	EXCL01	Pregnant or lactating	EXCLUSION	2.2

2. EC CDEs – Supplemental Qualifiers Domain Questions

- NCI CBIIT team will curate CDEs for the LPOs requesting assistance please submit request(s) to the NCI CDISC Mailbox
- Will curate only EC Forms where questions are not already curated CDASH or Inclusion/Exclusion
- For curation we will need CDISC details – LPOs provide map to SDTM:
  - Variable label
  - Variable name
  - Data format
  - List of values (if enumerated)

CDISC Implementation Workflow



CDISC Rave Global Library: CTSU Change Matrix

Rave Attribute	caDSR/ OCI Attribute	CTSU CDISC Global library	Change Management
Form Name	caDSR Form Long Name *(e.g., DEMOGRAPHICS)*	CDASH Domain Name *(e.g., DEMOGRAPHICS)*	*<no change: same as caDSR>*
Form OID	caDSR Form Long Name *(e.g., DEMOGRAPHICS)*	2 letter CDASH Domain Name *(e.g., DM)*	**Update by CTSU**
Variable OID	Short Name of the CDE *(e.g., RACE)*	CDASH/SDTM Variable Name *(e.g., RACE)*	*<no change: same as caDSR>*
Field Name	caDSR CDE Long Name + PID + MajorV + MinorV *(e.g., Race PID6343384_V1_0)*	Variable Label + PID + MajorV + MinorV *(e.g., Race PID6343384_V1_0)*	*<no change: same as caDSR>*
Field OID	CDASH variable *(e.g., RACE)*	CDASH/SDTM Variable Name (e.g., RACE)	*<no change: same as caDSR>*
Field Label	Question Text *(e.g., Which of the following five racial designations best describes you? (More than one choice is acceptable.))*	Question Text or Variable label *(e.g., Which of the following five racial designations best describes you? (More than one choice is acceptable.))*	**Update by CTSU** ***Specify the Variable label for Field Labels without question text ***
Data Dictionary Name	Defined by CBIIT *(e.g., CDISC_SDTM_RACE_PID6343345_V1_0F)*	CDISC Codelist Submission value	**Update by CTSU** ***Specify the dictionary name as per CDISC submission value***
Data Dictionary Values- User Data String	Permissible Value Meaning (PVM) *(e.g., Native Hawaiian or Other Pacific Islander)*	NCI preferred term *(e.g., Native Hawaiian or Other Pacific Islander)*	*<no change: same as caDSR>*
Data Dictionary Values- Coded Data String	Permissible Value (PV) *(e.g., NATIVE HAWAIIAN OR OTHER PACIFIC ISLANDER)*	Submission Value *(e.g., NATIVE HAWAIIAN OR OTHER PACIFIC ISLANDER)*	*<no change: same as caDSR>*
Format- Date	char - $11	dd MMM yyyy	**Update by CTSU**
Format- Time	char - $10	24 hr HH:nn:ss	**Update by CTSU**
Format- Char (W/ Dictionary)	char *(may be either the max length of dictionary value or extendable value;; e.g., $100)*	char *(may be either the max length of dictionary value or extendable value; e.g., $100)*	*<no change: same as caDSR>*
Format- Char (W/O Dictionary)	char -$200	char -$200	*<no change: same as caDSR>*
Format- Numeric	num	num	*<no change: same as caDSR>*
Control Type	*(e.g., DropDownList)*	DATE/TIME update	**Update by CTSU**
SAS Label	*n/a*	Variable Label from SDTM; if no SDTM variable, use CDASH variable label	**Update by CTSU**
Auto-Query for Required data entry	*n/a*	SDTM -Required & Expected Variables CDASH - HR and R/C	**Update by CTSU**
Auto-Query for future Date	*n/a*	Set flag in the CDISC Rave GLIB ALS	**Update by CTSU** ***Applicable to all dates***

eCRFs Build Scenarios *(1)*

Rave eCRFs may have different build scenarios:

1) Variables from a **single domain**
2) Variables from **multiple domains**
3) Custom variables that do not map to CDASH/SDTM

1) Single Domain: All variables in an eCRF are from a single domain.

Example: Concomitant Medication (CM) domain

- Set the CDISC Global library as the copy source for the study.
- Identify the Domain variables needed for the CRF.
- Use copy wizard to select and copy the variables from the domain identified for eCRF build.
- Multiple eCRFs can be built using the same domain by changing the FORM OID during the copy process.

eCRFs Build Scenarios *(2)*

2) Multiple Domains: Variables in an eCRF are from more than one domain.

Example: an Exposure form with variables from the EX and AE domains

- Follow the steps specified in the *"eCRFs Build Scenarios: Single Domain"* slide to build an Exposure form with variables from the EX domain.
- Using the copy wizard, copy the required variables from the AE domain to the study draft.
- Add AE domain variables to an Exposure form:

Option 1

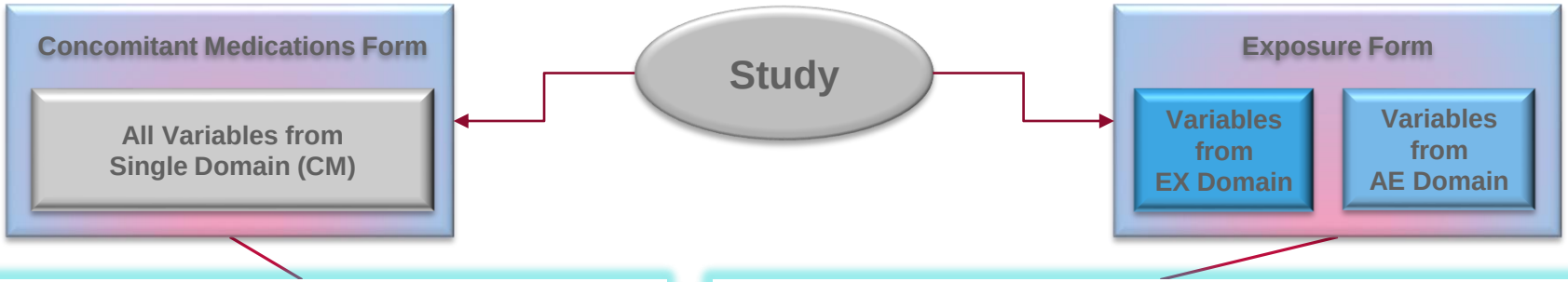
- Open the Exposure form; click add a new variable.
- Use the "Find" link on the variable definition screen to select the AE variable; click Apply to copy.

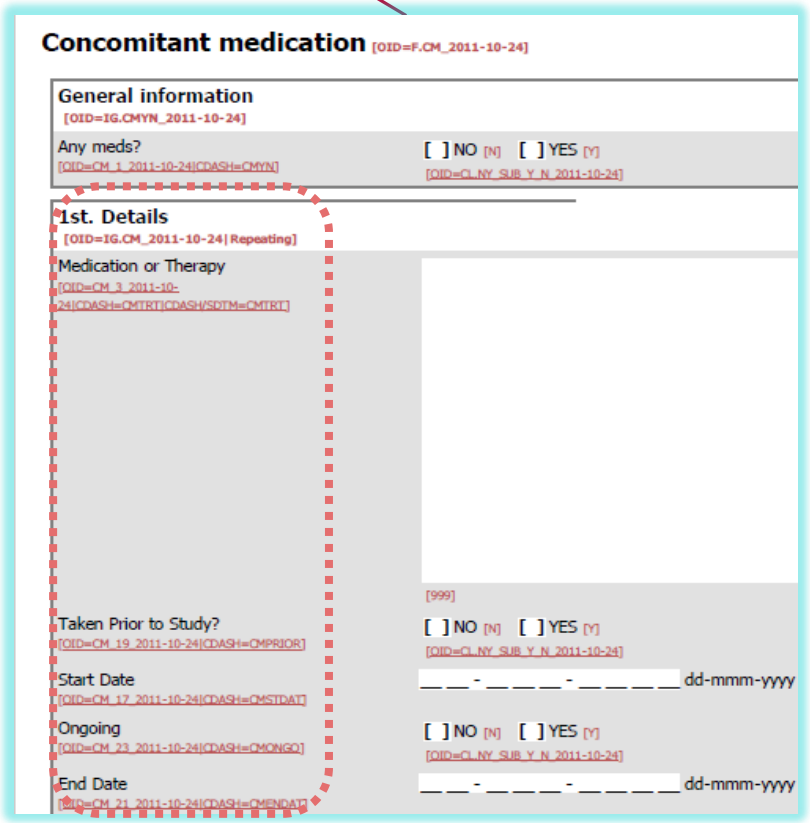
Note: the "Find" link brings only the Variable OID, Format and Data/Unit dictionary, not the associated Field attributes.

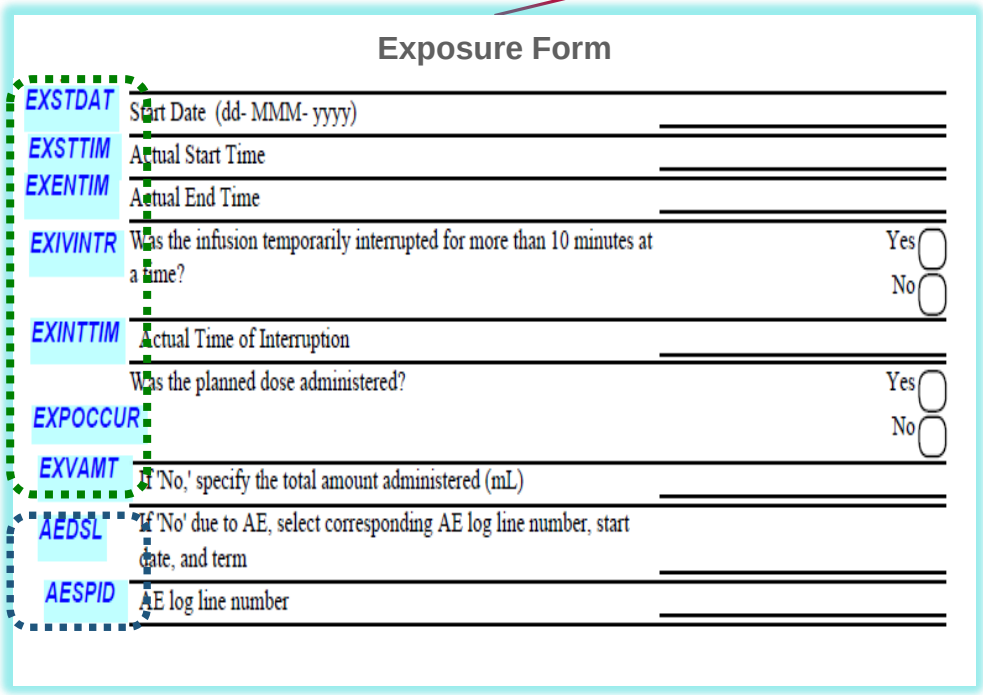
Option 2

- Download the ALS for the selected domains.
- Copy and paste the AE domain variables to the Exposure Form.
- Upload the ALS into Rave.

eCRFs Build Scenarios *(3-Examples)*







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