

CDISC Implementation Committee

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Agenda

- NCI Requirement
- Project Updates
- LPO
 - Status Updates
 - Support Requests
- Supplemental Question Workflow
- 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**

- CDISC Webinars and slides have been posted
- LPO FAQs have been updated
- CDISC LPO Impact Analysis Feedback and Support Request responses will be posted to the CDISC Wiki and the CTSU Collaboration Portal by July 1st

Project Updates (2)

■ LPO Support Webinars/Sessions

- July 9th, 2019 **CDISC SME Session**
 - Session: 2:00-4:00pm ET
 - Pre-planned agendas based on requests the week before
- July 23rd, 2019 **CDISC SME Session**
 - Session: 2:00-4:00pm ET
 - Pre-planned agendas based on requests the week before
- August 12th, 2019 **CDISC SME Session**
 - Session: 2:00-4:00pm ET
 - Pre-planned agendas based on requests the week before

LPO Status Updates

LPO	LPO 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> NCI Update: Addressed comments, formal response pending.	<Submitted> NCI Update: review/ response completed.
Theradex	Planning phase Waiting for tools; DMU is ready for CDISC implementation	<Submitted> NCI Update: Complete (Pre-Text will be in July Release)	<Submitted> NCI Update: review/ response completed.
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 GLIB	<Submitted> NCI Update: Pending initial review *Confirm if comments affect CDISC Implementations	<Submitted> NCI Update: items CTSU Integration-related, no CDISC action required; CTSU responded directly to NRG.
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	<Submitted> NCI Update: under review. A few items are CTSU Integration-related, no CDISC action required; CTSU will respond directly to EA.
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> NCI Update: Pending additional follow-up but no impact on delivery	<Submitted> NCI Update: Review completed, no action items, no impact to ALS.
COG	Initial planning phase	Not far enough to go through the GLIB	

LPO Support Questions (1)

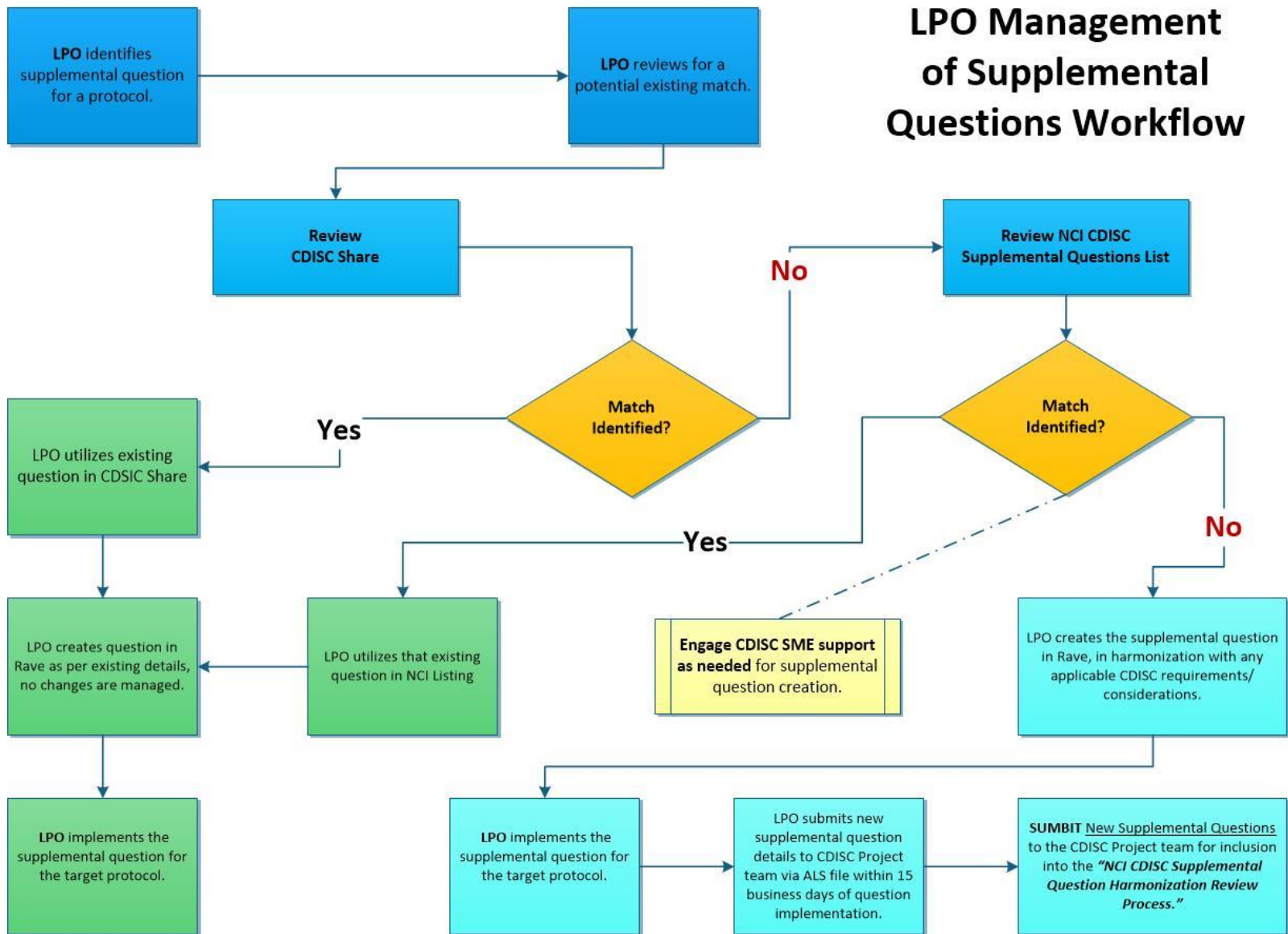
LPO Question	Response	LPO Feedback
<p>We are realizing that the CDASH standards clash in some ways with our previous NCI or NRDS standards as well as some required CDUS reporting questions that we've been mandated to use. Are we going to receive any guidance about how to resolve these? Or are we supposed to disregard the old mandates in favor of CDASH? We seem to be in a situation where we have two mandates that are in conflict and we're not sure what we're supposed to be following.</p>	<p>NCI: CDASH/CDISC overrides all standards.</p> <p>Applying CDASH standards to CDUS, CDUS will not be applicable to new studies, we will have DMU by then so we can accept it.</p>	<p>Groups would like to kick off the harmonization effort sooner than later.</p> <p>DMU Mapping?</p>
<p>I've been thinking more about the discussion from the last SME call to add the domain code to the beginning of all variables in the GLIB ALS. I'm still struggling with why this is something that is beneficial or helpful for our programmers.</p>	<p>CDISC SME: The first two characters of most SDTMIG variables is the domain code, but not all variables follow that convention:</p> <ul style="list-style-type: none"> • Most DM variables do not have the domain code prefix • Associated Persons variables do not have the AP-- domain code as a variable prefix • Supplemental Qualifier variables are normalized and do not have the domain-specific code prefix • Other study-level timing and identifier variables (e.g., SPDEVID) do not have domain codes <p>For the above commonly used fields/variables, a specific mapping would have to be created for the SDTM programmer to know where they are intended to go in the SDTM domains.</p> <p>Consistent, pattern-based variables is only one example of how to build efficiency into the process, it is primarily for the benefit of the SDTM programmers (to be able to write standard pattern-based programming</p>	

LPO Support Questions (2)

LPO Question	Response	LPO Feedback
<p>We do have a question on the QRS supplements. The CBIIT team provided support to curate validated tool in caDSR. Will there be any kind of support to create validated tools in CDASH?</p>	<p>NCI: The NCI is reviewing this item and working to find a process to curate these, especially those that are validated tools.</p>	

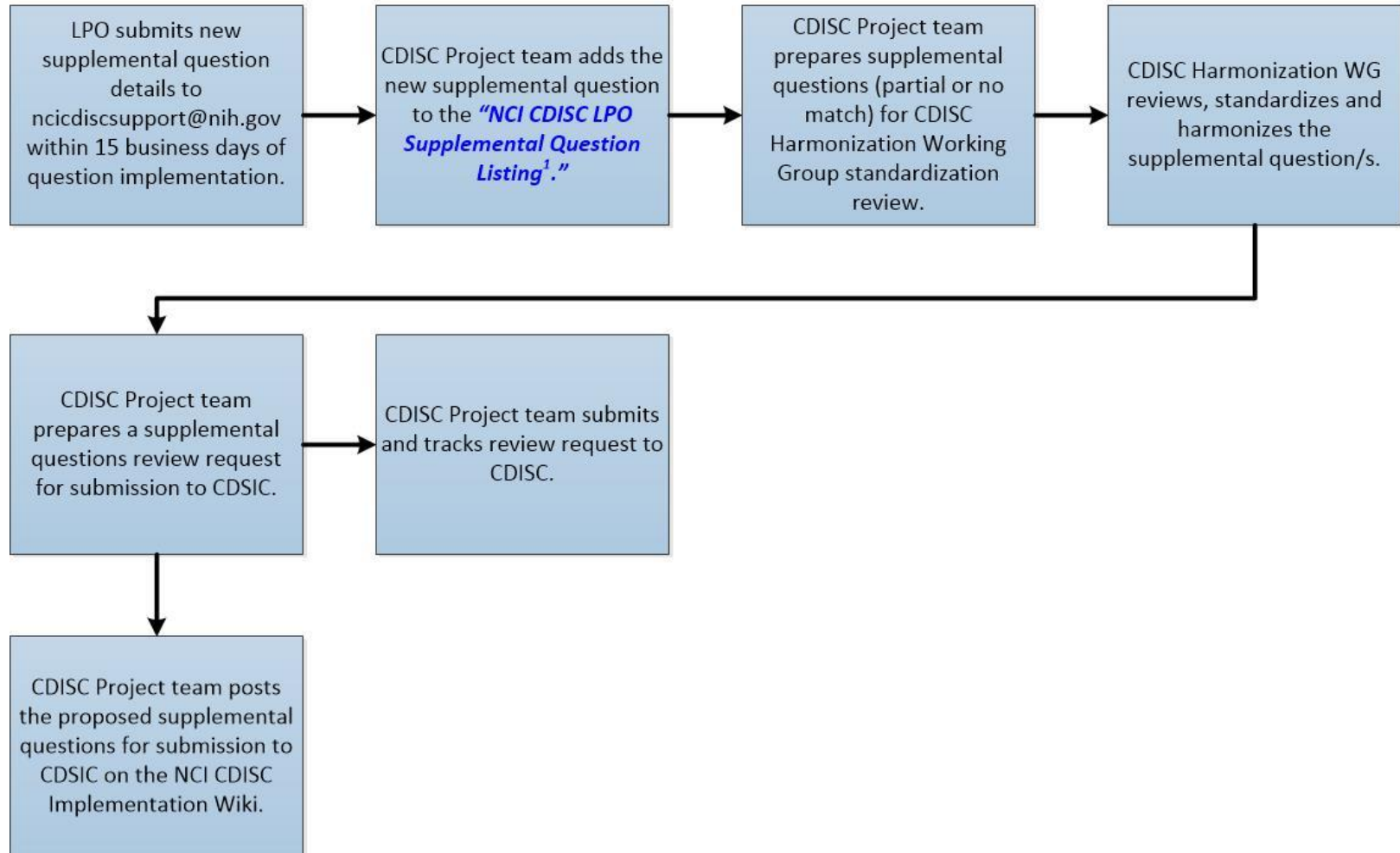
Supplemental Question Workflow (1)

LPO Management of Supplemental Questions Workflow



Supplemental Question Workflow (2)

NCI CDISC Supplemental Question Harmonization Review



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	July 2nd, 2019
6) CTSU send CDISC Rave GLIB to CDISC SME	July 8, 2019
7) Complete Internal Compliance review on Final ALS	July 15, 2019
8) Release Final CDISC Rave GLIB ALS to LPOs	July 31, 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**	June 7 th , 2019
▶ 5) Manage Impact Analysis updates	July 2nd, 2019
6) CTSU send CDISC CTSU Standard Forms ALS v7.0 to CDISC SME	July 8, 2019
6) Complete Internal Compliance review on Final ALS	July 15, 2019
7) Release Final CDISC CTSU Standard Forms ALS v7.0 to LPOs	July 31, 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 July 24th, 2019

Appendix

CDISC Harmonization Workflow
NCI CBIIT: CDEs for EC Forms Overview
CDISC Implementation Workflow
CTSU Change Matrix
eCRF Build Scenarios
LPO I.A.: Beta Release GLIB

CDISC Harmonization Workflow v06/25/2019

- 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 and submits the supplemental questions (partial or no match) to the CDISC Harmonization Working Group (WG).
- 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.

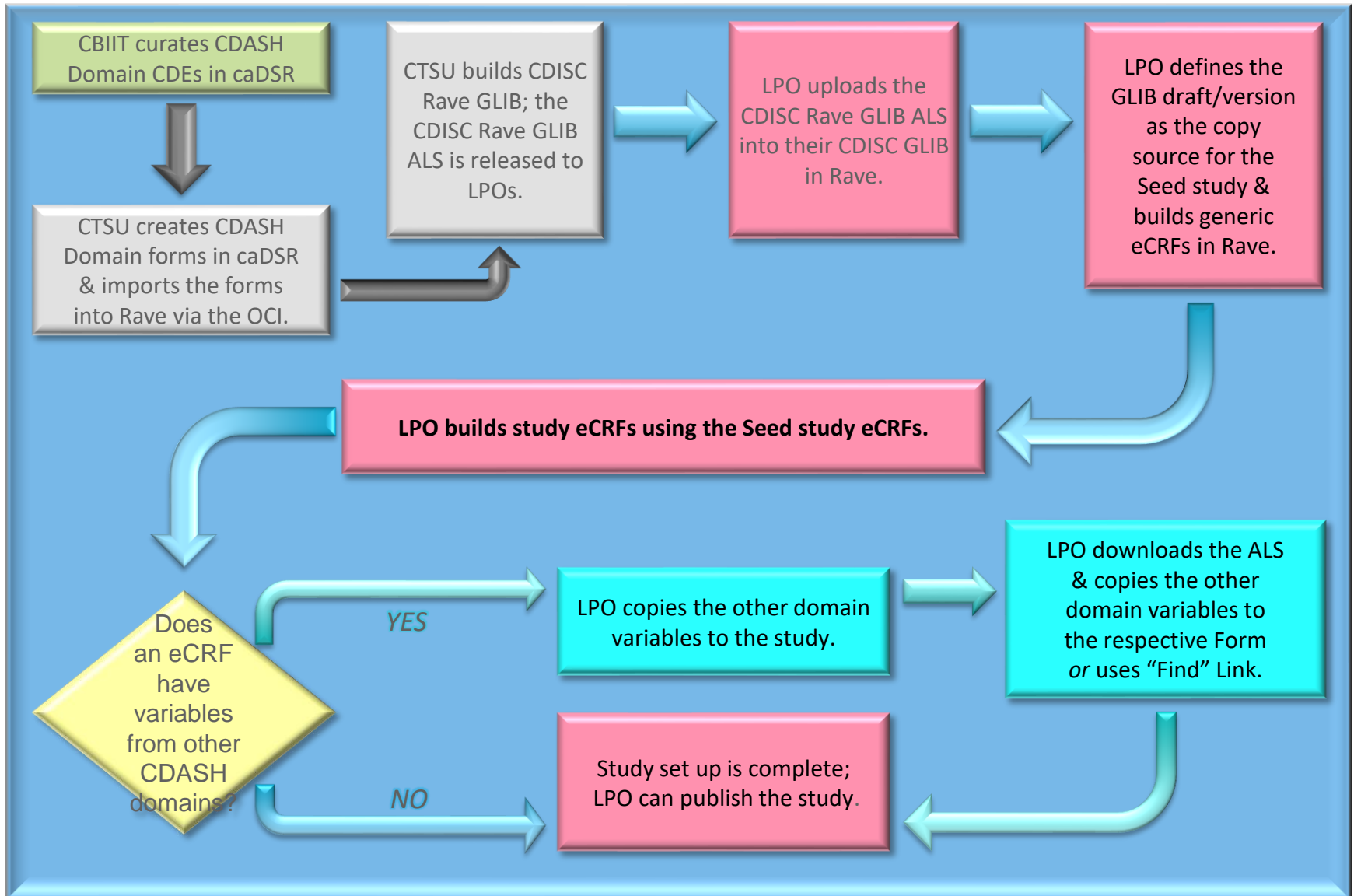
Rows 4-6 show the inclusion/exclusion criteria for version 2.2 of the protocol, which changed the minimum age entry from 21 to 18.

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 <i>(e.g., DEMOGRAPHICS)</i>	CDASH Domain Name <i>(e.g., DEMOGRAPHICS)</i>	<no change: same as caDSR>
Form OID	caDSR Form Long Name <i>(e.g., DEMOGRAPHICS)</i>	2 letter CDASH Domain Name <i>(e.g., DM)</i>	Update by CTSU
Variable OID	Short Name of the CDE <i>(e.g., RACE)</i>	CDASH/SDTM Variable Name <i>(e.g., RACE)</i>	<no change: same as caDSR>
Field Name	caDSR CDE Long Name + PID + MajorV + MinorV <i>(e.g., Race PID6343384_V1_0)</i>	Variable Label + PID + MajorV + MinorV <i>(e.g., Race PID6343384_V1_0)</i>	<no change: same as caDSR>
Field OID	2 letter domain prefix + CDASH/ SDTM variable <i>(e.g., RACE)</i>	2 letter domain prefix + CDASH/SDTM Variable Name <i>(e.g., RACE)</i>	<no change: same as caDSR>
Field Label	Question Text <i>(e.g., Which of the following five racial designations best describes you? (More than one choice is acceptable.))</i>	Question Text or Variable label <i>(e.g., Which of the following five racial designations best describes you? (More than one choice is acceptable.))</i>	Update by CTSU **Specify the Variable label for Field Labels without question text **
Data Dictionary Name	caDSR Value Domain Long Name + PID + MajorV <i>(e.g., CDISC_SDTM_RACE_PID6343345_V1_0F)</i>	caDSR Value Domain Long Name + PID + MajorV	<no change: same as caDSR>
Data Dictionary Values-User Data String	Permissible Value Meaning (PVM) <i>(e.g., Native Hawaiian or Other Pacific Islander)</i>	NCI preferred term <i>(e.g., Native Hawaiian or Other Pacific Islander)</i>	<no change: same as caDSR>
Data Dictionary Values-Coded Data String	Permissible Value (PV) <i>(e.g., NATIVE HAWAIIAN OR OTHER PACIFIC ISLANDER)</i>	Submission Value <i>(e.g., NATIVE HAWAIIAN OR OTHER PACIFIC ISLANDER)</i>	<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 <i>(may be either the max length of dictionary value or extendable value;; e.g., \$100)</i>	char <i>(may be either the max length of dictionary value or extendable value; e.g., \$100)</i>	<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	<i>(e.g., DropDownList)</i>	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

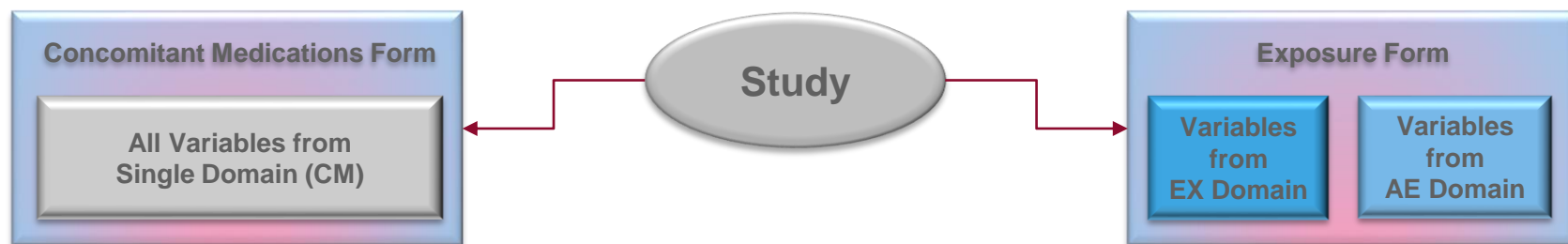
- Open the Exposure form; click add a new variable.
- Use the “**Find**” link on the variable definition screen to select the AE variable; click Apply to copy.

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)



Concomitant medication [OID=F.CM_2011-10-24]

General information
 [OID=IG.CMYN_2011-10-24]
 Any meds? NO [N] YES [Y]
[OID=CM_1_2011-10-24|CDASH=CMYN] [OID=CL.NY.SUB.Y.N_2011-10-24]

1st. Details
 [OID=IG.CM_2011-10-24|Repeating]
Medication or Therapy
[OID=CM_3_2011-10-24|CDASH=CMTRT|CDASH/SDTM=CMTRT]

Taken Prior to Study? NO [N] YES [Y]
[OID=CM_19_2011-10-24|CDASH=CMPRIOR] [OID=CL.NY.SUB.Y.N_2011-10-24]

Start Date _____ dd-mmm-yyyy
[OID=CM_17_2011-10-24|CDASH=CMSTDAT]

Ongoing NO [N] YES [Y]
[OID=CM_23_2011-10-24|CDASH=CMONGO] [OID=CL.NY.SUB.Y.N_2011-10-24]

End Date _____ dd-mmm-yyyy
[OID=CM_21_2011-10-24|CDASH=CMENDAT]

Exposure Form

EXSTDAT Start Date (dd- MMM- yyyy) _____

EXSTTIM Actual Start Time _____

EXENTIM Actual End Time _____

EXIVINTR Was the infusion temporarily interrupted for more than 10 minutes at a time? Yes No

EXINTTIM Actual Time of Interruption _____

Was the planned dose administered? Yes No

EXPOCCUR _____

EXVAMT If 'No,' specify the total amount administered (mL) _____

AEDSL If 'No' due to AE, select corresponding AE log line number, start date, and term _____

AESPID AE log line number _____

LPO Impact Analysis: Beta Release GLIB (1)

#	Document	LPO Findings/ Recommendations/ Questions	Decision
1	GLIB ALS	<p>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</p> <p>With the final release in July, will the GLIB and other ALS files be updated using the newest CDASH and SDTM controlled terminology packages?</p>	<p>CTSUS will do an impact analysis (for CT Packages released by CDISC before July 2019) to understand the potential impact to our timeline.</p> <p>The intent is to provide documents as current as possible to CDISC releases/ updates; this must be done in consideration of the implementation timelines.</p>
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.	<p>05/02/2019 CDISC Integration FG:</p> <ul style="list-style-type: none"> 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. <p>“Standard Forms” applies to the CTSU Standard Forms ALS, “Non-Standard Forms applies to variables not specified in the CTSU Standard Forms ALS.</p>
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?	<p>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”.</p> <p>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.</p>

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.	<p>LPOs can decide to use the NCI Preferred Term or the CDISC Synonym for the User Data String in Rave.</p> <p>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.</p> <p>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.</p>
6	GLIB ALS	<p>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.</p> <p>i. Rave does not allow us to subset the dictionary choices at the field level – a separate dictionary must be created</p> <p>ii. Example: NY. Sometimes Unknown or Not Applicable are not valid responses for a question</p> <p>iii. The Alliance would be open to a standard naming convention in this situation.</p>	<p>If authoring a form directly in Rave: LPOs should use the vetted CSC Data Elements Work Group (DEWG) naming convention (caDSR Value Domain Long Name + PID + MajorV + MinorV + _0F) for dictionary names and append an integer when subsetting a dictionary.</p> <p>If building a form in the caDSR/ using the OCI to import the form into Rave: the OCI will append an integer to the dictionary name when a dictionary is reused/ subsetted.</p> <p>Dictionary Names have been reverted to 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.</p>
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	<p>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</p> <p>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?)</p>	<p>All CTEP IND studies activated on/after 1/1/2020 must be CDISC compliant.</p> <p>CDISC Compliance is highly recommended for studies that activate prior to the 2020 date.</p> <p>The CDISC SME to provide additional feedback at the "Overview of FDA Submission Process" Webinar on May 22nd, 2019.</p>
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	<p>Throughout the entire document, please be specific what CDISC compliant means. CDASH, SDTM, ADaM, etc.</p> <p>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.</p>	<p>The FDA requirement is SDTM. the NCI requirement is CDASH.</p> <p>CDASH compliance makes SDTM compliance easier to achieve.</p> <p>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.</p>



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