

SDTM CRT Packages and the Submission Space

Created by Joshua Glaser and Charlotte Prior-Chappell

Abstract

The main goal of the Submission Space and generating Case Report Tabulation (CRT) Packages is to ensure “Submission Readiness” of a study by assessing the Datasets, Define and Validation Reports. For all current and ongoing studies, the aim should be for a high-quality package to be produced at a study level, with little to no changes required for the CRT Package.

This poster aims to outline what it means for the SDTM side of a CRT package to be “Submission Ready” and how this is achieved, whether we are starting from SDTMs created from the most recent Implementation Guide or from Datasets not in the format of SDTMs yet.

Determining “Submission Readiness”

Completeness

- All expected files complete and ready to send - the package should include the following:
 - SDTM Datasets
 - Annotated Case Report Form (aCRF)
 - SDTM Define.xml
 - Clinical Study Data Reviewer's Guide (cSDRG)

Quality and Consistency

- All Quality Checks complete per company and client standards
- Identify and check consistency if multiple studies are required under a single compound's submission

Validation

- Pinnacle 21 is a key resource in checking Submission Readiness
- No unexpected issues in Pinnacle 21
- All expected issues have valid explanations

Poster Focus

A CRT Package is submitted with a New Drug Application (NDA) to a Regulatory Agency in charge of Food and Medicine, most of the time being the FDA and PMDA. The complete package will consist of both the SDTMs and ADaMs, but we are going to focus on the SDTMs and how to get them “Submission Ready”.

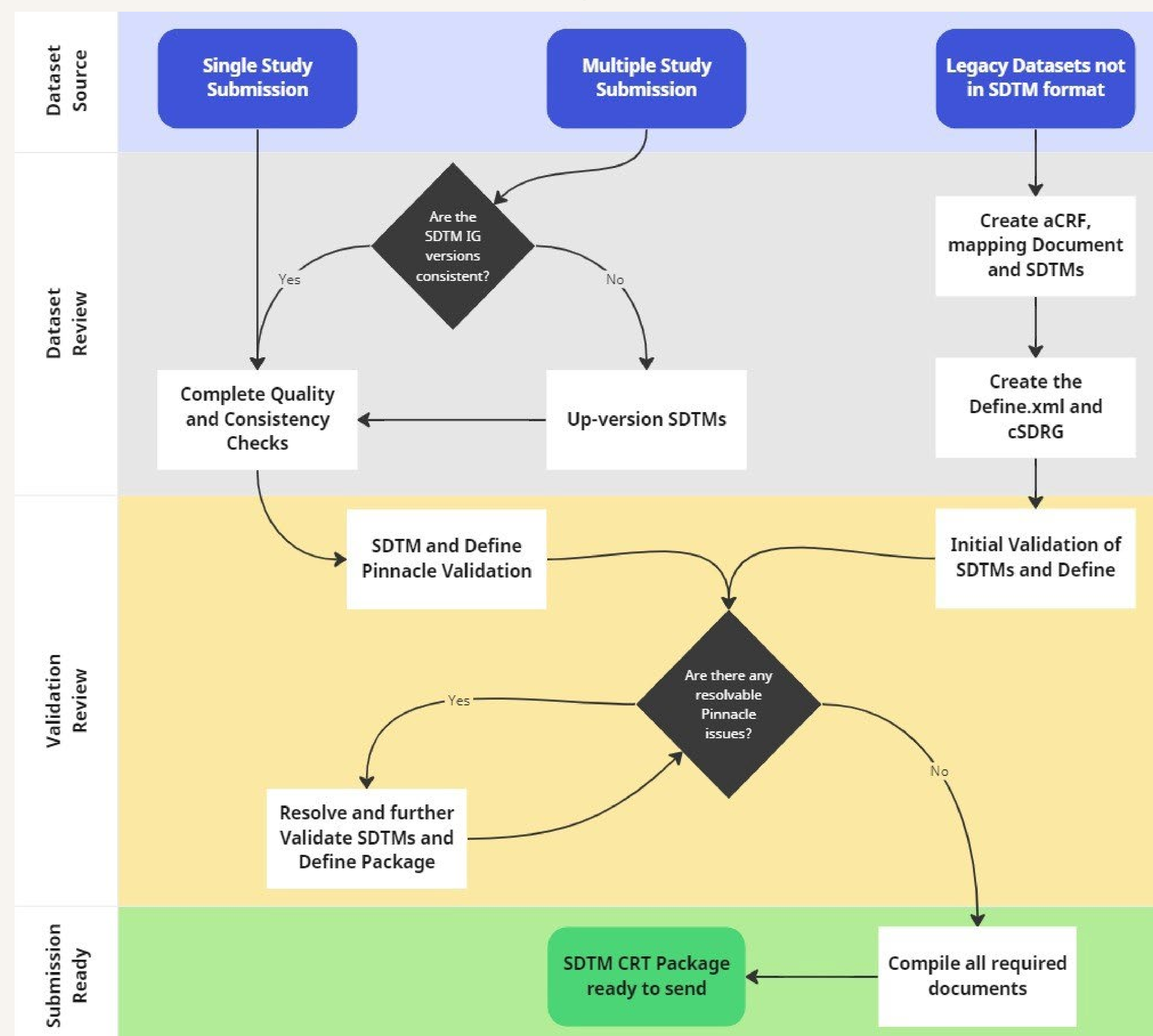
Spaces and Levels

When working in the Submission Space, it is key to understand the difference between the two below spaces/levels:

- **Study Level:** the space in which the programming package is initial created/finalised, during which the study was ongoing/active.
- **Submission Level:** the space in which the Submission CRT Package is updated/finalised.

If a study is ongoing/active and you are due to create the CRT Package, Submission reviews will be complete at the Study Level.

Consolidating the Process



Single Study Submission

The simplest situation where the Submission Package contains a single study, already in SDTM format. Once all Quality, Consistency and Validation checks are complete, this is good to go!

Study State

The Study that we create the SDTM CRT package can be in one of two states:

- Ongoing/Active (we produce a CRT at Study Level)
- A study that is finished and SDTMs Finalised (we produce a CRT at Submission Level)

Depending on which state the study is in will impact how we approach producing a “Submission Ready” Package and which space we produce it in.

Multiple Study Submission

It is possible for multiple studies investigating the same compound to be submitted together. In this case, each study will have their own CRT Package, but will have a few unique interactions to consider:

- We aim for consistency across studies where possible, one example being Trial Summary information like TRT and PCLAS being consistent
- The same SDTM Implementation Guide version should be used across all studies, this may lead to Up-versioning of some Studies to match the highest version study in the set

Legacy Study Submission

What happens if the datasets required for a submission are not contained in SDTMs? We would need to create the SDTMs ourselves, performing a Legacy conversion on the “Legacy” datasets.

The key aspect to a Legacy Conversion is Traceability between the “Legacy” datasets and the SDTMs through sufficient documentation – two important documents:

- A Mapping Document, clearly stating how data is translated from the “Legacy” datasets into the SDTMs
- The aCRF, identifying mapping between the original database and the newly created SDTMs