CRT for eCTD submission

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Introduction

CRTs are part of the electronic submission (US only at the moment)

From the 1\textsuperscript{st} of January 2008, CRT specifications are based on the eCTD (electronic Common Technical Document) rules.

These new rules implied a lot of changes.

How Novartis have changed processes to be compliant?

What have been encountered issues?
What is a CRT?

Definition

CRT = Case Report Tabulation

Required by the FDA

Package created in order to provide to the FDA the raw & derived datasets

2 CRTs for one study:

- One containing only raw datasets (if needed)
- One containing only derived datasets
What is a CRT?

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Package contains:

- SAS V5 Transport files of either the raw or derived datasets (<xxx>.xpt) Format required by FDA.
- The data definition table (Define.pdf), regrouping:
  - A table of content of the raw or derived datasets
  - Hyperlinks to the XPT files
  - An alphabetical list of variables and formats
- The CRF / eCRF for raw CRT
- the Data Derivation and handling Document (DDD) for derived CRT
What is a CRT?

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- The Data Derivation & Handling document is not required by the FDA but Novartis considers the DDD as mandatory.

- The DDD should contain details which are not self-explanatory by the variable label within the data definition table.

- It provides further detailed explanation to the reviewer on special data handling issues and other data imputation rules or convention that were used in the study.
What is a CRT?

Data Definition table

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**Annotated CRF/eCRF**

### Introduction

**What is a CRT?**

- **Definition**
- **Content**
- **Data Derivation & Handling document**
- **Data definition table**

**Annotated CRF/eCRF**

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**Name of the dataset**

**Label of the dataset**

**Name of the variable as it appears in the dataset**
What is a CRT?

Summary

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CRF

Define.pdf

CRT: Raw datasets

Raw datasets

Derived datasets

Transport files

CRF

Define.pdf

CRT: Derived datasets

Transport files

DDD
The new electronic format of CRT required additional rules to be followed:

- Naming conventions (no underscores)
- Specific folder structure (listings, analysis)

No underscores:

- Previous naming convention for a derived dataset was `a_dataset`.
- For old studies, underscores are removed at the moment of the translating in SAS version 5 transport files.

The specific folder structure implied to create two CRTs instead of one.
Novartis processes

Before: example

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Before:

Raw and derived datasets in the same define

Now:

Raw datasets
Derived datasets

Listings CRT
Analysis CRT
Novartis processes

Novartis tools

- A standard macro create the data definition table automatically.
- The macro works with a call of parameters, the update to be compliant with the new eCTD rules have added a new parameter.
- The different parameters are:
  - DVD or NODVD: to include the Data Derivation and Handling document or not.
  - RAW or NORAW: to include Listings datasets or not.
  - eCTD: to apply the new rules during the creation of the CRT.
Novartis processes

DRA and review

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- The programmer has to deliver the CRT to DRA (Drug Regulatory Affair) in Novartis.
- DRA is the primary contact with FDA. They have to transfer CRTs.
- As the CRT is the last part of the programming work, these have to be fully reviewed by both programmer and statistician.
- A final review is done by the project programmer and the global manager.
FDA have recently asked us to provide more information in the data definition table.

- Actually, because of the automation the only document we can modify is the DDD.
- The standard macro will be updated.
CRT: final step of Statistical Reporting work

CRT is one of the most important SR document but the working time to create a CRT is often underestimated…

Next step: CRT in XML format
Questions ?