**TROUBLE SHOOTINGS**

Throughout the years, we have accumulated experiences with hundreds of NDA and BLA submissions and close to a thousand clinical studies. The recurrent scenarios (e.g., study designs, data formats, and analysis methodologies) observed from these past submissions have been categorized in a knowledgebase, with the corresponding solutions built into PKView.

With the current version, there are still remaining issues that mostly associated with data contents and specific PKView requirement. The following table documents some common issues and possible solutions. We plan to develop a "Data Check" function to automatically detect the known issues during the "Data Management" step.

| Scenario | Issue/Error | Solution |
|---|---|---|
| PC:VISIT has blank records | Errors occurs during "Run Analysis" | 1. Add values to the blank records<br>2. Delete the row with blank VISIT |
| PC:PCTPT has duplicated values for the same set of USUBJID, PPCAT, VISIT. This might be caused by multiple assays performed for each sample. | Cause error during "NCA reporting" | Resolve the duplication |
| PC:PCTPT during washout have time points other than 'pre-dose' | Concentration time profiles cannot be properly deciphered, and cause error during "Run Analysis" | Remove the washout time points |
| Symbols used in PP:PPCAT or DM:ARM, such as #, * | Cause run time errors, during "Run Analysis" or "NCA reporting" | 1. Delete or replace the symbols in PP.xpt and PC.xpt<br>2. Use "EDIT" function to remove the symbols |
| Unique values in PP:VISIT and PC:VISIT does not match | Cause error during "NCA reporting" | Use "EDIT" function to revise the VISIT values |
| Missing records in PP or PC. All PERIOD specified in DM:ARM for each USUBJID | The "Study Design" and "Reference Trt/Grp" cannot be properly mapped | 1. Remove the missing PERIOD from the subjecs (for example: instead of AB use A for DM:ARMCD) |

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| must have corresponding records in PP.xpt and PC:xpt. | | 2. Add the missing records to PP and PC with values specified as  ".."<br>3. Remove the subjects with missing record from DM.xpt |
| When the study design are different among different cohorts in a study (e.g. mix of crossover, parallel, and sequential) | The "Study Design" and "Reference Trt/Grp" cannot be properly mapped | Break down the study data by cohort.  Revise only DM.xpt for each cohort, by deleting USUBJID from other cohorts).  PP.xpt and PC.xpt can remain the same as the original data. |
| For a sequential study, in DM:ARM, the alphabetically sorting order of PERIOD may not match the chronological sorting order.  This could due to the length of the PERIOD name. | When the alphabetically sorting order of PERIOD may not match the chronological sorting order. This will cause the reversal of point estimate of test/reference for a sequential study during "Statistical Reporting" and "Forest plot" | Make sure the alphabetical order match the chronological order of PERIOD in DM:ARM using the EDIT function. |
| The format (Character or Numeric) of VISIT, PPSTRESN, and PCSTRESN, PCTPT are incorrect. | The "Study Design" and "Reference Trt/Grp" cannot be properly mapped | Verify the required variable format in DM.xpt, PC.xpt, PP.xpt |
| Duplicated selection of same variable from a Domain on the Mapping page.  For examples: same variable is selected for "PP:Category" and "PP:TEST" | Cause run time error | Select different variables for different required fields. |