# Documentation of the Novartis Decision Support Tool Project

**Abstract**: This document records the steps taken to produce deliveries for the Novartis Decision Support Tool project. The objective is to make it easy to reproduce all the deliveries. The document includes the description of data, code, usage and results produced.

1. Descriptive Statistics
   1. Abstract: Two versions of descriptive statistics are delivered: the first with the IMS-proposed cut-offs for EDSS score variables while the second with the Novartis-proposed cut-offs.
   2. With IMS-proposed cut-offs
      1. Preprocess data for computing descriptive statistics
         1. Input: Raw data from Ray (path and file name needed, also need to copy the data)
         2. The code
            1. General flow: run ‘Jie’s version’ (see below) first using the raw data from Ray; then run ‘Lichao’s version’ using the results from ‘Jie’s version’. The output is ready for computing the descriptive statistics.
            2. ‘Jie’s version’

Code location: F:\Jie\MS\04\_Delivery\01\_DescriptiveStats\02 Code\01 prepare cohort for descriptive stats\createJie'sVersion

Code description: It performs initial processing of the data such as dichotomising variables and merging small categories. It can be used to produce two types of results: the first with continuous variables intact and the second with continuous variables dichotomised.

Input: raw data from Ray. The input path needs to be changed as follows: Change the input directory (i.e. line 12 of the code “F:\Jie\MS\04\_Delivery\01\_DescriptiveStats\02 Code\01 prepare cohort for descriptive stats\createJie'sVersion\Scripts\main\_createCohort.R”) into: “F:/Jie/MS/04\_Delivery/01\_DescriptiveStats/01 Data/”

How to run the code: execute script ‘./Scripts/main\_createCohort.R’ and it will produce the types of results described in 1.2.1.2.2.2.

Example results: F:\Jie\MS\04\_Delivery\01\_DescriptiveStats\03 Results\2016-07-08 02.16.04

* + - * 1. ‘Lichao’s version’

Note: There’s a bug in this code. The bug-fixed version can be found in the corresponding code in Section 1.3 below. The corresponding problem in this delivery was fixed manually.

Code location: F:\Jie\MS\04\_Delivery\01\_DescriptiveStats\02 Code\01 prepare cohort for descriptive stats\createLichao'sVersion

Code description: It performs variable-specific processing to the set of manually-selected ‘base variables’ which will be used for modelling.

Input: both types of results from ‘Jie’s version’ above. The input path needs to be changed as follows: Change the input directory (i.e. line 7 of the code “F:\Jie\MS\04\_Delivery\01\_DescriptiveStats\02 Code\01 prepare cohort for descriptive stats\createLichao'sVersion\main\_lichao\_Jul06.R”) into: "F:/Jie/MS/04\_Delivery/01\_DescriptiveStats/03 Results/2016-07-08 02.16.04/"

How to run the code: execute script ‘main\_lichao\_Jul06.R’

Example results: F:\Jie\MS\04\_Delivery\01\_DescriptiveStats\03 Results\2016-07-11 20.01.50

* + 1. Compute descriptive statistics
       1. Code location: F:\Jie\MS\04\_Delivery\01\_DescriptiveStats\02 Code\02 create descriptive stats
       2. Input: the results from ‘Lichao’s version’ above. The input paths need to be changed (Lines 17 and 19 of the pyspark code).
       3. How to run the code: execute script ‘run\_DS.sh’ (in the script the Spark code is run 5 times with different arguments, each for a cohort. )
       4. Example results: F:/Jie/MS/04\_Delivery/01\_DescriptiveStats/03 Results/DS\_20160713
    2. Delivery: Lichao to add
  1. With Novartis-proposed cut-offs
     1. The input and code are the same as 1.2 except two changes are made to ‘Lichao’s version’ in 1.2.1.2.3. (the updated code is under F:\Jie\MS\04\_Delivery\11\_DescriptiveStats\02 Code\01 prepare cohort for descriptive stats\02 createLichao'sVersion). The two changes are:
        1. A bug is fixed about dichotomising missing values in continuous variables;
        2. The cut-offs are updated using Novartis-proposed values.
     2. Example results: F:\Jie\MS\04\_Delivery\11\_DescriptiveStats\03 Results\
        1. Data for computing descriptive statistics: 2016-07-19 01.52.43 (Lichao to clarify: results from Jie’s version, 2016-07-08 02.16.04, is the same as the one used in 1.2. But here it’s copied again into the folder 11\_DescriptiveStats)
        2. Descriptive statistics: DS\_20160719
     3. Delivery: Lichao to add

1. Modelling
   1. Abstract: This includes the modelling of the 5 cohorts by 6 outcomes. It also includes different models, namely logistic-regression with the elastic-net penalty and extensive base variables, logistic-regression with the elastic-net penalty and the top 10 most important variables and the standard logistic-regression with the top 10 most important variables. The following divides the documentation into two parts: one for the composite cohort and the other for the other 4 cohorts. This is because firstly the result of the composite cohort is delivered separately prior to that of the other 4 cohorts, and secondly the code for the other 4 cohorts is slightly different for the purpose of incorporating multiple cohorts.
   2. The Composite cohort
      1. Prepare the modelling data
         1. Code location: F:\Jie\MS\04\_Delivery\02\_ExtractModelData4Cmp\02 Code\main.R
         2. Code description: The code further pre-processes the data generated for descriptive statistics to generate data ready for modelling (e.g., removing reference categories, etc.)
         3. Input: Outputs from both 1.2.1.2.2 and 1.2.1.2.3 (pre-processed data from Jie’s version and Lichao’s version, respectively). The input paths need to be changed as the following: change the directory of input file (i.e. line 99 of the code "F:\Jie\MS\04\_Delivery\02\_ExtractModelData4Cmp\02 Code\main.R") into: "F:\Jie\MS\04\_Delivery\01\_DescriptiveStats\03 Results\2016-07-11 20.01.50\", and then change the directory of the input file (i.e. line 118 of the code "F:\Jie\MS\04\_Delivery\02\_ExtractModelData4Cmp\02 Code\main.R") into: "F:\Jie\MS\04\_Delivery\01\_DescriptiveStats\03 Results\2016-07-08 02.16.04"
         4. How to run the code: execute the main.R above.
         5. Code for QC the above result:
            1. Location: F:\Jie\MS\04\_Delivery\02\_ExtractModelData4Cmp\02 Code\QcData4ModelUsingRayData.R
            2. Input: the raw data and an already-generated dataset for modelling. The input paths need to be changed as follows: Change the already-generated data directory (i.e. line 5 of the code "F:\Jie\MS\04\_Delivery\02\_ExtractModelData4Cmp\02 Code\QcData4ModelUsingRayData.R") into: "F:\Jie\MS\04\_Delivery\02\_ExtractModelData4Cmp\03 Results" and then change the raw data directory (i.e. line 7 of the code "F:\Jie\MS\04\_Delivery\02\_ExtractModelData4Cmp\02 Code \QcData4ModelUsingRayData.R") into: "F:\Jie\MS\04\_Delivery\01\_DescriptiveStats\01 Data"
            3. Usage: Execute the file ‘QcData4ModelUsingRayData.R’. Check the resultant value of bEq at Line 359 and unmatchedRows at Line 366. In order to pass the QC, bEq needs to be True AND unmatchedRows needs to be empty.
         6. Additional variable-merging: after initial modelling, some categories of the baseline EDSS score are merged because they have very similar coefficients and confidence intervals. The code used to do this is in F:\Jie\MS\04\_Delivery\12\_ExtractModelData4Cmp\02 Code\mergeCategory.R. The input to this code is the data generated using code described from 2.2.1.1 to 2.2.1.4.
         7. Example results:
            1. Using IMS-proposed EDSS cut-offs and with additional variable-merging: F:\Jie\MS\04\_Delivery\02\_ExtractModelData4Cmp\03 Results
            2. Using Novartis-proposed EDSS cut-offs and without additional variable merging: F:\Jie\MS\04\_Delivery\12\_ExtractModelData4Cmp\03 Results\2016-07-20 06.58.41
      2. Modelling
         1. Modelling is done by two pieces of code: InitModel and NonRegularizedGLM. (Lichao to do: remove unused files in the folders of these two piece of code)
         2. InitModel
            1. Code location: F:\Jie\MS\04\_Delivery\13\_InitModel\_NonRegulatizedGLM\02 Code\InitModel
            2. Code description: The code can be used to produce two types of results: the logistic regression with the elastic-net penalty and the extensive set of variables and the logistic regression with the elastic-net penalty and the top 10 most important variables.
            3. Input: data for model produced in 2.2.1. If it is for logistic regression with the elastic-net penalty and the top 10 most important variables, the result from logistic regression with the elastic-net penalty and the extensive set of variables is also needed.
            4. How to run the code:

To run the logistic regression with the elastic-net penalty and the extensive set of variables, execute “./scripts/run\_\_ BooleanPredictor.R” with the variable ‘main.arglist$bTopVarsOnly’ set to ‘FALSE’ and use the corresponding input in 2.2.2.2.3.

To run the logistic regression with the elastic-net penalty and the top 10 most important variables, execute “./scripts/run\_\_BooleanPredictor.R” with the variable ‘main.arglist$bTopVarsOnly’ set to ‘TRUE and use the corresponding input in 2.2.2.2.3.

* + - * 1. Example results: F:\Jie\MS\04\_Delivery\03\_InitModel\_NonRegulatizedGLM\03 Results

Logistic regression with the elastic-net penalty and the extensive set of variables: 2016-07-14 20.48.49 (Lichao to add: explain the results)

logistic regression with the elastic-net penalty and the top 10 most important variables: 2016-07-15 02.33.5 (Lichao to add: explain the results)

* + - 1. NonRegularizedGLM
         1. Code location: F:\Jie\MS\04\_Delivery\03\_InitModel\_NonRegulatizedGLM\02 Code\Non\_RegularizedGLM
         2. Code description: This code is used to produce the AUC of the standard logistic regression with the top 10 most important variables and the confidence intervals and p-values of those variables.
         3. Input: the output of 2.2.2.2.4.2 above (logistic-regression with the top 10 most important variables)
         4. How to run the code: execute “./scripts/main.R”
         5. Example results: 2016-07-14 22.26.06 (Lichao to add: explain the results)
    1. Collecting results
       1. Code location: F:\Jie\MS\04\_Delivery\14\_FinalTables\02 Code
       2. Code description: The code collects results from 2.2.2 above into tables.
       3. Input: results from 2.2.2.2 and 2.2.2.3 above (i.e., logistic regression with the elastic-net penalty and the extensive set of variables, logistic regression with the elastic-net penalty and the top 10 most important variables and standard logistic regression with the top 10 most important variables). The input paths need to be changed (the following is an example):
          1. change the input directory (i.e. line 4 of the code “F:\Jie\MS\04\_Delivery\14\_FinalTables\02 Code\createFinalTables.R) into: “F:\Jie\MS\04\_Delivery\04\_FinalTables\01 Data\2016-07-14 20.48.49” (logistic regression with the elastic-net penalty and the extensive set of variables),
          2. change the input directory (i.e. line 7 of the code “F:\Jie\MS\04\_Delivery\14\_FinalTables\02 Code\createFinalTables.R” ) into: “F:\Jie\MS\04\_Delivery\04\_FinalTables\01 Data\2016-07-15 02.33.53” (logistic regression with the elastic-net penalty and the top 10 most important variables )
          3. change the input directory(i.e. line 9 of the code “F:\Jie\MS\04\_Delivery\14\_FinalTables\02 Code\createFinalTables.R”) into: “F:\Jie\MS\04\_Delivery\04\_FinalTables\01 Data\2016-07-14 22.26.06” (standard logistic regression with the top 10 most important variables)
       4. How to run the code: execute createFinalTables.R.
       5. Example results:
          1. With IMS-proposed EDSS cut-offs: F:\Jie\MS\04\_Delivery\04\_FinalTables\03 Results\2016-07-14 05.33.59
          2. With Novartis-proposed EDSS cut-offs: F:\Jie\MS\04\_Delivery\14\_FinalTables\03 Results\2016-07-27 06.10.28
    2. Delivery: (Lichao to add) (Also note that the tables are not in the same order / indices as those in 2.2.3 above)
  1. All other cohorts
     1. Code for all other cohorts is similar to that in 2.2.1, 2.2.2 and 2.2.3 for the composite cohort, with small changes to adapt to produce results for multiple cohorts. Note that modelling results for the other cohorts are only from using the Novartis-proposed EDSS cut-offs.
     2. Prepare the modelling data
        1. Code location: F:\Jie\MS\04\_Delivery\22\_ExtractModelData4Cmp\02 Code\scripts\GenDataFromRaw.R
        2. Code description: Prepare modelling data for the other 4 cohorts using those for computing descriptive statistics
        3. Input: Raw data from Ray and information about which records are to be randomly selected if they are from the same patients in the same cohort (an example dataset containing this information is “F:\Jie\MS\04\_Delivery\11\_DescriptiveStats\03 Results\2016-07-19 01.52.43\”). The input paths could be changed in Lines 5 and 35 for the raw dataset and the record-selection information, respectively.
        4. How to run the code: execute the script.
        5. Code for QC:
           1. Code location: Under F:\Jie\MS\04\_Delivery\22\_ExtractModelData4Cmp\02 Code\scripts , ./scripts/main.R and QC4ModelDataBasedOnGenDataFromRaw.R are for QC
           2. Code description: ./scripts/main.R generates the data for model using a different method. QC4ModelDataBasedOnGenDataFromRaw.R compares results from ./scripts/main.R and ./scripts/GenDataFromRaw.R mentioned in 2.3.2.1 above.
           3. Input:

./scripts/main.R: the output from Lichao’s version for descriptive statistics (the corresponding line to change in ./scripts/main.R is Line 120), the raw data from Ray (the corresponding line to change in ./scripts/main.R is Line 128) and the output from Jie’s version for descriptive statistics (the corresponding line to change in ./scripts/main.R is Line 140).

QC4ModelDataBasedOnGenDataFromRaw.R: the output from ./scripts/GenDataFromRaw.R in 2.3.2.1 above (the corresponding line to change in QC4ModelDataBasedOnGenDataFromRaw.R is Line 4) and the output from ./scripts/main.R in 2.3.2.5.3.1 above (the corresponding line to change in QC4ModelDataBasedOnGenDataFromRaw.R is Line 5)

* + - * 1. How to justify whether the QC is successfully passed: execute QC4ModelDataBasedOnGenDataFromRaw.R, and if it prints 5 ‘TRUE’ on the screen, the QC is passed successfully.
      1. Example results: F:\Jie\MS\04\_Delivery\22\_ExtractModelData4Cmp\03 Results (the two result folders are generated by 2.3.2.1 and 2.3.2.5)
    1. Modelling
       1. InitModel
          1. Code location: F:\Jie\MS\04\_Delivery\23\_InitModel\_NonRegulatizedGLM\02 Code\InitModel (Lichao to do: remove unused files)
          2. Code description: It runs the logistic-regression with the elastic-net penalty. There are 4 different situations: whether to use the extensive set of variables or the top 10 most important variables, and whether it is for the BRACE to BRACE cohort or the other three cohorts (BRACE to Firstline Oral, BRACE to Secondline and BRACE Continuation). The reason that BRACE to BRACE is separated is that there are some complete-separation variables. The following subsections will be described according to these 4 situations.
          3. Input: Similar to what is described in 2.2.2.2.3, if it is for the extensive variables, the input is the model for data obtained from 2.3.2; if it is for the top 10 most important variables, the input is the model for data from 2.3.2 and the corresponding modelling result from using the extensive set of variables. The corresponding lines to change are Lines 25, 26 and 36.
          4. How to run the code: the same as 2.2.2.2.4
          5. Example results: Under F:\Jie\MS\04\_Delivery\23\_InitModel\_NonRegulatizedGLM\03 Results

Extensive variables and BRACE to BRACE: 2016-08-11 12.00.36

Extensive variables and BRACE to Firstline Oral, BRACE to Secondline and BRACE Continuation: 2016-08-08 08.19.05

Top 10 variables and BRACE to BRACE: 2016-08-11 12.09.42

Top 10 variables and BRACE to Firstline Oral, BRACE to Secondline and BRACE Continuation: 2016-08-08 09.24.44

* + - 1. NonRegularizedGLM
         1. Code location: F:\Jie\MS\04\_Delivery\03\_InitModel\_NonRegulatizedGLM\02 Code\Non\_RegularizedGLM\
         2. Code description: see 2.2.2.3.2.
         3. Input: similar to 2.2.2.3.3, the output of logistic regression with the elastic-net penalty and the top 10 most important variables described in 2.3.3.1 above.
         4. How to run the code: similar to 2.2.2.3.4.
         5. Example results:

BRACE to BRACE: 2016-08-11 12.14.54

BRACE to Firstline Oral, BRACE to Secondline and BRACE Continuation: 2016-08-11 06.51.00.

* + 1. Collecting results: Everything is similar to 2.2.3 but with slightly changed code (e.g., changing the table names, etc.).
    2. Delivery (Lichao to add)

1. Exploring New Variables (Not Delivered)
2. Raw data and dictionary (Lichao to add)
3. The latest variable lookup table (Lichao to add)