## Coding assessment for the Van Allen Laboratory

Complete the following programming assignment using this dataset. If you are familiar with Python or R, please use one of these languages; otherwise, use a language of your choice. This exercise should not take more than a couple of hours. Please send your source code with your conclusion to Jihye Park (jpark@broadinstitute.org).

In this exercise you will explore cancer genomic data (tumor/normal whole exome sequencing) from 50 patients that received the same type of treatment, half of whom responded.

Overall question: Can you discover any mutations that are associated with treatment response?

- Download the dataset linked to above and load the <u>Mutation Annotation Format (MAF)</u> files found in the data/mafs/ folder. Each of these 50 files contains the genomic mutations observed in a different patient's tumor, obtained by biopsy and sequenced with <u>whole-exome sequencing</u>. Each row in a MAF file corresponds to a different mutation.
- 2. Subset for mutations that are not of the Variant Classification "Silent". For the purposes of this analysis, we will restrict ourselves to substitutions which result in changes to the produced protein ("nonsynonymous mutations").
- **3. Find the 10 most common mutations.** Genes names are included in the column Hugo\_Symbol and protein changes are stored in the column Protein\_Change.
- 4. Perform a statistical test to explore if any mutated genes are enriched in patients who either responded or not. Response labels for individual patients are found in the file data/sample-information.tsv.
- 5. Create a scatter plot of genes with the number of mutated patients on the x-axis and your results from question 4 on the y-axis. Can the axes be scaled or transformed in any way to improve readability? If so, recreate the plot using your suggestion(s).
- 6. How many samples are wild-type versus mutant with respect to the most significantly enriched gene from Question 4? Plot the number of nonsynonymous mutations per megabase in the mutant vs. wild-type samples. Is there a significant difference in the number of mutations between the two groups? Information on the number of nonsynonymous mutations per megabase for each patient can be found in the file data/sample-information.tsv.
- 7. Write conclusions have made based on your analysis. How might this analysis be improved or expanded upon? Please include all requested figures in your report.