

Genetic Risk Scores for Type 2 Diabetes Using Python

1 Background:

Type 2 Diabetes (T2D) is a metabolic disorder with a significant genetic component. This project focuses on calculating genetic risk scores (GRS) based on identified SNPs associated with T2D from GWAS studies.

2 Objective:

To write a Python program to calculate GRS for T2D for 1000 individuals, utilizing provided genotypic data and GWAS information. This includes data handling, computation of risk scores, visualization, and identification of high-risk individuals.

3 Genetic Risk Score:

The formula for calculating the Genetic Risk Score (GRS):

$$GRS = \sum (\log(\text{Odds Ratio}) \times \text{Allele Count})$$

In this formula:

- \sum denotes the summation across all SNPs.
- $\log(\text{Odds Ratio})$ is the natural logarithm of the odds ratio associated with each SNP's risk allele.
- Allele Count is the number of risk alleles (0, 1, or 2) present in an individual's genotype for each SNP.

4 Provided Data:

1. GWAS Information for T2D:

rsID	Risk Allele	Odds Ratio
rs7903146	T	1.4
rs1801282	G	1.3
rs5219	A	1.2
rs4402960	G	1.5
rs13266634	C	1.6

2. Genotypic Data for 1000 Individuals:

The [genotypes.tsv](#) table of 1000 individuals, where each row represents an individual, and each column represents a SNP.

Here are the top five rows of the genotypic data table:

individual_id	rs13266634	rs1801282	rs4402960	rs5219	rs7903146
1	CA	AA	AA	TT	AA
2	CA	GA	AA	TT	AA
3	CC	AA	AA	TT	AA
4	AA	GG	AA	TT	TT
5	CA	AA	AA	AT	AA

3. Example Calculation for First Individual:

For individual_id 1, suppose the genotypes are:

- rs13266634: CA
- rs1801282: AA
- rs4402960: AA
- rs5219: TT
- rs7903146: AA

The GRS is calculated as follows:

- rs13266634 (C, 1.6): 1 risk allele, score contribution = $\log(1.6)$
- rs1801282 (G, 1.3): 0 risk alleles, score contribution = 0
- rs4402960 (G, 1.5): 0 risk alleles, score contribution = 0
- rs5219 (A, 1.2): 0 risk alleles, score contribution = 0
- rs7903146 (T, 1.4): 0 risk alleles, score contribution = 0

Thus, the total GRS = $0 + 0 + 0 + 0 + \log(1.6) \approx 0.47$

5 Tasks:

- Data Preprocessing
- Risk Score Calculation
- Statistical Analysis and Visualization
- Identification of High-Risk Individuals

6 Deliverables:

1. Google Colab Notebook: Create a Google Colab notebook that includes all parts of the project:
 - Data Loading and Preprocessing.
 - Risk Score Calculation.
 - Statistical Analysis and Visualization. Use Python libraries like Matplotlib and Seaborn to visualize the distribution of genetic risk scores. Include histograms, density plots, or other relevant visualizations.
 - Identification of High-Risk Individuals: Write code to identify individuals within the top 5% of GRS.
 - Discussion: Use markdown cells to interpret your findings, discuss potential implications for T2D risk prediction, and possible limitations of this approach.
2. Comprehensive Report within the Notebook: Alongside the code, your Colab notebook should include a detailed report. Use markdown cells to document:
 - The methodology used.
 - The results obtained from the analysis.
 - Visualizations of the data.
 - A discussion section covering the interpretation of results, implications, and limitations.
 - Any conclusions drawn from the project.
3. Shareable Link to the Notebook: Once your project is complete, ensure that the Google Colab notebook is shareable and accessible. Provide a link to the notebook as part of your submission.