# AI Colab Group 1 - Clinical Data Science & Modeling

Yara Yaghi, Naod Dawit, Kareem Aly, & Indira Kuppa

# Proposed Research Question

How does kidney function, as measured by creatinine clearance (CCR) and blood urea nitrogen (BUN), relate to β-cell dysfunction and type 2 diabetes risk, and does this relationship differ by smoking status and alcohol consumption among U.S. adults?

# Potential Hypothesis

Reduced kidney function, indicated by lower creatinine clearance and higher BUN, is associated with impaired  $\beta$ -cell function and increased risk of type 2 diabetes, especially among current smokers and drinkers.

# Research Scope and Objectives

# Scope:

**Population**: U.S. Adults from NHANES

# Key Explanatory Variables:

- Kidney function: Creatinine Clearance (CCR), Blood Urea Nitrogen (BUN)
- Behavioral factors: Smoking status, alcohol consumption

# Mediating/Target Outcome:

- $\beta$ -cell dysfunction (estimated via HOMA-B)
- Type 2 diabetes status (diagnosed or based on biomarkers)

# Stratification/Interaction: Smoking and alcohol status groups

# **Objectives:**

**Primary**: Assess how kidney function markers (CCR and BUN) are associated with  $\beta$ -cell dysfunction and type 2 diabetes

# **Secondary**: Support early identification of high-risk groups based on renal-endocrine interactions. Determine whether impaired renal function precedes or parallels endocrine dysfunction in T2D

# **Research Potential**

## **Clinical Insight**

- Improves early diabetes screening by flagging high-risk individuals with both metabolic red flags and family history
- Promotes personalized prevention strategies

#### **Research Contribution**

- Explores an under-researched physiological pathway linking kidney and endocrine health
- Moves beyond traditional T2D predictors (e.g., BMI, glucose) toward organ-level systems modeling

#### <u>Public Health Relevance</u>

- Offers insights for screening and risk stratification based on routine kidney function labs
- Informs prevention strategies for smokers and drinkers, who may be more vulnerable to combined renal and endocrine stress

# 1.Data Source - National Health and Nutrition Examination Survey)

<u>Source:</u> NHANES (National Health and Nutrition Examination Survey); large, nationally representative U.S. adult sample. Data collected through interviews, physical examinations, and laboratory tests.

• Dataset Size: ~30,000+ participants per combined multi-year cycles.

#### **Target Outcomes** → **Binary Variables**

• Diabetes Status: Self-reported diagnosis and laboratory-confirmed diabetes (Fasting Plasma Glucose ≥ 126 mg/dL or HbA1c ≥ 6.5%).

#### **Predictors Included: Kidney Function:**

- Blood Urea Nitrogen (BUN)
- Creatinine Clearance Rate (CCR)

#### β-cell Function:

Fasting Insulin & Fasting Glucose (HOMA-β calculation)

#### **Family History:**

• Family history of diabetes (parent, sibling) collected via questionnaire.

#### **Physical Behaviors:**

- Smoking status, alcohol use, physical activity levels.
- Link to the Dataset -

https://wwwn.cdc.gov/nchs/nhanes/continuousnhanes/default.aspx?BeginYear=1999

# **Machine Learning Potential**

#### **Predictive Modeling:**

Logistic Regression, Random Forest, XGBoost, LightGBM for type 2 diabetes classification.

## **Moderation and Interaction Analysis:**

Machine learning to detect interactions between metabolic biomarkers, family history, and lifestyle.

#### **Risk Stratification:**

Identifying high-risk subgroups using combined NHANES and Kaggle datasets.

## Feature Importance & Explainability:

SHAP (SHapley Additive exPlanations), feature ranking, and visualization to improve clinical interpretability.

## **Unsupervised Learning:**

Clustering to explore hidden diabetes phenotypes.

# **Web Scraping Integration**

#### **Supplementary Data Collection:**

- Scrape region-specific diabetes prevalence rates, lifestyle trends, and public health reports from CDC, WHO, and other health portals.
- Extract recent research abstracts and news updates from PubMed to track emerging diabetes risk factors.

#### **Added Value:**

- Enriches model context with real-time, location-based, and literature-based insights.
- Allows integration of external environmental or policy-level factors.

#### **Tools:**

- Python (BeautifulSoup, Requests, Pandas)
- Target websites: CDC Diabetes Atlas, BRFSS, PubMed

# Literature Review

#### **Metabolic & Behavioral factors:**

- Individuals with high-normal fasting glucose (91–99 mg/dL) have a greater risk of developing diabetes compared to those with lower normal levels (Brambilla et al., 2011)
- Triglycerides are an independent and early predictor of type 2 diabetes (Zhao et al., 2019)
- High BMI amplifies risk across all metabolic risk markers (Zhao et al., 2019)
- Smoking and alcohol use worsen metabolic regulation (Akhuemonkhan & Lazo, 2017)

# Literature Review

## **Beta Cell Dysfunction**

- Beta cells located in the pancreas produce and secrete insulin (Dludla et al., 2023).
- Beta cell dysfunction indicates impaired insulin secretion, contributing to T2DM (Dludla et al., 2023).
- The Homeostatic Model Assessment of Beta-cell Function (HOMA-B) and the Insulinogenic Index can be used to indicate beta cell function (Kim et al., 2024; Sung et al., 2009)
  - These indicators can be calculated with fasting and post-load glucose and insulin values.

# Literature Review

#### **Type II Diabetes:**

- Type II diabetes, also known as adult-onset diabetes, occurs when the body is not able to utilize insulin correctly and sugar builds up in the blood (Mayo Clinic, 2025).
  - Type II diabetes is more common in older adults (hence adult-onset), however, more and more children are being diagnosed with the rise of childhood obesity (Mayo Clinic, 2025).
- As of 2024, more than 38 million Americans have diabetes, with close to 95% of diagnoses being for Type II diabetes (CDC, 2024)
  - Mostly in adults over 45 years old, but more and more children are getting diagnosed.

# References

- Akhuemonkhan, E. & Lazo, M. (2017). Association between family history of diabetes and cardiovascular disease and lifestyle risk factors in the U.S. population (NHANES 2009–2012).

  \*Preventive Medicine, 96, 129–134. https://doi.org/10.1016/j.ypmed.2016.12.015
- Baliunas, A., Taylor, B., Irving, H., Roerecke, M., Patra, J., Mohapatra, S., & Rehm, J. (2009). Alcohol as a Risk Factor for Type 2 Diabetes. *Diabetes Care, 32*(11), 2123-2132. https://doi.org/10.2337/dc09-0227
- Brambilla, P., La Valle, E., Falbo, R., Limonta, G., Signorini, S., Cappellini, F., & Mocarelli, P. (2011). Normal fasting plasma glucose and risk of type 2 diabetes. *Diabetes Care*, 34(6), 1372–1374. https://doi.org/10.2337/dc10-2263
- CDC. (2024, May 15). About Type 2 Diabetes. Diabetes. https://www.cdc.gov/diabetes/about/about-type-2-diabetes.html
- Dludla, P. V., Mabhida, S. E., Ziqubu, K., Nkambule, B. B., Mazibuko-Mbeje, S. E., Hanser, S., Basson, A. K., Pheiffer, C., & Kengne, A. P. (2023). Pancreatic β-cell dysfunction in type 2 diabetes: Implications of inflammation and oxidative stress. *World Journal of Diabetes*, 14(3), 130–146. https://doi.org/10.4239/wjd.v14.i3.130
- Kim, J. Y., Lee, J., Kim, S. G., & Kim, N. H. (2024). Recent Glycemia Is a Major Determinant of β-Cell Function in Type 2 Diabetes Mellitus. *Diabetes & Metabolism Journal*, 48(6), 1135-1146. https://doi.org/10.4093/dmj.2023.0359
- Mayo Clinic. (2025, February 27). Type 2 diabetes. Mayo Clinic. https://www.mayoclinic.org/diseases-conditions/type-2-diabetes/symptoms-causes/syc-20351193
- Sung, K. C., Reaven, G., & Kim, S. (2009). Utility of Homeostasis Model Assessment of β-Cell Function in Predicting Diabetes in 12,924 Healthy Koreans. *Diabetes Care*, *33*(1), 200-202. https://doi.org/10.2337/dc09-1070
- Zhao, J., Zhang, Y., Wei, F., Song, J., Li, W.-D., Chen, C., Zhang, K., & Feng, S. (2019). Triglyceride is an independent predictor of type 2 diabetes among middle-aged and older adults: A prospective study with 8-year follow-ups in two cohorts. *Journal of Translational Medicine*, 17, 354. https://doi.org/10.1186/s12967-019-02156-3

## **DATA FILES AND SOURCES**

https://www.cdc.gov/brfss/annual\_data/annual\_2015.html

https://www.kaggle.com/datasets/cdc/behavioral-risk-factor-surveillance-system/data

https://www.kaggle.com/datasets/alexteboul/heart-disease-health-indicators-dataset/data