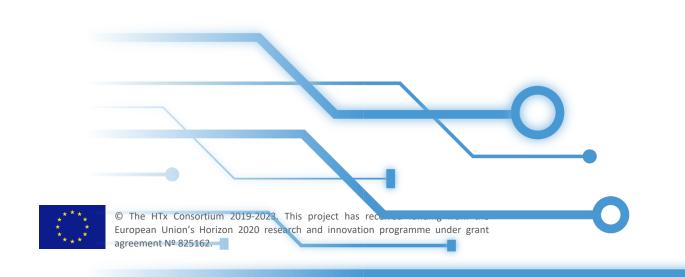


# Prediction modeling for HTA using Explainable AI (XAI)



**Gunjan Chandra, BISG, University of Oulu** 



## Goal





Create a Clinical Decision Support System (CDSS) to support clinicians make better decisions

- XAI ensures that the system's recommendations are understandable and can be explained.
- Addressing challenges associated with the complexity and interpretability of AI-driven clinical decision-making.



# Predicting clinical outcomes



ORIGINAL RESEARCH

## Data-Driven Identification of Long-Term Glycemia Clusters and Their Individualized Predictors in Finnish Patients with Type 2 Diabetes

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#### **Objectives:**

- Identify patients with homogeneous long-term HbA1c trajectories.
- Predict trajectory membership using explainable machine learning and various predictors (clinical, treatment, socio-economic).





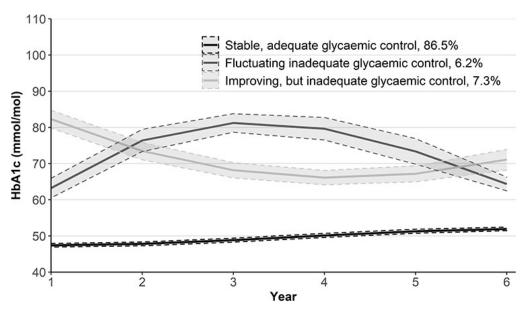
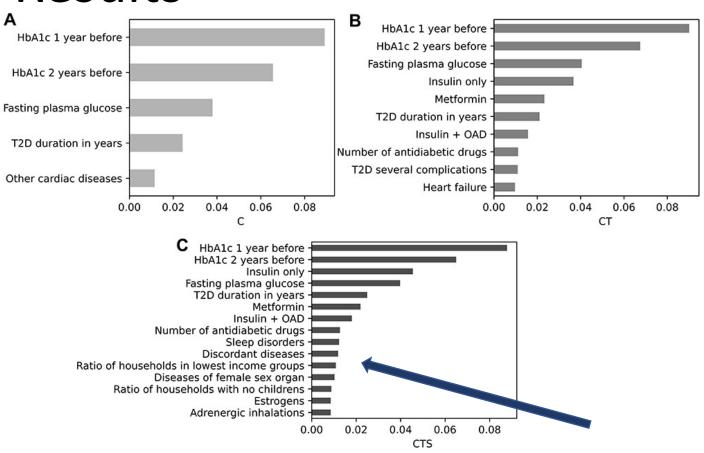
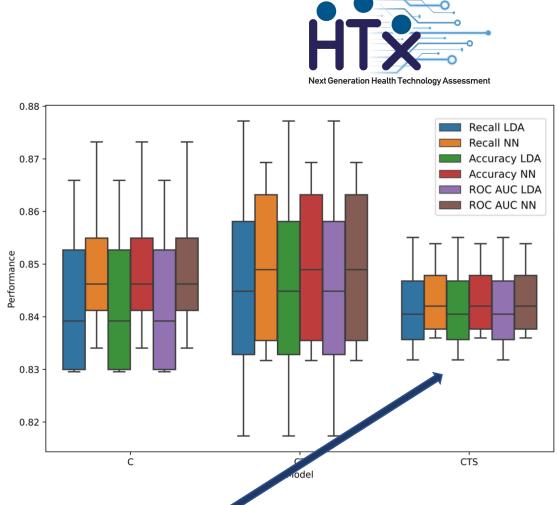


Figure 1. Estimated HbA1c trajectories.



**Figure 2.** Feature importance plot for (**A**) Clinical (C), (**B**) Clinical + Treatment (CT), (**C**) and Clinical + Treatment + SES (CTS) models.





**Figure 3.** Performance of models over different splits in 4-fold cross-validation.

# Global and local explaination (SHAP)

High



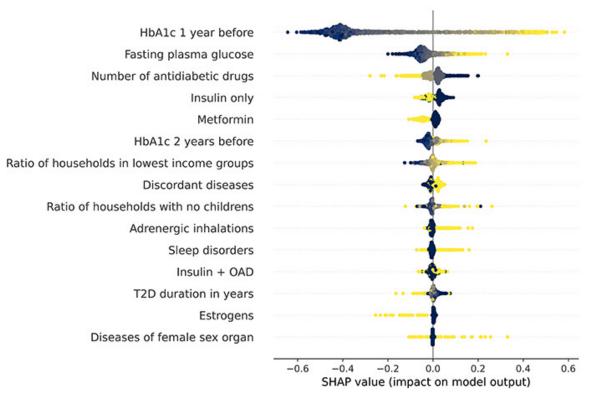
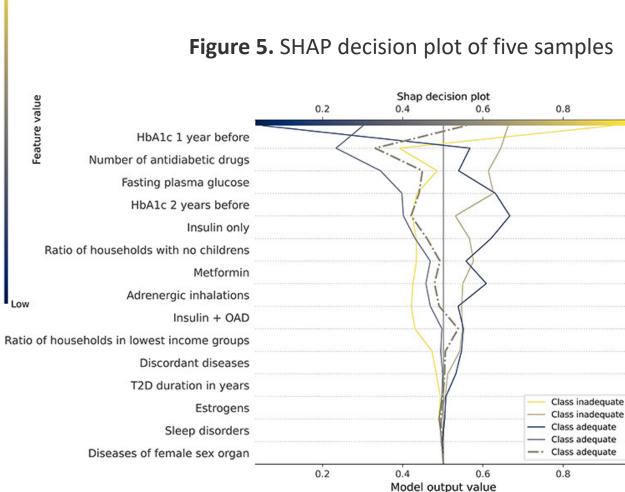


Figure 4. SHAP summary plot





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#### Informatics in Medicine Unlocked

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# Explainable Artificial Intelligence to predict clinical outcomes in type 1 diabetes and relapsing-remitting multiple sclerosis adult patients

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- e Research Center for Clinical Neuroimmunology and Neuroscience (RC2NB), University Hospital and University of Basel, Basel, 4001, Switzerland

## **Objectives:**



Predict clinical outcomes in type 1 diabetes and relapsing-remitting multiple sclerosis adult patients



Compare machine learning and statistical methods



Outcome	Models built using statistically identified prognostic / risk factors	Models built using features selected through ML methods	Statistical model
Relapses (MS)	AUC – 0,67 BA – 0,66 F1 score – 0,71	Male AUC – 0,70 BA – 0,70 F1 score – 0,84	AUC – 0,65
		Female AUC – 0,69 BA – 0,68 F1 score – 0,76	
Severe hypoglycemia (T1D)	AUC – 0.65 BA – 0,66 F1 score – 0,65	Male AUC – 0,88 BA – 0,85 F1 score – 0,84	-
		Female AUC – 0,82 BA – 0,79 F1 score – 0,84	
Diabetic Ketoacidosis (T1D)	AUC – 0,69 BA – 0,68 F1 score – 0,78	AUC – 0,85 BA – 0,83 F1 score – 0,78	-



- Machine learning models that rely only on known risk factors yield moderate prediction accuracy.
- Feature selection methods have the potential to improve the prediction of medical outcomes.
- Socioeconomic factors, physical health, and mental health impact the prediction of medical outcomes.



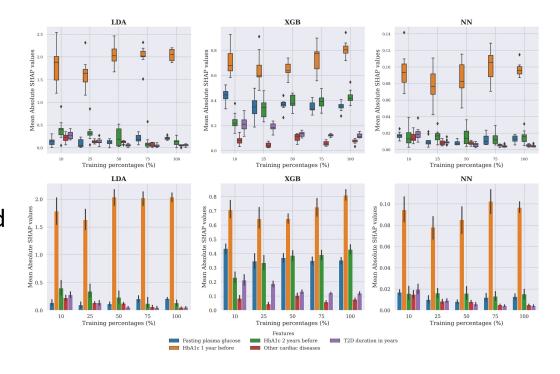
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# **Objective:** Influence of Data Size and Class Balance on Machine Learning Classification Performance and SHAP explanations



#### **Results:**

- Various machine learning models work best with different amounts of training data, and the effect of imbalanced data on performance depends on the metrics used.
- SHAP explanations are more effective when there is balanced background data, and their stability improves with larger background datasets.



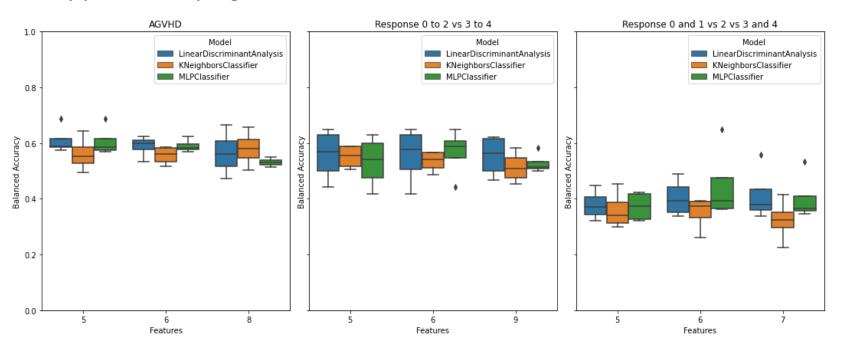
# Orphan diseases



**Objective:** Al for Predicting Acute Graft versus-Host Disease and Subtypes in Allogeneic Hematopoietic Cell Transplantation for T-cell Prolymphocytic Leukaemia

#### **Methods:**

- Open data set from Centre for International Bone and Marror Transplant Research (CIBMTR)
- Only predefined prognostic features were used



#### **Results:**

- Models predict the occurance of aGVHD and its sub-types with moderate to low accuracy.
- The performance of the models could be impacted by the data size or the absence of comprehensive data.



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## In progress







Predict the success of ESAs in EUMDS patients and time to response.

Investigating ITE using
Causal ML with Time-toEvent Data in AML Patients
Undergoing Allo-HCT with
Different Treatment
Regimens.





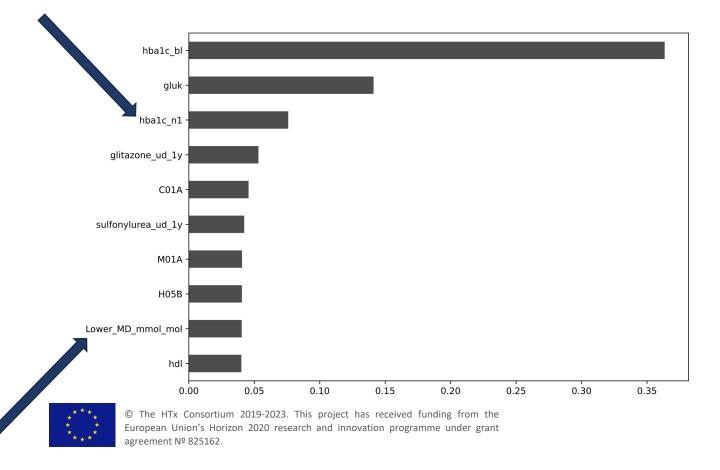
# Predicting treatment outcomes

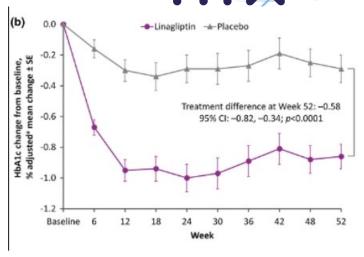
## Offset model

**Objective:** Predicting Change in HbA1c Values Following Initiation of

Antidiabetic Drugs in Type 2 Diabetes using XAI

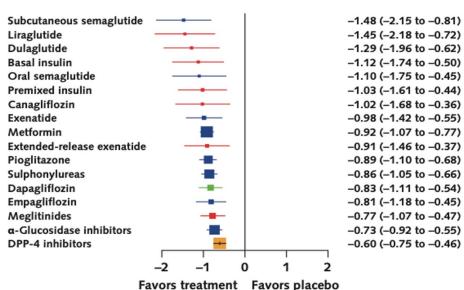
- Distance from baseline to target: Minimum: 80, Median: 280.0, Maximum: 364
- Added expected HbA1c changes from RCT as predictors.
- Added HbA1c follow up value.



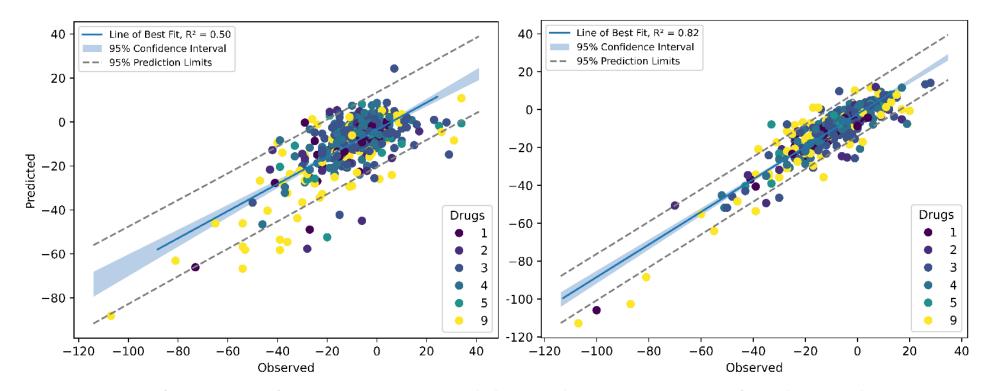


#### A. Change in Hemoglobin A<sub>1c</sub> Level in Drug-Naive Patients

#### MD (95% CI)







Code Name

1 Metformin

2 GLP-1 analogues

3 DPP-4 inhibitors

4 SGLT2 inhibitors

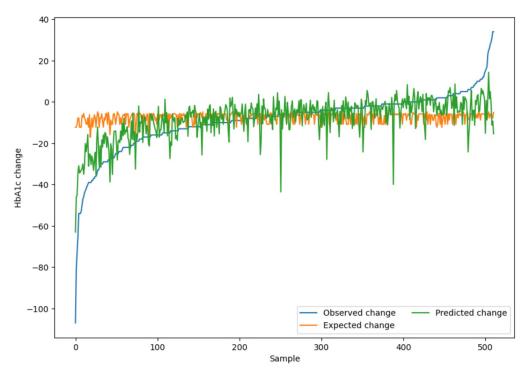
5 Combinations of oral blood glucose lowering drugs

9 Insulin

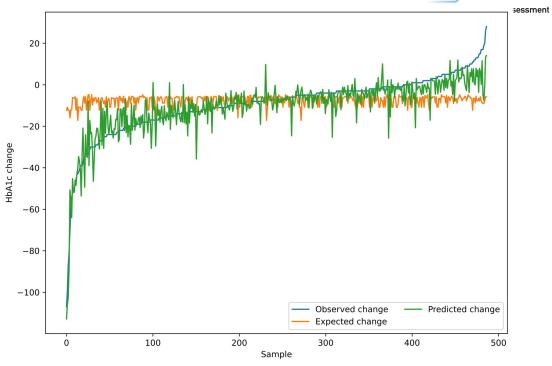
Figure 1. Performance of MLPRegressor Model: Fitted Regression Line for HbA1c Change Before and After Drug Initiation. Base model on the left and model using follow-up HbA1c value after drug initiation on the right.







Base model: Al predicted better in 287 cases, RCT 223 cases



Follow-up HbA1c value: Al predicted better in 290 cases, RCT 196 cases

- All outperforms RCT values in predicting individualized treatment responses in both cases.
- The occurrence of positive changes following drug initiation raises questions.



# Next steps







Multi-target regression modeling for tretament effect calculation and optimal treatment selection.

XAI-based clinical decision support system (<u>Demo-CDSS</u>)

### Welcome to HTx Website!

Please select a disease to study from the dropdown menu: Disease

Please select a model to study the disease from the menu: Model 

Model

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## Conclusion

- When employing more holistic modelling approaches, AI demonstrates heightened efficacy in predicting both clinical and treatment outcomes.
- The magnitude of the dataset significantly influences both the performance and explainability of the model.
- Artificial intelligence exhibits the potential to enhance predictive performance specifically for orphan diseases.



# Thank you!

