Empowering Clinicians to Manage Chronic-comorbid Conditions by Predicting Adverse Outcomes with Transparent AI at the Point of Care

Dillon Tracy, MS Physics – Principal Data Scientist, Amy Sheide, BSN, MPH, MS Biomedical informatics – VP Data Platform and Partnerships

Challenge

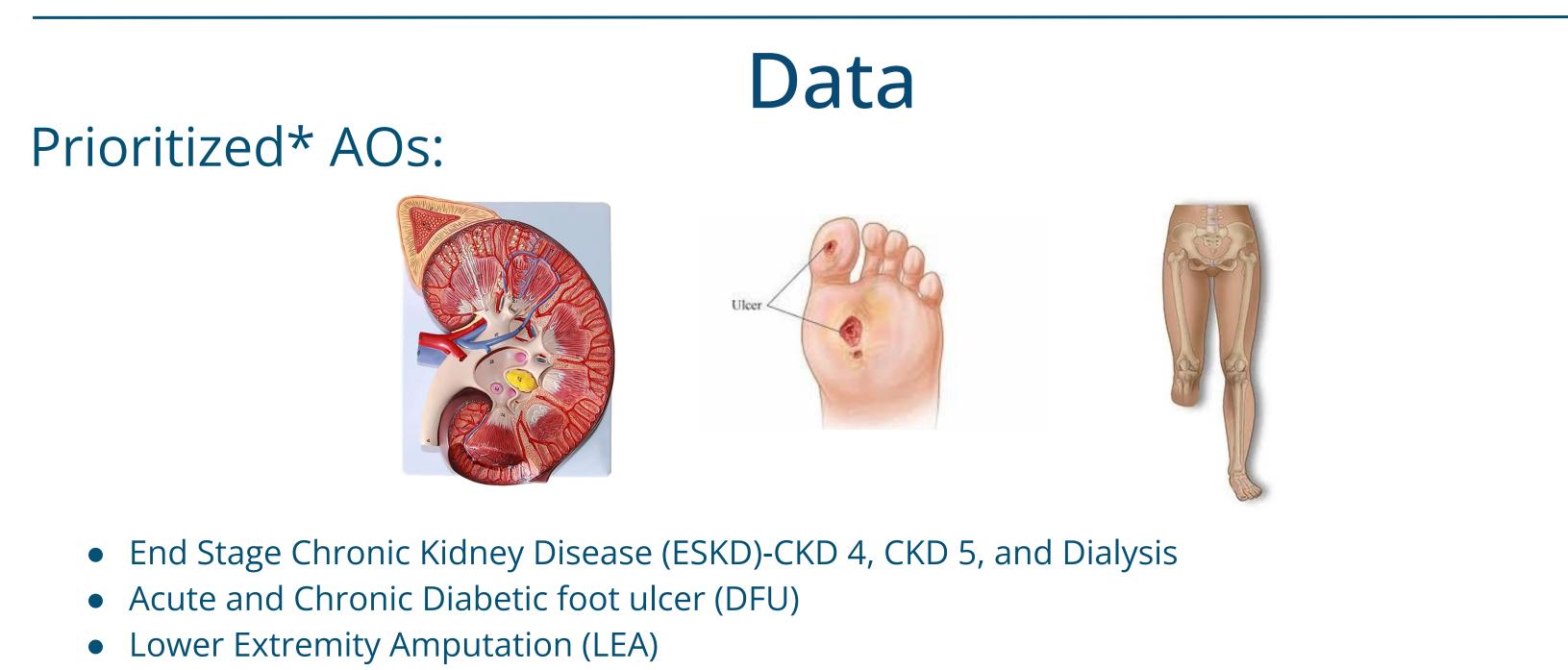
The interconnectivity of factors contributing to adverse outcomes (AOs) in type 2 diabetes (T2D) makes it difficult to predict when complications will occur for individual patients. While population stratification of T2D AOs has been studied^{1,2,5,6,7,8}, predicting when a specific patient will experience explicit outcomes and deploying person-specific interventions to address those outcomes have not been integrated into clinical workflows.

Objective

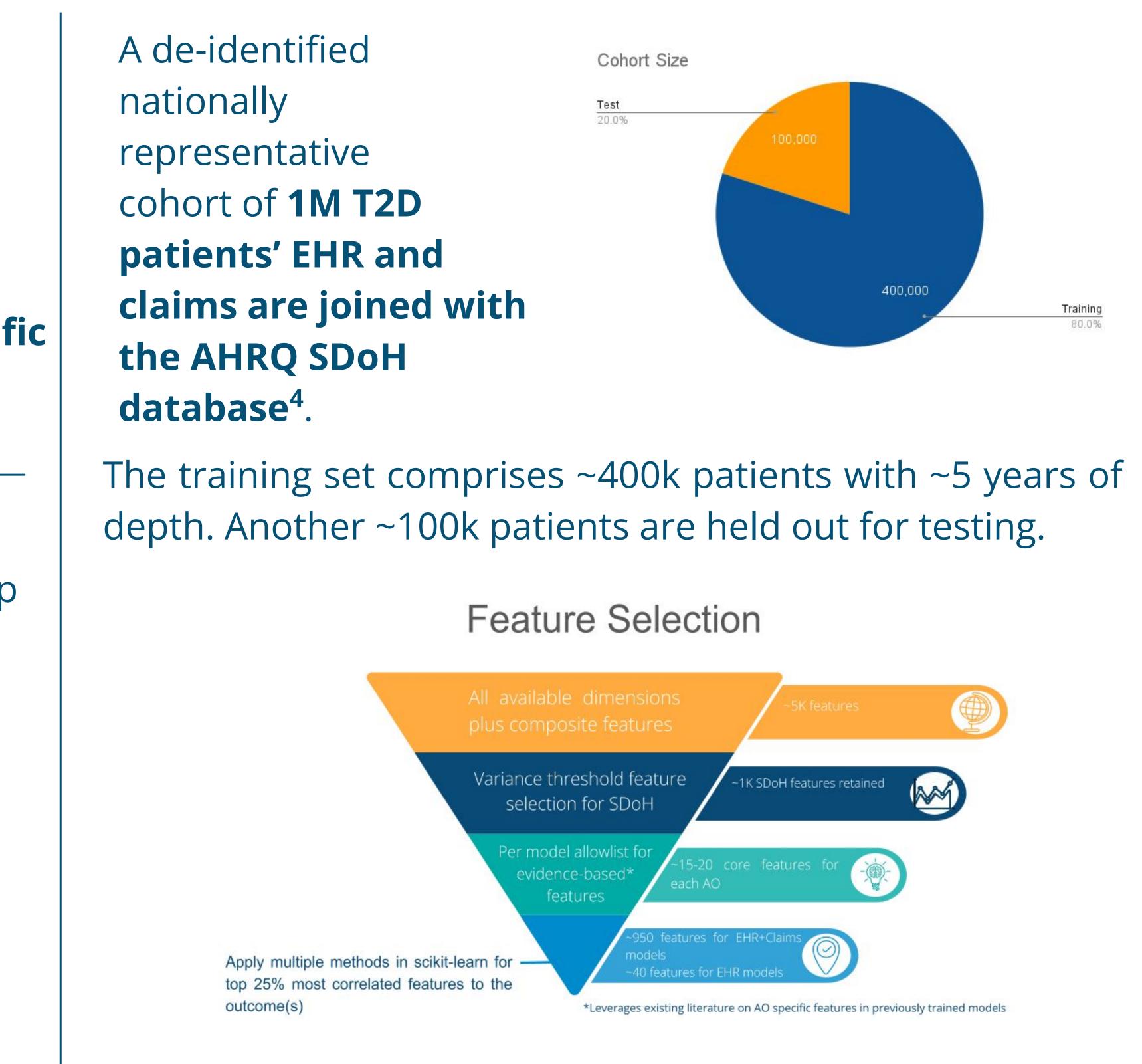
Zephyr Al[™] and MedStar Health® have partnered to develop and deploy **interpretable AI/ML models** into the workflow.

sk Timeline				Filter ≢
CKD Stage 4				
	6 - 12 months	12 - 24 months 24 - 36	months	
	45% Moderate		0% High	
Contributing Factor	Date	Value	Influence	
Non Diabetic Kidney Disease	5/21/2021	N20.0-Kidney stone	37%	
Non Diabetic Kidney Disease DCSI Risk	5/21/2021 1/9/2021	N20.0-Kidney stone	37% 13%	
	16 - 15 -			
DCSI Risk	1/9/2021	10	13%	

Patient AO predictions for 0-5 years are returned as probability measures with individualized contributing factors to assist clinicians in the delay or prevention of AOs.







We compute ~5k features and **use the highest quality** ~1k features in training. Feature provenance is tracked to enable smart imputation when limited patient data is available.

Methods

Multiple types of survival models (RSF, AFT, MTLR, PLSR, CCA) are trained for each AO from first interaction to the date of T2D diagnosis plus a fixed observation period (6 or 12 months). We have generally obtained the best performance, in metrics and scalability, from MTLR.

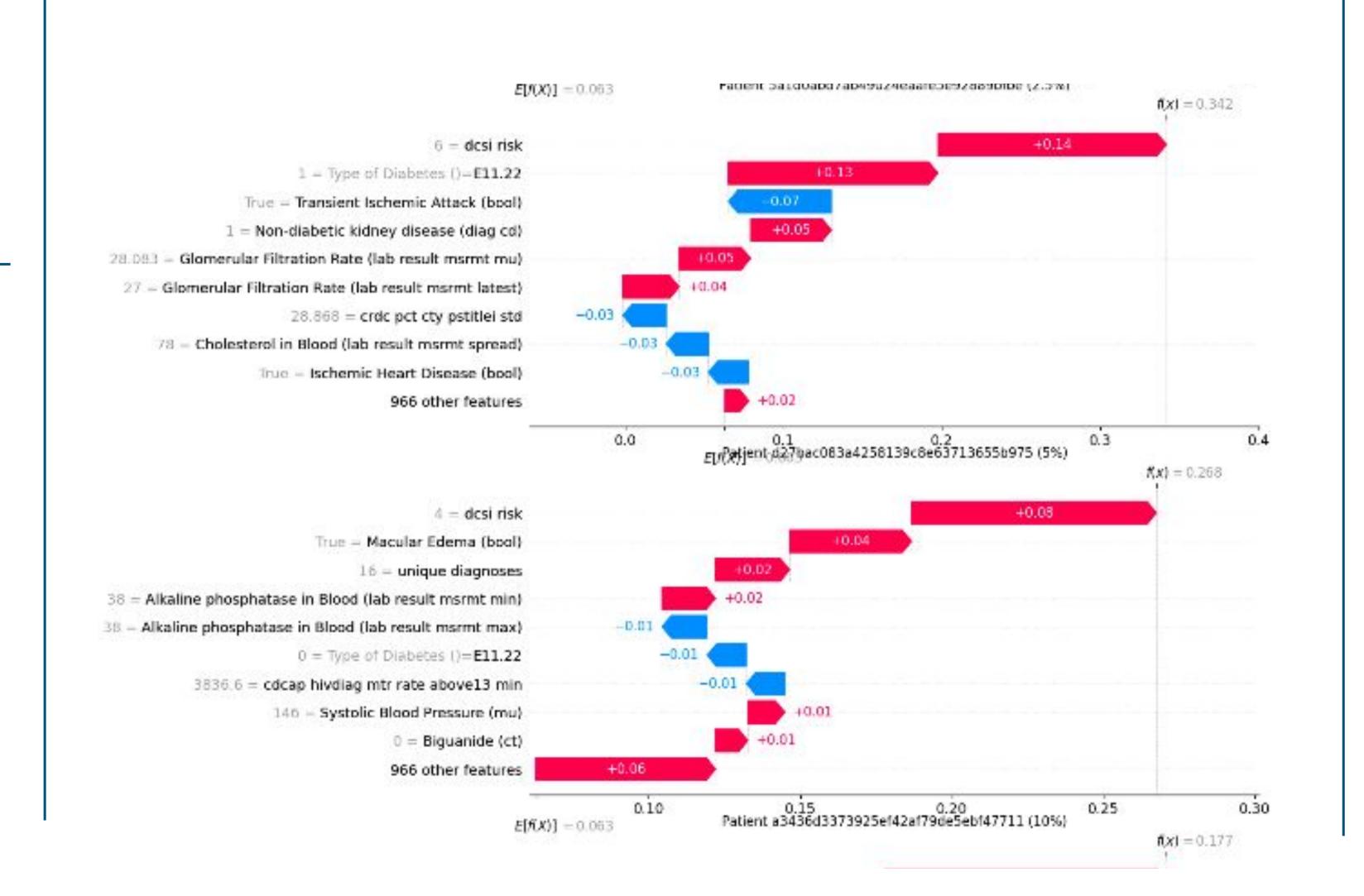
*considered clinically relevant and actionable

Evaluating predictive capability, generalizability and clinical utility of Zephyr models on 300K T2D MedStar patients is underway

Stratification of a homogenous cohort with explainability is achieved using SHapley Additive **exPlanations (SHAP)**¹¹ and aids in establishing model trust.

Type of Diabetes ()=E11.22 nique diagnoses Biguanide (ct) Non-diabetic kidney disease (diag cd) age in days Transient Ischemic Attack (bool)=True Glomerular Filtration Rate (lab result msrmt min) Hypertension (bool)=True Depression Diagnosis (bool)=True Diabetic Medications (ct) Urea nitrogen in Blood (BUN) (lab result msrmt mu) Glomerular Filtration Rate (lab result msrmt mu) Coronary Artery Disease (CAD) (bool)=True Systolic Blood Pressure (mu) Glomerular Filtration Rate (lab result msrmt earliest) Urea nitrogen in Blood (BUN) (lab result msrmt max) Peripheral artery disease (bool)=True Loop Diuretic (ct)

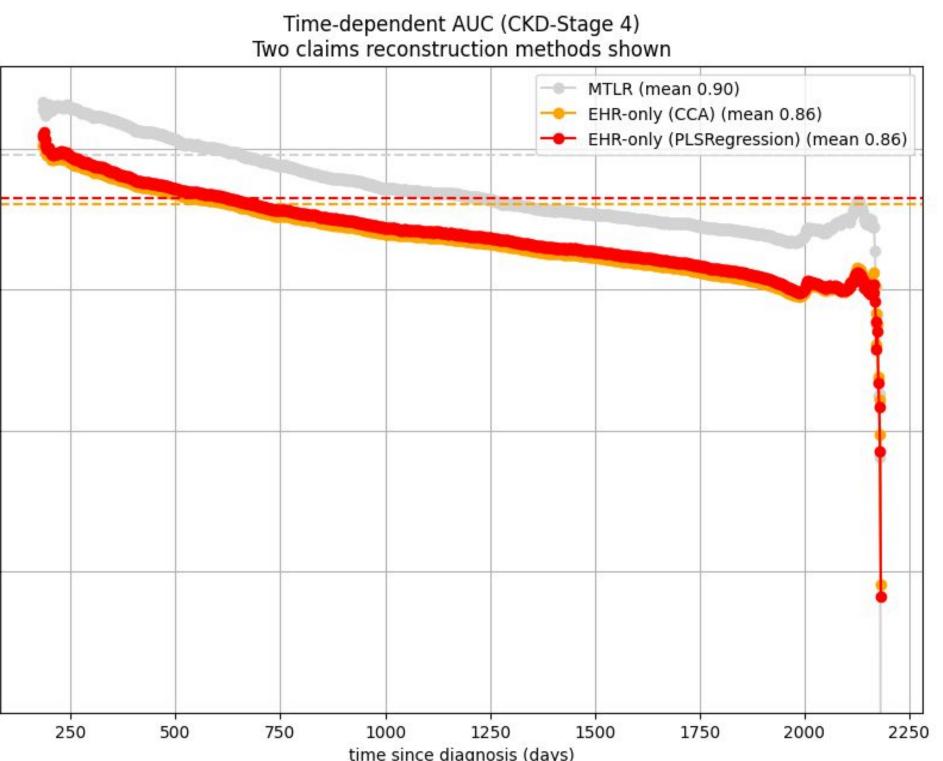
A SHAP "waterfall plot" shows the 10 most influential features for a given patient. SHAP values are additive, their sum gives the individual risk vs. population prevalence of an adverse outcome within in the prediction window. Feature-level contributions to risk are critical to clinicians' interpretation of models^{12,13} and prioritization of interventions.



Results

AO	Lit. AUC	Zephyr AUC
ESKD	0.81 ⁵ -0.84 ⁶	0.80-0.93
DFU	0.86 ⁷ -0.88 ⁸	0.78-0.86
LEA	0.72 ⁹ -0.84 ¹⁰	0.83-0.88

Most models compare favorably to metrics published in the space^{14, 18, 19, 20} and show an **average AUC of 0.85**, concordance indices > 0.76, and Brier scores < 0.07 across the AOs.

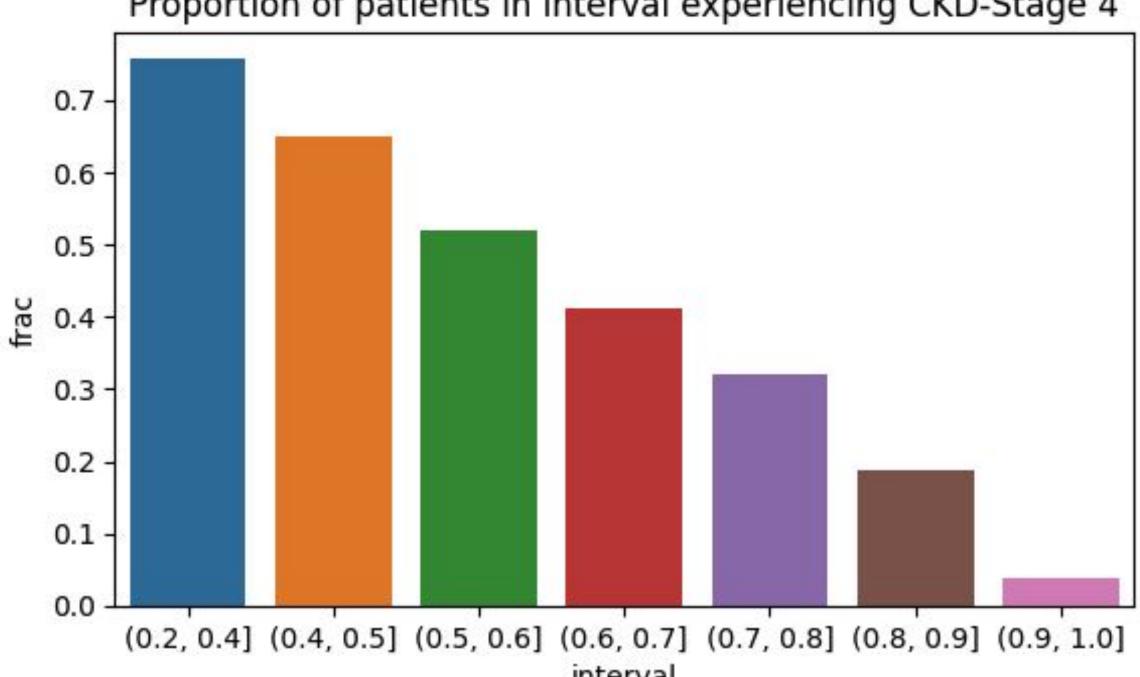


Matrix reconstruction techniques allow use of partial patient data at a modest cost in AUC.

Backtesting: **the** proportion of patients experiencing a given adverse outcome correlates with the predicted risk.

1007/s 101 98-01 2-04 10-

Proportion of patients in interval experiencing CKD-Stage 4



References

vakowska, Magdalena, et al. "The comorbidity burden of type 2 diabetes mellitus: patterns, clusters and predictions from a large English primary care cohort." BMC medicine 17.1 (2019): 1-10.

Wang, Shiqi et al. "Machine Learning Models for Predicting the Risk of Hard-to-Heal Diabetic Foot Ulcers in a Chinese Population." Diabetes, metabolic syndrome and obesity : targets and therapy vol. 15 3347-3359. 29 Oct. 2022,

on risk in patients undergoing lower extremity amputation due to the

16. Yang, Jiaxi, et al. "Predicting Risk of Hypoglycemia in Patients With Type 2 Diabetes by Electronic Health Record–Based Machine Learning: Development and Validation." JMIR Medical Informatics 10(6):e36958 (2022). do 10.2196/36958 17. Ravaut, Sedeghi, et al. "Predicting adverse outcomes due to diabetes complications with machine learning using administrative health data." npj Digital Medicine (2021)4:24. doi: 10.1038/s41746-021-00394-8

18. Lo-Ciganic, Donohue, et al. "Using Machine Learning to Examine Medication Adherence Thresholds and Risk of Hospitalization." Med Care. 2015 August ; 53(8): 720–7 20. Sarah Clifford, Magaly Perez-Nieves, Anne M. Skalicky, Matthew Reaney and Karin S. Coyne (2014) "A systematic literature review of methodologies used to assess medication adherence in patien

Research and Opinion, 30:6, 1071-1085, DOI: 10.1185/03007995.2014.884491 Amanda Momenzadeh, Ali Shamsa, Jesse G Meyer, "Bias or biology? Importance of model interpretation in machine learning studies from electronic health records," JAMIA Open, Volume 5, Issue 3, October 2022, ooac063

22. Perveen S, Shahbaz M, Ansari MS, Keshavjee K and Guergachi A (2020). "A Hybrid Approach for Modeling Type 2 Diabetes Mellitus Progression." Front. Genet. 10:1076. doi: 10.3389/fgene.2019.01076