



CLEF 2022

Notebook for the iDPP Lab on Intelligent Disease Progression Prediction

Baseline Machine Learning Approaches to Predict Amyotrophic Lateral Sclerosis Disease Progression

Isotta Trescato, PhD student
isotta.trescato@unipd.it

SBB team, UNIPD

Barbara Di Camillo
Alessandro Guazzo
Enidia Hazizaj
Enrico Longato
Chiara Roversi
Erica Tavazzi
Isotta Trescato
Martina Vettoretti





Task 1

Rank subjects based on the risk of early occurrence of clinical events for ALS

Task 2

Predict time of occurrence of clinical events for ALS

SURVIVAL ANALYSIS (model time-to-event directly)

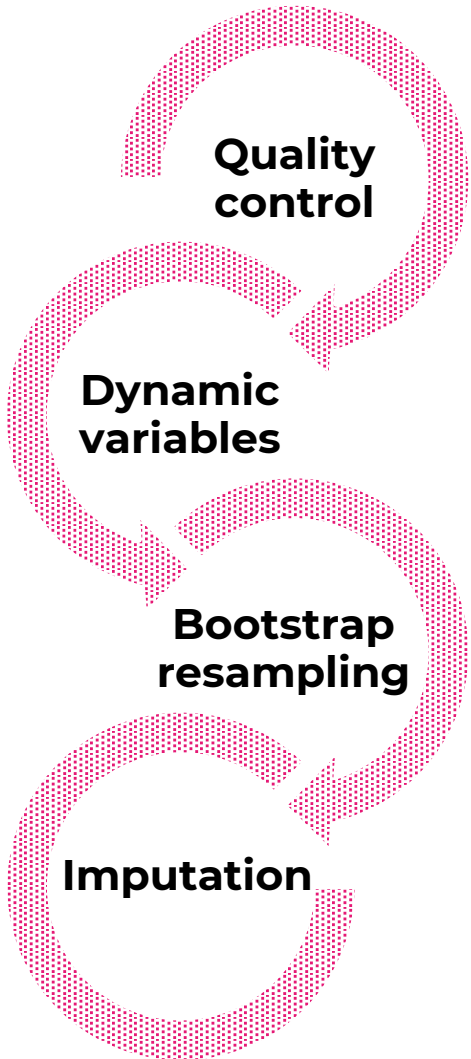
- ▷ Cox proportional-hazards model (Cox)
- ▷ Survival Support Vector Machines (SSVM)
- ▷ Random Survival Forest (RSF)



- ▷ **Preprocessing**
- ▷ **Model training pipelines**
- ▷ **Outcomes**
 - Models' outcomes
 - Task 1 outcomes
 - Task 2 outcomes
- ▷ **Results**
 - Task 1 results
 - Task 2 results
- ▷ **Conclusion**



Preprocessing



- ▶ No subject with more than 20% missing
- ▶ One subject was removed, positive onset date

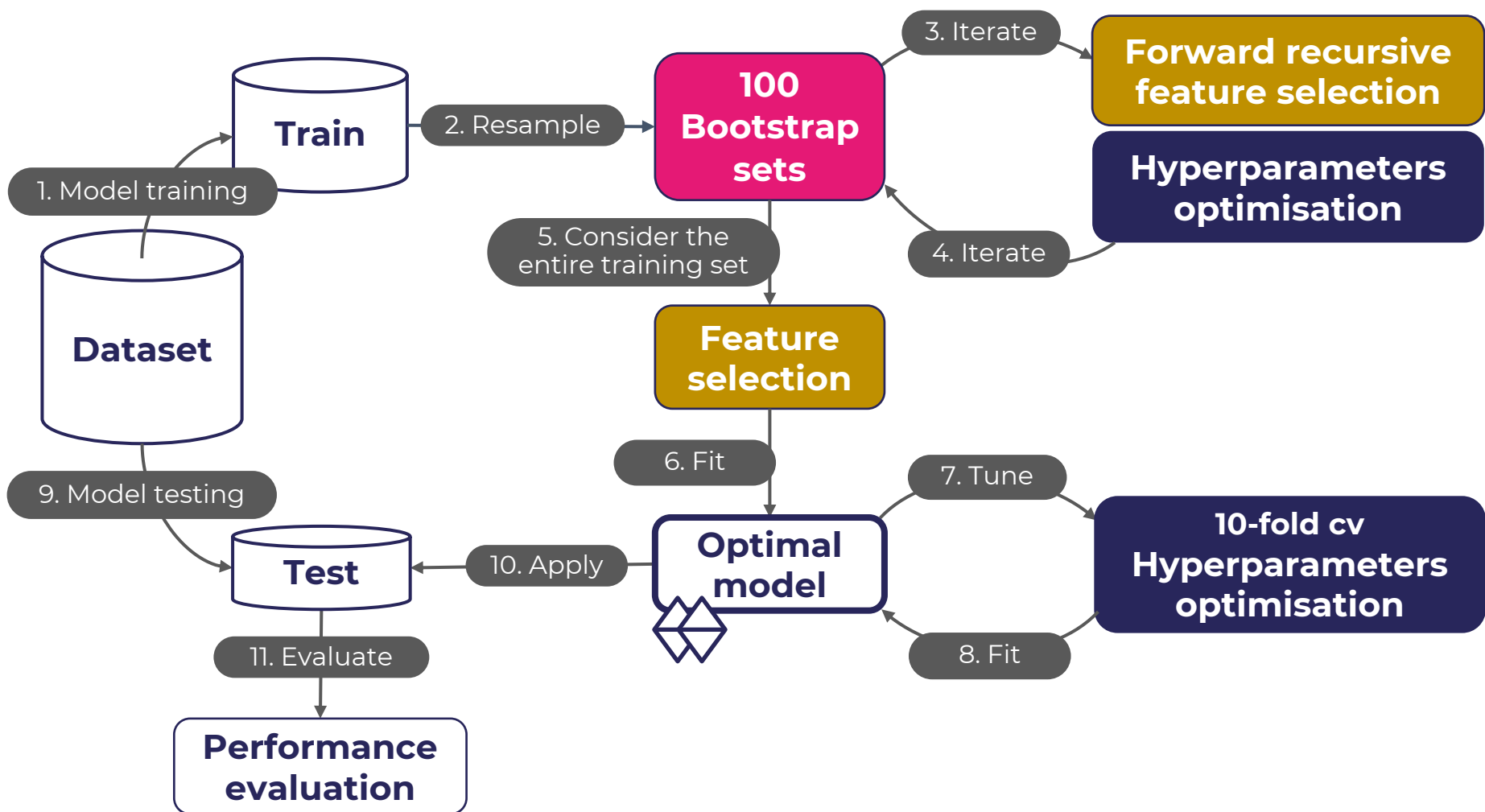
- ▶ Considered as slopes to capture the variations over time

- ▶ Hyperparameters tuning and feature selection
- ▶ 100 sets: internal training set + validation set

- ▶ All variables with more than 70% missing removed
- ▶ 4 variables needed imputation: FVC, instruction level, BMI, weight slope

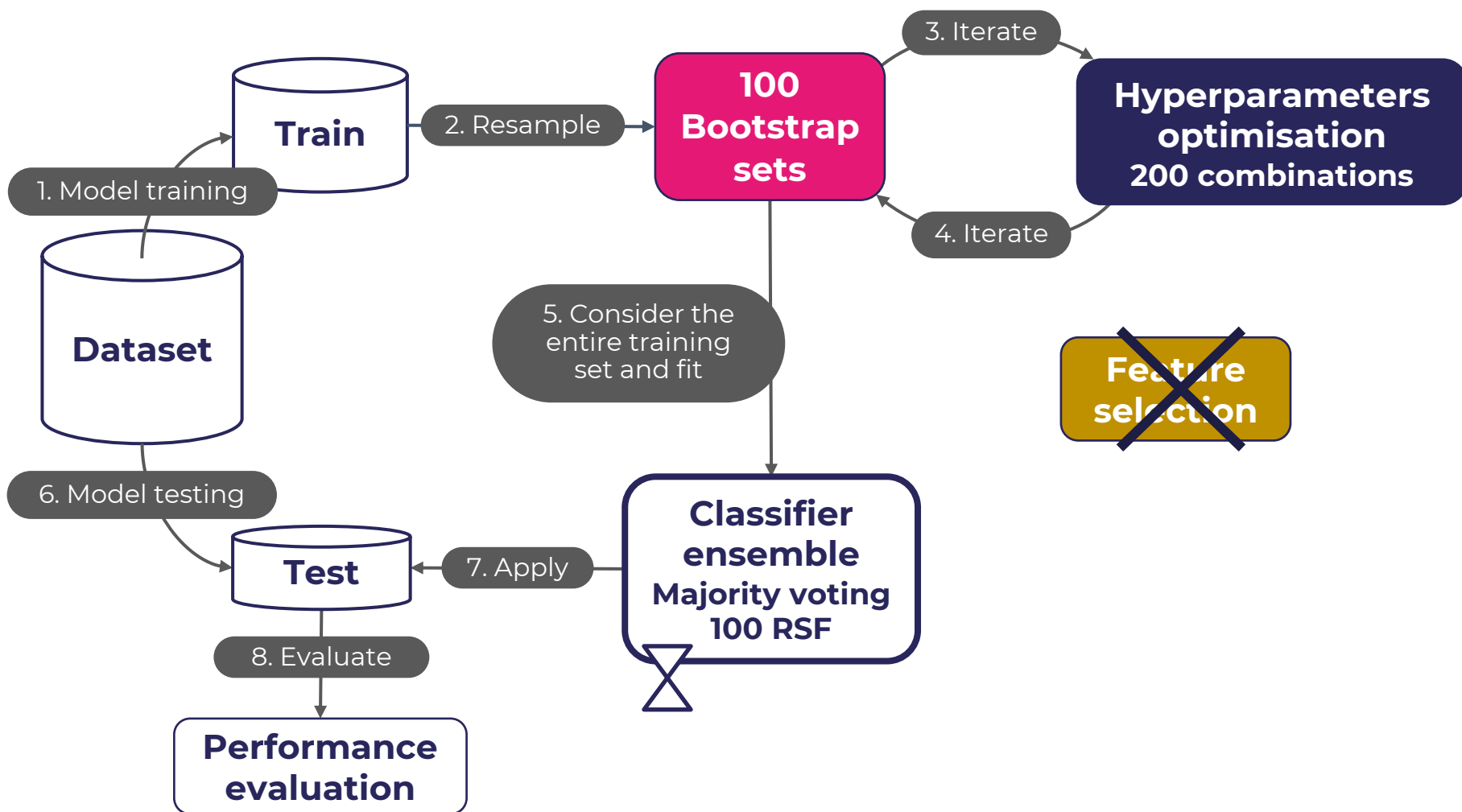


Cox and SSVM pipeline





RSF pipeline





Model outcome

Cox proportional-hazards model



Survival Support Vector Machines



Random Survival Forest





Model outcome (1 of 2)

Cox proportional-hazards model



▶ **Model outcome: risk scores**

Survival Support Vector Machines



▶ **Model outcome: risk scores**

Random Survival Forest





Model outcome (2 of 2)

Cox proportional-hazards model



▶ Model outcome: risk scores

Survival Support Vector Machines



▶ **Model outcome: time to event**

Random Survival Forest



▶ Model outcome: risk scores



Task 1 – Risk prediction

Cox proportional-hazards model



▶ Model outcome: risk scores

Survival Support Vector Machines



▶ Model outcome: time to event

Random Survival Forest



▶ Model outcome: risk scores



Task 1 – Risk prediction

Cox proportional-hazards model



- ▶ Model outcome: risk scores
- ▶ **Task 1: straightforward from the model**

Survival Support Vector Machines



- ▶ Model outcome: time to event

Random Survival Forest



- ▶ Model outcome: risk scores
- ▶ **Task 1: straightforward from the model**



Task 1 – Risk prediction

Cox proportional-hazards model



- ▶ Model outcome: risk scores
- ▶ Task 1: straightforward from the model

Survival Support Vector Machines



- ▶ Model outcome: time to event
- ▶ **Task 1: derived via Platt scaling**

Random Survival Forest



- ▶ Model outcome: risk scores
- ▶ Task 1: straightforward from the model



Task 2 – Time to event prediction

Cox proportional-hazards model



- ▶ Model outcome: risk scores
- ▶ Task 1: straightforward from the model

Survival Support Vector Machines



- ▶ Model outcome: time to event
- ▶ Task 1: derived via Platt scaling

Random Survival Forest



- ▶ Model outcome: risk scores
- ▶ Task 1: straightforward from the model



Task 2 – Time to event prediction

Cox proportional-hazards model



- ▶ Model outcome: risk scores
- ▶ Task 1: straightforward from the model
- ▶ **Task 2: predicted survival = 0.5**

Survival Support Vector Machines



- ▶ Model outcome: time to event
- ▶ Task 1: derived via Platt scaling

Random Survival Forest



- ▶ Model outcome: risk scores
- ▶ Task 1: straightforward from the model
- ▶ **Task 2: predicted survival = 0.5**



Task 2 – Time to event prediction

Cox proportional-hazards model



- ▶ Model outcome: risk scores
- ▶ Task 1: straightforward from the model
- ▶ Task 2: predicted survival = 0.5

Survival Support Vector Machines



- ▶ Model outcome: time to event
- ▶ Task 1: derived via Platt scaling
- ▶ **Task 2: straightforward from the model**

Random Survival Forest



- ▶ Model outcome: risk scores
- ▶ Task 1: straightforward from the model
- ▶ Task 2: predicted survival = 0.5

Results





Task 1: submission selection

- ▶ For each subtask (a, b, c) and each dataset version (M0 and M6), the two best-performing models according to the average C-index on the 100 bootstrap sets were selected.
- ▶ 12 submitted runs for Task 1
 - 3 subtasks
 - 2 dataset versions

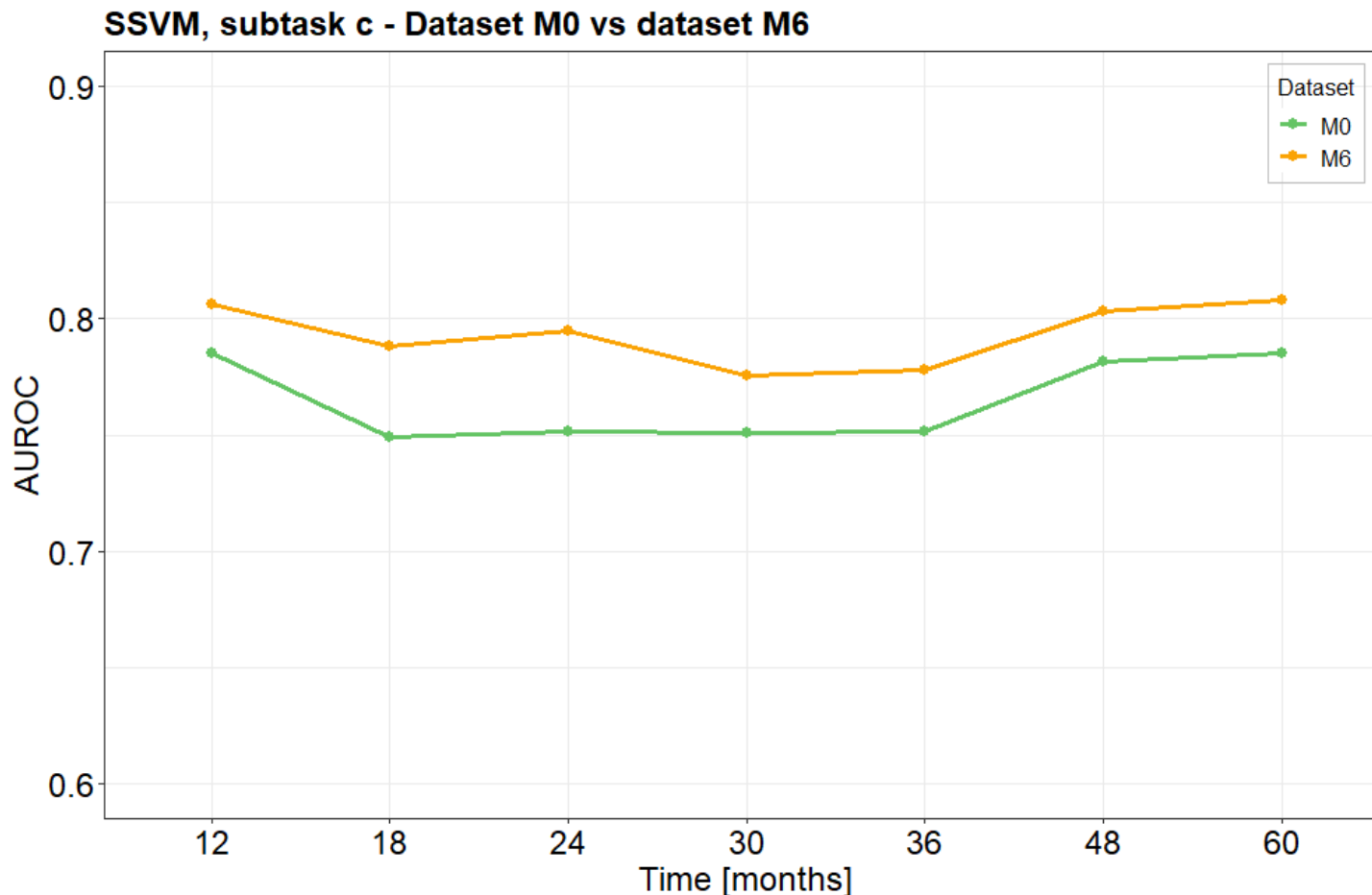
	Subtask a: PEG or death		Subtask b: NIV or death		Subtask c: death	
	M0	M6	M0	M6	M0	M6
Cox	0.678 ± 0.009	0.717 ± 0.010	0.684 ± 0.016	0.726 ± 0.015	0.693 ± 0.014	0.726 ± 0.016
SSVM	0.679 ± 0.010	0.715 ± 0.011	0.695 ± 0.013	0.735 ± 0.011	0.698 ± 0.011	0.733 ± 0.012
RSF	0.697 ± 0.020	0.703 ± 0.009	0.690 ± 0.010	0.729 ± 0.008	0.687 ± 0.009	0.722 ± 0.013

In the table: C-index [mean ± sd]



Task 1: results (1 of 3)

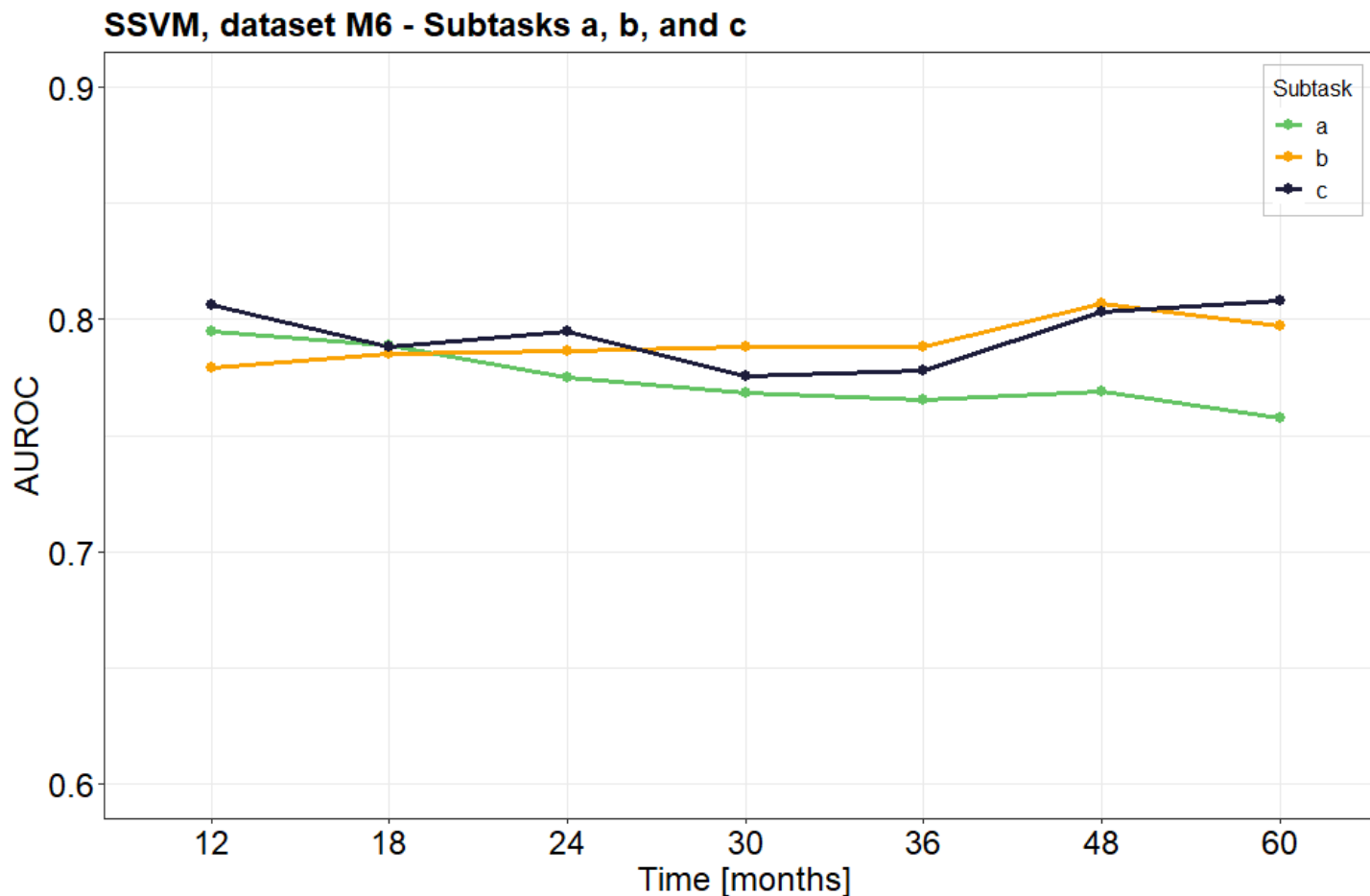
Model discrimination improves when adding 6 months of data to the baseline (M6 vs. M0)





Task 1: results (2 of 3)

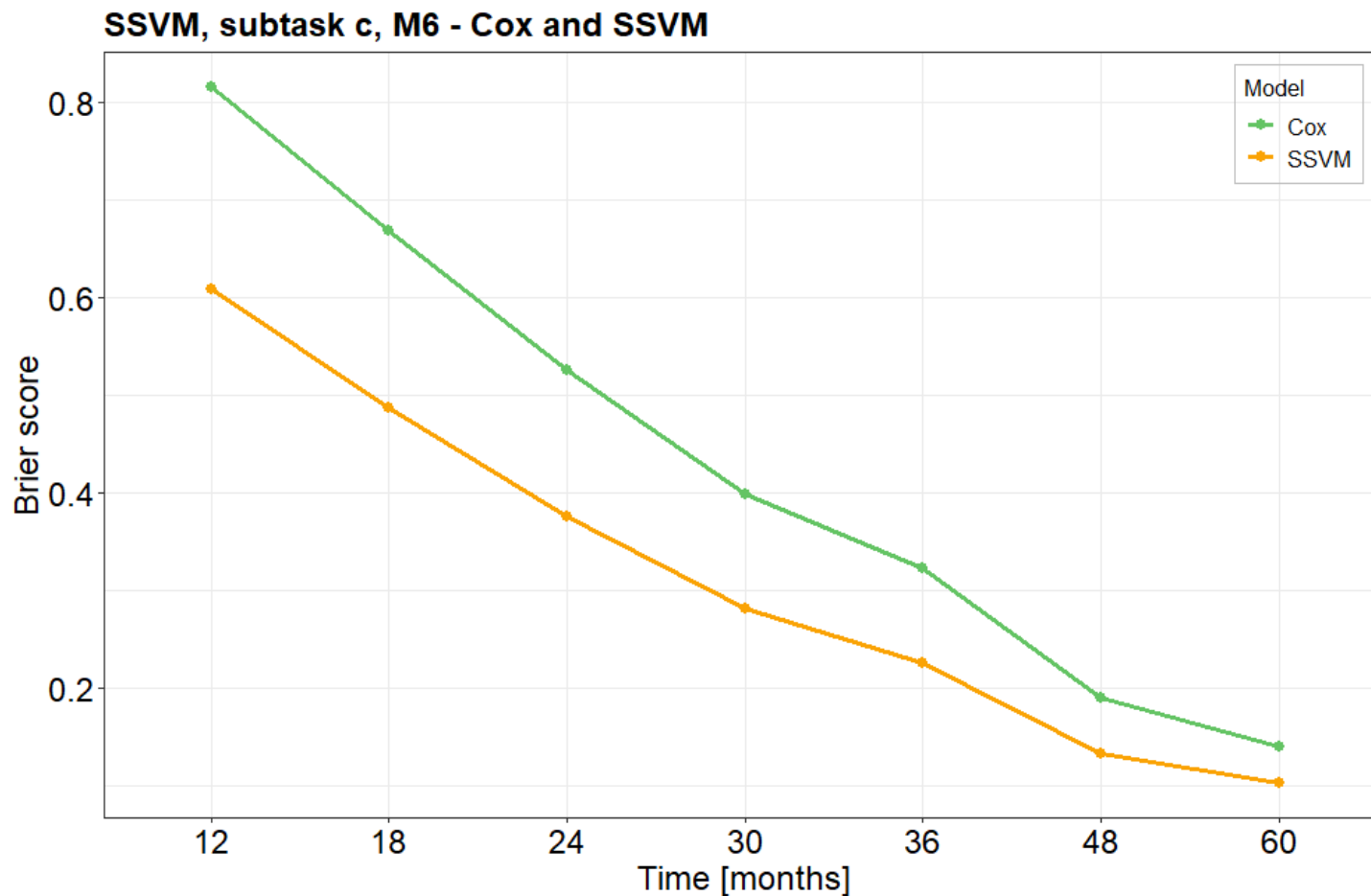
Better performance in subtasks b (PEG) and c (death)





Task 1: results (3 of 3)

Better model calibration with longer prediction horizons





Task 2: submission selection

- ▶ For each subtask (a, b, c) and each dataset version (M0 and M6), the two best-performing models according to the average C-index on the 100 bootstrap sets were selected.
- ▶ 12 submitted runs for Task 2 (same methods as task 1)
 - 3 subtasks
 - 2 dataset versions

	Subtask a: PEG or death		Subtask b: NIV or death		Subtask c: death	
	M0	M6	M0	M6	M0	M6
Cox	--	7.796	--	--	8.187	7.683
SSVM	8.847	7.662	7.714	7.358	8.115	7.582
RSF	8.599	--	7.407	6.857	--	--

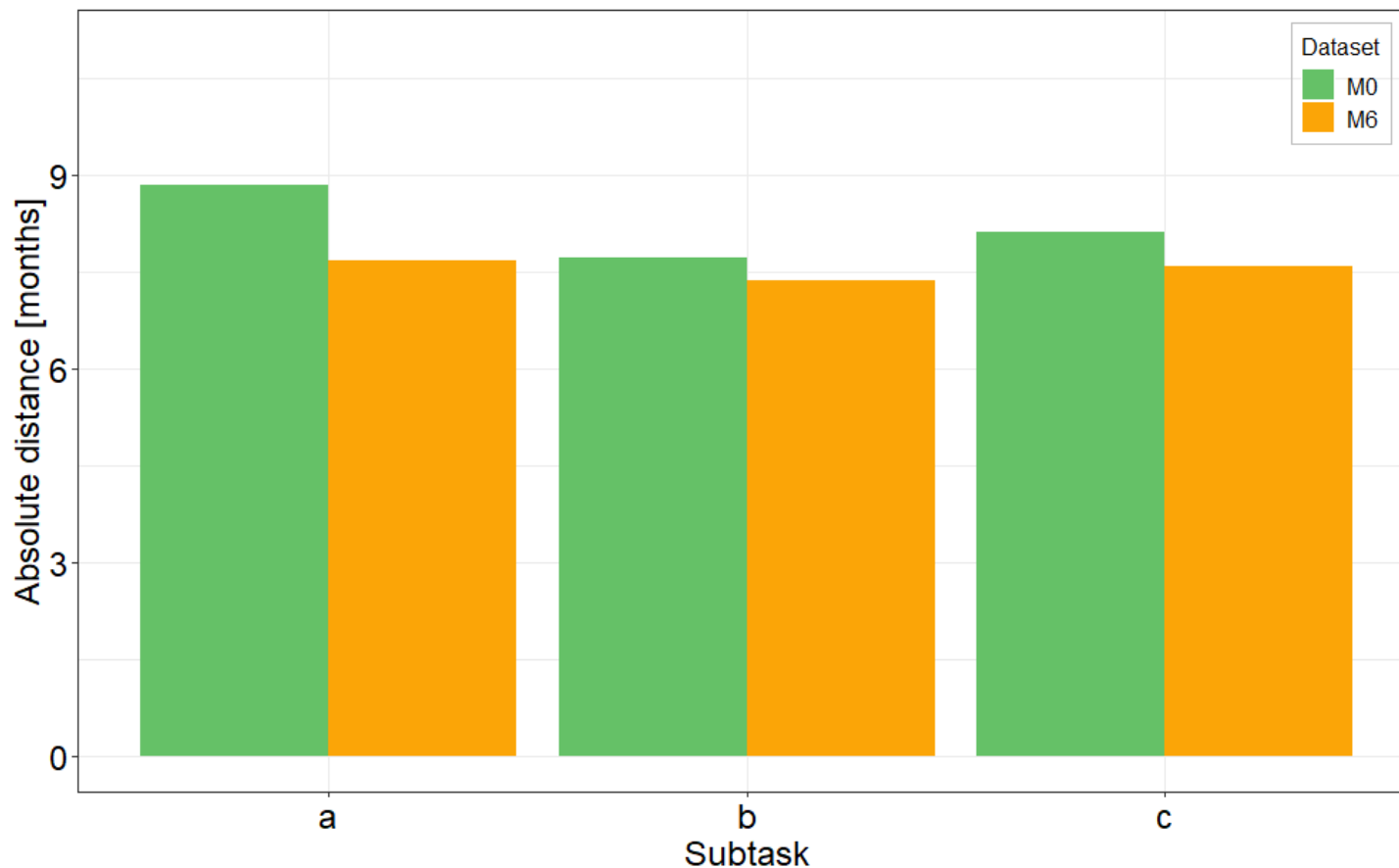
In the table: absolute distance [in months]



Task 2: results (1 of 3)

Absolute distance between real and predicted times is smaller when adding 6 months of follow-up

SSVM, subtask c - Dataset M0 vs dataset M6

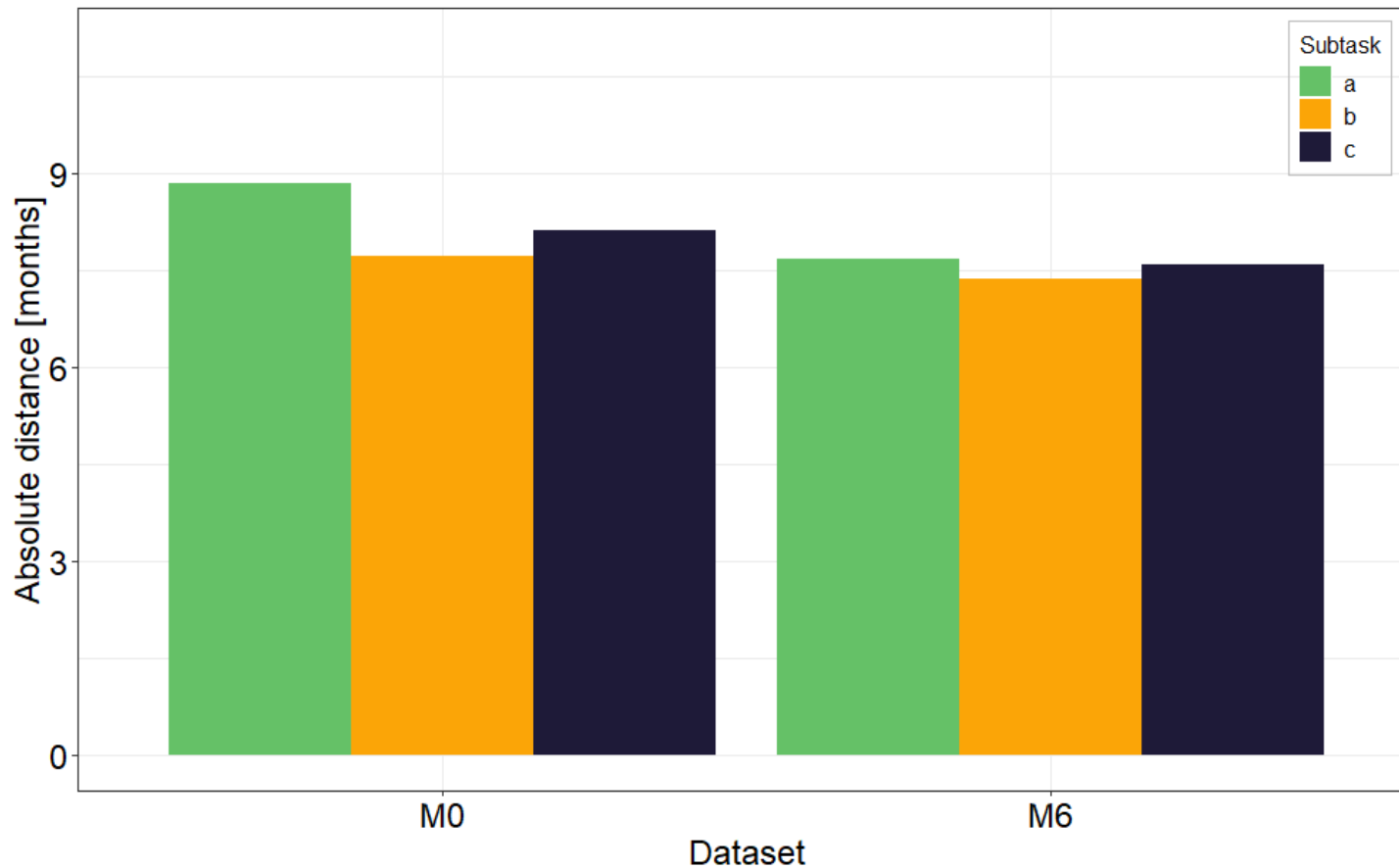




Task 2: results (2 of 3)

Better performance in subtasks b (PEG) and c (death)

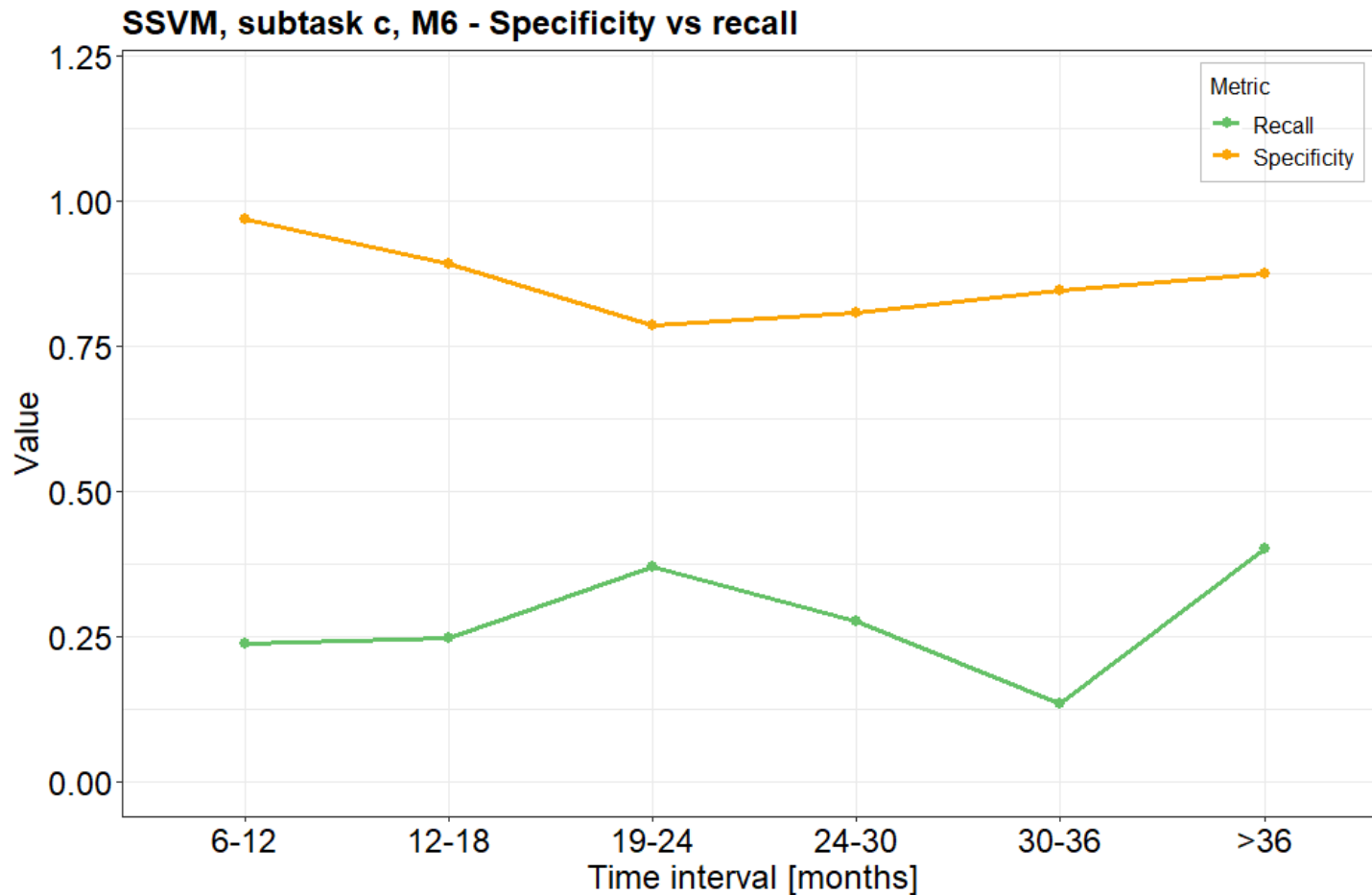
SSVM - Subtasks a, b, and c





Task 2: results (3 of 3)

High specificity, recall needs to be improved

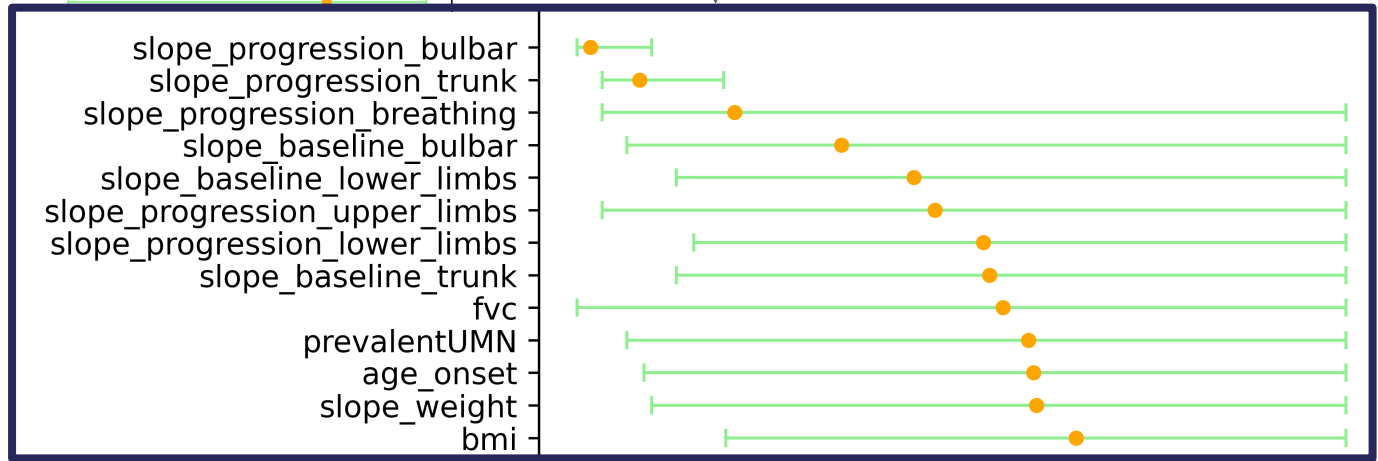
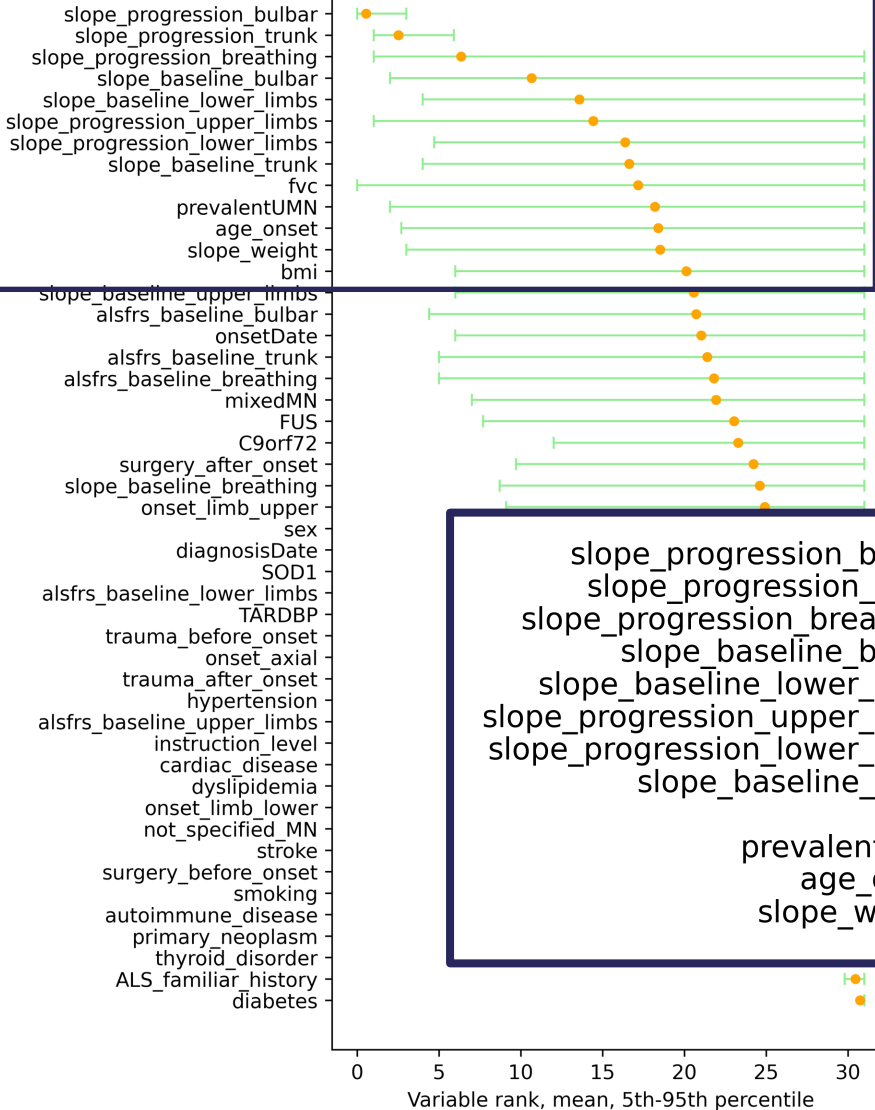




Variables ranking

Strongest predictors:

- ▷ ALSFRS-R slopes
- ▷ FVC
- ▷ Impairment in upper motor neuron
- ▷ Age at onset
- ▷ Weight variations
- ▷ High BMI





Baseline Machine Learning Approaches to Predict Amyotrophic Lateral Sclerosis Disease Progression **SBB Team**

- ▶ For each subtask and model used, performance improved when dynamic features were added as predictors.
- ▶ Predicting NIV seems to be more difficult than predicting PEG or death.
- ▶ Patient's stratification may lead to more accurate predictions.
- ▶ ALSFRS-R slopes, FVC values, and age at onset are among the strongest predictors of ALS clinical events.

Thank You



Isotta Trescato, PhD student
isotta.trescato@unipd.it

SBB team, UNIPD

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 [@sysbiobigunipd](https://twitter.com/sysbiobigunipd)