

## **Disentangling Blink Reflexes in Multiple Sclerosis with machine learning** techniques

## M. Biggio<sup>1</sup>, D. Caligiore<sup>2,3</sup>, M. Merone<sup>4</sup>, F. D'Antoni<sup>4</sup>, M. Bove<sup>5-6</sup>, L. Bonzano<sup>1-6</sup>

<sup>1</sup>Department of Neuroscience, Rehabilitation, Ophthalmology, Genetics, Maternal and Child Health, University of Genoa, Genoa, Italy <sup>2</sup>Computational and Translational Neuroscience Laboratory, Institute of Cognitive Sciences and Technologies, National Research Council (CTNLab-ISTC-CNR), Rome, Italy. <sup>3</sup>AI2Life s.r.l., Innovative Start-Up, ISTC-CNR Spin-Off, Rome, Italy. <sup>4</sup>Research Unit of Computer Systems and Bioinformatics, Department of Engineering, Università Campus Bio-Medico di Roma, Rome, Italy. <sup>5</sup>Department of Experimental Medicine, Section of Human Physiology and Centro Polifunzionale di Scienze Motorie, University of Genoa, Genoa, Italy. <sup>6</sup>IRCCS Ospedale Policlinico San Martino, Genoa, Italy.



22/09 09:00 - 10:30 Artificial intelligence and Machine learning in digital health



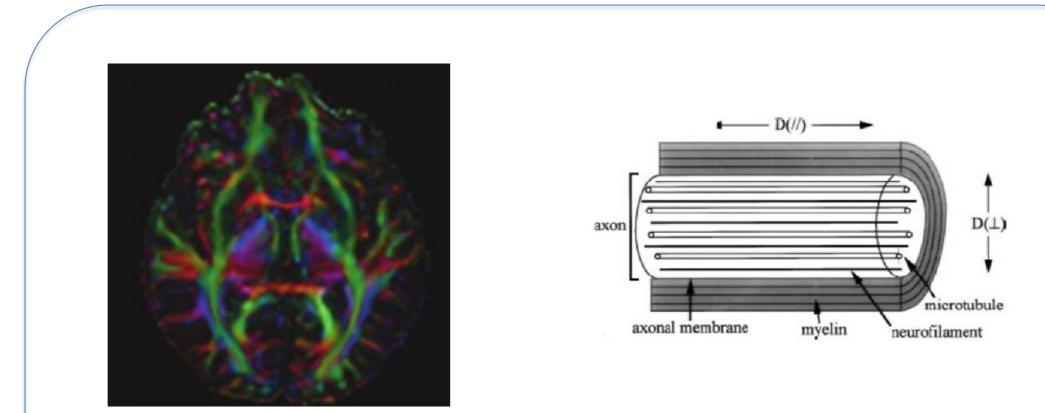
Brainstem dysfunctions are common in Multiple Sclerosis (MS) and are a critical predictive factor for future disability<sup>1</sup>. Brainstem functionality can be explored with blink reflexes, i.e., subcortical responses consisting in a blink following peripheral stimulation. Hand blink reflex (HBR)<sup>2,3</sup>, differently from other reflexes like the trigeminofacial blink reflex (TBR)<sup>4</sup>, is modulated by the proximity of the stimulated hand to the face. Despite brainstem dysfunctions are very common in MS, HBR was never been investigated in people with MS (PwMS). Due to its sensitivity to a top-down modulation, the study of HBR in PwMS could allow investigating the activity of the cortico-bulbar circuits and provide information related to functionality of the corticalbrainstem pathways.

AIM: Main goal was to investigate HBR response, its relationship with the motor and associative cortical regions in PwMS with a relapsing-remitting multiple sclerosis course and its possible alteration due to the disease.

- 20 PwMS

- 20 Healthy Controls

Reflexes were recorded by means of two MP100 BIOPAC EMG channels from the orbicularis oculi



We explored neurophysiological data with Machine Learning (ML) techniques.

- Datasets based on HBR and TBR features.

- Dichotomic classification(PwMS/Control)

muscles bilaterally.

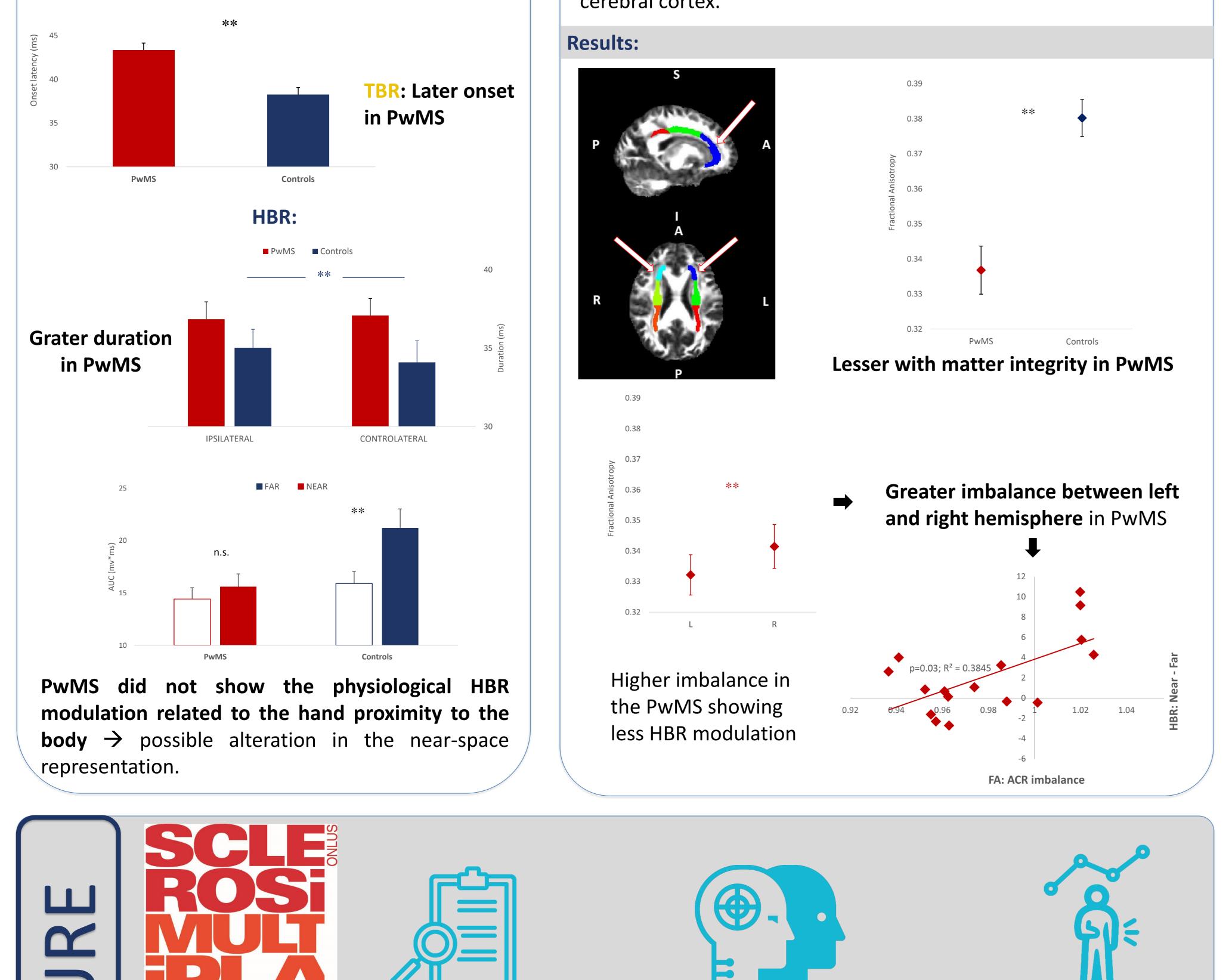
TBR was evoked by administering transcutaneous electrical stimuli to the supraorbital branch of the trigeminal nerve.





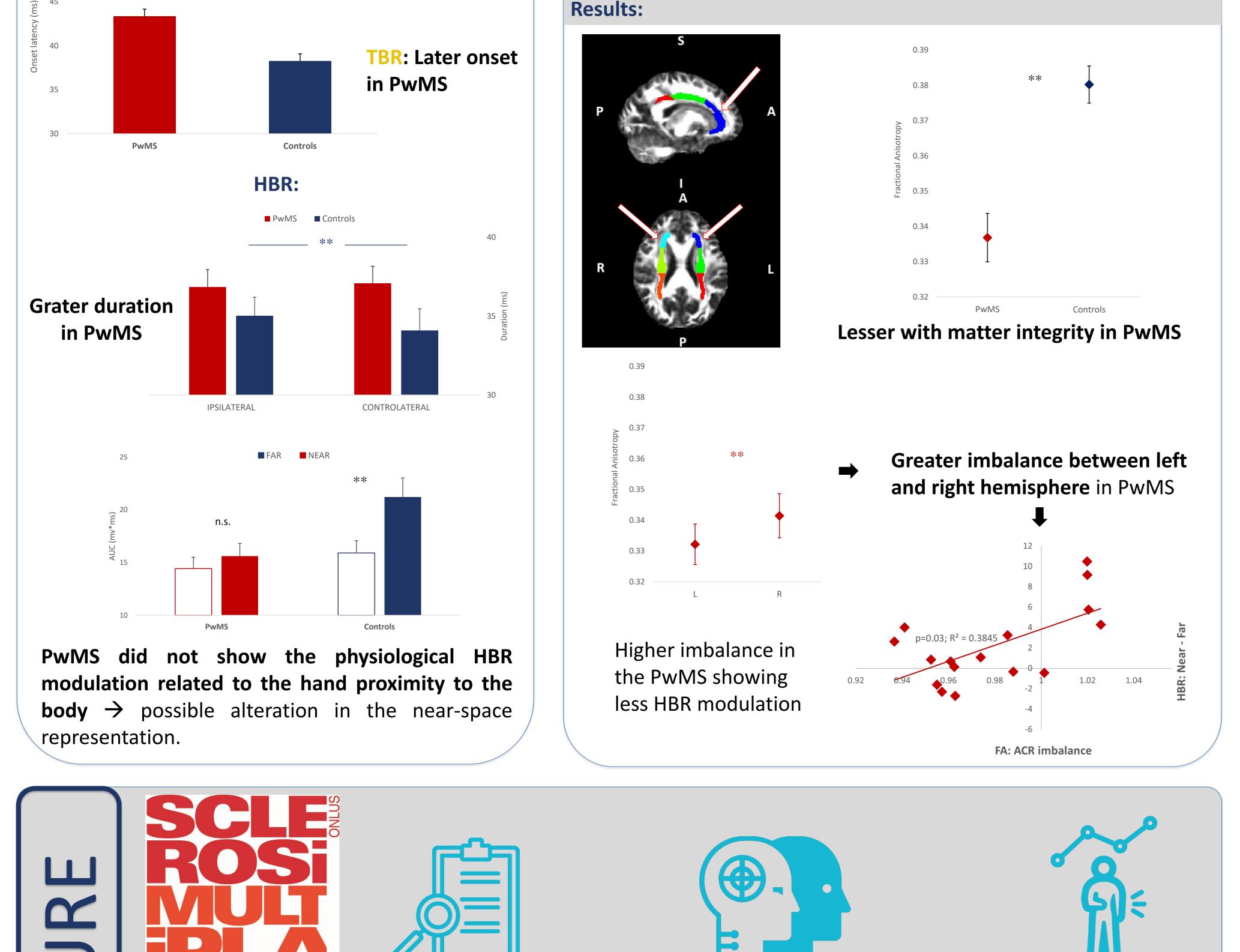
HBR was evoked by administering transcutaneous electrical stimuli to the median nerve. HBR was elicited bilaterally in two conditions, NEAR and FAR, with the stimulated hand of the subject respectively near and far from the face.

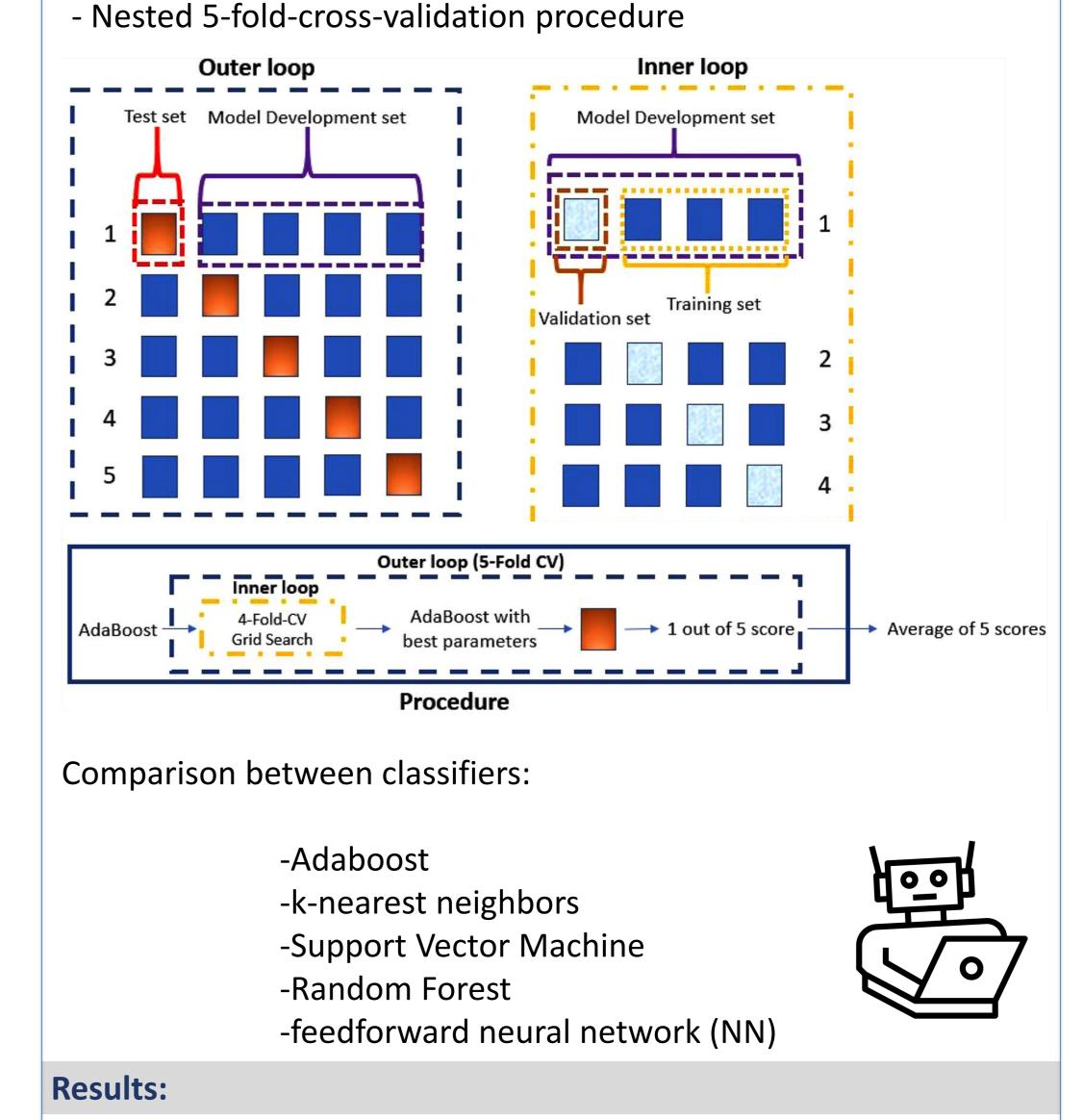
**Results:** 



To investigate microstructural integrity of the cortico-brainstem pathway in association with reflexes features, participants underwent Diffusion Tensor Imaging (DTI: 3 Tesla, 64 non collinear directions; b=1000 s/mm<sup>2</sup>).

By using FSL-FDT we obtained parametric maps of fractional anisotropy, mean diffusivity, axial diffusivity, and radial diffusivity. The mean value of each parameter was calculated on each scan as indication of tissue integrity in the whole white matter (**WM**) and in different white matter tracts, with special focus on the corona radiata (**CR** – anterior, posterior and superior), a bundle of ascending and descending fibers connecting the brainstem to the cerebral cortex.





AdaBoost classifiers showed the highest accuracy for both datasets.

Class	Metrics	HBR	TBR
	Accuracy [%]	86.7	73.3
	Recall [%]	88.3	71.7
PwMS	Precision [%]	91.0	82.7
	F1-Score [%]	89.6	76.8
	Recall [%]	83.3	76.7
Control	Precision [%]	88.3	74.7
	F1-Score [%]	85.8	75.7
mua nacitina truca	nagatina	<b>D</b>	true po
rue positive+true 1 N Recall · Precision		Docall	true po ie positive + true positi
		tri Docelli	ie positive +



specific DESCRIBE HBR in alteration of PwMS with ML in relation with clinical parameters on 100 PwMS and 100

Controls.

Extract the most relevant features of the reflex and virtually **REPLICATE** subject responses.

patient's MONITOR after 18 response months, with the goal to identify biomarkers reflecting pathology.



Machine learning for exploring neurophysiological functionality in multiple sclerosis based on trigeminal and hand blink reflexes.<sup>5</sup>

## Bibliography

- Habek, M. (2014). Evaluation of brainstem involvement in multiple sclerosis. *Expert Review of Neurotherapeutics*
- Sambo C.F., Forster B., Williams S.C., Iannetti G.D. (2012). To Blink or Not to Blink: Fine Cognitive Tuning of the Defensive Peripersonal Space. Journal of Neuroscience.
- Bisio A.\*, Garbarini F.\*, Biggio M., Fossataro C., Ruggeri P., Bove M. (2017). Dynamic shaping of the defensive peripersonal space through predictive motor mechanisms: when the "near" becomes "far". Journal of Neuroscience
- Magnano I, Pes G.M., Pilurzi G., Cabboi M.P., Ginatempo F., Giaconi E., et al. (2014) Exploring brainstem reflexes, evoked potentials, clinical and MRI investigations. International Federation of Clinical Neurophysiology
- Biggio M., Caligiore D., D'Antoni F. Bove M., Merone M. (2022). Machine learning for exploring neurophysiological functionality in multiple sclerosis based on trigeminal and hand blink reflexes. Scientific Reports

