

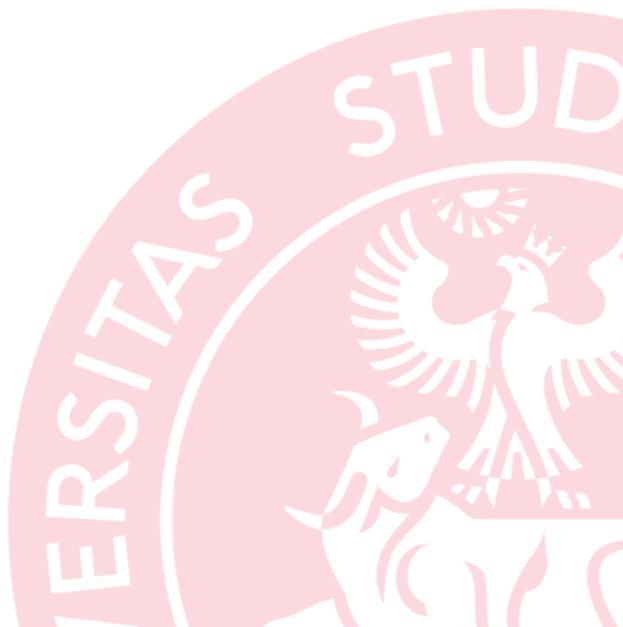


# Integrating Anatomical Information in Weakly Contrastive Learning for Neuroimaging

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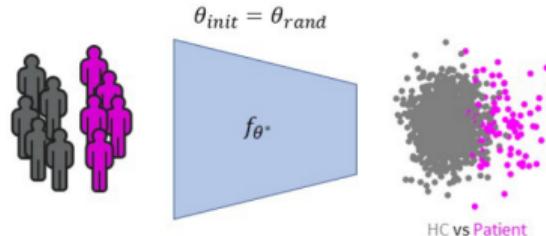
# Deep Learning in Neuroimaging



- ▶ Deep learning has demonstrated remarkable results across various fields that benefit from **automated image analysis**, including neuroimaging
- ▶ Neuroimaging concerns the acquisition and analysis of **brain images**
- ▶ Neurobiological characterization of psychiatric and neurological disorders encompasses clinical, biological and environmental factors
- ▶ **Limited** relative size of neuroimaging datasets, especially related to specific neurological conditions

# Transfer Learning

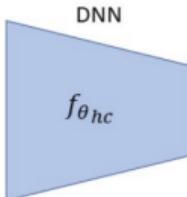
## Old Paradigm: supervised learning from scratch



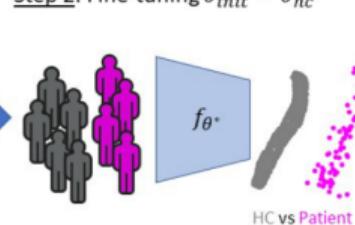
## Our New Paradigm

### Step 1: Pre-training

$N = 10K$



### Step 2: Fine-tuning $\theta_{init} = \theta_{hc}$

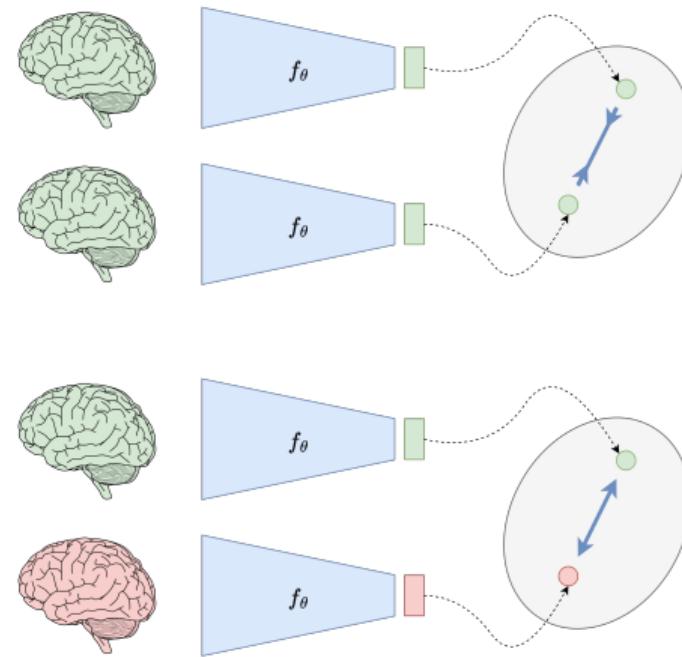


Credits: Dufumier et. al, 2024

# Contrastive Learning

- ▶ **Minimize** the distance between positive pairs of embeddings
- ▶ **Maximize** the distance between negative pairs of embeddings

$$\mathcal{L}_{i,j} = -\log \left( \frac{\exp(s_{i,j}/\tau)}{\sum_{a \in A(i)} \exp(s_{i,a}/\tau)} \right)$$



# Weakly Supervised Contrastive Learning



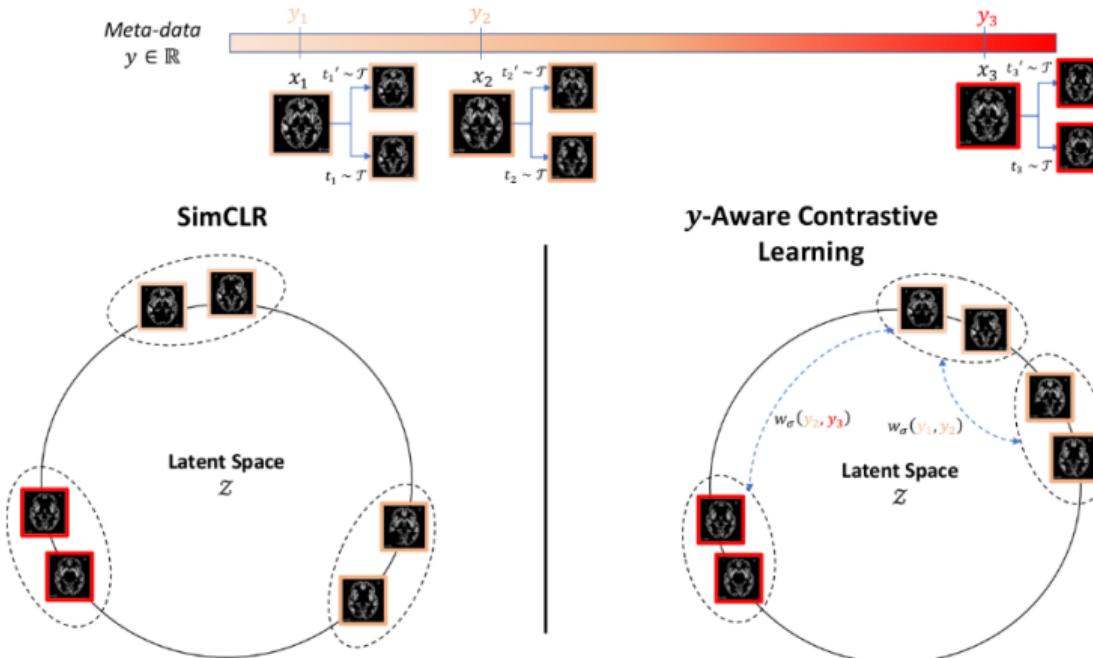
- ▶ Existing weakly contrastive learning methods augment the learning process with the **age attribute**
- ▶ These methods works by computing a **weight term** that increase the pushing/pulling strength between latent representations based on the closeness of the age attribute

$$\mathcal{L}^{\text{y-aware}} = - \sum_{k \in A(i)} \frac{w_k}{\sum_t w_t} \log \left( \frac{\exp(s_k/\tau)}{\sum_{a \in A(i)} \exp(s_a/\tau)} \right)$$

Where:

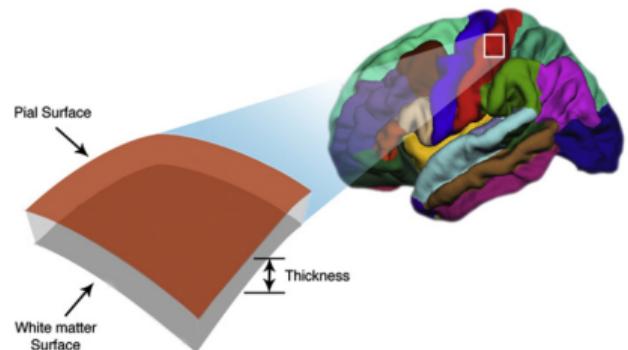
$$w_k = K_\sigma(y_i - y_k)$$

# Weakly Supervised Contrastive Learning

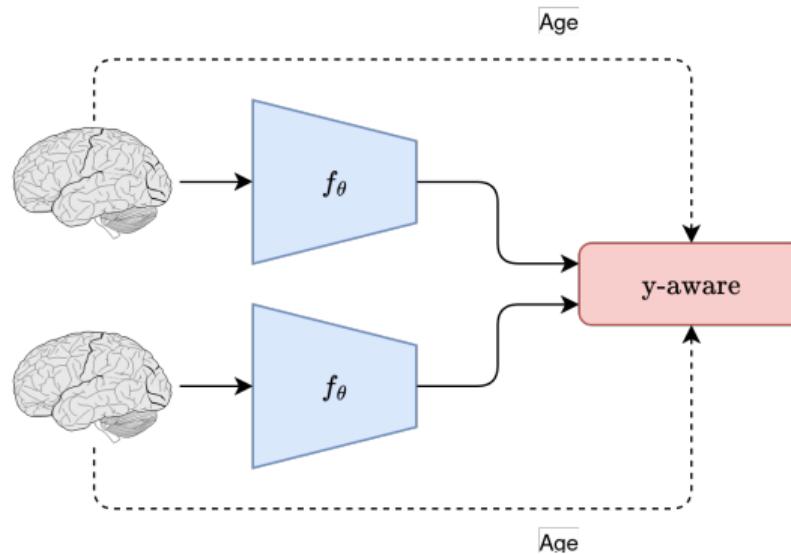


# Goal of this Thesis

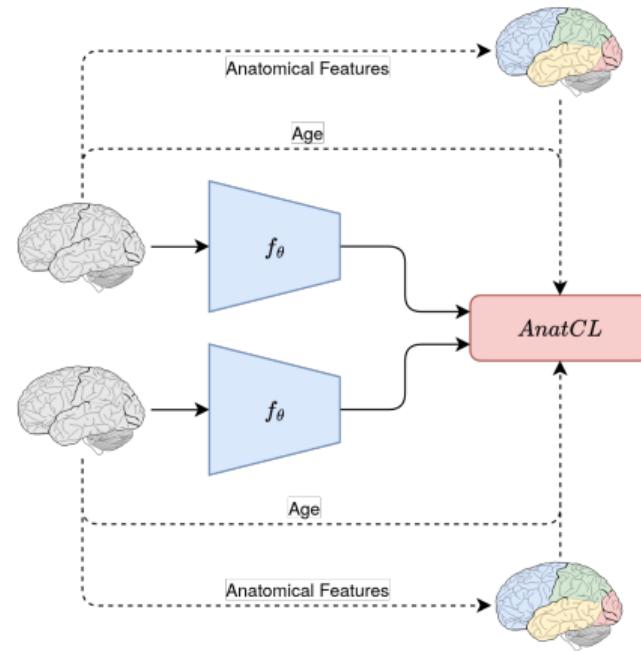
- ▶ Neuroimaging datasets commonly include other **anatomical measures** along with **patient-specific informations** (age, sex, etc.)
- ▶ Leverage this data during the **pre-training** process to further enhance the learned representations
- ▶ This may in turn enhance the performance and **generalizability** of the model in various downstream tasks



# Current Approach

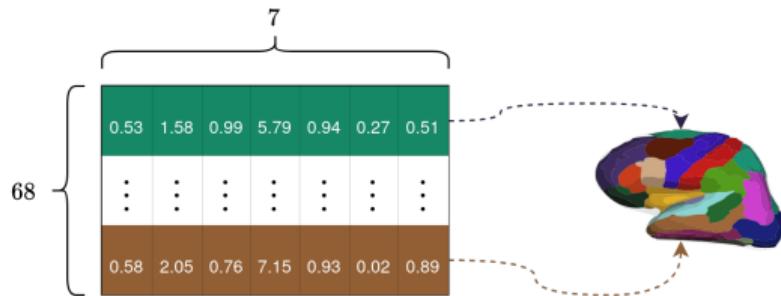


# Proposed Approach



# Integrating Anatomical Information

- ▶ For each  $i^{th}$  patient, its **anatomical measures** are stored in a matrix  $\mathcal{D}^i \in \mathbb{R}^{68 \times 7}$
- ▶ Each **row vector** contain 7 anatomical measures, corresponding to a specific **region of interest** out of the total 68



## Anatomical Measures:

- ▶ Average Cortical Thickness
- ▶ Standard Deviation Cortical Thickness
- ▶ Gray Matter Volume
- ▶ Total Surface Area
- ▶ Integrated Mean Curvature
- ▶ Gaussian Curvature
- ▶ Intrinsic Curvature Index

# Proposed Method: Local Descriptor

- ▶ Define the **similarity** between two subjects as the expected cross-similarity between local measurements vectors
- ▶ A **normalization**  $\gamma(x)$  must be applied to normalize **each component** of the vectors to a common range

$$w_k = \frac{1}{68} \sum_{n=1}^{68} sim(\gamma(\mathcal{D}_n^i), \gamma(\mathcal{D}_n^k))$$

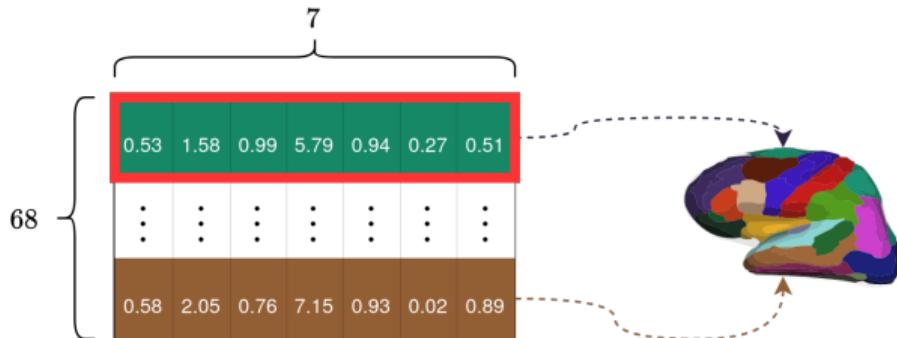
# Proposed Method: Global Descriptor



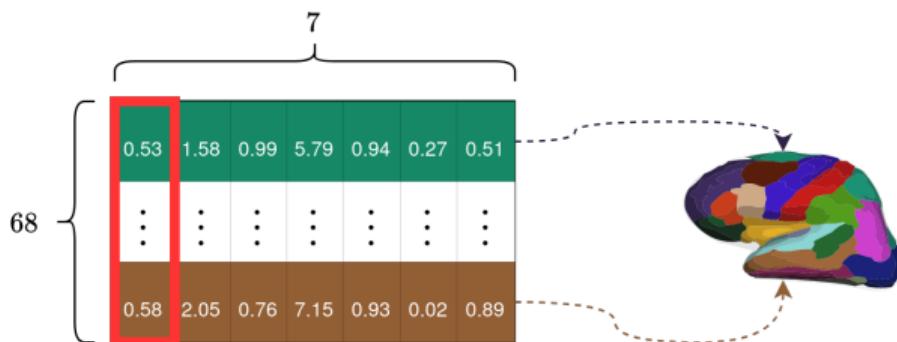
- ▶ Rather than viewing the data as 68 measurement vectors one can consider the format as 7 **feature vectors**, each composed of 68 measurement values
- ▶ Let  $\omega_n^i = (\mathcal{D}^i)_n^T$  be the  $n^{th}$  row vector of the **transposed measurements matrix** associated to the  $i^{th}$  patient

$$w_k = \frac{1}{7} \sum_{n=1}^7 sim(\omega_n^i, \omega_n^k)$$

# Descriptors Summary



Local Descriptor



Global Descriptor

# Final Formulation

- ▶ To incorporate also the age attribute into the final objective loss, the resulting function is structured as a **weighted sum**
- ▶ This sum combines a weakly supervised loss that is augmented with the **age** attribute ( $\mathcal{L}^{\text{age}}$ ) and another weakly supervised loss that utilizes the **anatomical information** ( $\mathcal{L}^{\text{AnatCL}}$ )

$$L = \lambda_1 \mathcal{L}^{\text{age}} + \lambda_2 \mathcal{L}^{\text{AnatCL}}$$

# Experiments Overview

- ▶ The proposed method was compared with other **baselines** such as:
  - L1: Mean Absolute error using the age attribute
  - **SimCLR**: Unsupervised Contrastive (*Chen et. al, 2020*)
  - **y-aware**: Weakly Supervised Contrastive (*Dufumier et. al, 2021*)
  - **ExpW**: Weakly Supervised Contrastive (*Barbano et. al, 2023*)
- ▶ The **experimental setting** consisted in training a ResNet18 model for each evaluated method on a dataset of **healthy individuals**
- ▶ Each trained model was then tested on 20 different **downstream tasks** across 5 **different datasets** in a 5 fold cross-validation
- ▶ Experiments were run using the Leonardo CINECA HPC cluster on Nvidia A100 GPUs, with each experiment taking approximately 24 hours

# Experiments

Dataset	Task / Condition
<b>OpenBHB</b>	Age (HC) Sex
<b>ADNI</b>	Alzheimer's Disease sMCI vs pMCI
<b>OASIS-3</b>	Alzheimer's Disease
<b>SchizConnect</b>	Schizophrenia Broad Schizophrenia Strict Bipolar Disorder Schizoaffective
<b>ABIDE I</b>	Autism Aspergers PDD-NOS

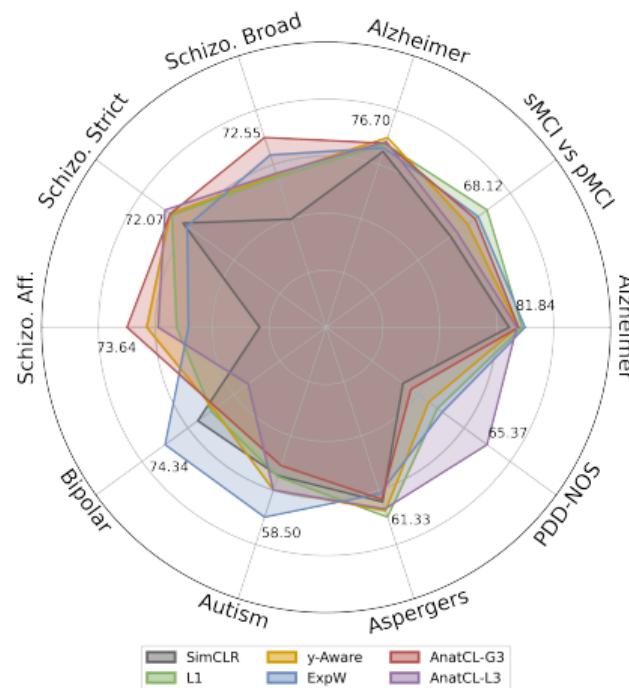
Dataset	Phenotype
<b>SchizConnect</b>	AIMS Overall Severity AIMS Upper Body AIMS Lower Body Depression Handedness SAS GAIT
<b>ABIDE I</b>	Handedness FIQ (WASI) VIQ (WASI) PIQ (WASI)

# Results

## Brain Age Prediction

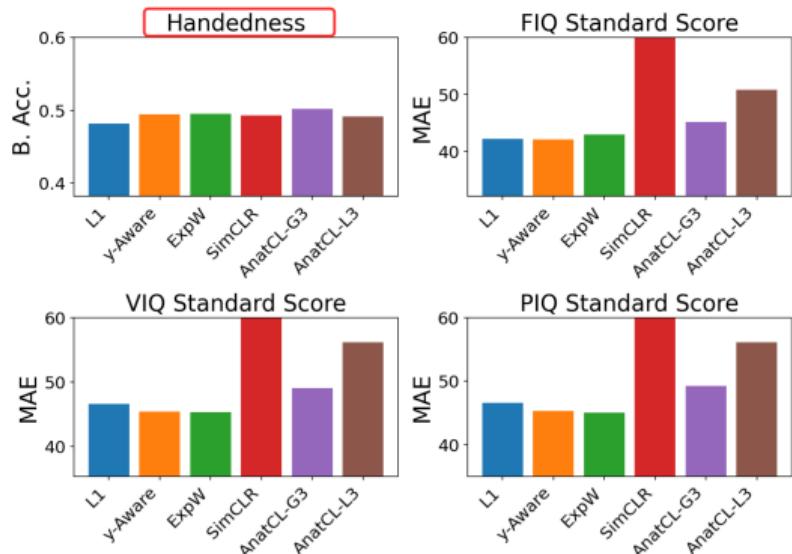
Method	Age MAE	Sex
SimCLR	$5.58 \pm 0.53$	$76.7 \pm 1.67$
L1 (age sup.)	$2.73 \pm 0.14$	$76.7 \pm 0.67$
y-Aware	$2.66 \pm 0.06$	$79.6 \pm 1.13$
ExpW	$2.70 \pm 0.06$	<b><math>80.3 \pm 1.7</math></b>
AnatCL-G3	<b><math>2.61 \pm 0.08</math></b>	$78.2 \pm 1.25$
AnatCL-L3	$2.64 \pm 0.07$	$78.2 \pm 0.7$

## Clinical Downstream Tasks

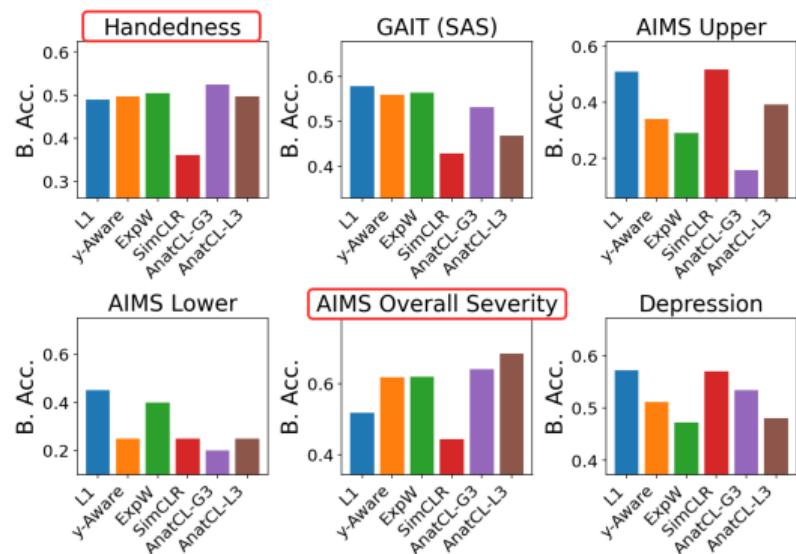


# Results: Phenotypes and Clinical Scores

Abide



SchizConnect



# Conclusions

- ▶ The proposed method consistently **outperforms unsupervised methods**, while reaching SOTA performances in some tasks
- ▶ This suggests that adding anatomical information may actually improve the **generalizability** of the learned representations
- ▶ This work has been **submitted** to the NeurIPS conference in May 2024 (*Barbano et. al, Anatomical Foundation Models for Brain MRIs, 2024*)
- ▶ Further research work is needed to enhance and refine these methods

# Limitations and Future Developments



- ▶ The age attribute **may not** be always present in every neuroimaging dataset
- ▶ Novel methods may integrate **only** anatomical measures without using the age. Those measures can be obtained from pre-processing pipelines common in neuroimaging, effectively making the methods fully unsupervised
- ▶ Additional methods to integrate **other anatomical measures** can be explored
- ▶ Develop **multi-modal models** to integrate other **acquisition methods** as well as data of different nature such as **textual data** coming from clinical records/assessments

# Thanks for your attention

*Any questions?*