

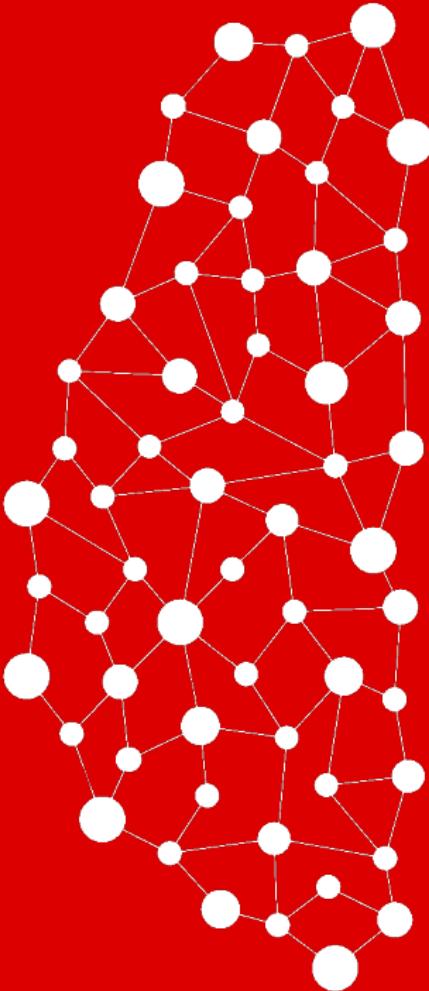
Exploring brain heterogeneity in neuropsychiatric patients with explainable artificial intelligence



Esten H. Leonardsen

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Outline

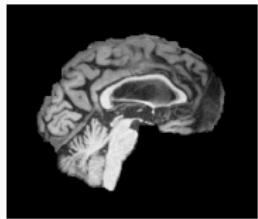
1. Background and motivation
2. Explainable dementia classification
3. Explainable brain age predictions



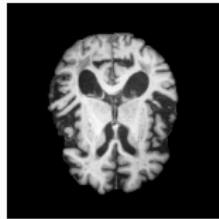
Background: Predictive clinical neuroscience



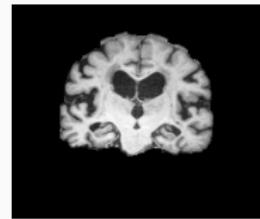
Background: Predictive clinical neuroscience



Side

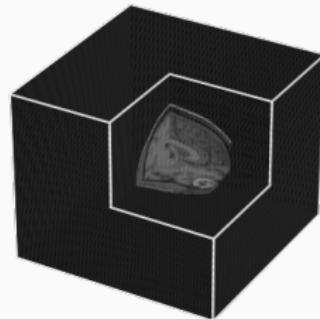


Above



Front

Structural Magnetic
Resonance Imaging (MRI) scans



Background: Predictive clinical neuroscience



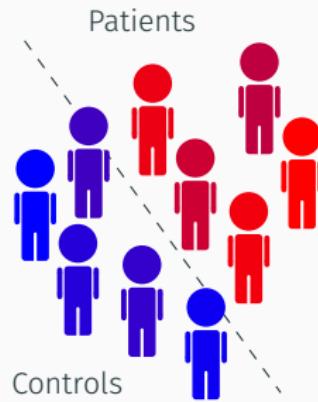
Background: Predictive clinical neuroscience



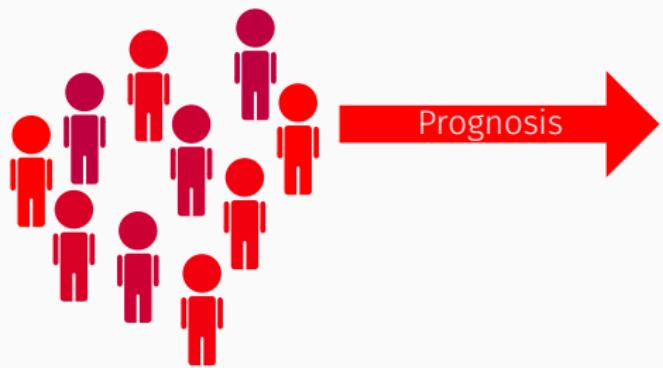
Background: Predictive clinical neuroscience



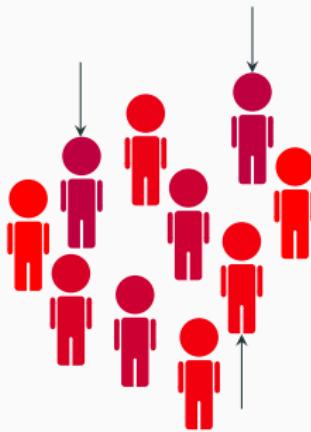
Background: Predictive clinical neuroscience



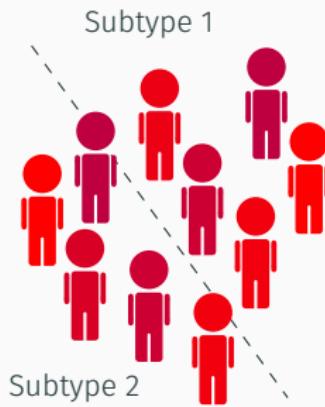
Background: Predictive clinical neuroscience



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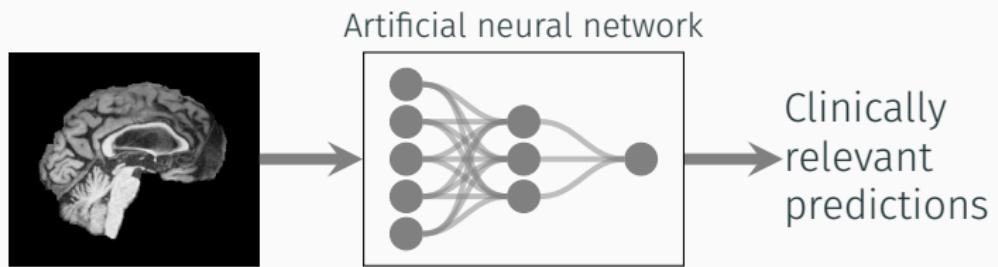
Background: Predictive clinical neuroscience



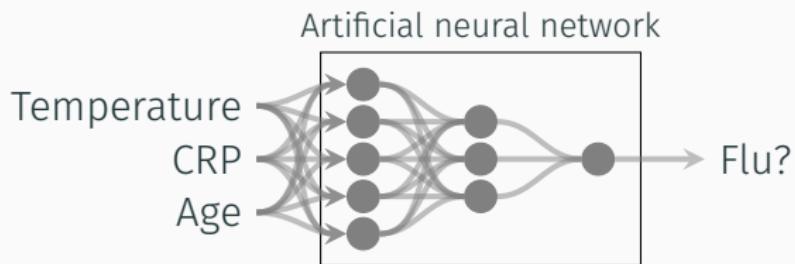
Background: Explainable artificial intelligence



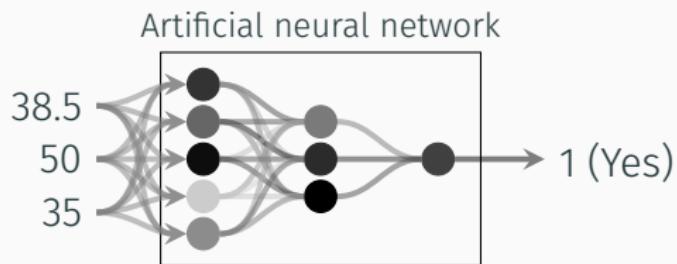
Background: Explainable artificial intelligence



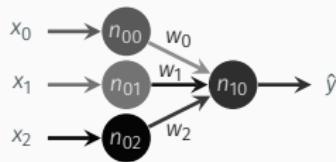
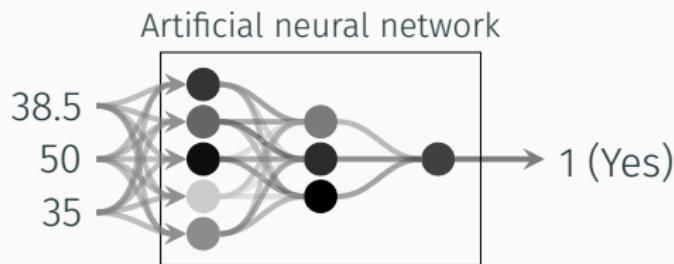
Background: Explainable artificial intelligence



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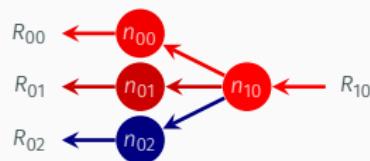
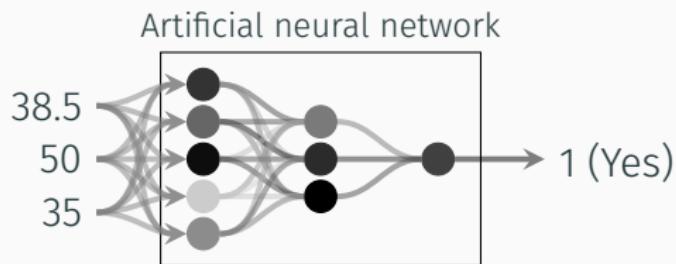
Background: Explainable artificial intelligence



$$\hat{y} = f \left(\sum_i^N w_i \cdot n_{0i} \right)$$



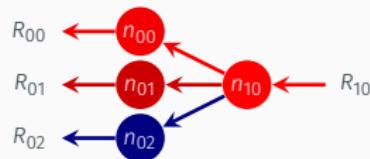
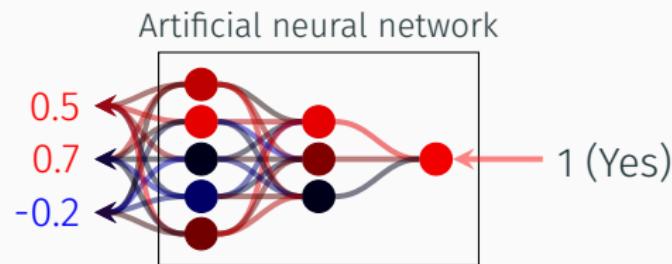
Background: Explainable artificial intelligence



$$R_{0i} = \sum_j \frac{n_{0i} w_i}{\sum_k n_{0k} w_k} R_{1j}$$



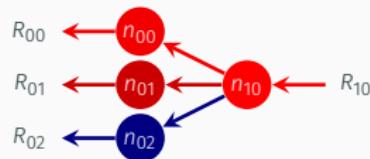
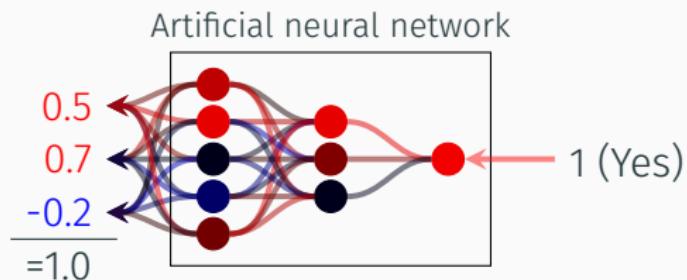
Background: Explainable artificial intelligence



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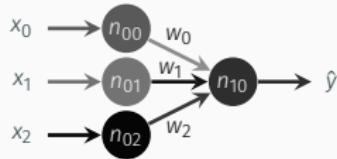
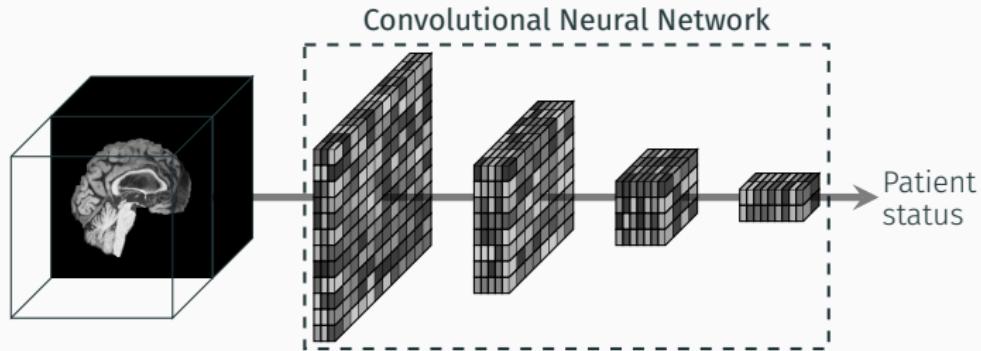
Background: Explainable artificial intelligence



$$R_{0i} = \sum_j \frac{n_{0i} w_i}{\sum_k n_{0k} w_k} R_{1j}$$



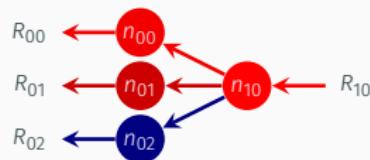
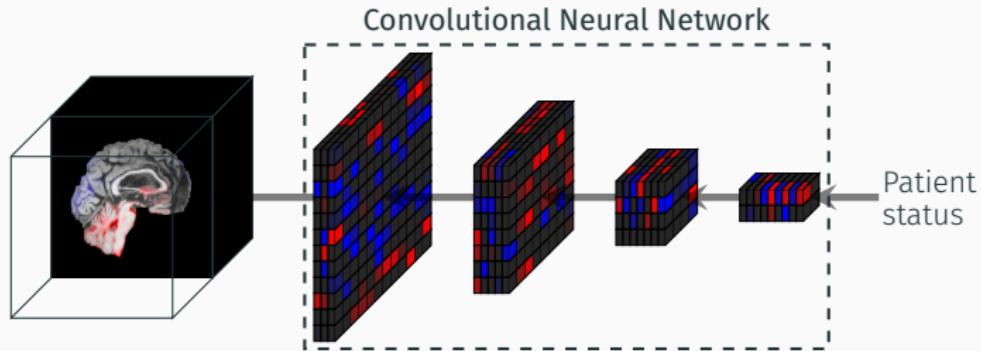
Background: Explainable artificial intelligence



$$\hat{y} = f \left(\sum_i^N w_i \cdot n_{0i} \right)$$



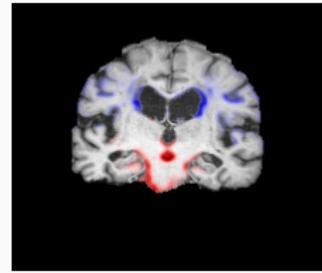
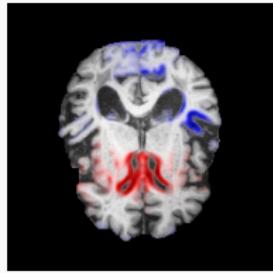
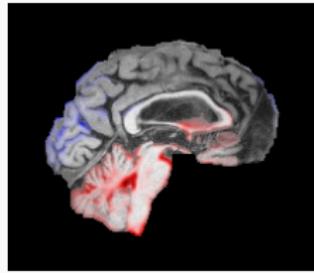
Background: Explainable artificial intelligence



$$R_{0i} = \sum_j \frac{n_{0i} w_i}{\sum_k n_{0k} w_k} R_{1j}$$



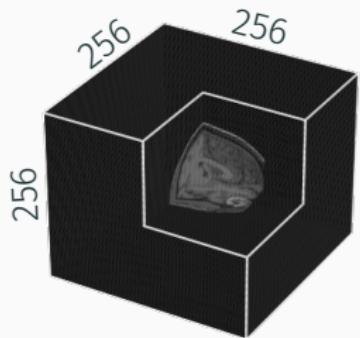
Background: Explainable artificial intelligence



Background: Conceptualization



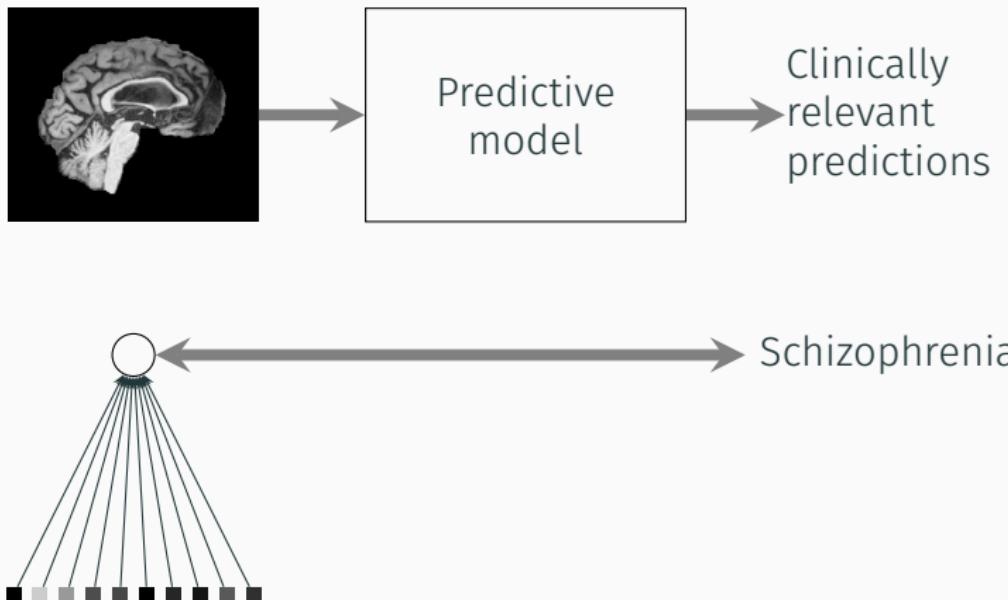
Background: Conceptualization



Background: Conceptualization



Background: Conceptualization



Background: Conceptualization

"Despite considerable heterogeneity at the regional level, deviations were often coupled to common functional circuits and networks"

Regional, circuit and network heterogeneity of brain abnormalities in psychiatric disorders
Segal, A. et al., *Nature Neuroscience* (2023)



Background: Conceptualization



Background: Conceptualization



Background: Conceptualization



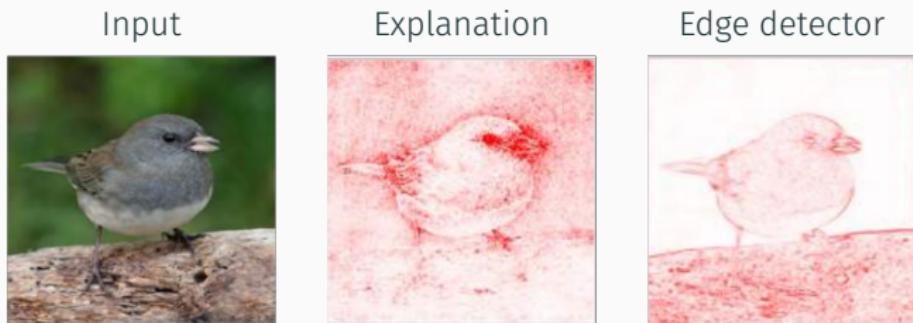
Background: Conceptualization

"We propose explanation space as a powerful embedding to facilitate detection of latent disease subtypes, [...]. Transforming from feature space to explanation space should not be seen in competition to dimensionality reduction, but rather as a complimentary processing step"

Inferring disease subtypes from clusters in explanation space
Schulz, MA. et al., *Scientific Reports* (2020)



Background: Conceptualization



Sanity Checks for Saliency Maps
Adebayo, J. et al., *preprint at arXiv* (2018)

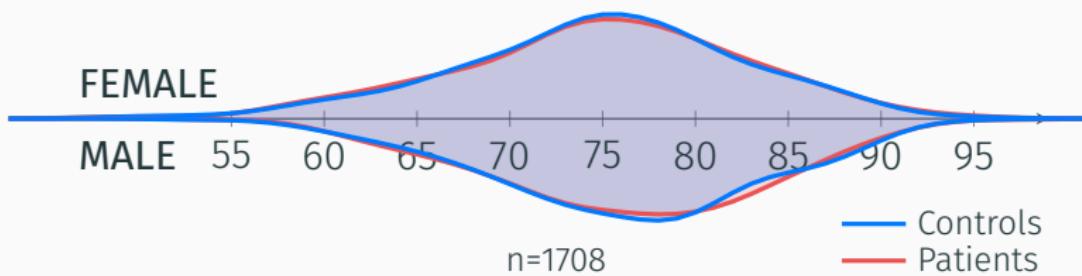


Explainable dementia classification

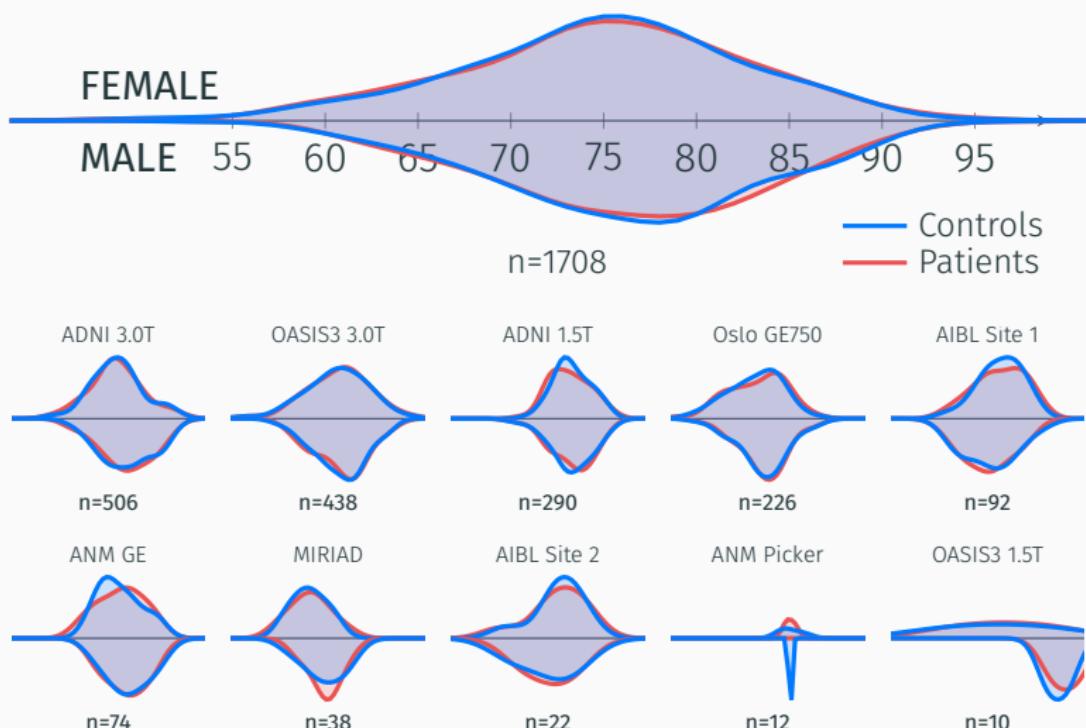


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Explainable dementia classification: Dataset



Explainable dementia classification: Dataset

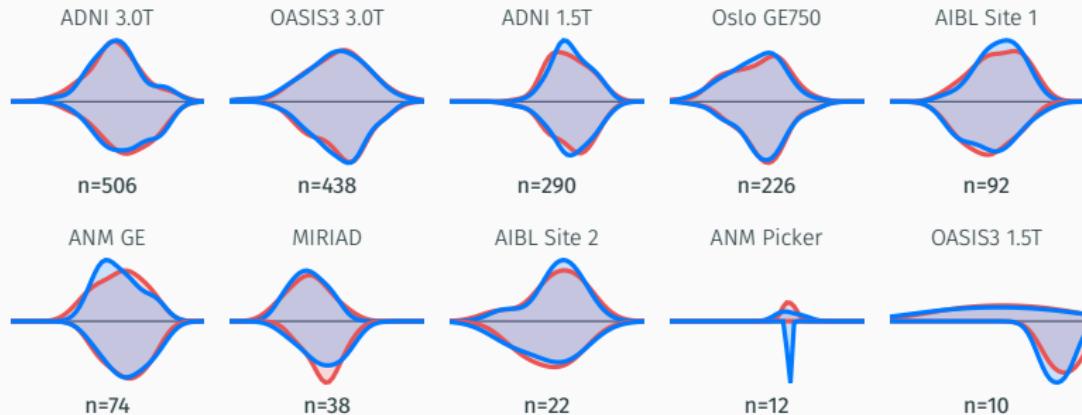


Explainable dementia classification: Dataset

ADNI:
Probable
Alzheimer's disease

OASIS3:
Dementia

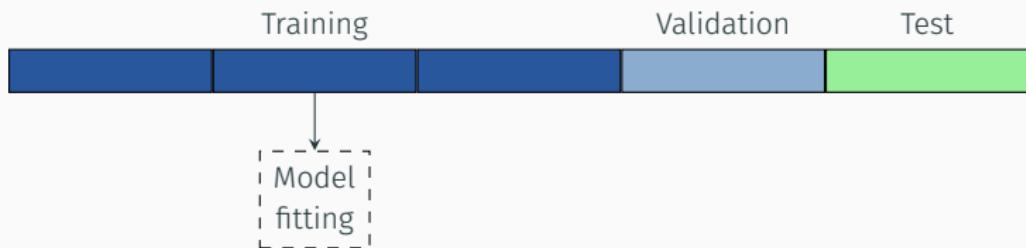
Oslo:
Alzheimer's disease,
Vascular dementia,
Unspecified dementia



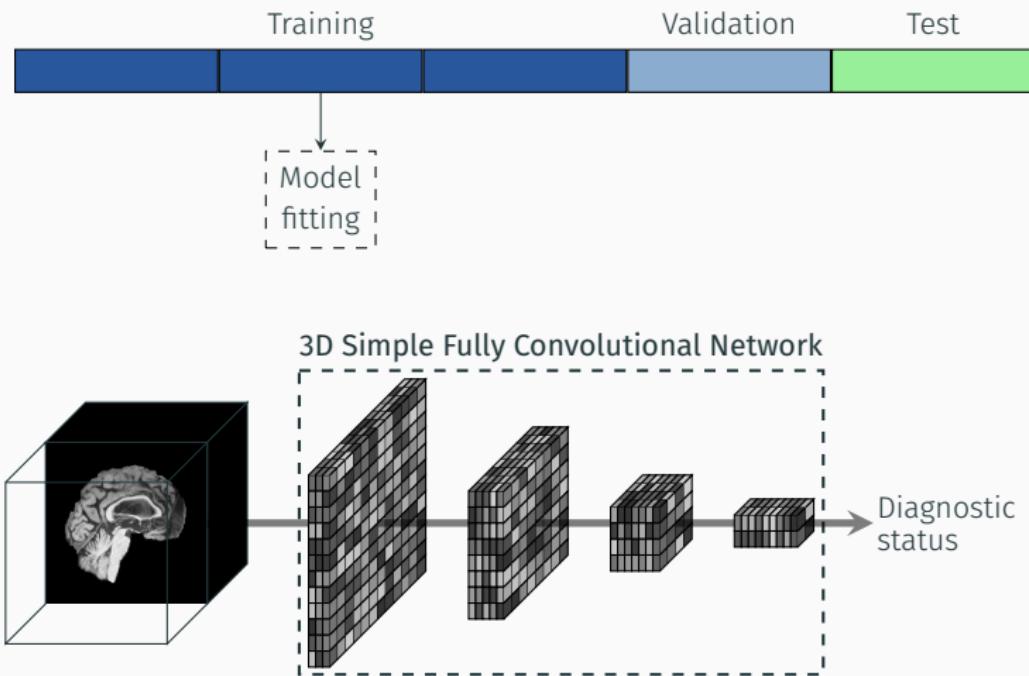
Explainable dementia classification: Modelling



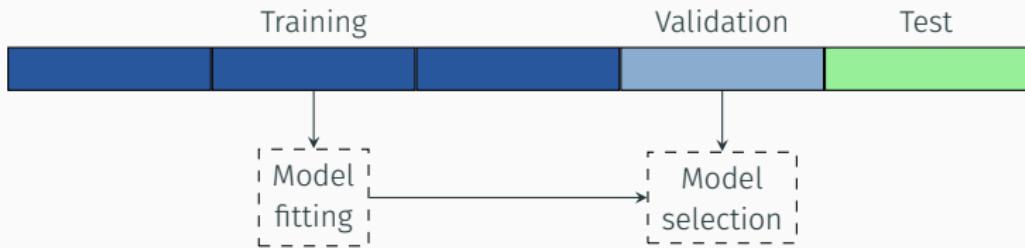
Explainable dementia classification: Modelling



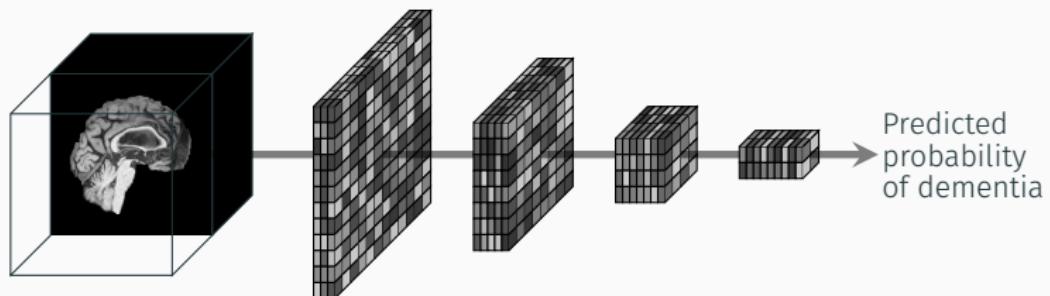
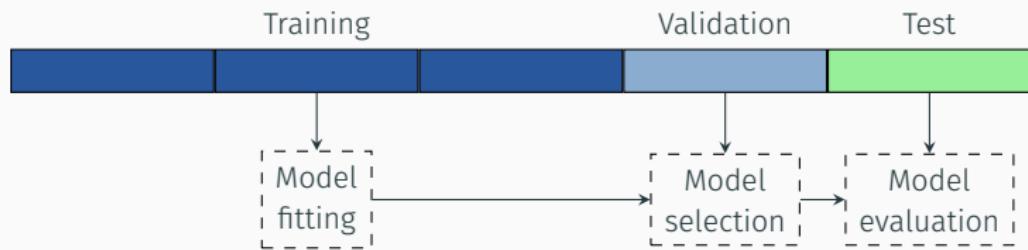
Explainable dementia classification: Modelling



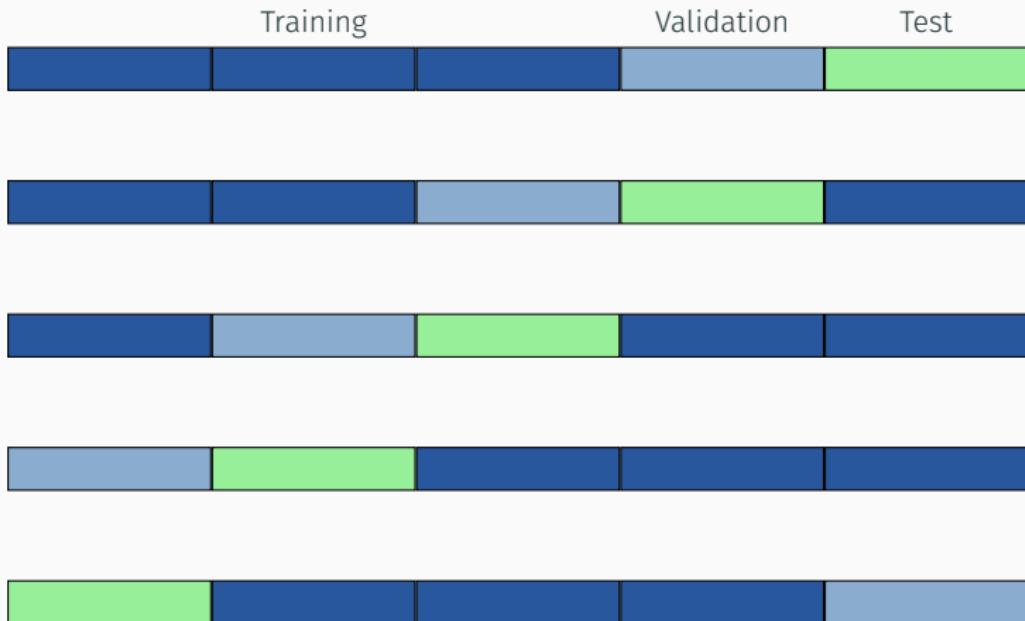
Explainable dementia classification: Modelling



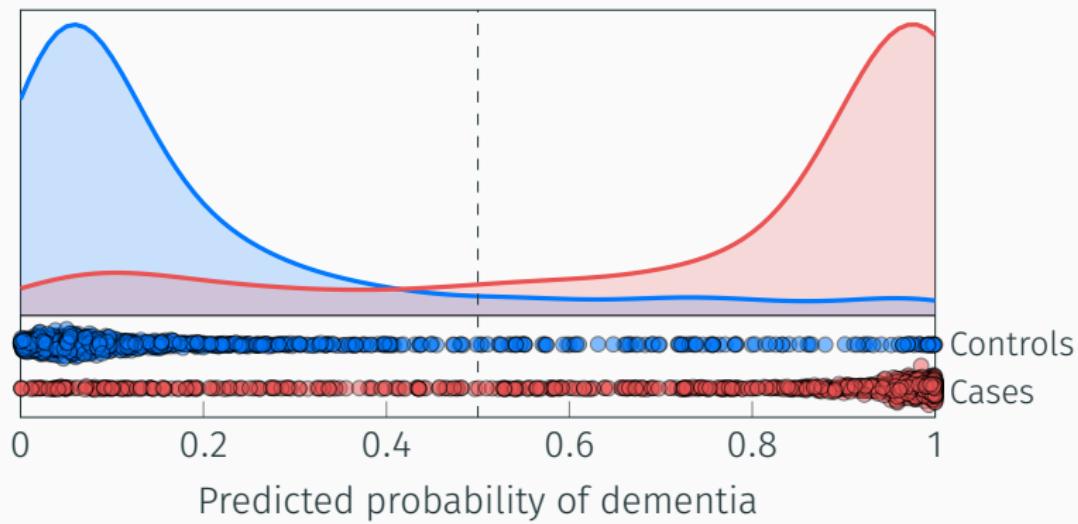
Explainable dementia classification: Modelling



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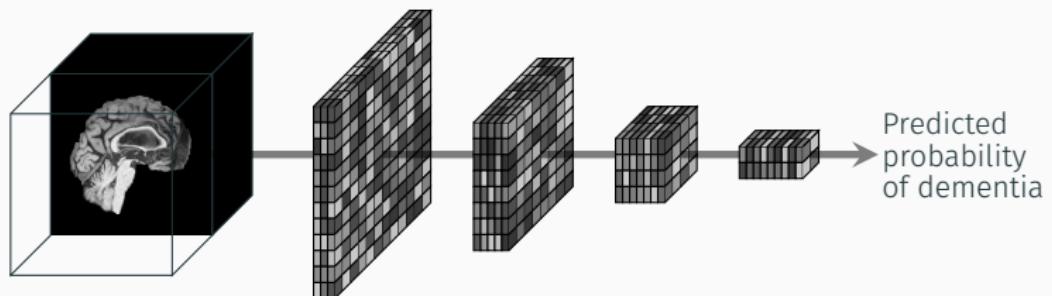
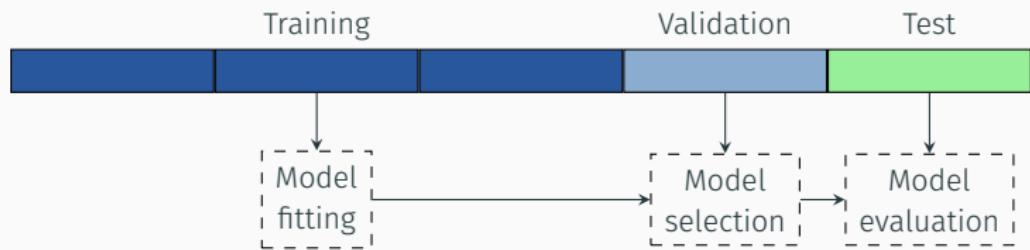


Explainable dementia classification: Modelling

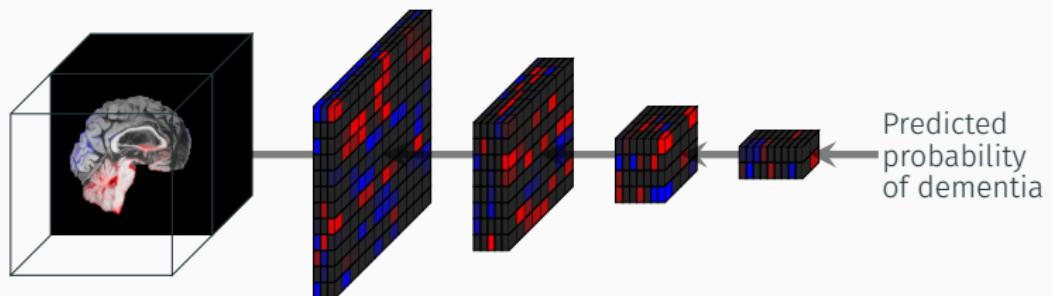
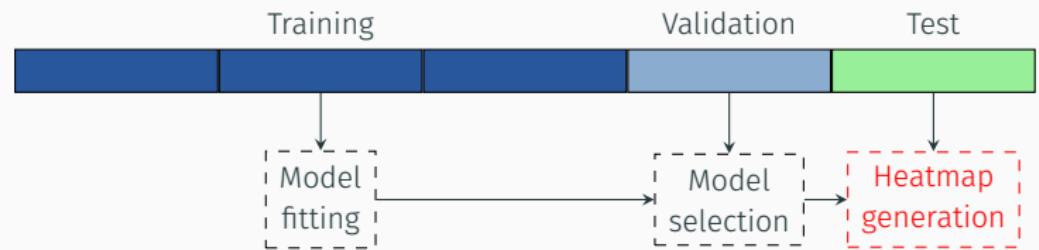


AUC=0.91, balanced accuracy=85%

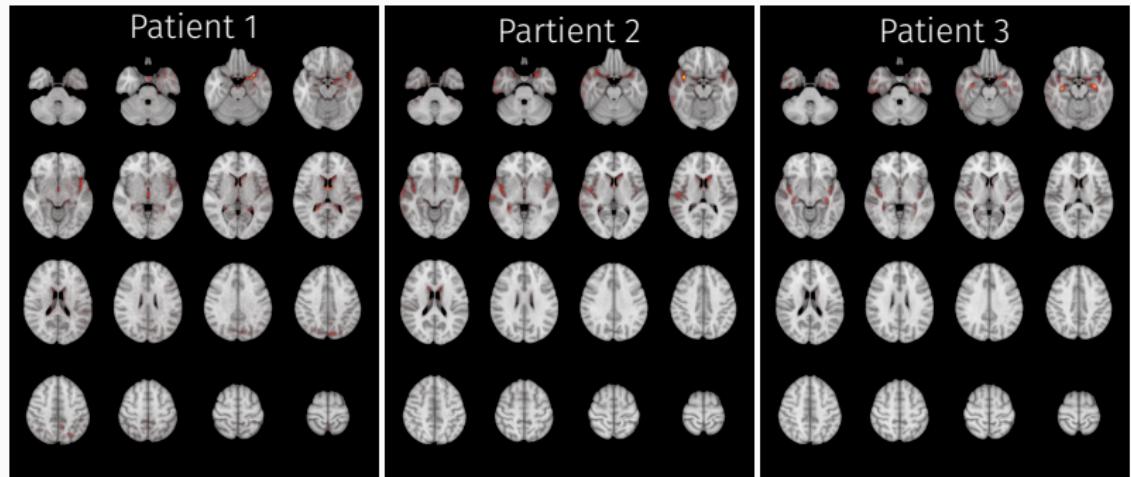
Explainable dementia classification: Modelling



Explainable dementia classification: Modelling

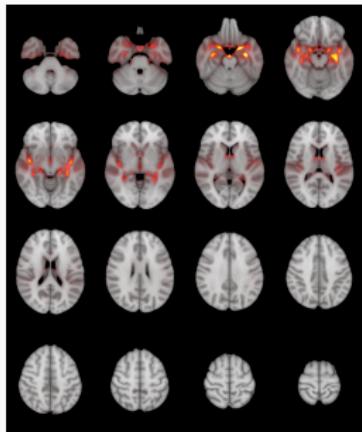


Explainable dementia classification: Validation



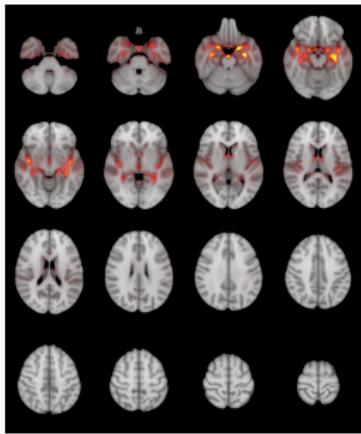
Explainable dementia classification: Validation

LRP

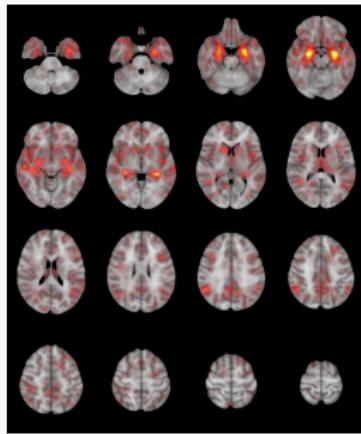


Explainable dementia classification: Validation

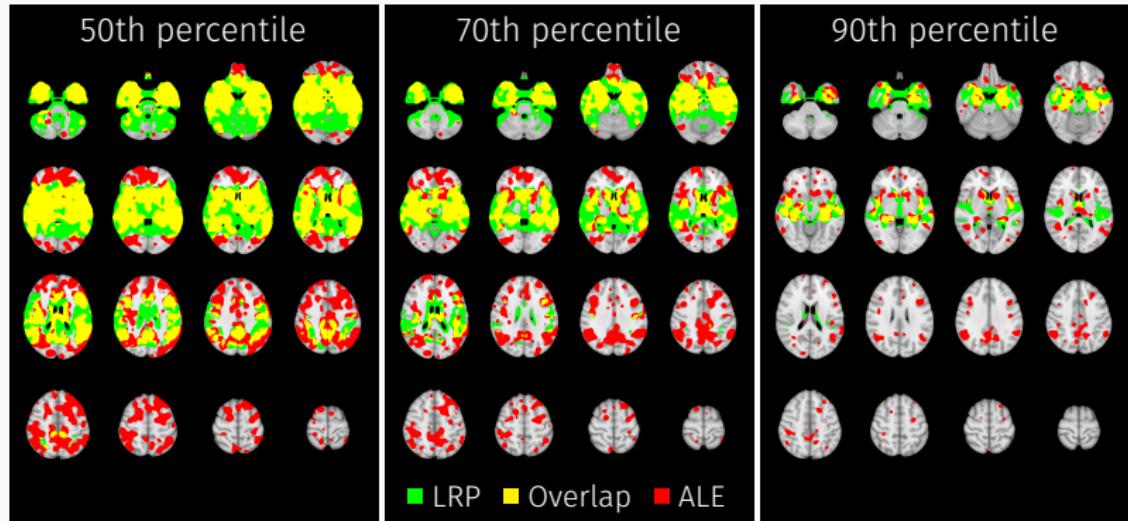
LRP



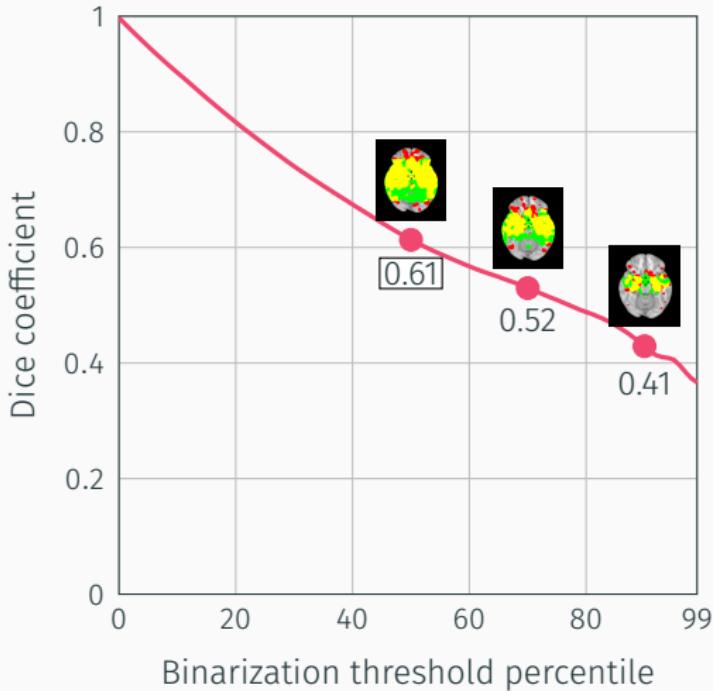
GingerALE



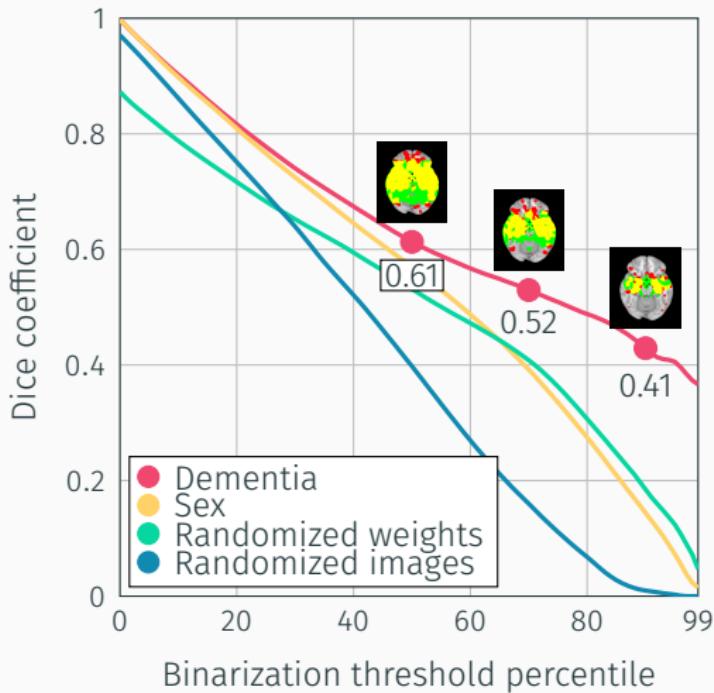
Explainable dementia classification: Validation



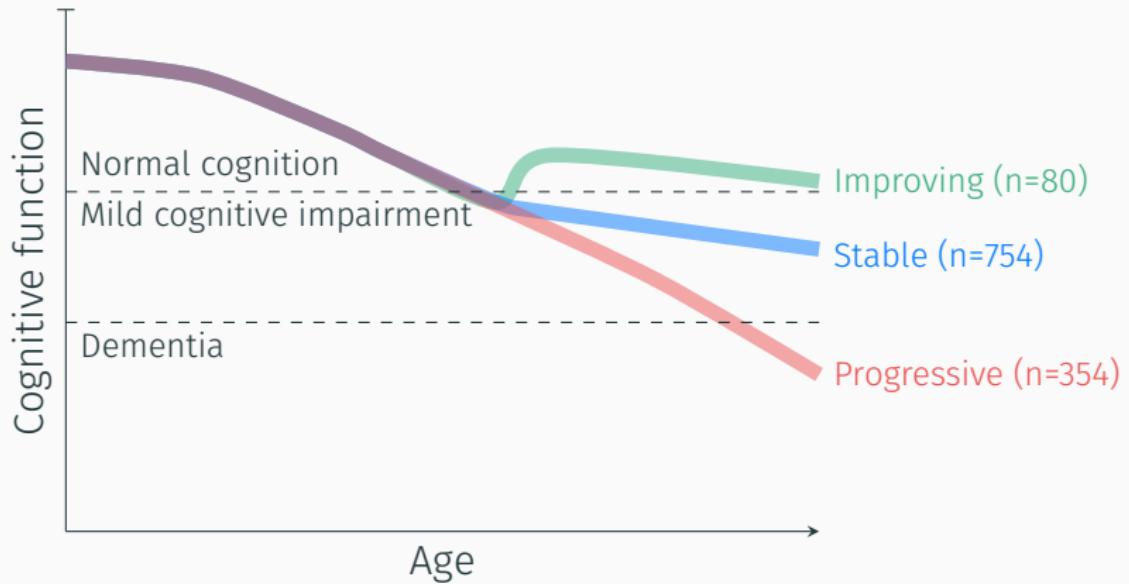
Explainable dementia classification: Validation



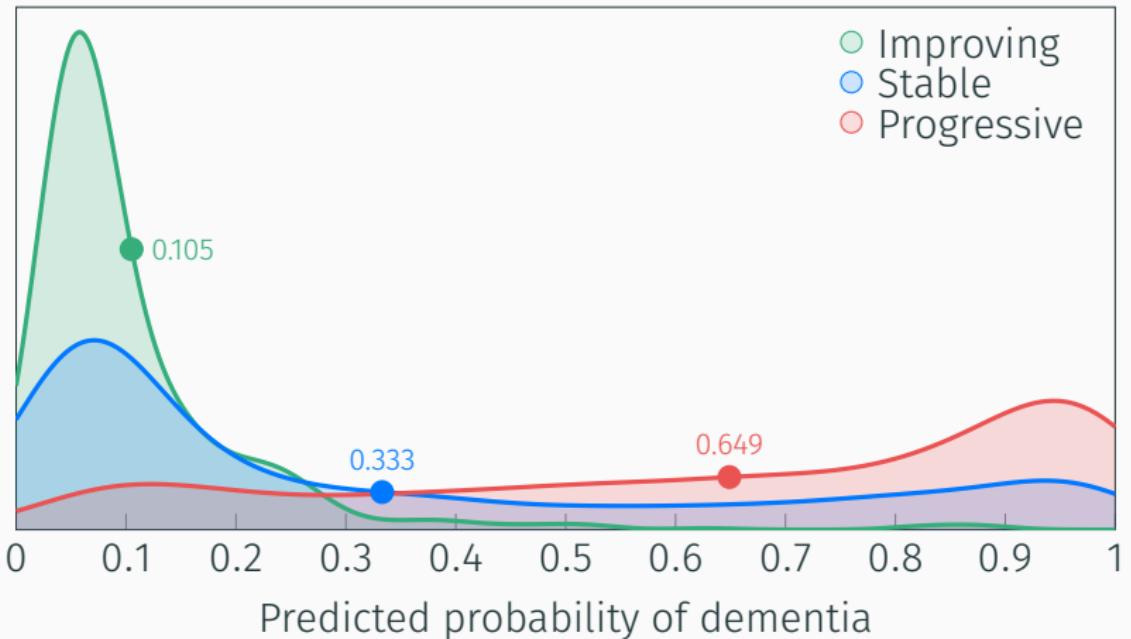
Explainable dementia classification: Validation



Explainable dementia classification: Application

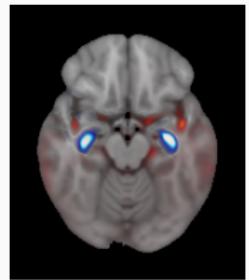
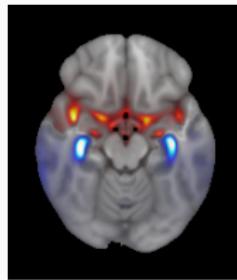
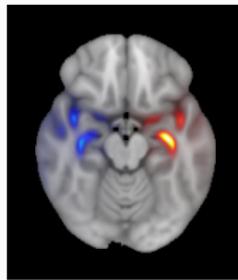
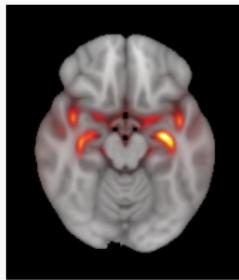


Explainable dementia classification: Application



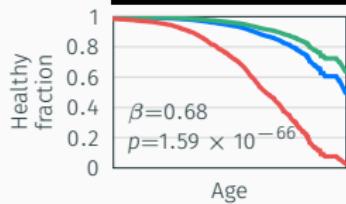
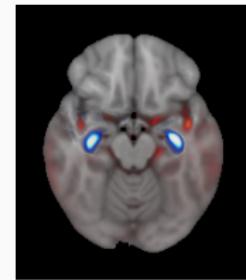
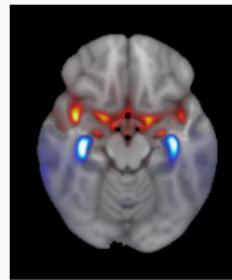
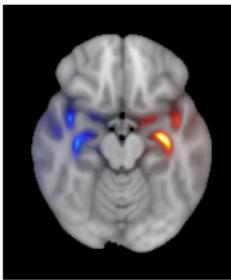
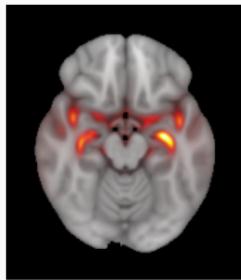
Explainable dementia classification: Application

Component 0 Component 1 Component 2 Component 3

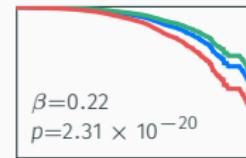
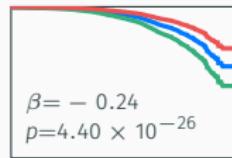


Explainable dementia classification: Application

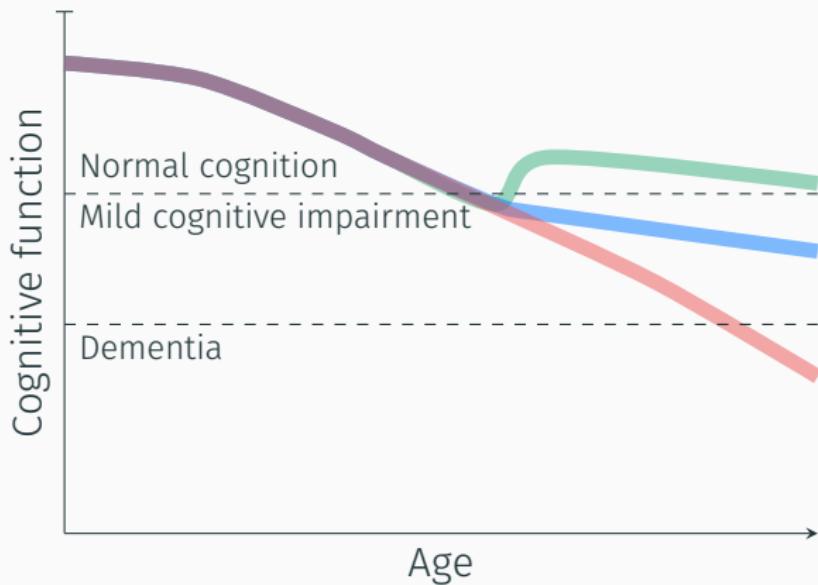
Component 0 Component 1 Component 2 Component 3



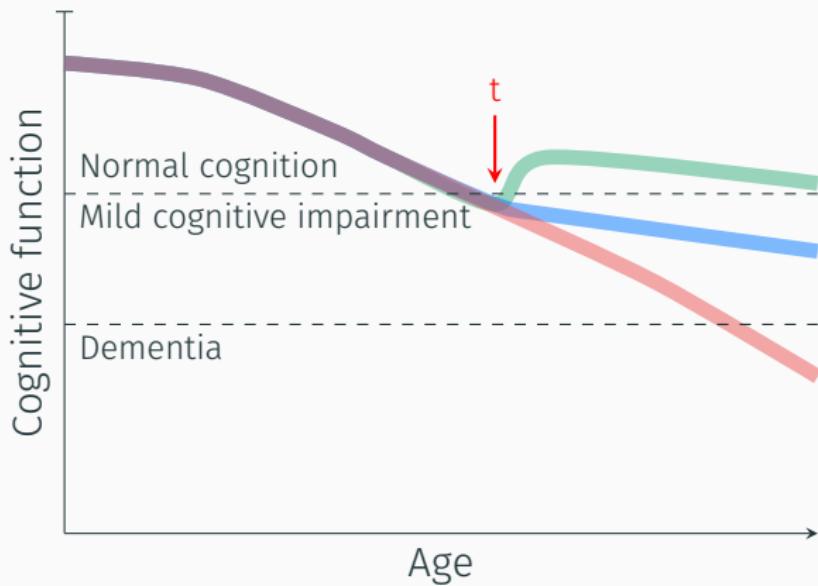
Percentiles
— 5th
— 50th
— 95th



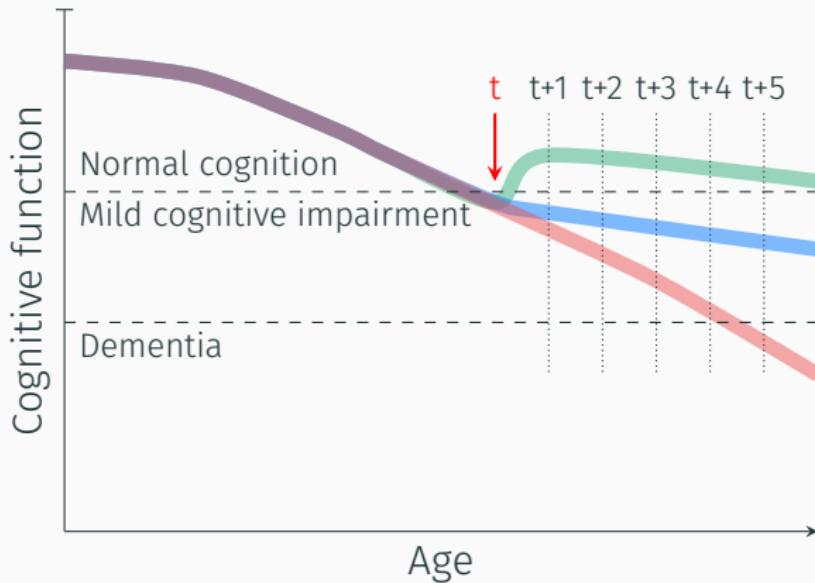
Explainable dementia classification: Application



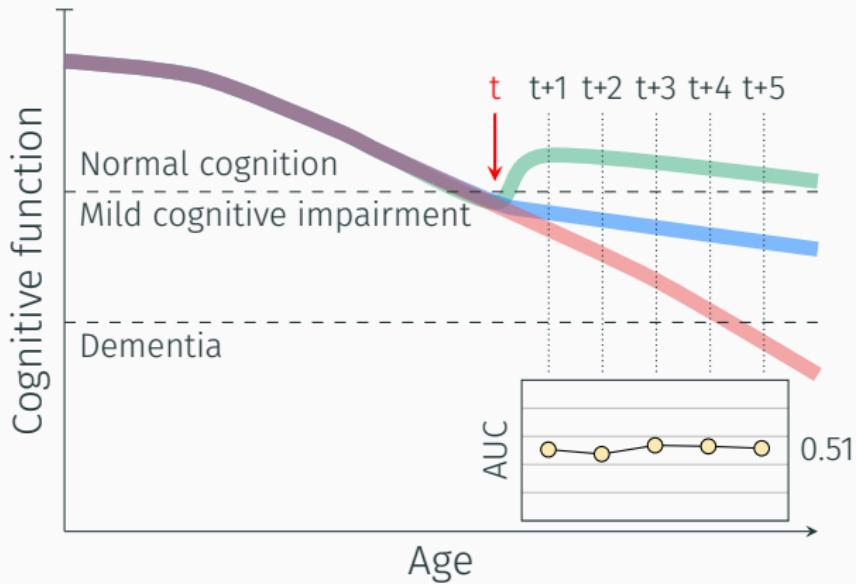
Explainable dementia classification: Application



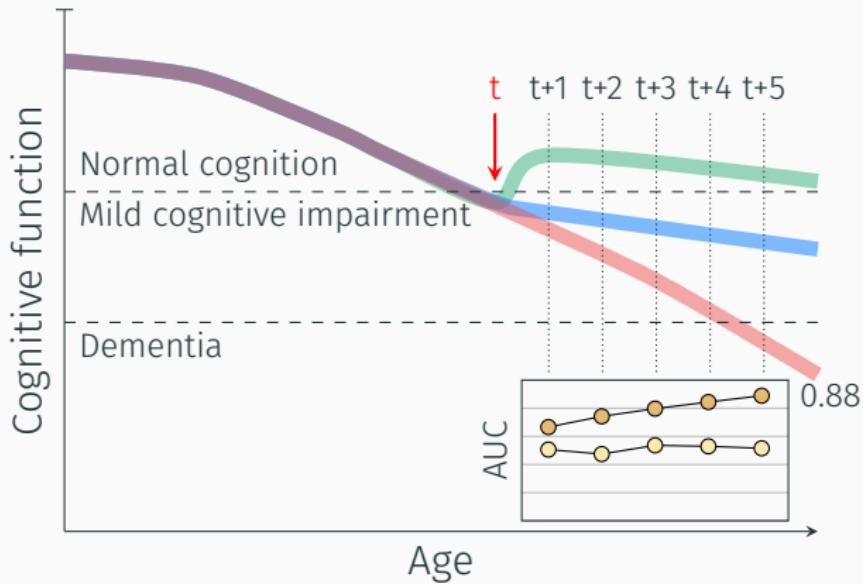
Explainable dementia classification: Application



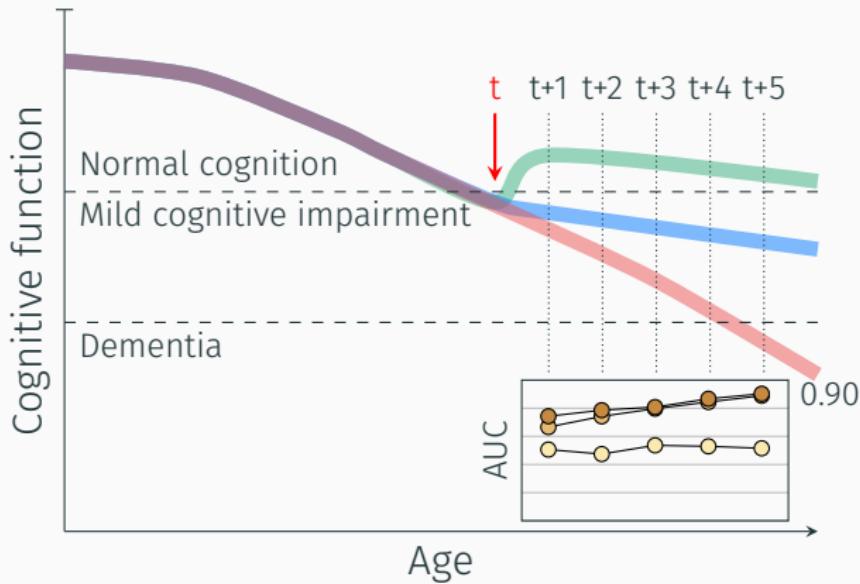
Explainable dementia classification: Application



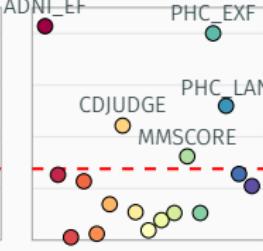
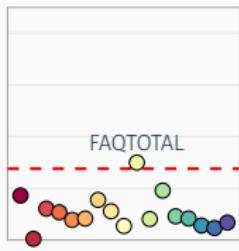
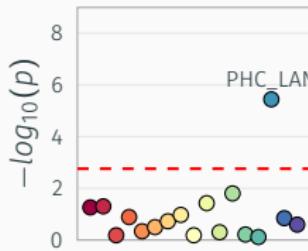
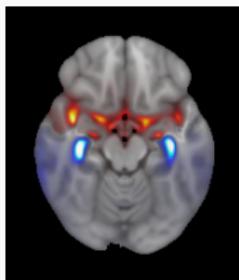
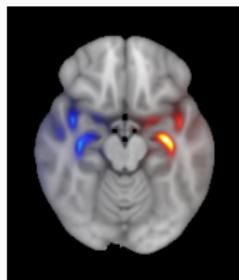
Explainable dementia classification: Application



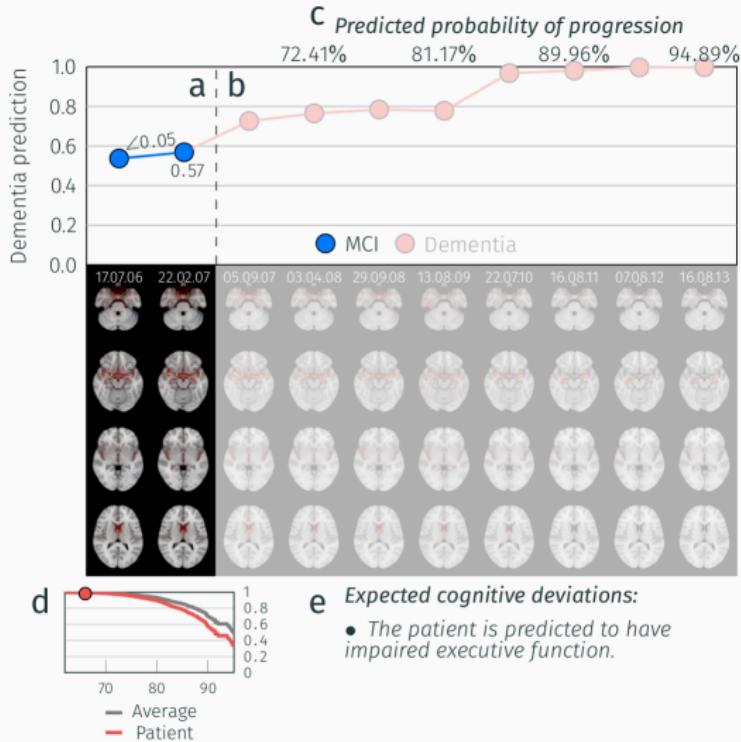
Explainable dementia classification: Application



Explainable dementia classification: Application



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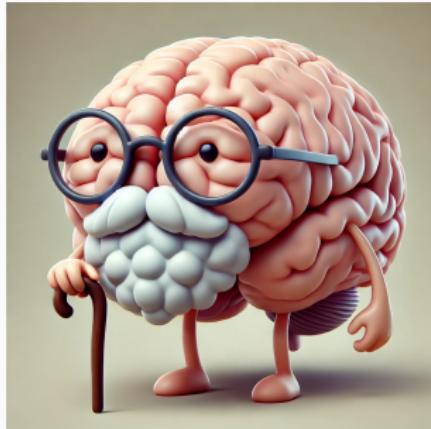


Explainable brain age



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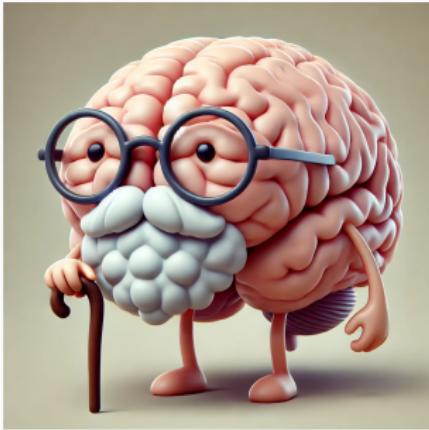
Explainable brain age: Motivation



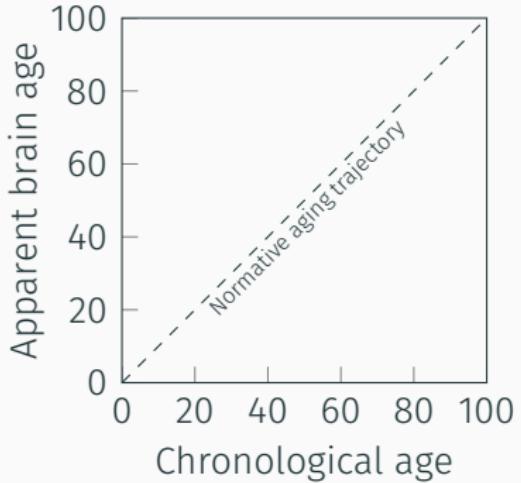
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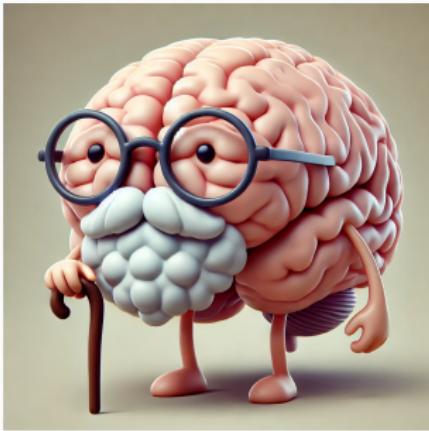
Explainable brain age: Motivation



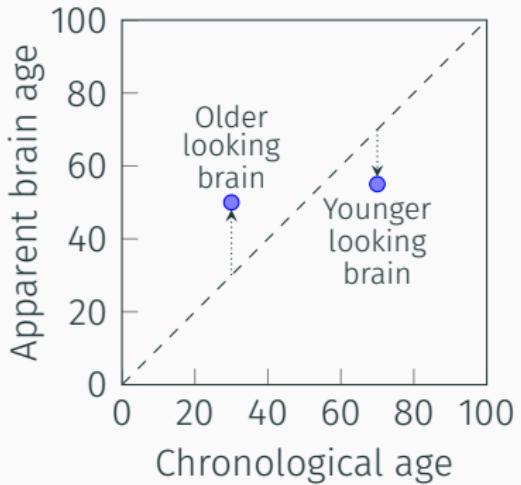
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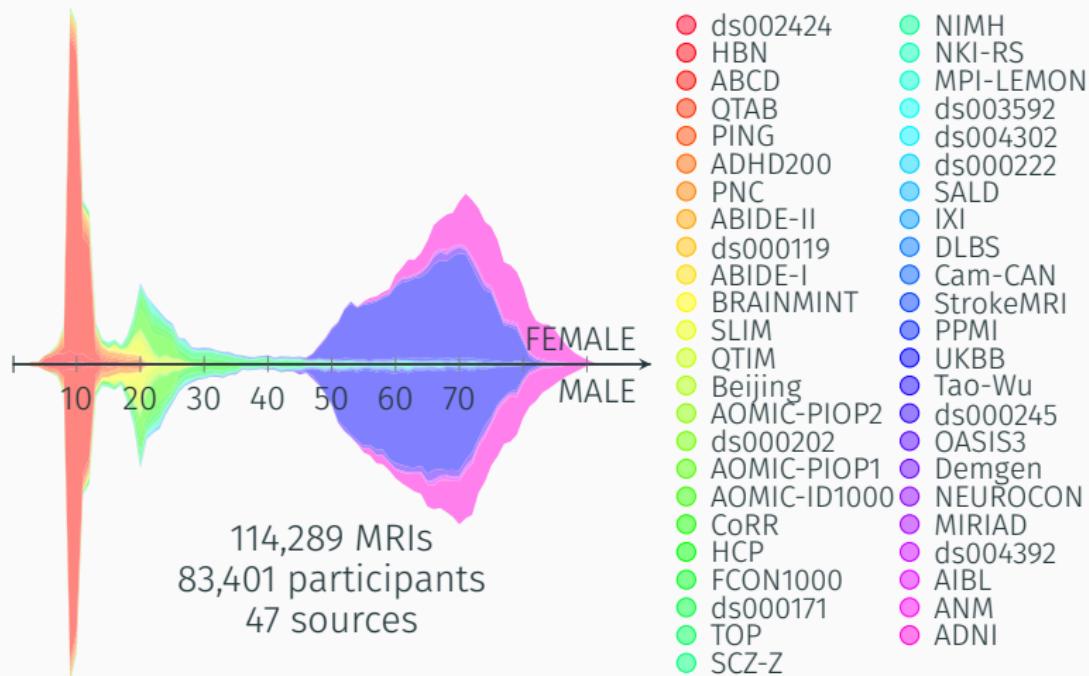
Explainable brain age: Motivation



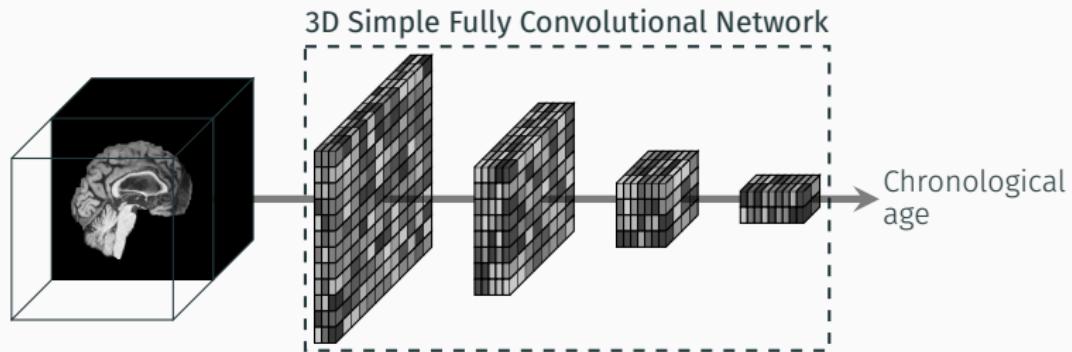
Generated by Dall-E 3



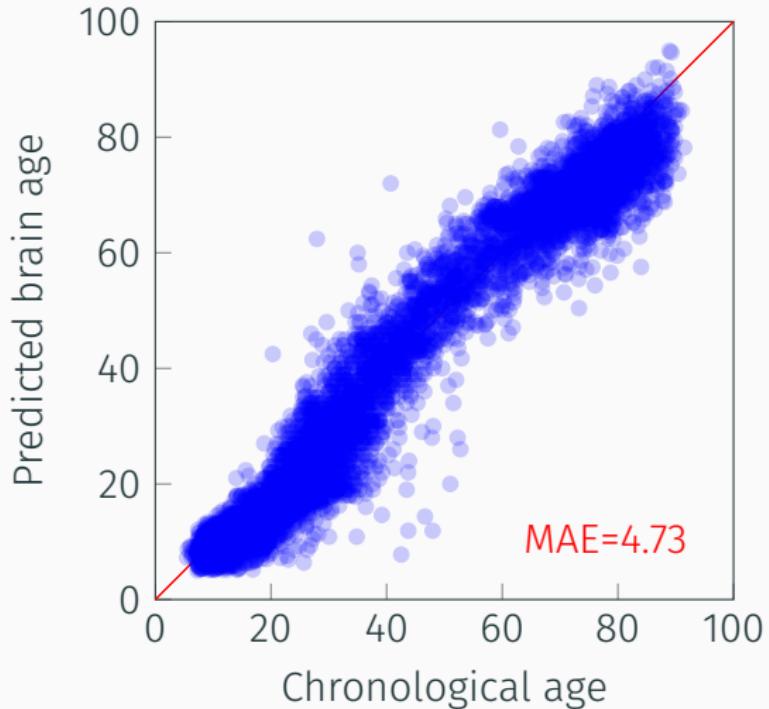
Explainable brain age: Motivation



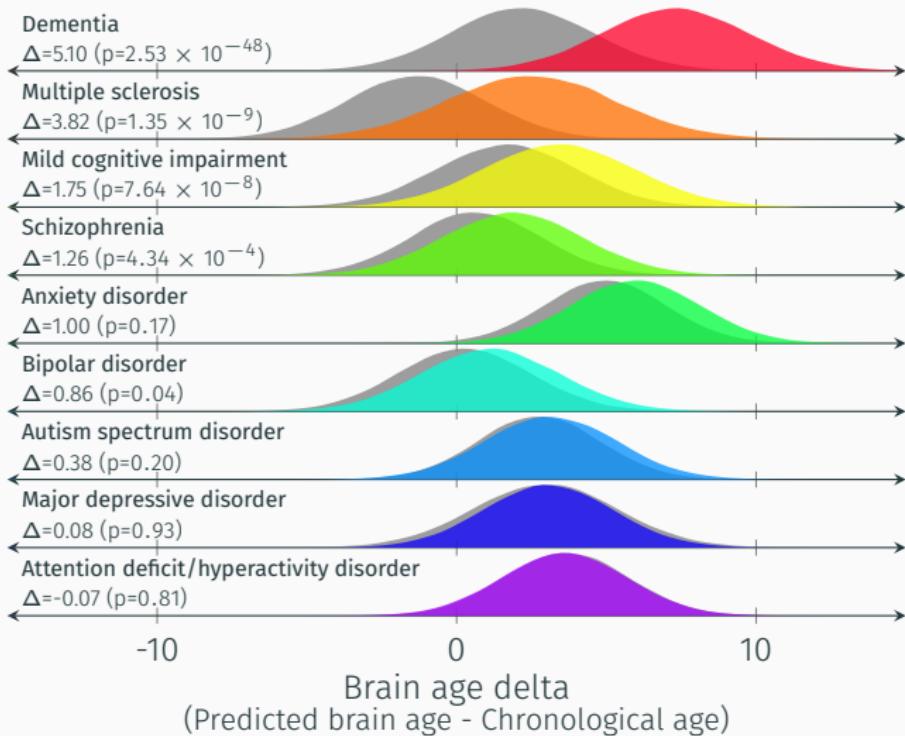
Explainable brain age: Motivation



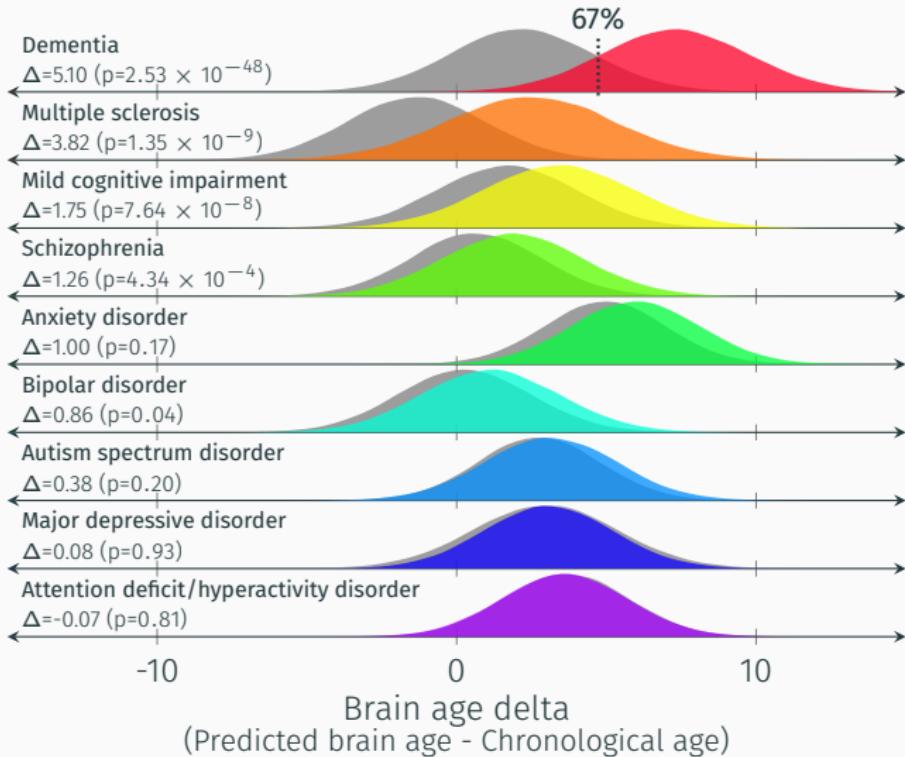
Explainable brain age: Motivation



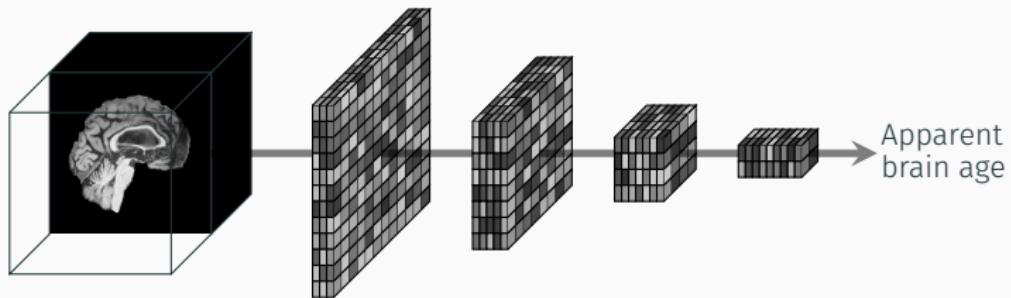
Explainable brain age: Motivation



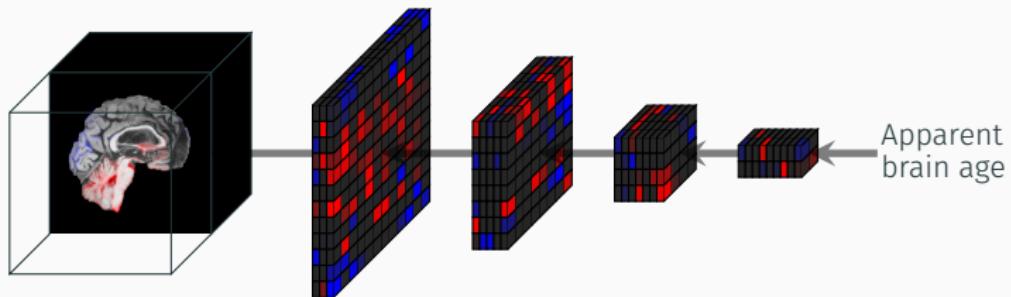
Explainable brain age: Motivation



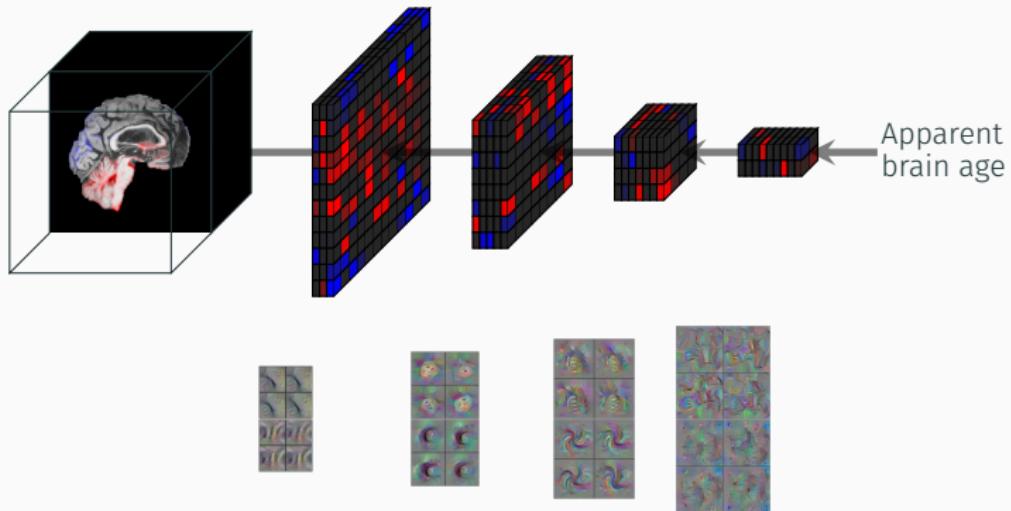
Explainable brain age: Methods



Explainable brain age: Methods



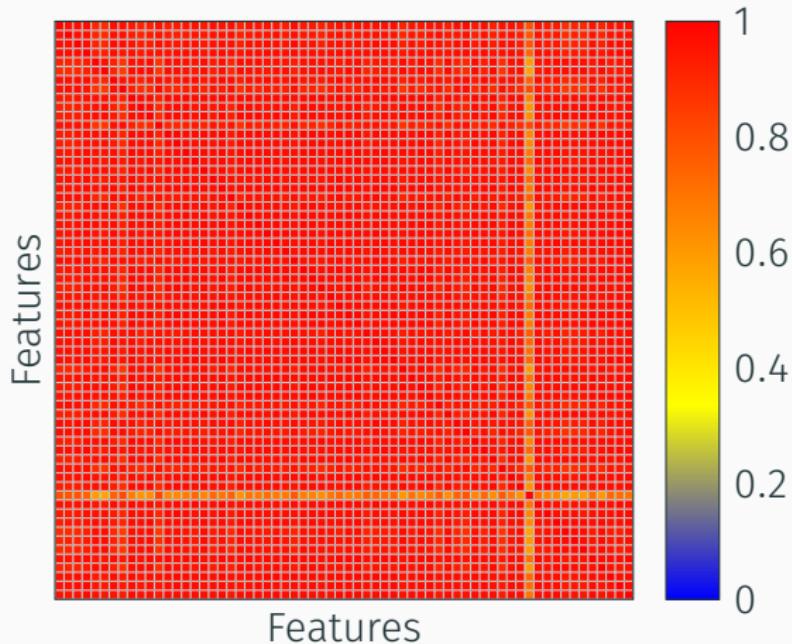
Explainable brain age: Methods



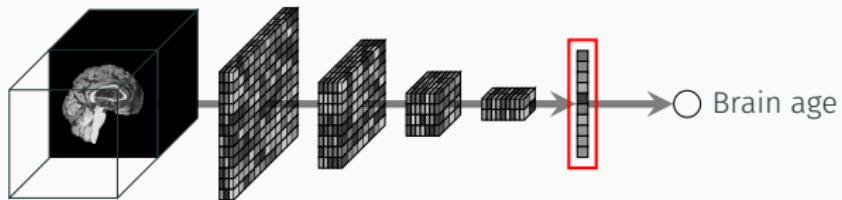
Understanding Neural Networks Through Deep Visualization
Yosinski, J. et al., preprint at arXiv (2015)



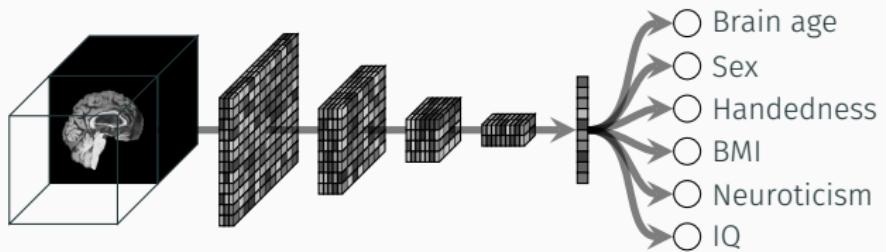
Explainable brain age: Methods



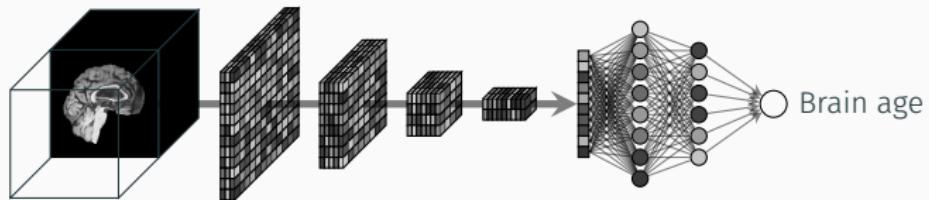
Explainable brain age: Methods



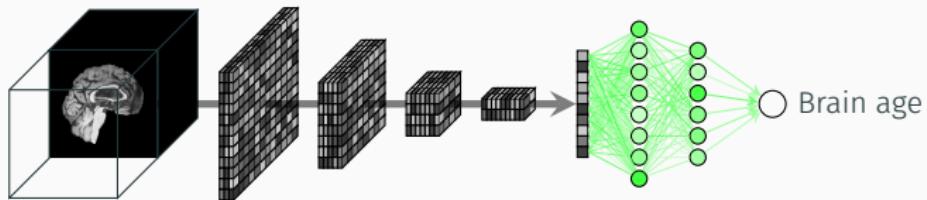
Explainable brain age: Methods



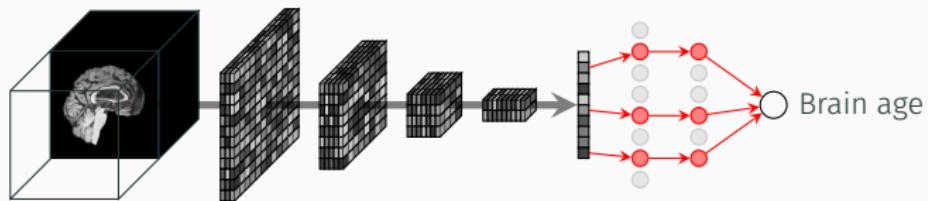
Explainable brain age: Methods



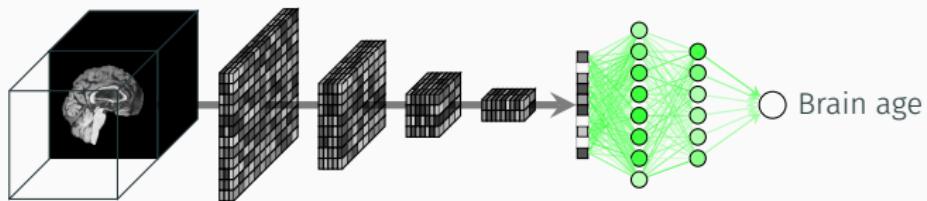
Explainable brain age: Methods



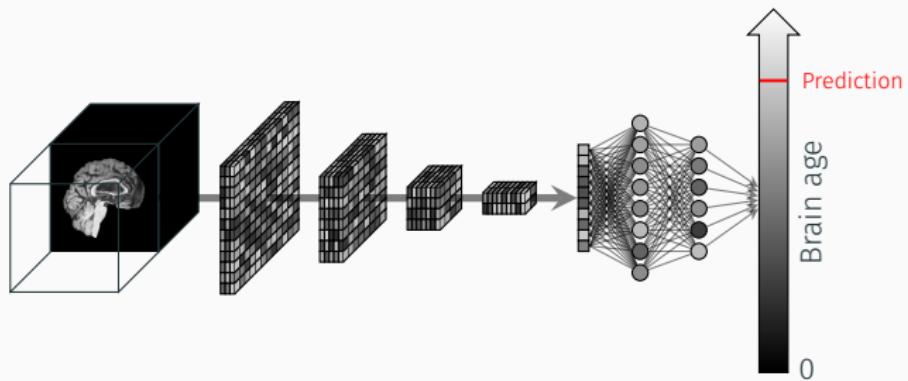
Explainable brain age: Methods



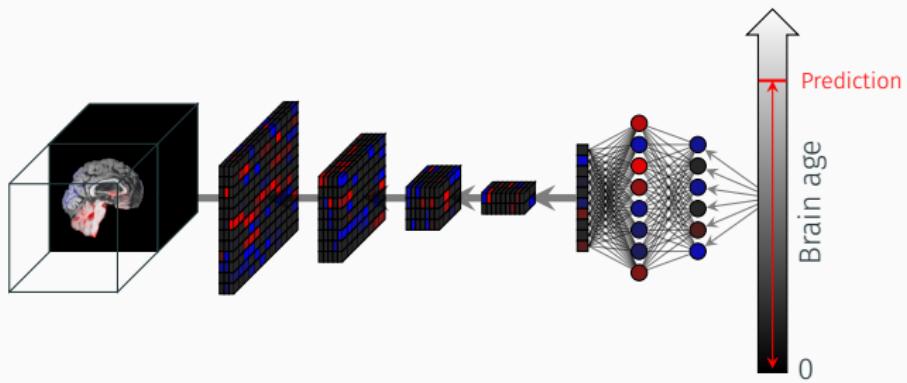
Explainable brain age: Methods



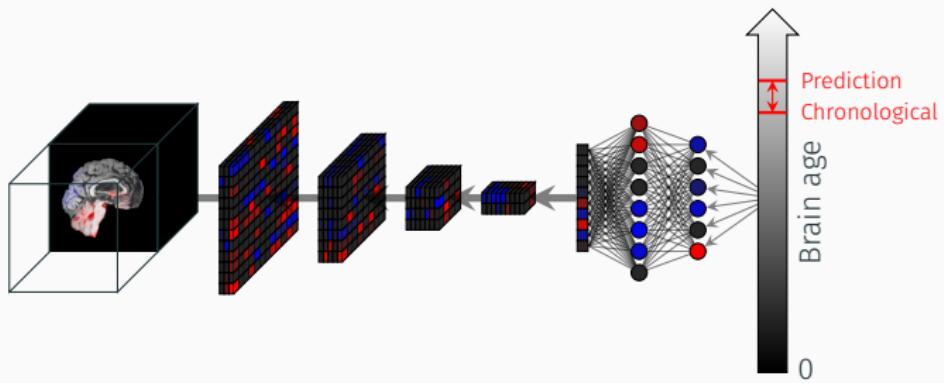
Explainable brain age: Methods



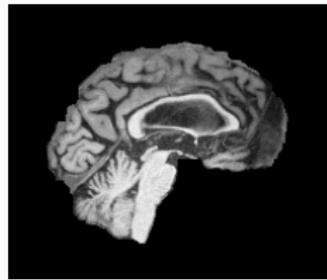
Explainable brain age: Methods



Explainable brain age: Methods



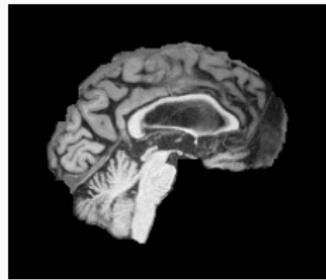
Explainable brain age: Methods



Age
62

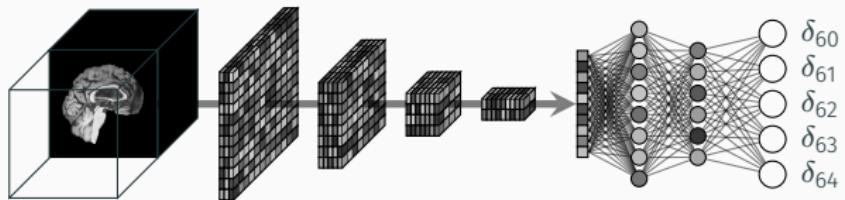


Explainable brain age: Methods

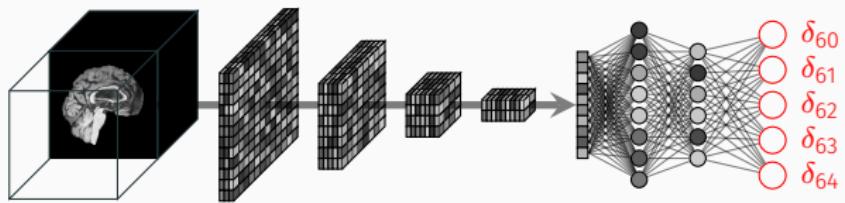


δ_{60}	δ_{61}	δ_{62}	δ_{63}	δ_{64}
2	1	0	-1	-2

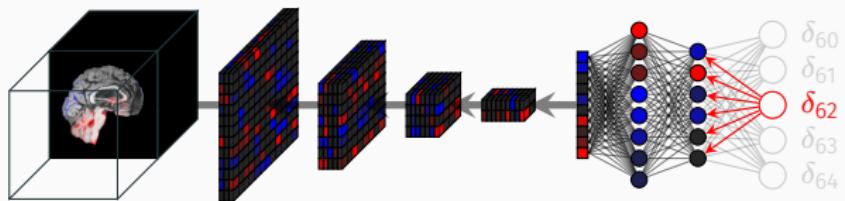
Explainable brain age: Methods



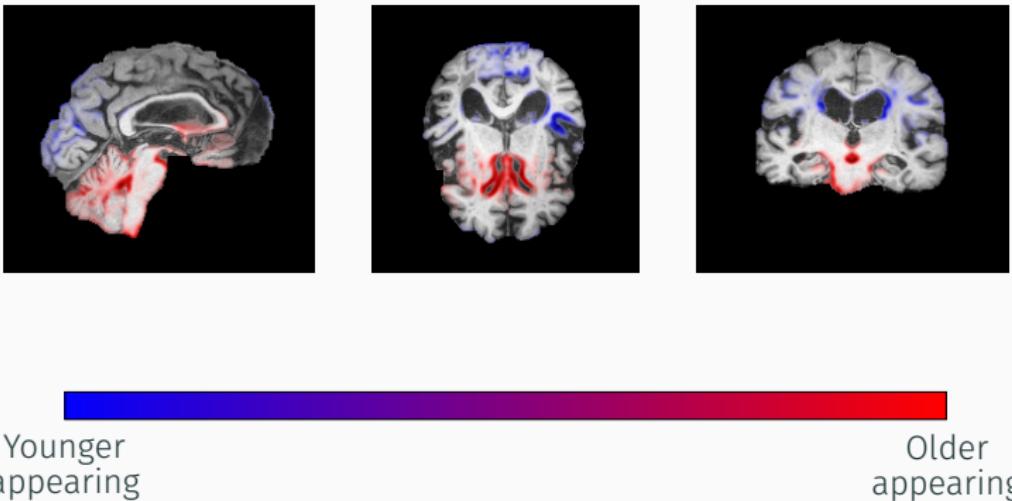
Explainable brain age: Methods



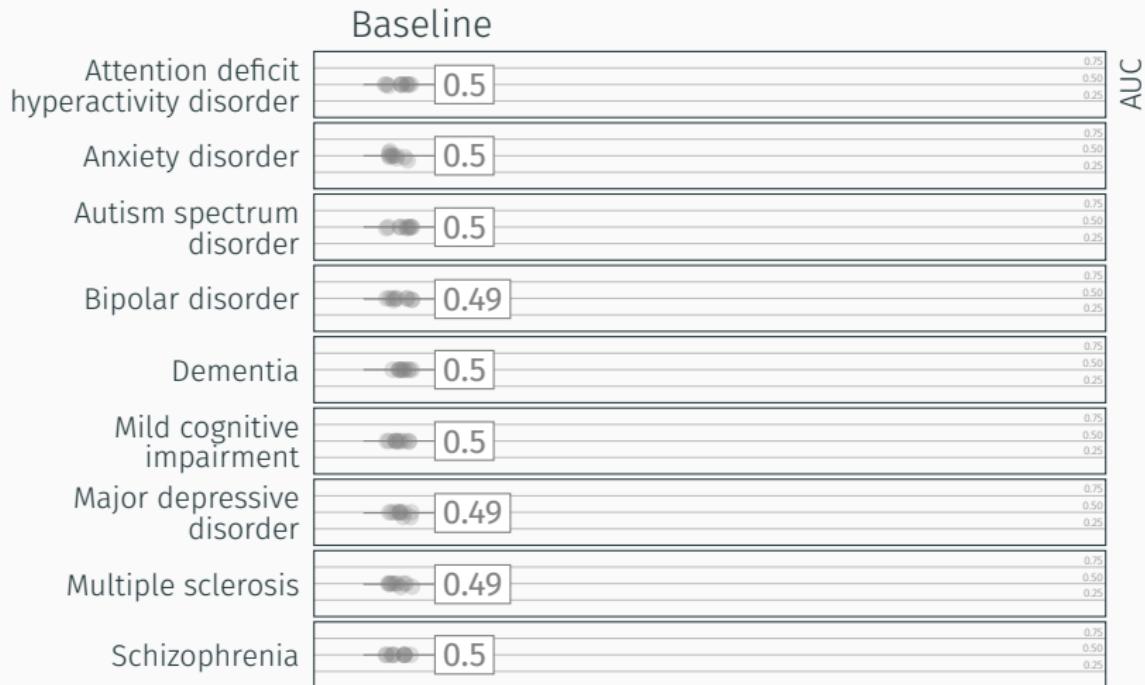
Explainable brain age: Methods



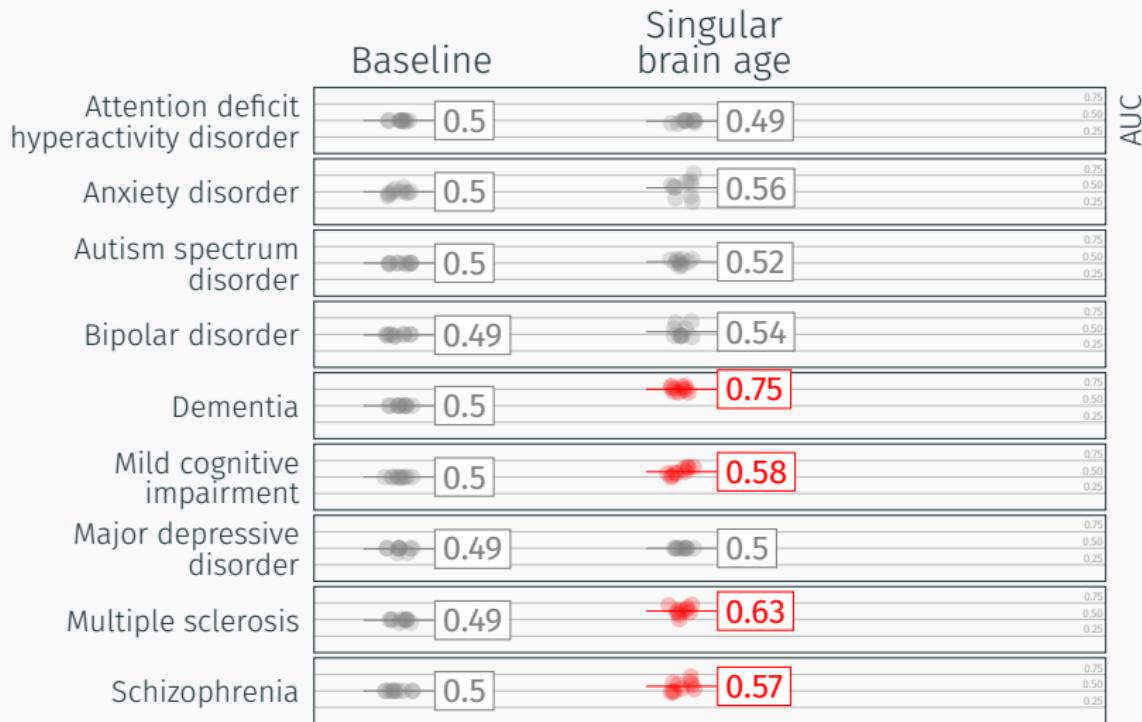
Explainable brain age: Results



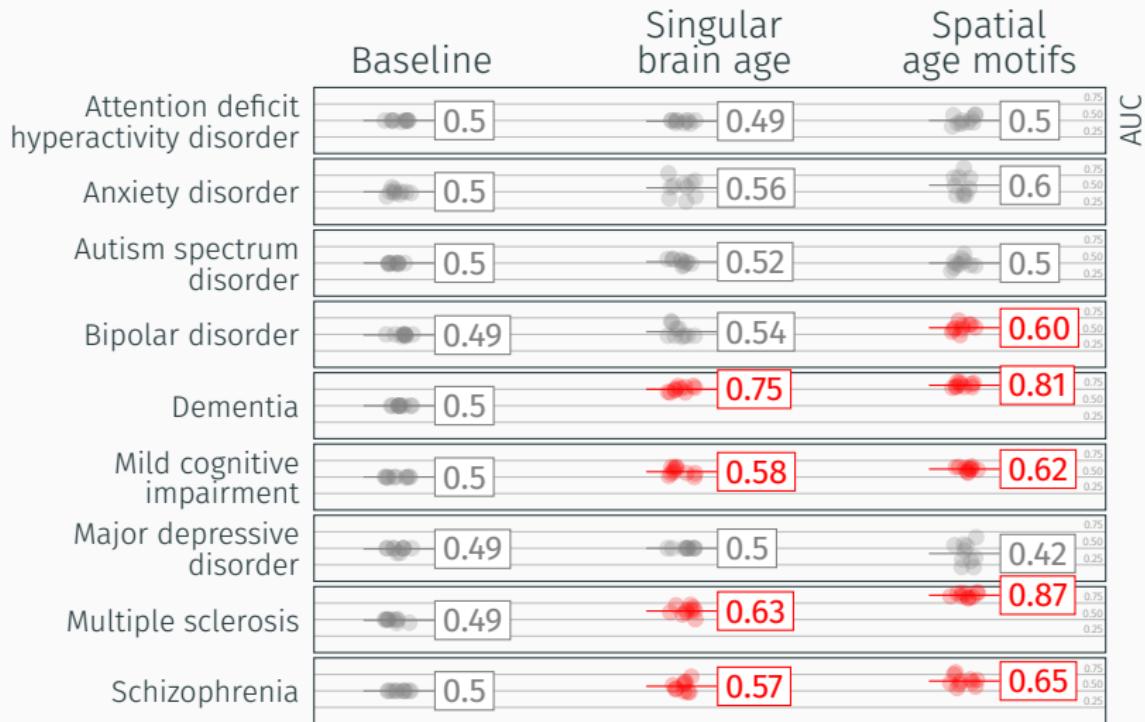
Explainable brain age: Results



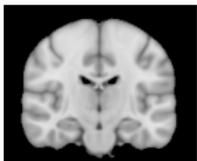
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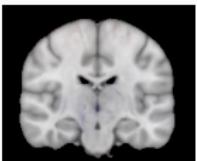
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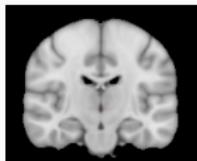
Explainable brain age: Results



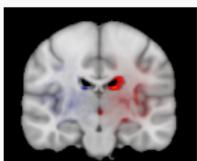
Attention deficit
hyperactivity disorder



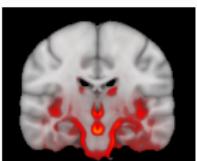
Anxiety
disorder



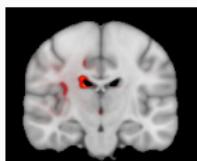
Autism spectrum
disorder



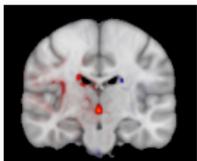
Bipolar
disorder



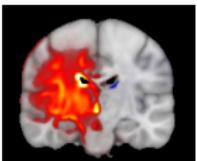
Dementia



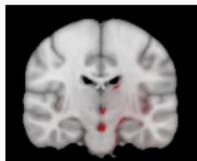
Schizophrenia



Major depressive
disorder



Multiple
sclerosis



Mild cognitive
impairment

Explainable brain age: Results



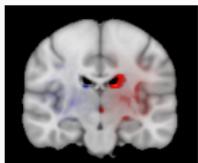
Attention deficit
hyperactivity disorder



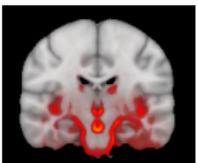
Anxiety
disorder



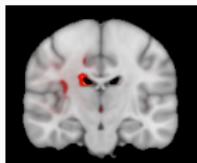
Autism spectrum
disorder



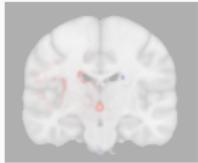
Bipolar
disorder



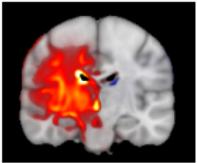
Dementia



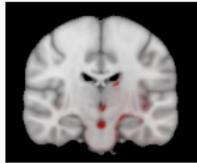
Schizophrenia



Major depressive
disorder

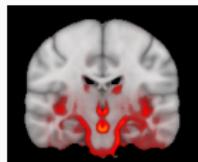


Multiple
sclerosis



Mild cognitive
impairment

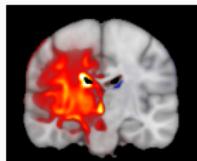
Explainable brain age: Results



Dementia

- Temporal pole
- Left amygdala
- Parahippocampal gyrus

Explainable brain age: Results

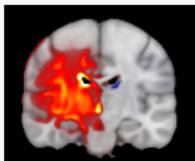


Multiple
sclerosis

- Right pallidum
- Right thalamus
- Right putamen



Explainable brain age: Results

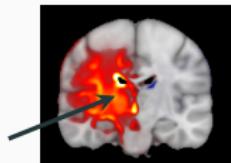


Multiple
sclerosis

- Right pallidum
- Right thalamus
- Right putamen



Explainable brain age: Results

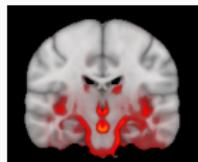


Multiple
sclerosis

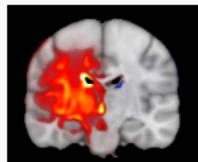
- Right pallidum
- Right thalamus
- Right putamen



Explainable brain age: Results



Dementia

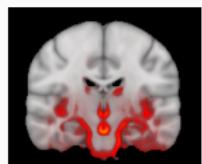


Multiple
sclerosis



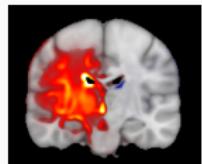
Explainable brain age: Results

AUC



Dementia

Singular brain age	Spatial age motifs
0.75	0.81



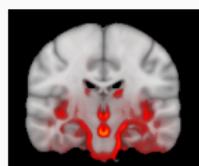
Multiple
sclerosis

0.63	0.87
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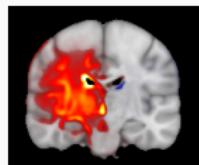
Explainable brain age: Results

AUC



Dementia

Singular brain age	Spatial age motifs
0.75 0.06	0.81

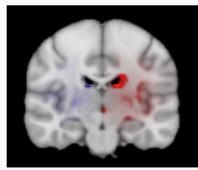


Multiple
sclerosis

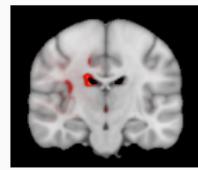
0.63 0.24	0.87
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Explainable brain age: Results



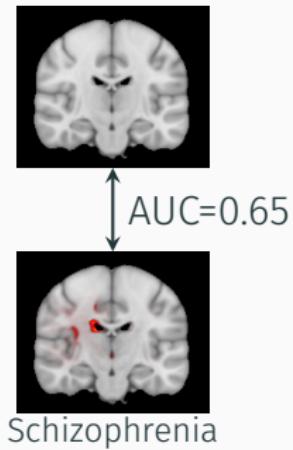
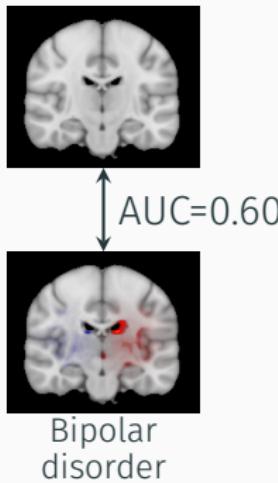
Bipolar
disorder



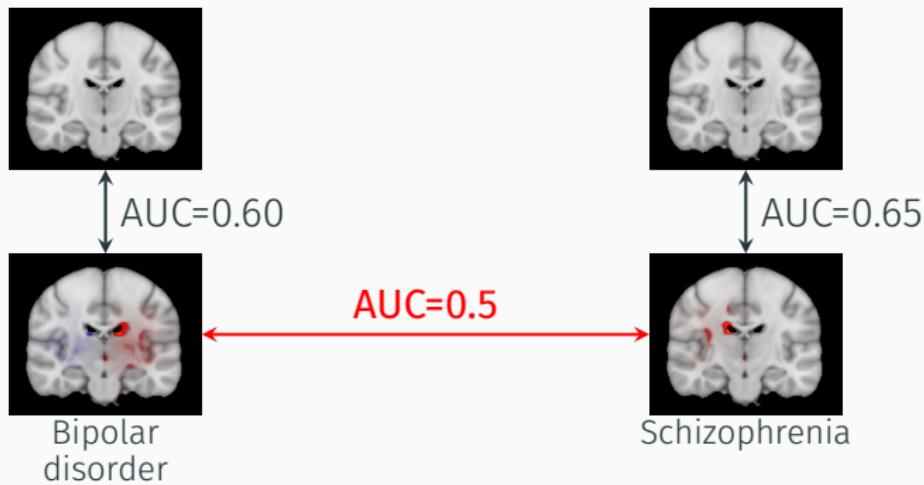
Schizophrenia



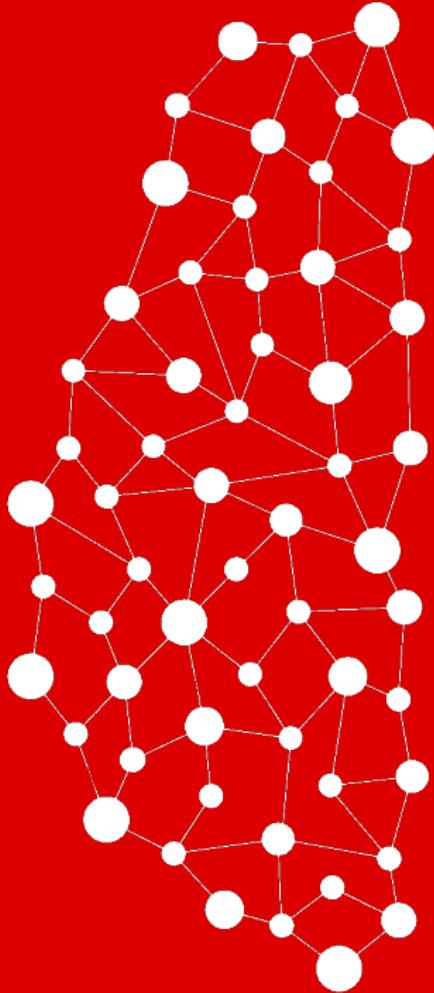
Explainable brain age: Results



Explainable brain age: Results



Thank you for your attention!
estenhl@ui.no



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