

Learning How to See the Invisible

A data-driven approach to finding underlying patterns of abnormality in visually normal MR brain images from patients with epilepsy

Oscar Bennett¹ John Duncan^{2,3} M. Jorge Cardoso¹ Gavin Winston^{2,3}
Sebastien Ourselin¹

1. Translational Imaging Group, Centre for Medical Image Computing, University College London, London, UK

2. Department of Clinical and Experimental Epilepsy, UCL Institute of Neurology, London, UK

3. Epilepsy Society MRI Unit, Chalfont St Peter, UK

Summary

Epilepsy

The Problem

Methods

Results

Discussion

Further Work

Epilepsy

The Problem

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Epilepsy

- Common and Serious
- Characterised by recurrent seizures
- First-line treatment with oral medication
- Medication ineffective in 1/3 of cases

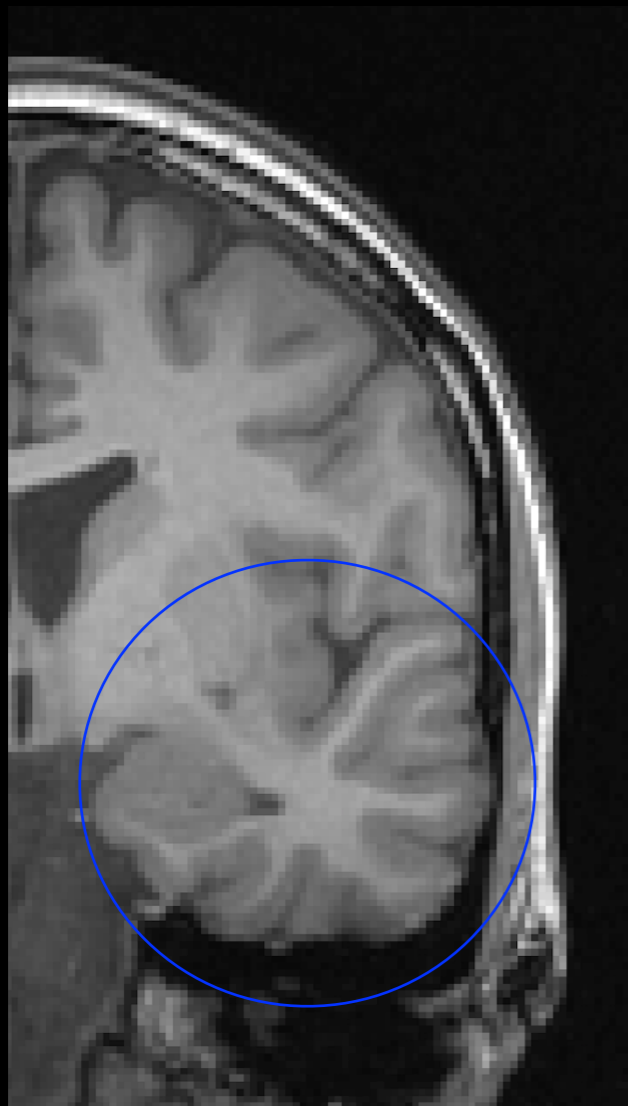
Epilepsy

- Drug resistant focal epilepsy is a challenging problem
- Neurosurgery is often considered in these cases
- Aim to remove the 'epileptogenic' region of the brain
- Have to find it first...

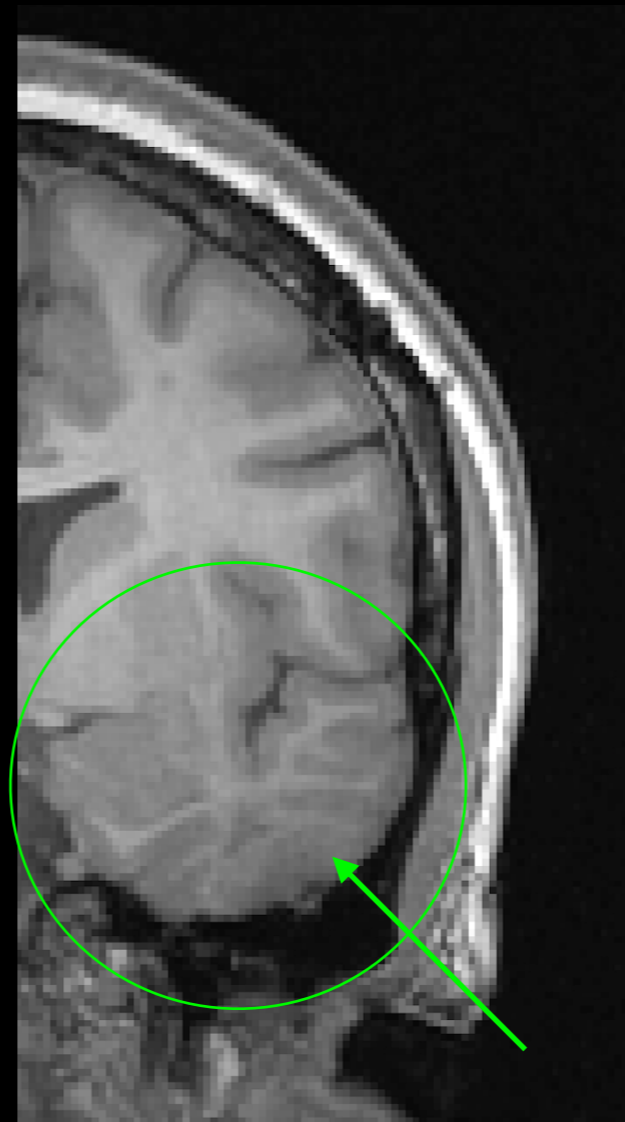
The Problem

- MRI is normally used to look for these abnormalities
- Abnormality found by a radiologist 2/3 of the time
- ...leaving 1/3 of patients 'MR negative'

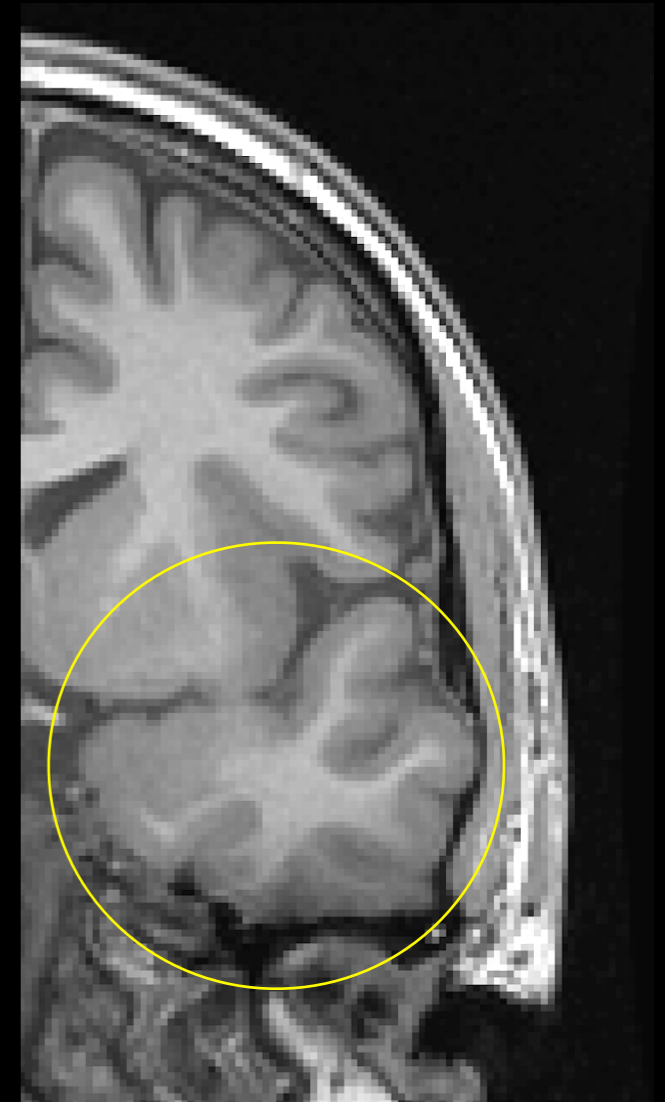
The Problem



Healthy



Visible Disease
'MR positive'



Invisible Disease
'MR negative'

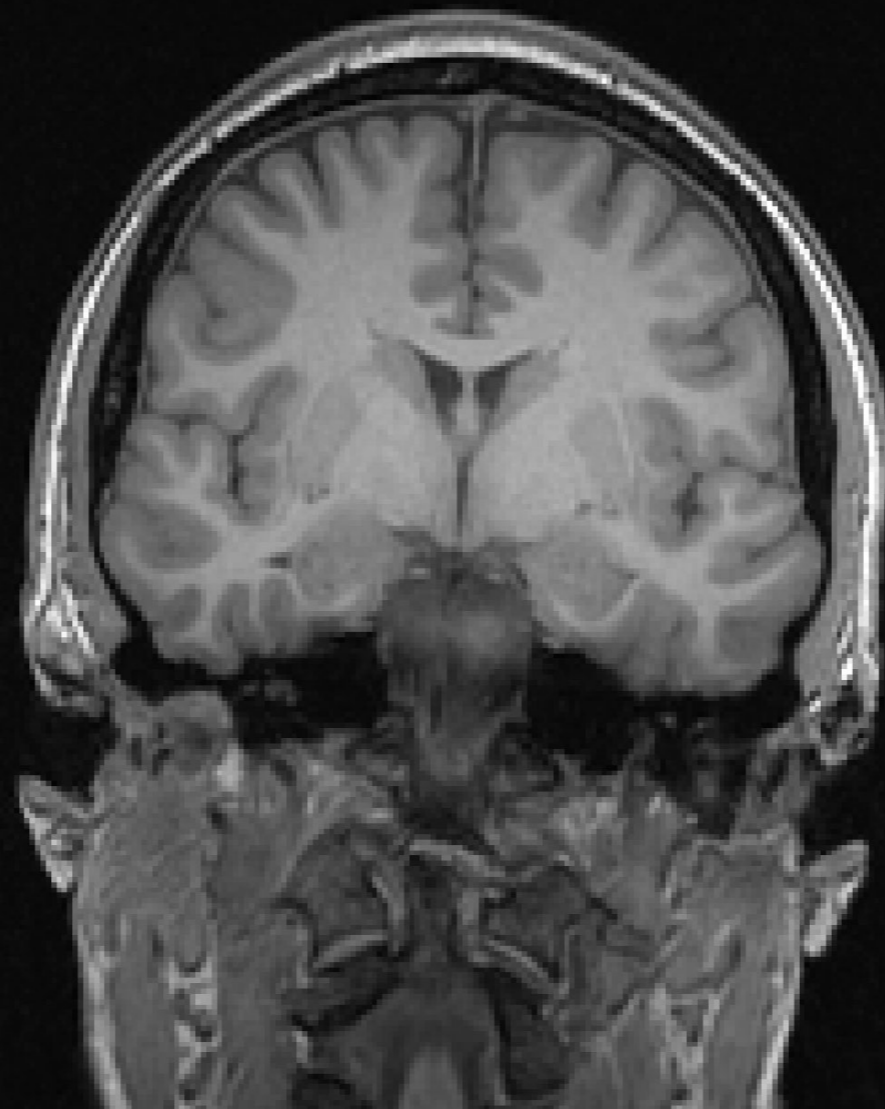
The Problem

- Are these visually normal MR negative images truly normal?
- Or are the abnormalities within them simply too subtle to see?
- Or in a different pattern from what we expect?



Invisible Disease
'MR negative'

Our Approach



Our Approach

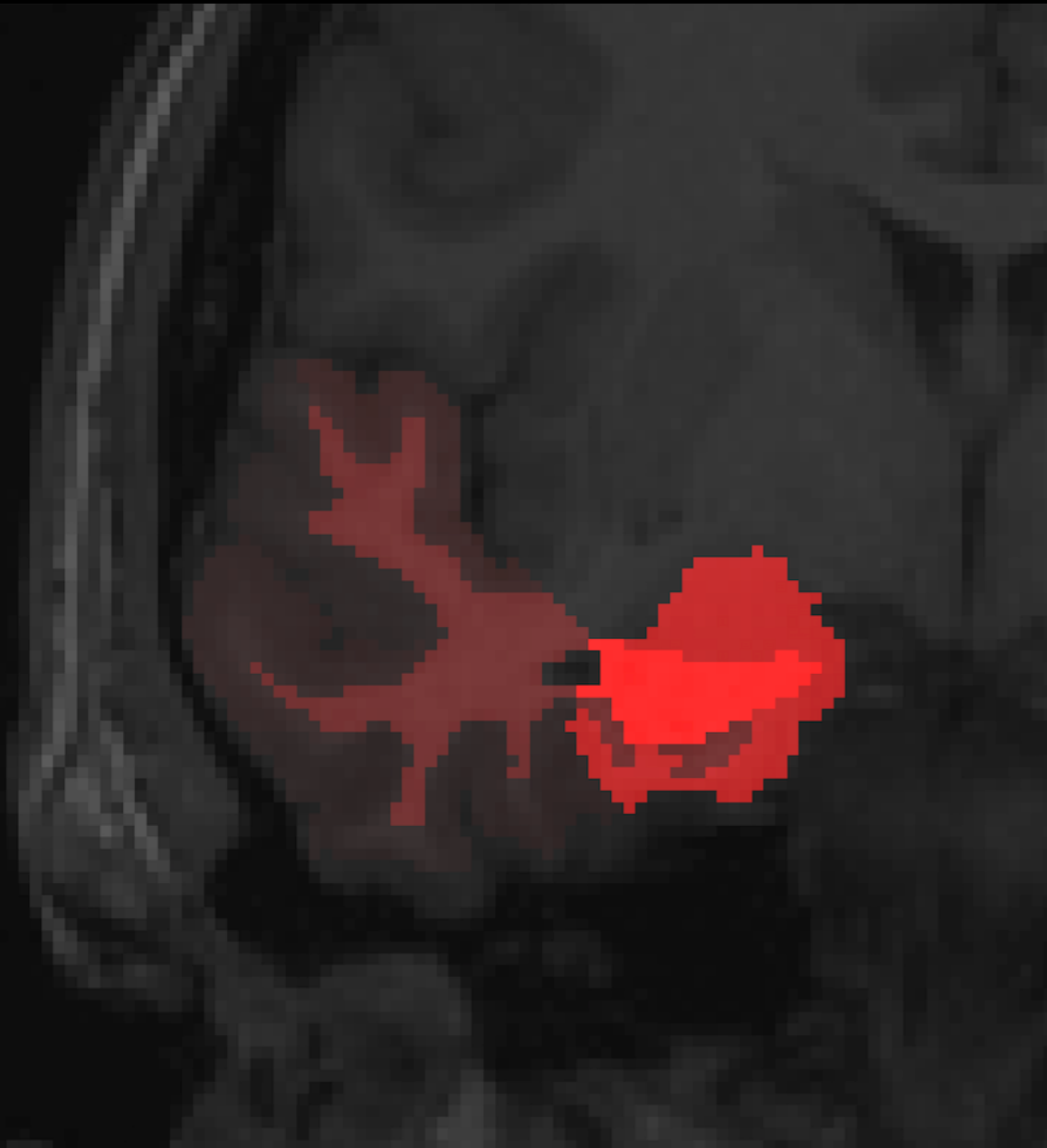


Our Approach

MR positive



Our Approach



MR positive



MR negative

Methods

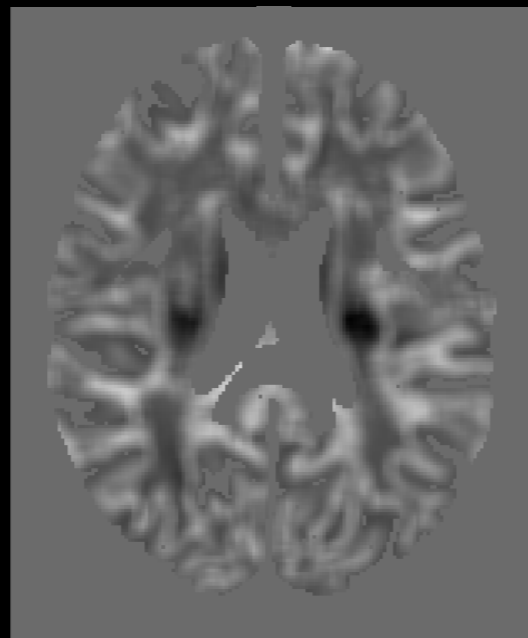
- 82 MR +ve and 26 MR -ve subjects
- All with temporal lobe epilepsy
- Seizure lateralisation known for each subject
- 4 image modalities/types available for each subject
- T1, T2, FLAIR, Junction Map

Methods

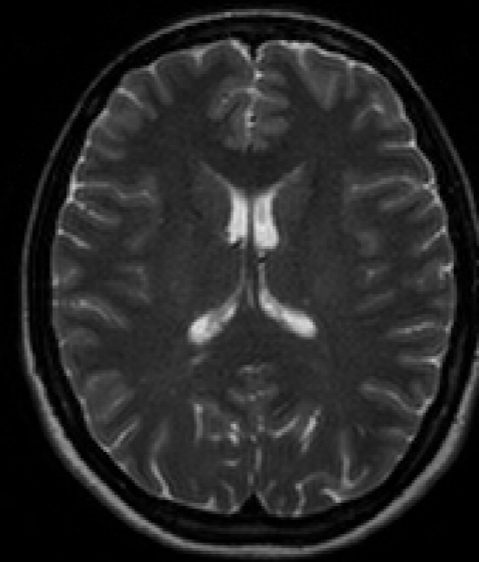
1. Start with the 4 image volumes from each subject



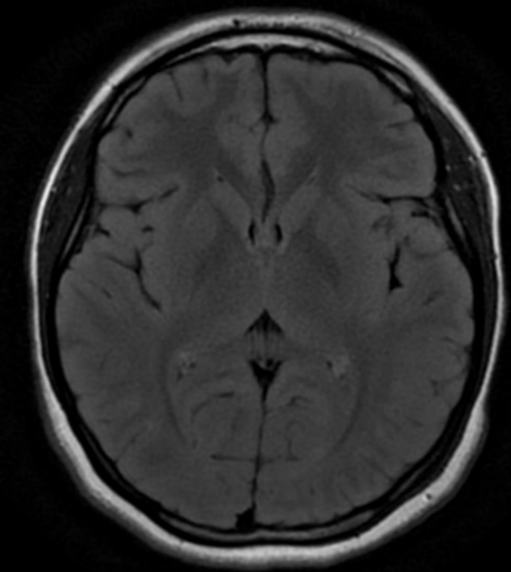
T1



Junction Map



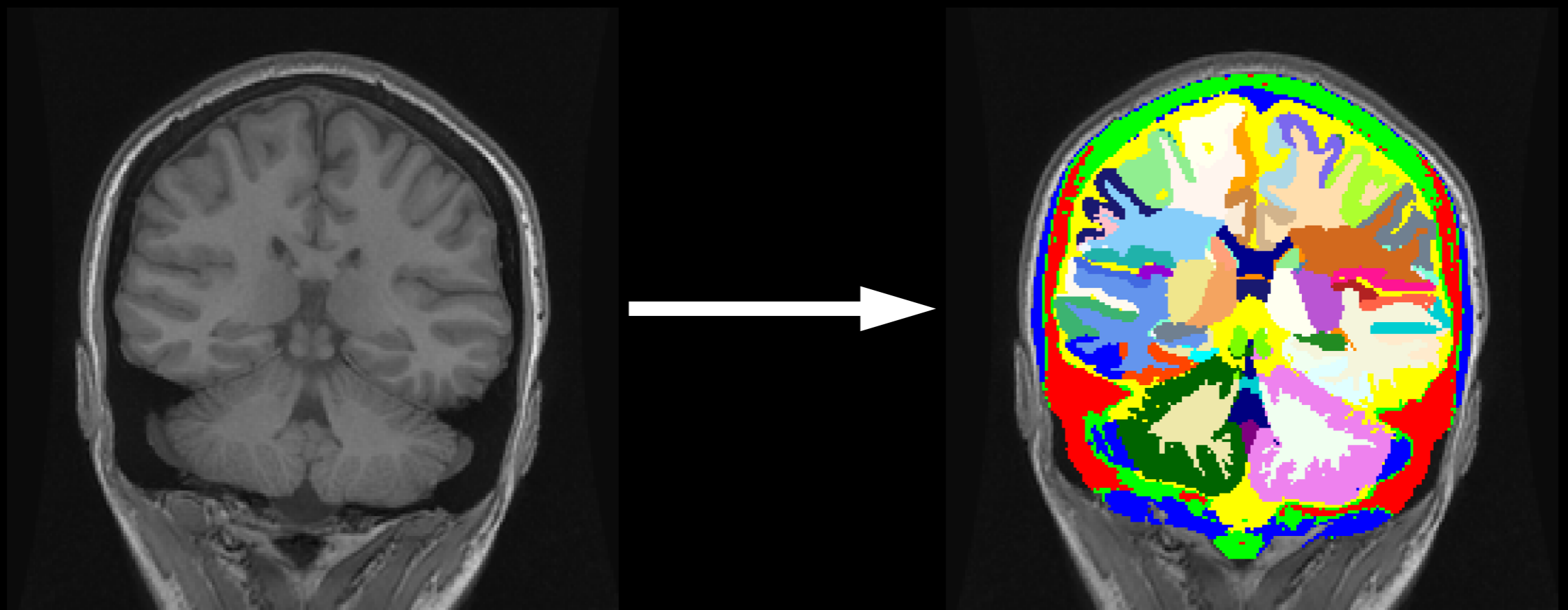
T2



FLAIR

Methods

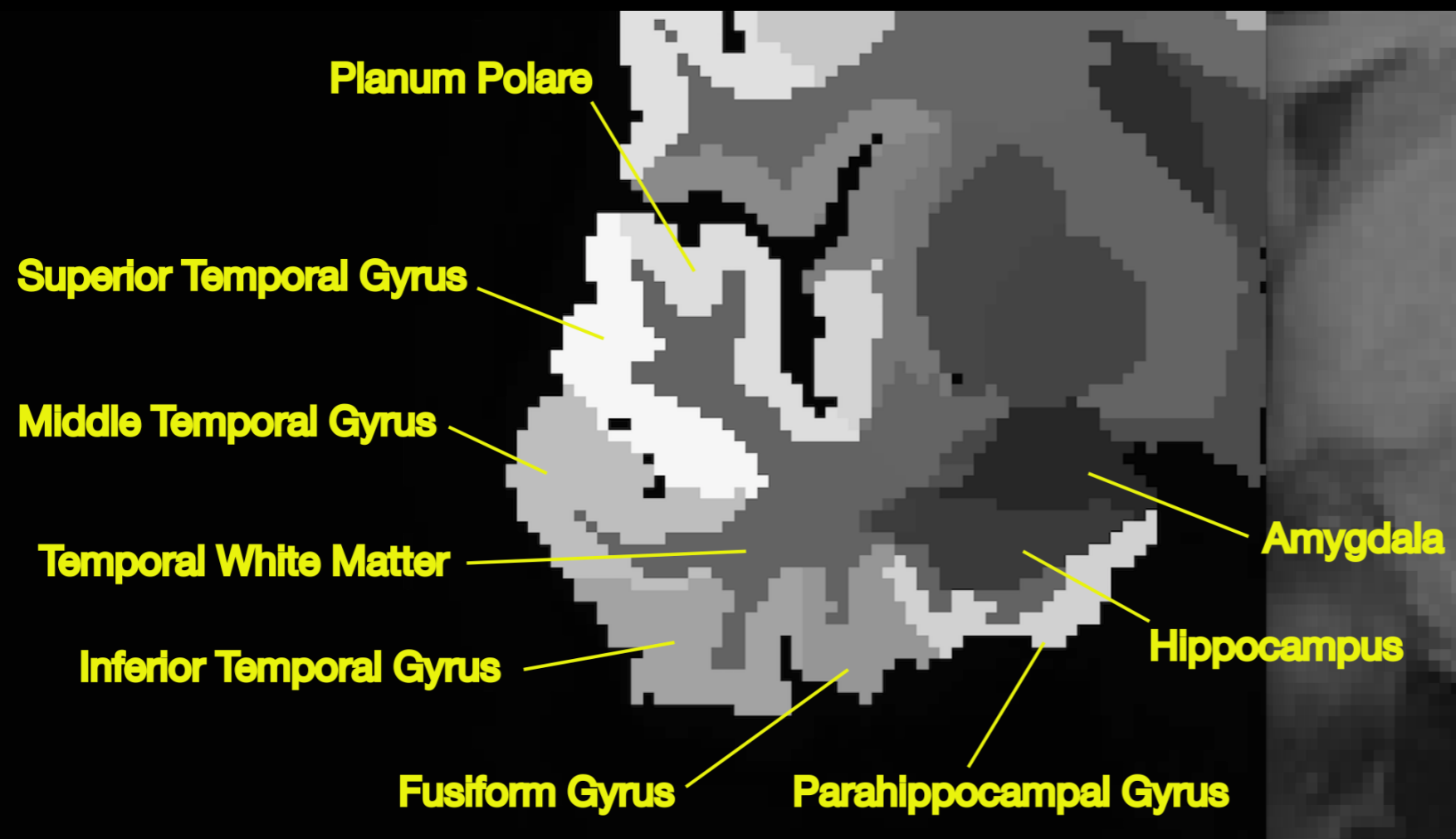
2. Segment out all anatomical regions within images (GIF)



[http://cmictig.cs.ucl.ac.uk/wiki/index.php/
Full brain parcellation using Geodesic Information Flow](http://cmictig.cs.ucl.ac.uk/wiki/index.php/Full_brain_parcellation_using_Geodesic_Information_Flow)

Methods

3. Look specifically at the regions within the temporal lobe



Also:

Entorhinal area

Planum Temorale

Transverse Temporal Gyrus

Temporal Pole

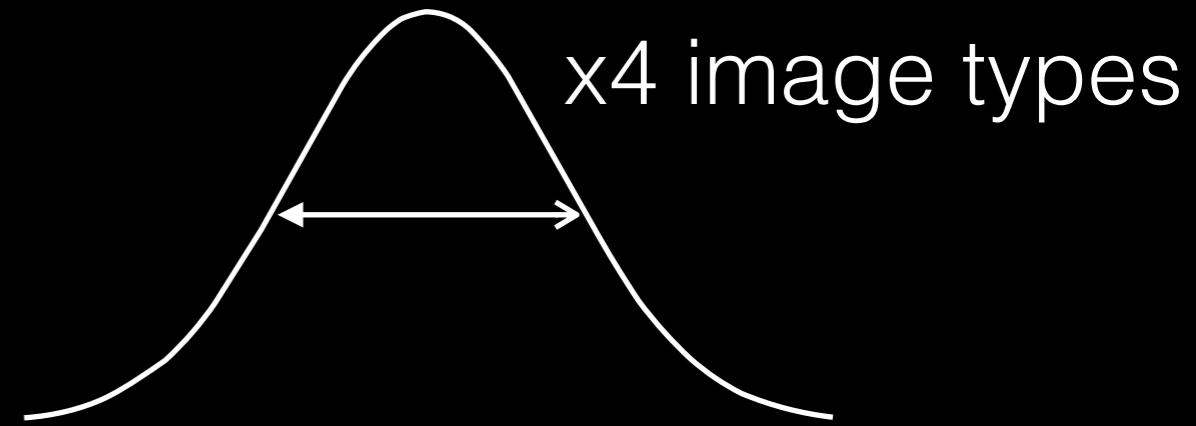
Methods

4. Compute image features from each region within the 13 temporal lobes of each subject

a. **Mean Intensity**
left/right difference



b. Intensity **Standard Deviation**
left/right difference



c. Region **volume**
left/right ratio

Also: Total intracranial
volume

Methods

4. Compute image features from each region within the 13 temporal lobes of each subject

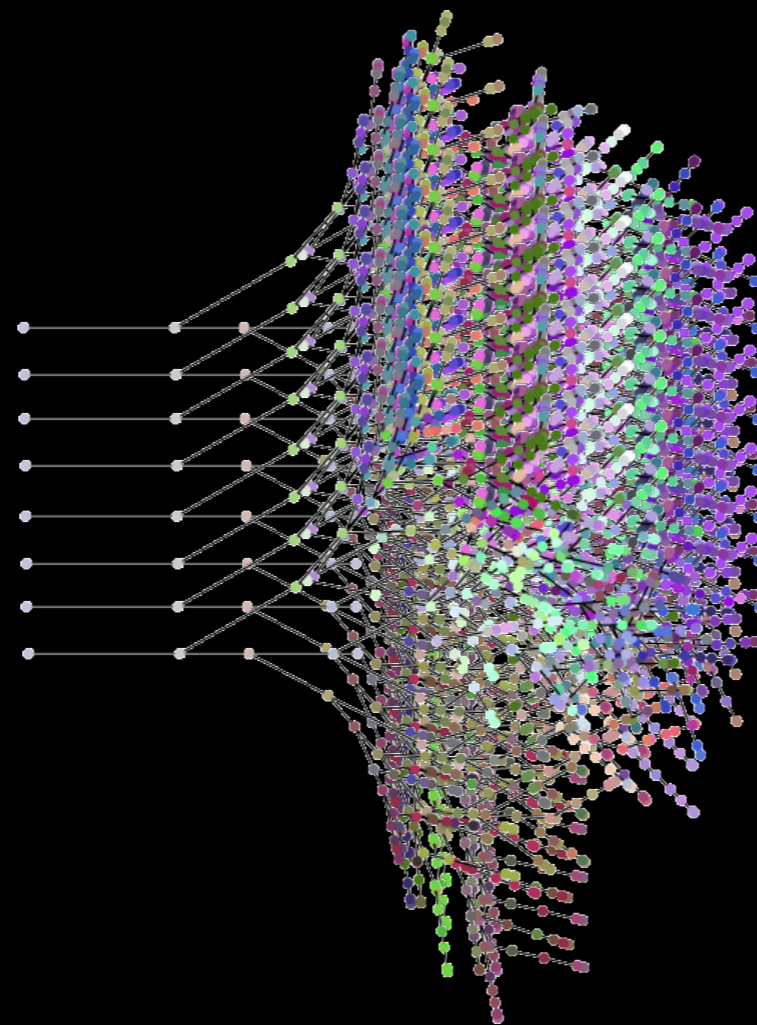
118 features generated per subject

Methods

5. Train a random forest classifier on the image features to predict seizure lateralisation

Random Forest

Image features
from within
temporal lobes



Left temporal lobe

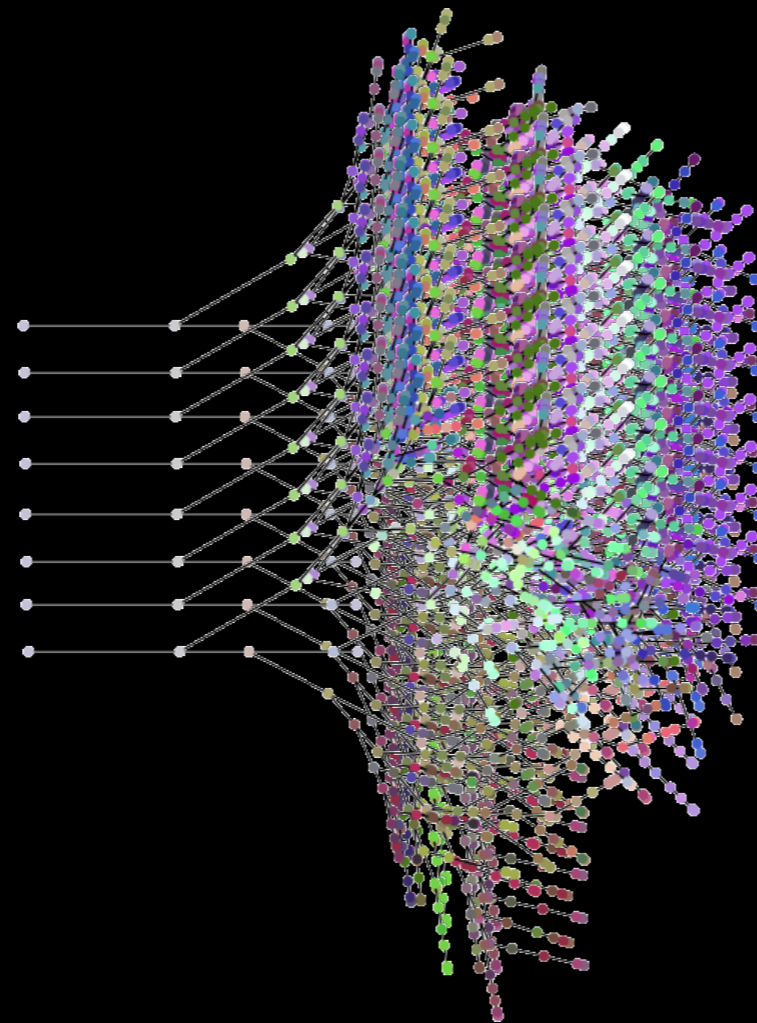
Right temporal lobe

Methods

6. Feature importance measurements

Random Forest

Image features
from within
temporal lobes



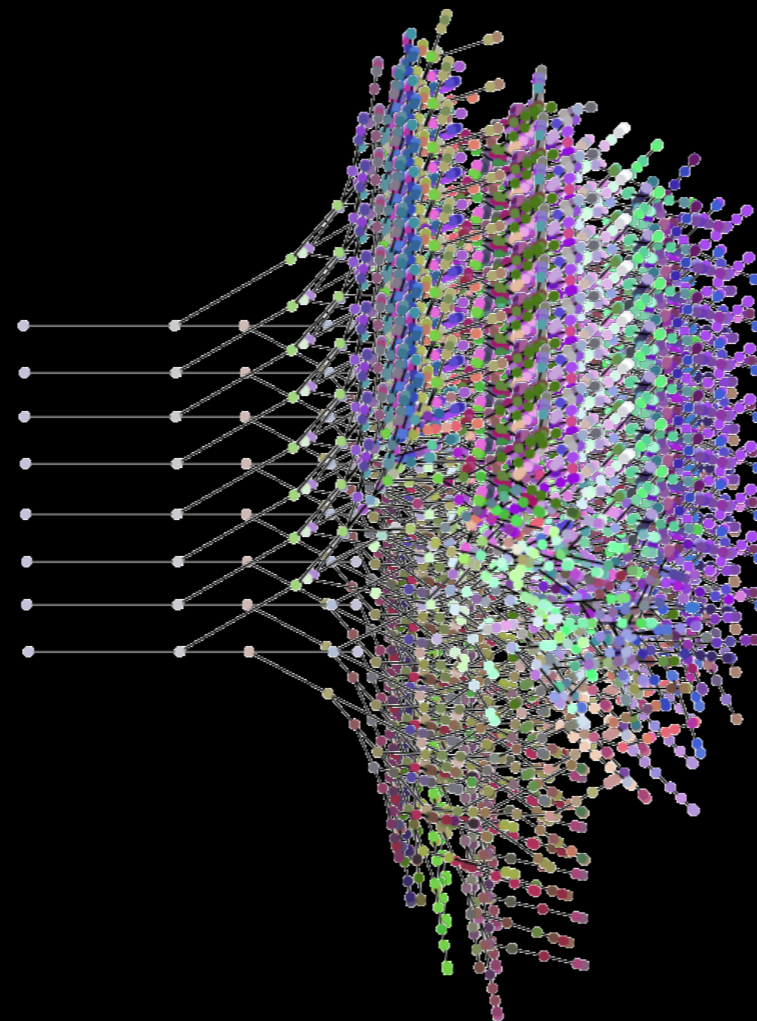
Left temporal lobe

Right temporal lobe

Methods

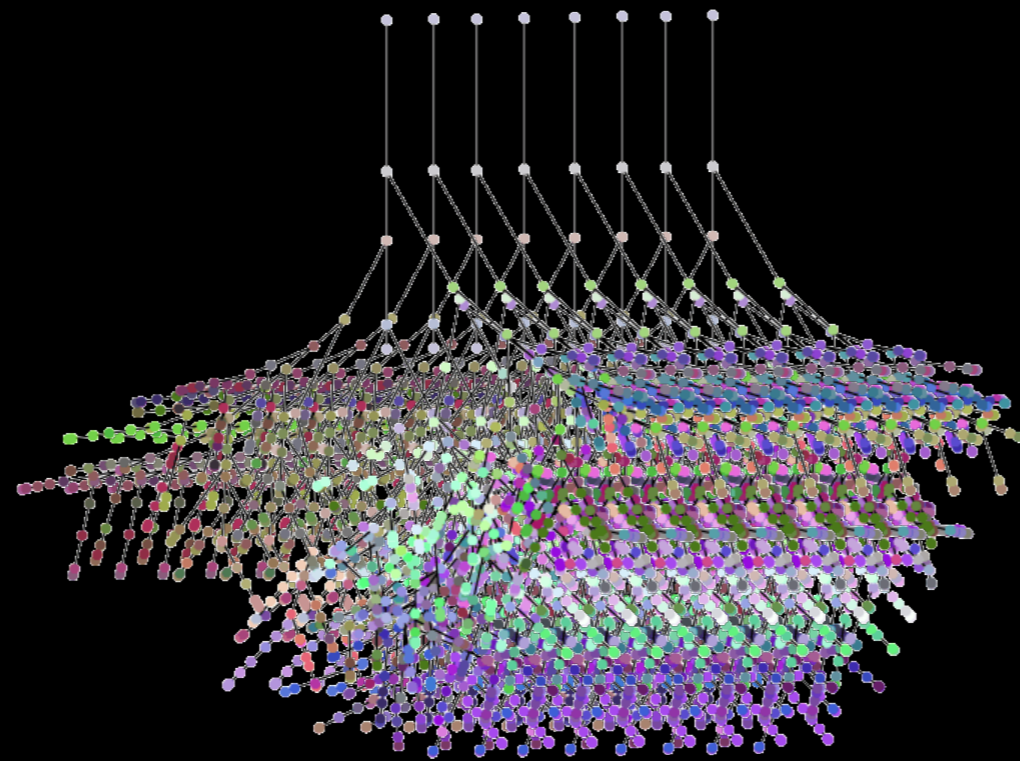
6. Feature importance measurements

Random Forest



Methods

6. Feature importance measurements



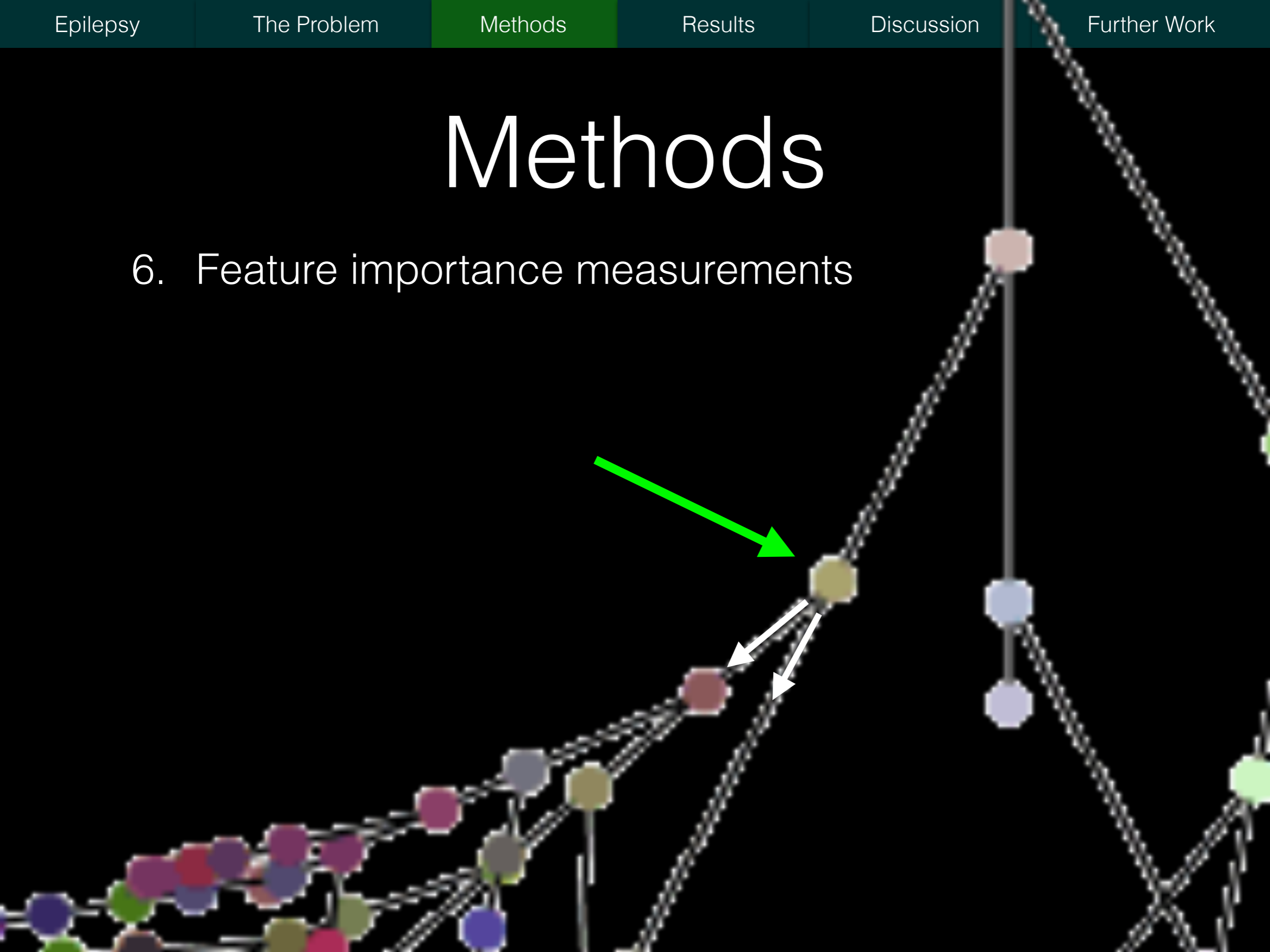
Methods

6. Feature importance measurements



Methods

6. Feature importance measurements



Methods

6. Feature importance measurements

A feature's importance is: the drop in Gini impurity it provides weighted by the chance of reaching that decision in the tree, averaged across all trees in the forest

Provides a quantitative measure of the 'usefulness' of each feature for distinguishing between classes

Methods

7. Generate Importance Maps

Visualisations of feature importances created in the form of Importance Maps

Can be thought of as 'heat maps' of abnormality

Results

SVM Classification accuracy using features:

MR **positive**:

Accuracy: 94% (CI 86-98%) using top 3 features

MR **negative**:

Accuracy: 82% (CI 63-93%) using top 38 features

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Results

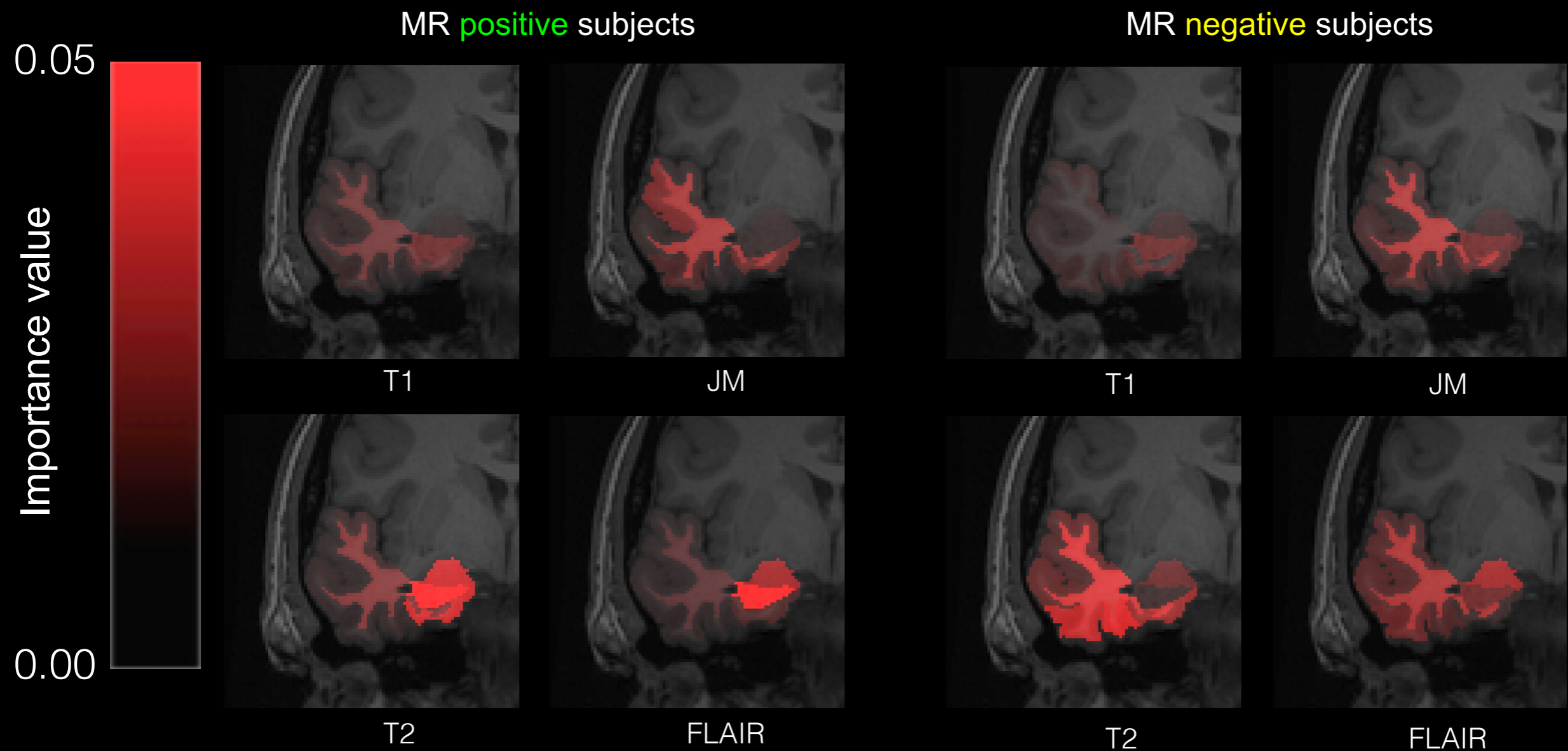
MR **positive** subjects

MR **negative** subjects

<u>Region</u>	<u>Feature</u>	<u>Region</u>	<u>Feature</u>
1) Hippocampus	vol	1) Amygdala	std T1
2) Hippocampus	mean T2	2) Fusiform Gyrus	mean T2
3) Hippocampus	std FLAIR	3) Temporal White Matter	mean T2
4) Temporal White Matter	vol	4) Amygdala	std T2
5) Hippocampus	mean FLAIR	5) Inferior Temporal Gyrus	mean T2
6) Hippocampus	std T1	6) Planum Temporale	std T2
7) Temporal Pole	mean T2	7) Hippocampus	std T2
8) Parahippocampal Gyrus	mean T2	8) Temporal White Matter	mean JM
9) Amygdala	mean T2	9) Hippocampus	std T1
10) Hippocampus	std T2	10) Amygdala	mean FLAIR

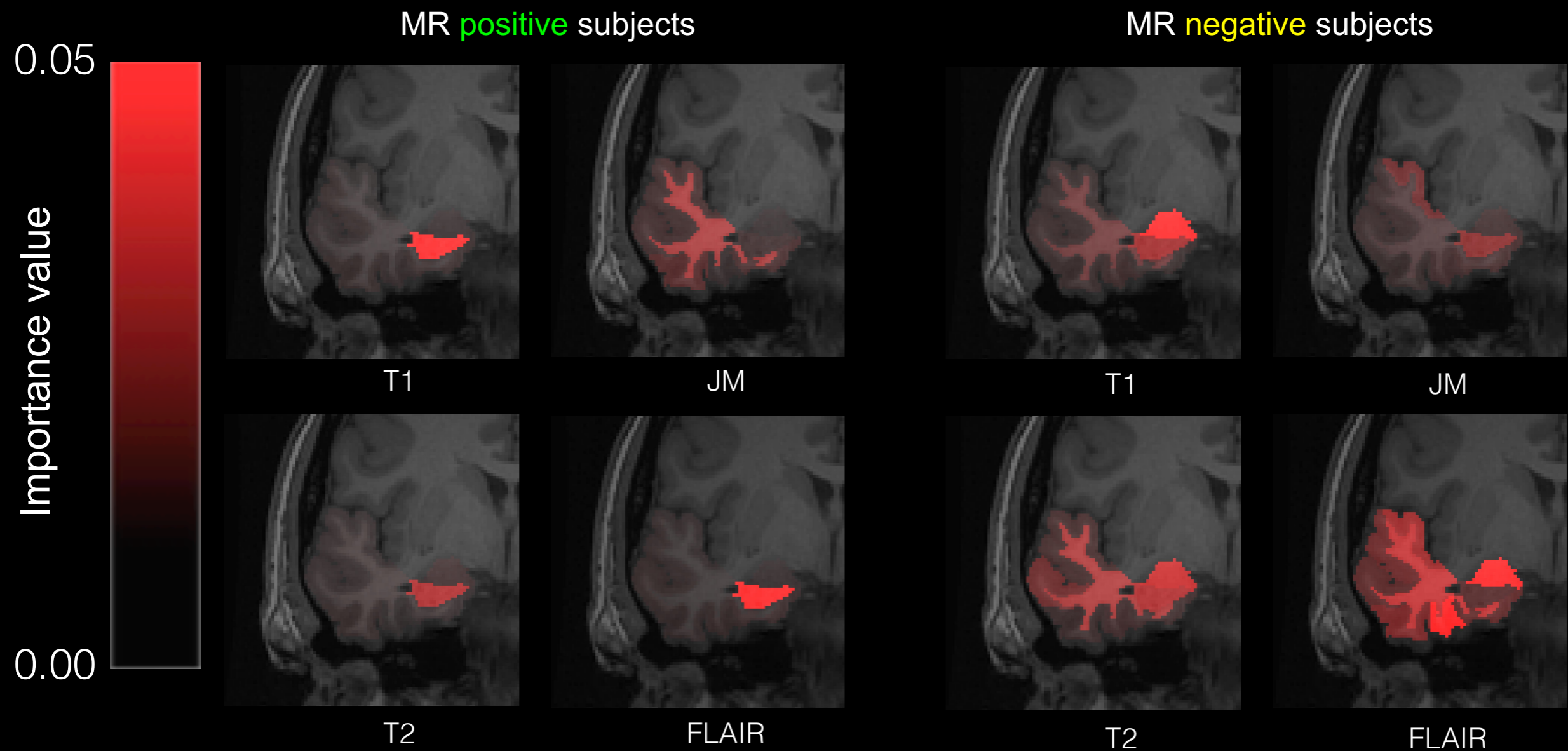
Results

Mean Intensity R - L difference Importance Maps



Results

Intensity standard deviation R - L difference Importance Maps



Results

R/L volume ratio Importance Maps



Discussion

We have demonstrated that:

- MR negative images do contain abnormalities
- These abnormalities seem to lie in a different pattern from those seen in MR positive (visible disease) cases

Discussion

More generally:

- Generates a disease's 'abnormality signature' in imaging investigations
- A training tool for radiologists?

Further Work

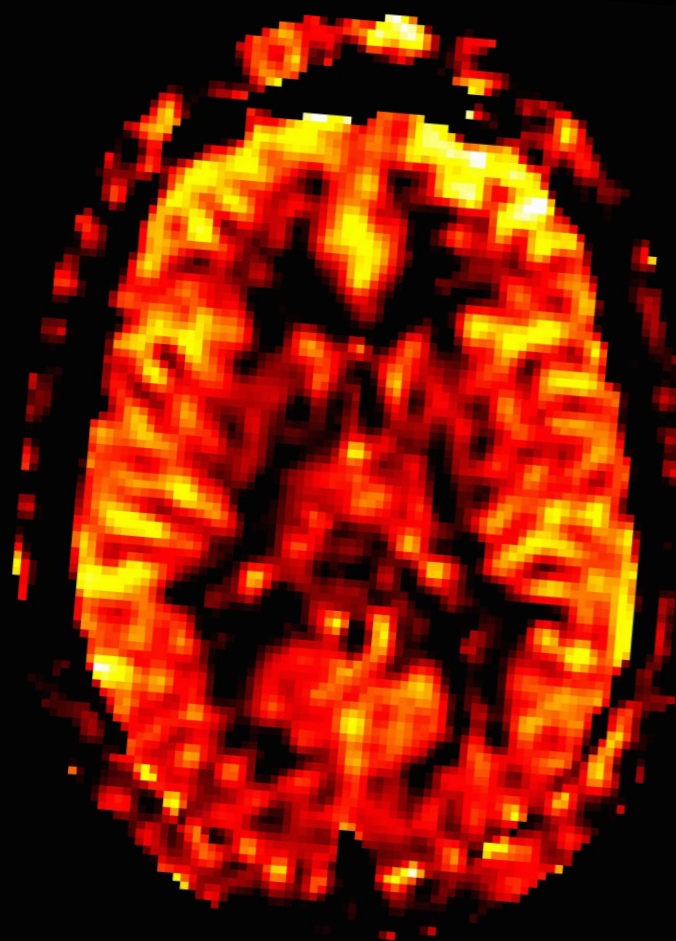
- Could be a way of characterising the amount of extra information novel imaging modalities provide
- The methodology could be applied to other disease processes with subtle or poorly understood visual appearances

Questions

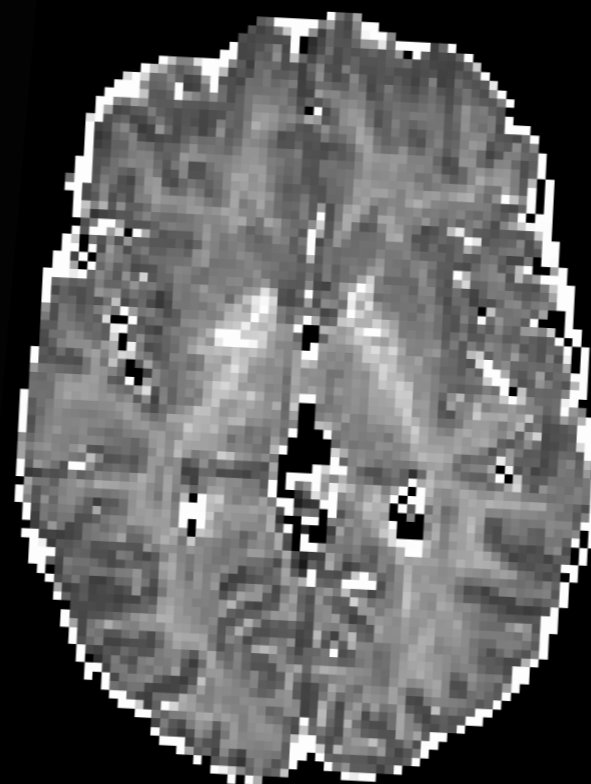
Further Work

Could be a way of characterising the amount of extra information novel imaging modalities provide

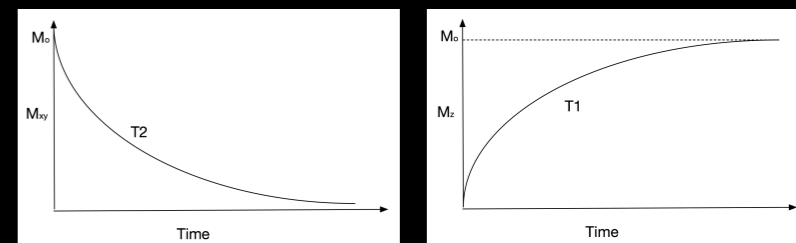
Arterial Spin Labelling



Multi-Compartment Diffusion Modelling



T1, T2 Relaxometry



Sodium Imaging

