

# Neuroimaging and mathematical modelling

## Lesson 2: Voxel Based Morphometry

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# Volume and surface morphometry

Brain volume

White matter

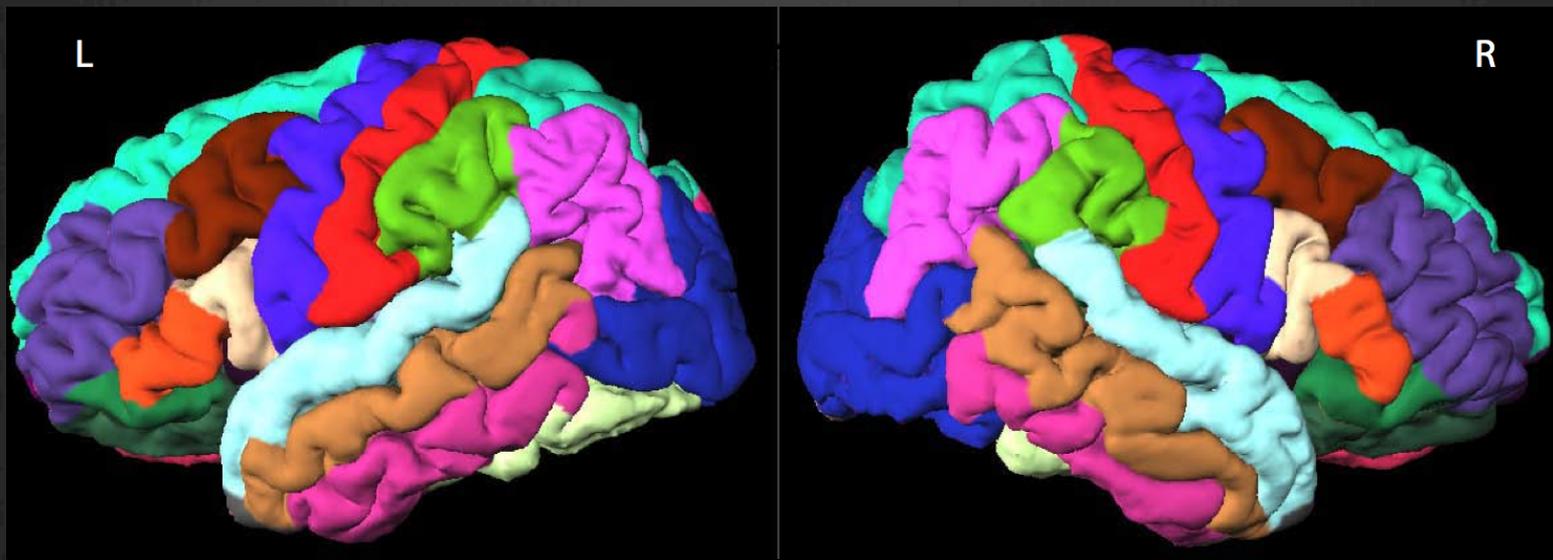
Grey matter

Volume of specific brain structures

hippocampus (Alzheimer's disease, memory loss states)

frontal lobe (intelligence!, analytical brain)

basal ganglia (Parkinson's disease, metabolic disorders)



# Examples applications of VBM

- ⊗ Many scientifically or clinically interesting questions might relate to the local volume of regions of the brain
- ⊗ For example, whether (and where) local patterns of brain morphometry help to:
  - ⊗ Distinguish groups (e.g. schizophrenics and healthy controls)
  - ⊗ Explain the changes seen in development and aging
  - ⊗ Understand plasticity, e.g. when learning new skills
  - ⊗ Find structural correlates (scores, traits, genetics, etc.)

# Volume and surface morphometry

This is where mathematicians and physicists come in to the scene!

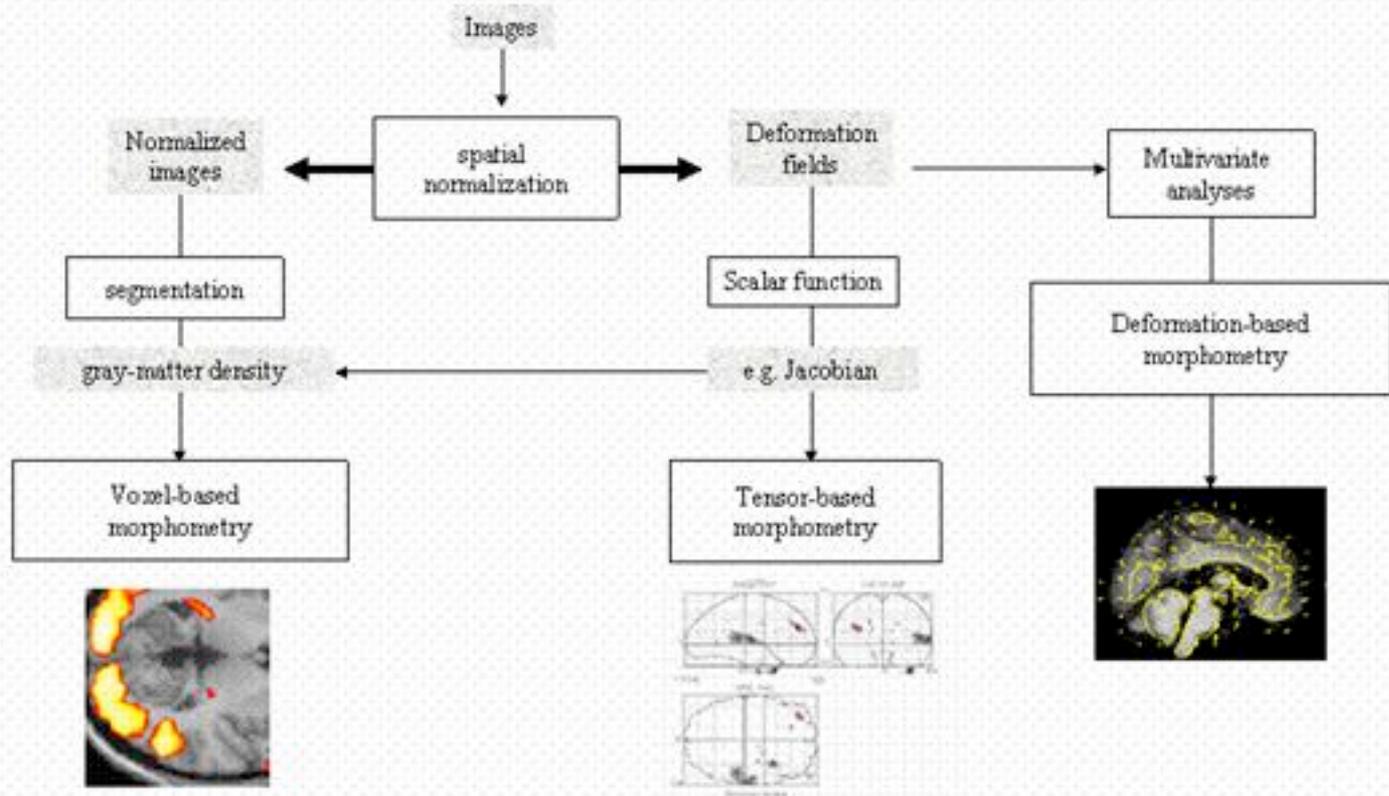
Lots of online free tools to study volume of brain structures and identify surface anatomy

<http://surfer.nmr.mgh.harvard.edu/fswiki/FreeSurferAnalysisPipelineOverview>

<http://surfer.nmr.mgh.harvard.edu/fswiki/Slicer>

<http://afni.nimh.nih.gov/>

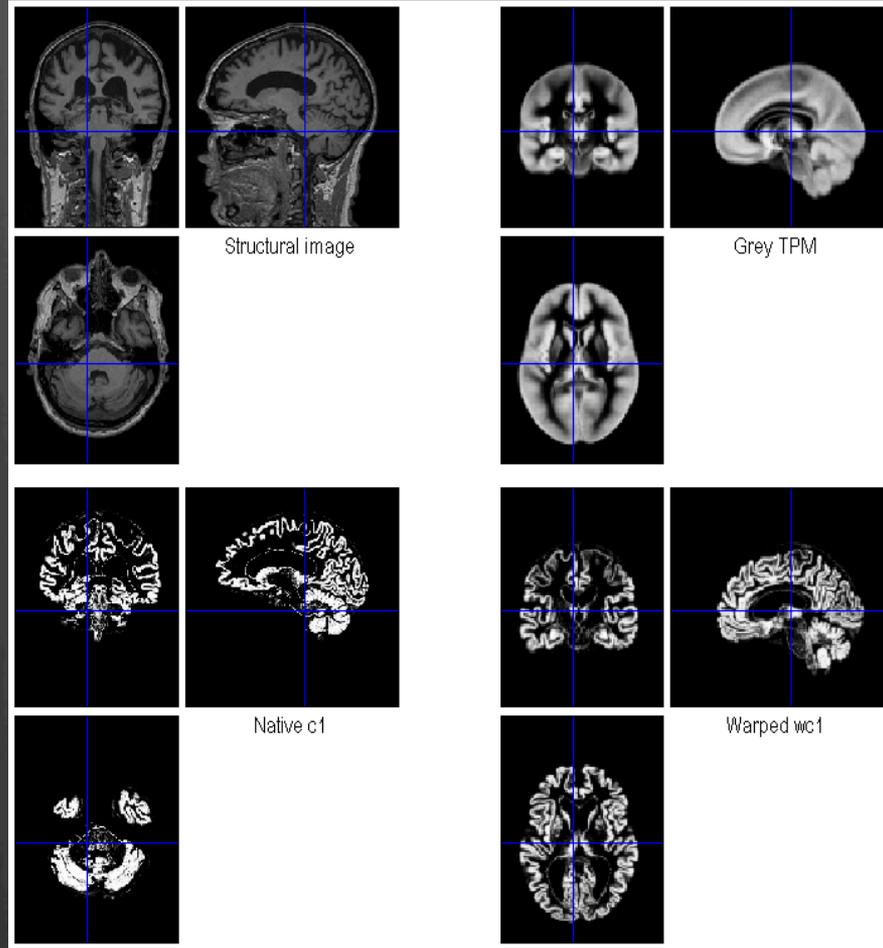
# Computational neuroanatomy



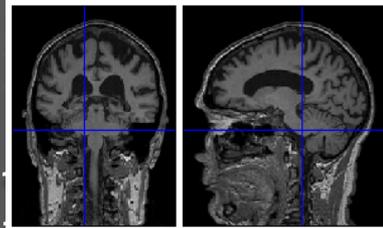
# VBM in pictures

Segment

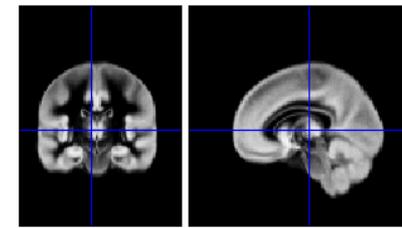
Normalise



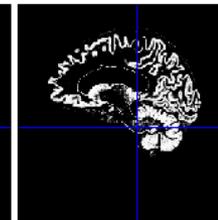
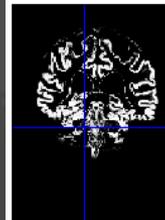
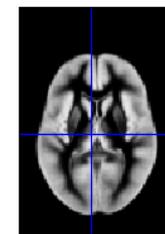
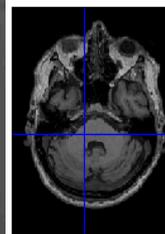
# VBM in



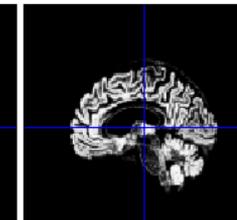
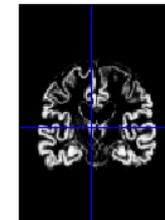
Structural image



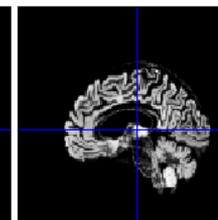
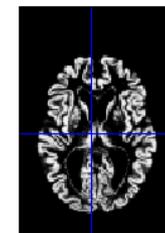
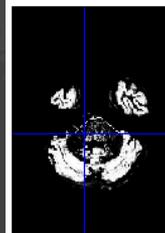
Grey TPM



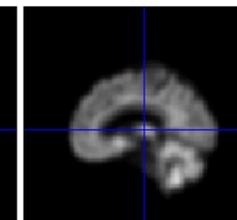
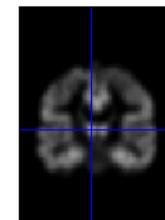
Native c1



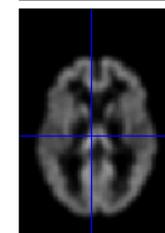
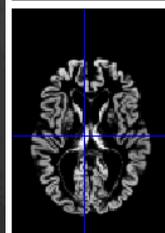
Warped wc1



Modulated mwc1



Smoothed smwc1



segment

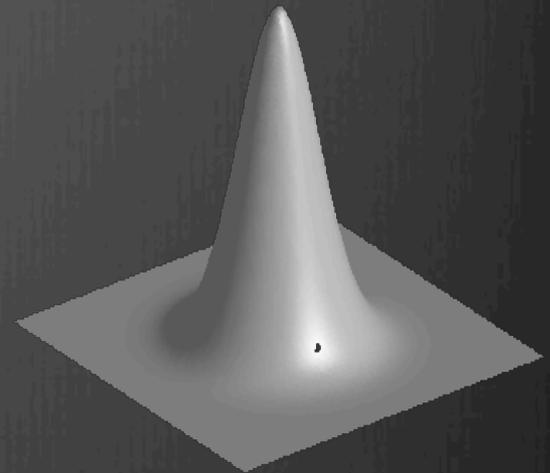
normalise

**Modulate**

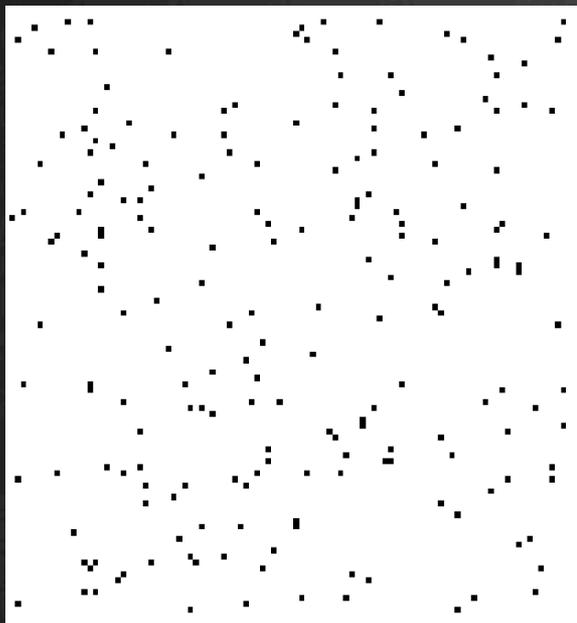
**Smooth**

# Smoothing

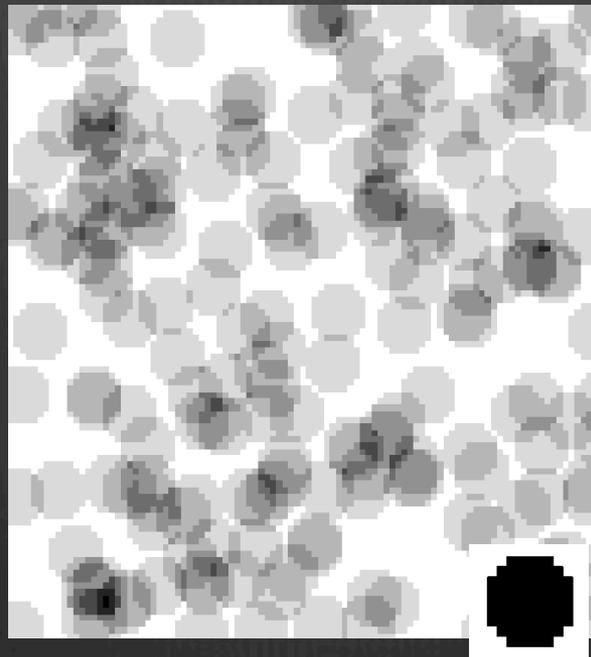
Each voxel after smoothing effectively becomes the result of applying a weighted region of interest (ROI).



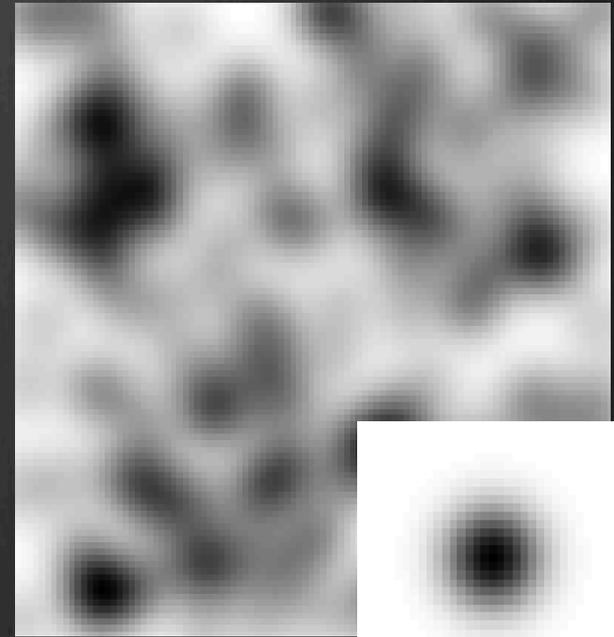
Before convolution



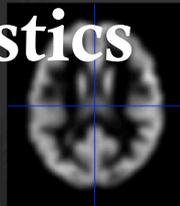
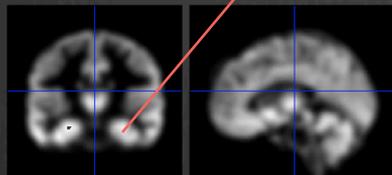
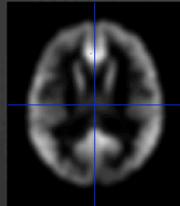
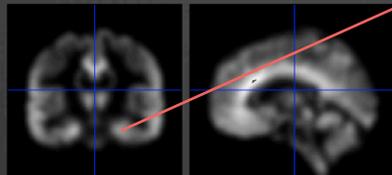
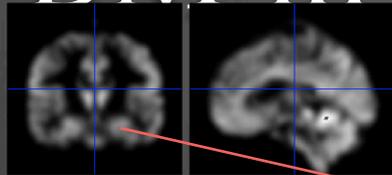
Convolved with a circle



Convolved with a Gaussian



# VBM in pictures



$$\begin{bmatrix} a1xyz \\ a2xyz \\ \vdots \\ aNxyz \end{bmatrix} = Y = X\beta_{xyz} + e_{xyz}$$

$e_{xyz} \sim N(0, \sigma_{xyz}^2 V)$

$$X = \begin{bmatrix} 1 & 0 \\ 1 & 0 \\ \vdots & \vdots \\ 0 & 1 \\ 0 & 1 \end{bmatrix}$$

**Voxel-wise statistics**

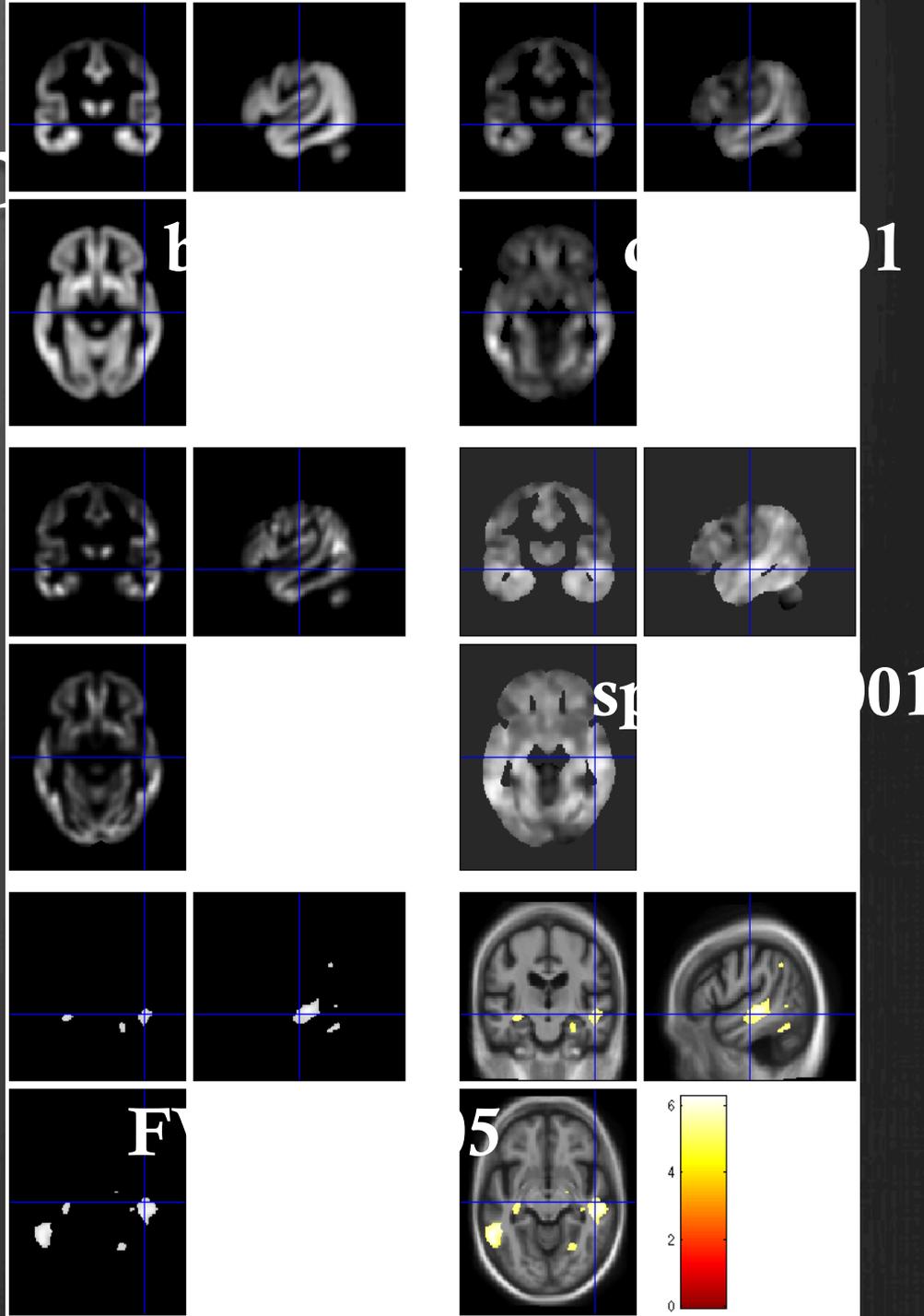
segment

normalise

modulate

smooth

# VBM in

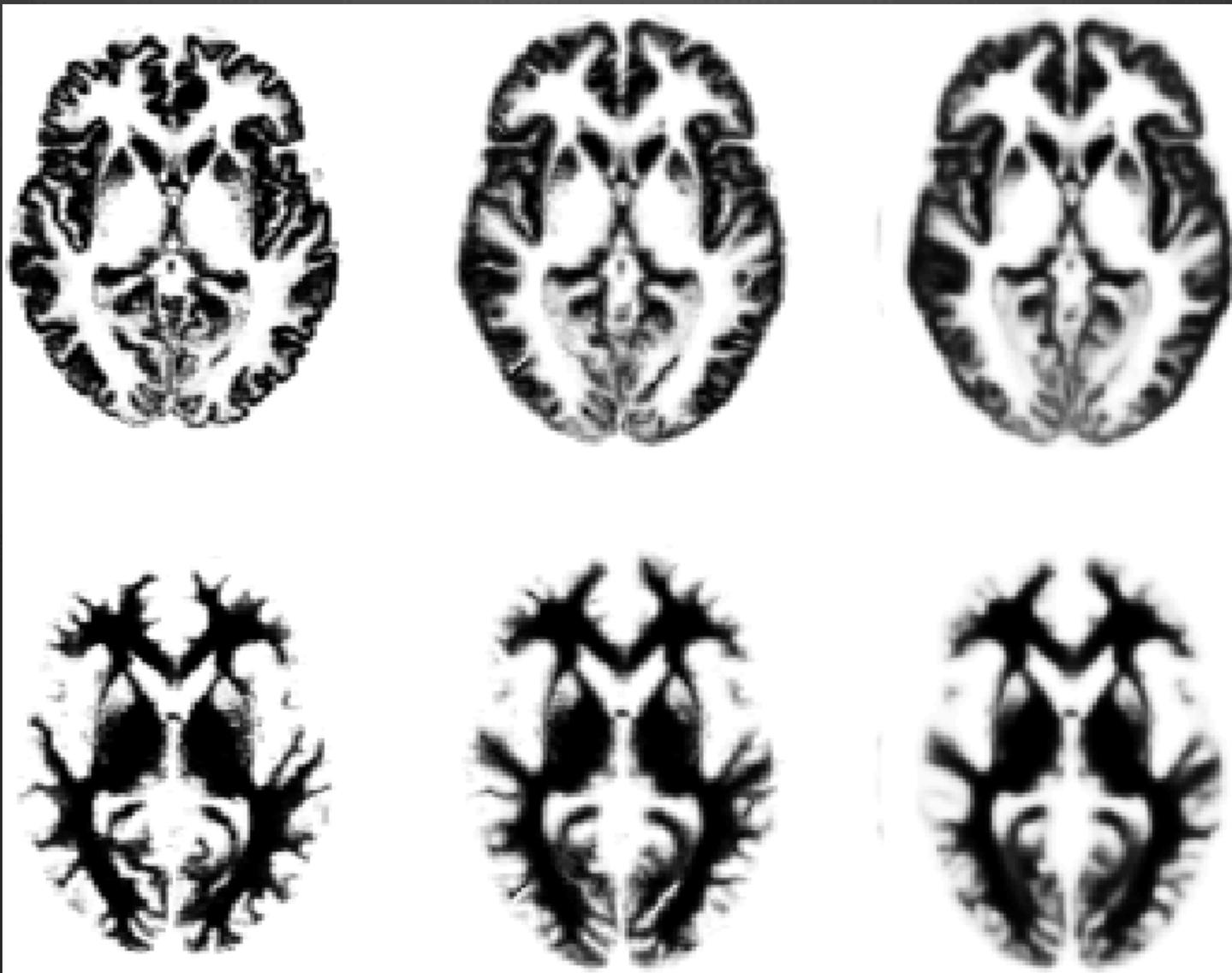


## Voxel-wise statistics

Original

Warped

Template



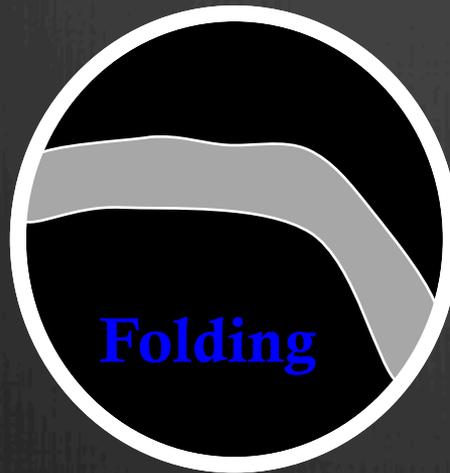
# Smoothing as a locally weighted

ROI

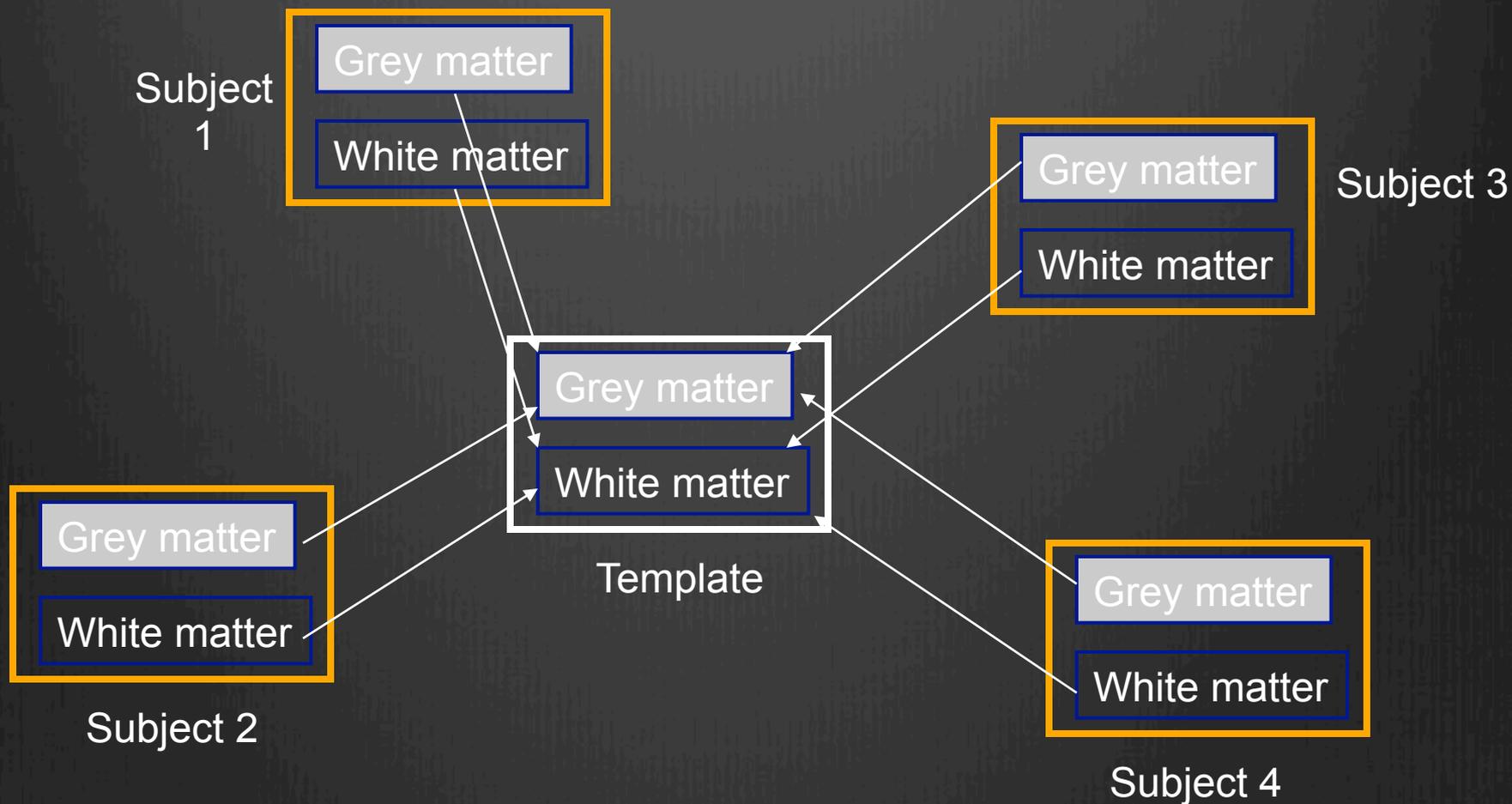


- ⊗ VBM > ROI: no subjective (or arbitrary) boundaries
- ⊗ VBM < ROI: harder to interpret blobs & characterise error

# Interpreting findings



# Simultaneous registration of GM to GM and WM to WM



# Template

Iteratively  
generated from  
471 subjects

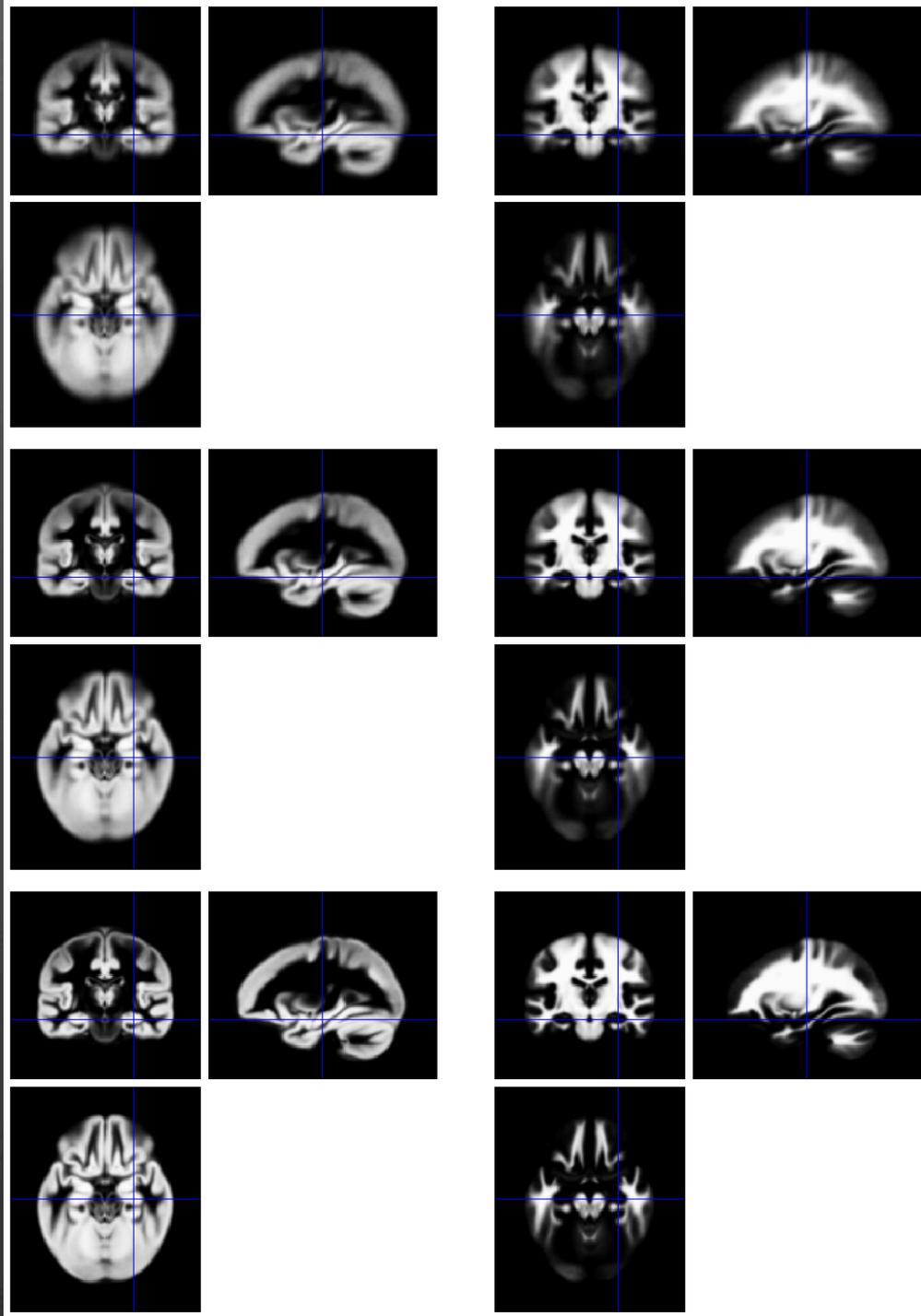
Began with rigidly  
aligned tissue  
probability maps

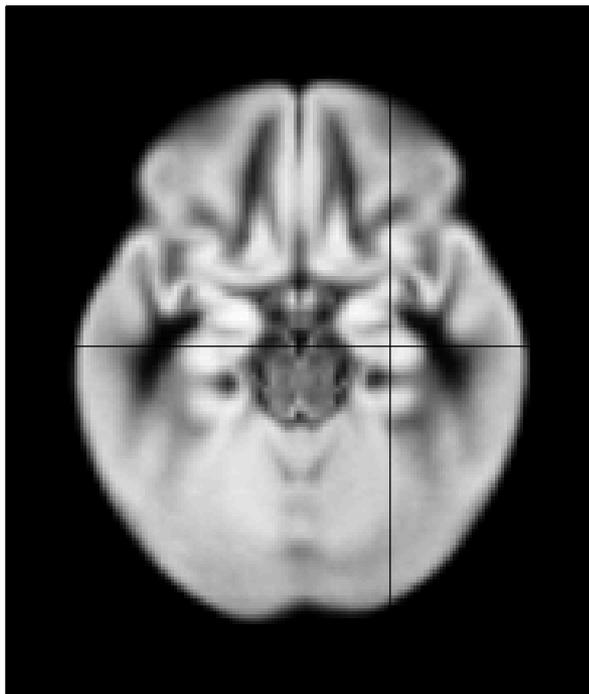
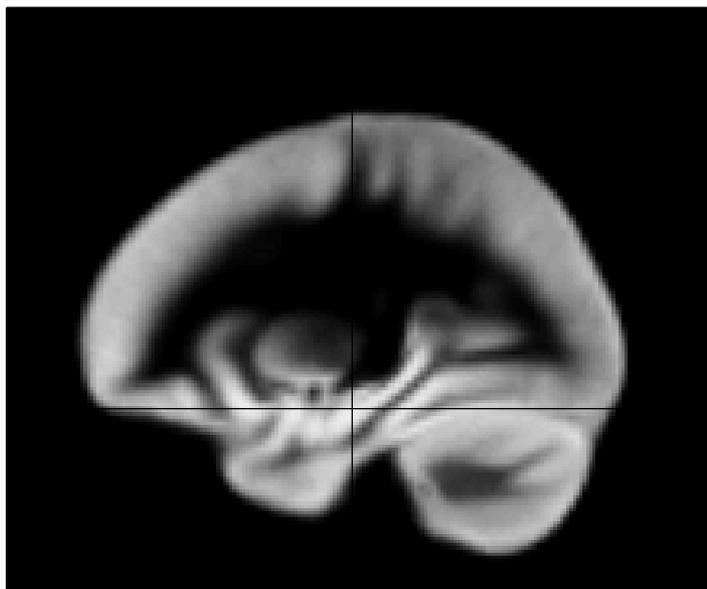
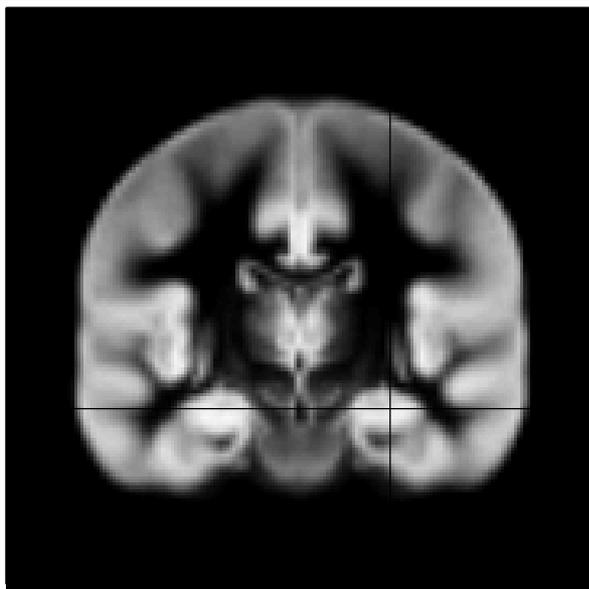
Used an inverse  
consistent  
formulation

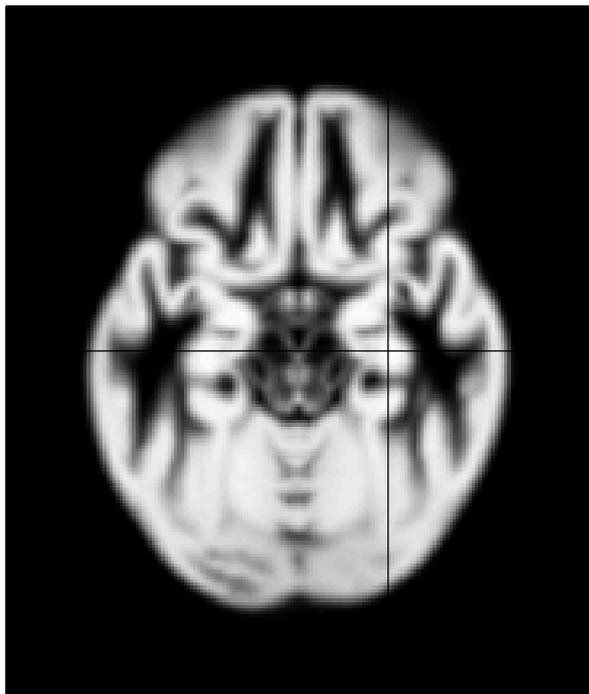
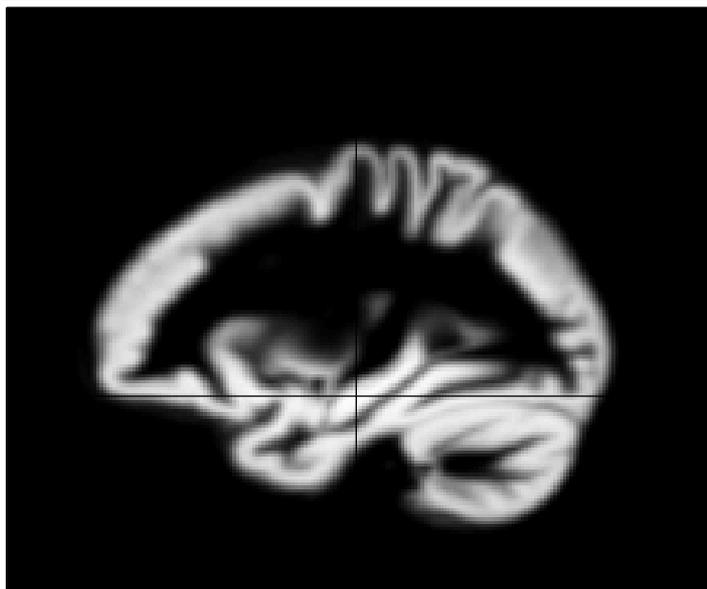
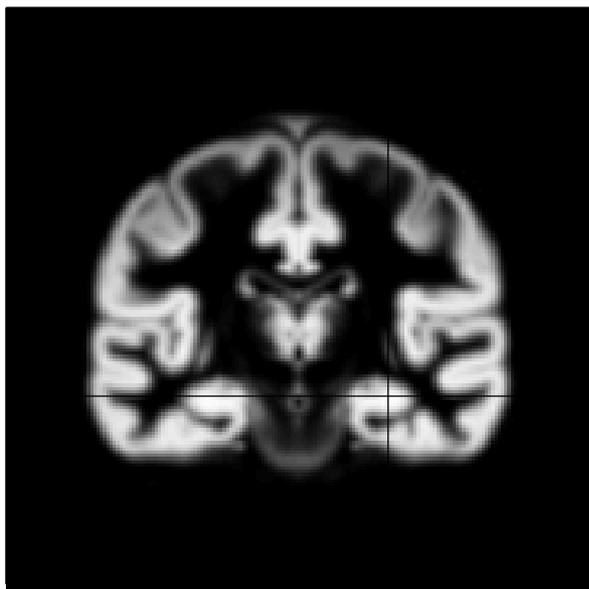
Initial  
Average

After a few  
iterations

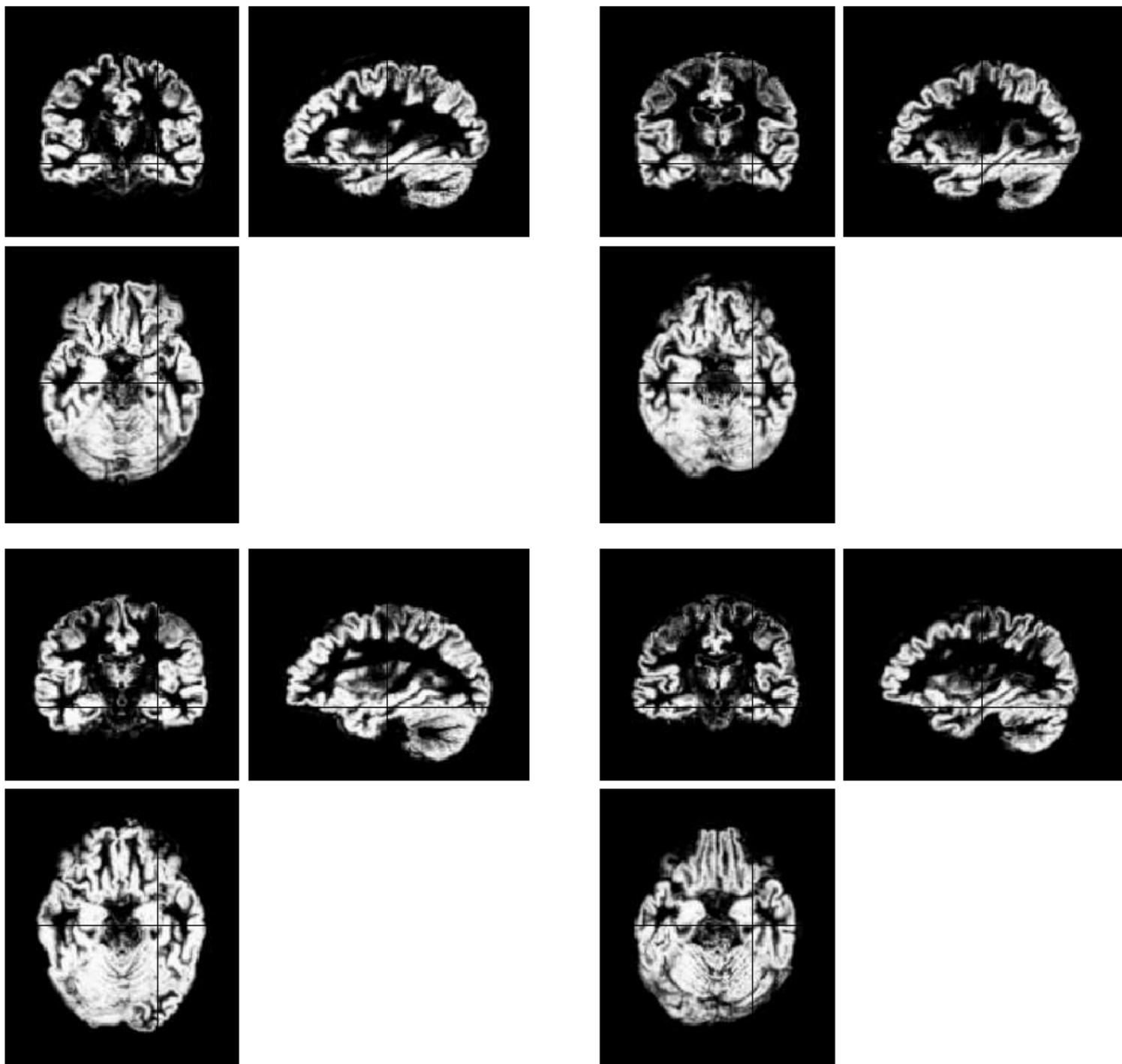
Final  
template



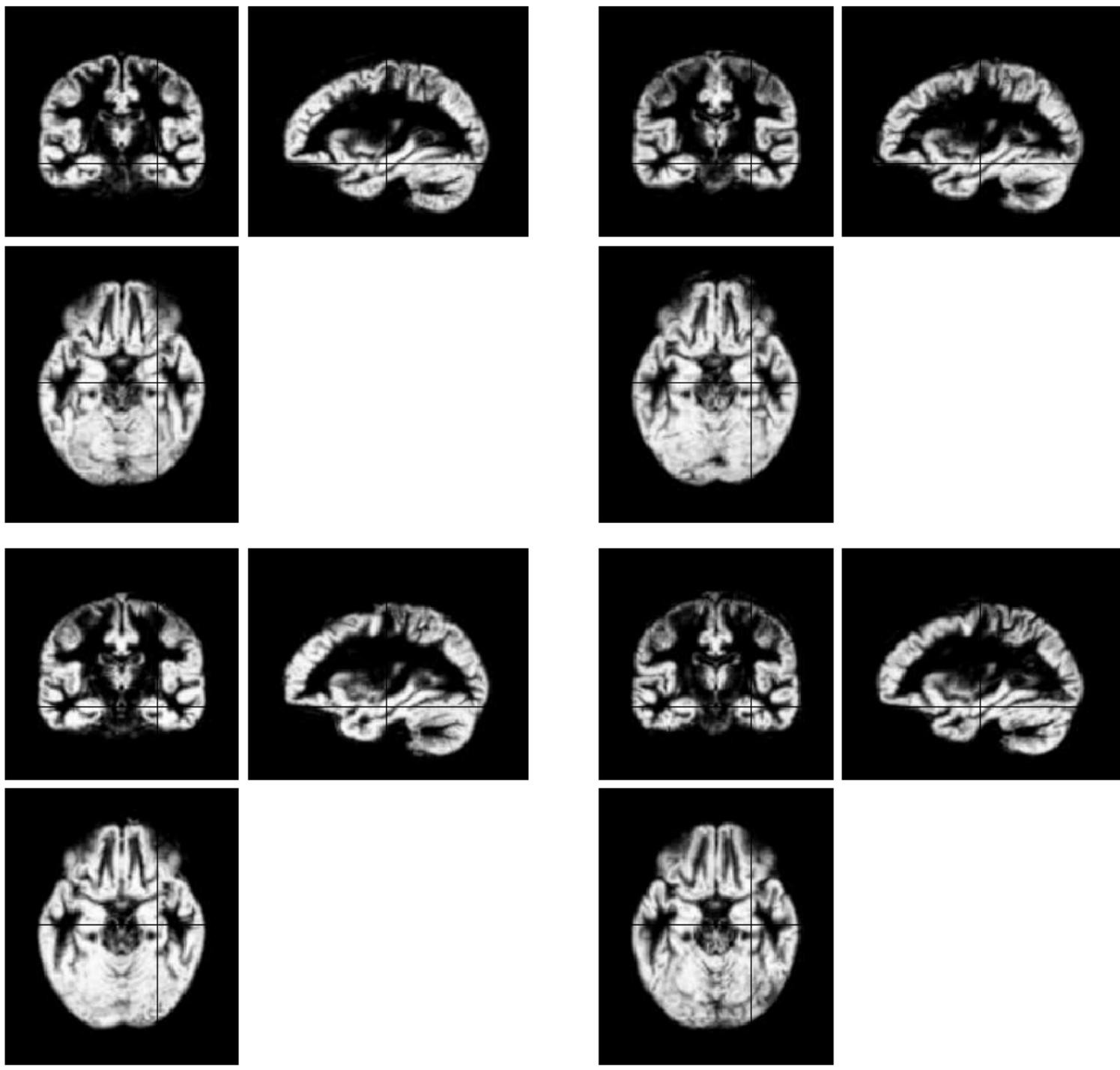


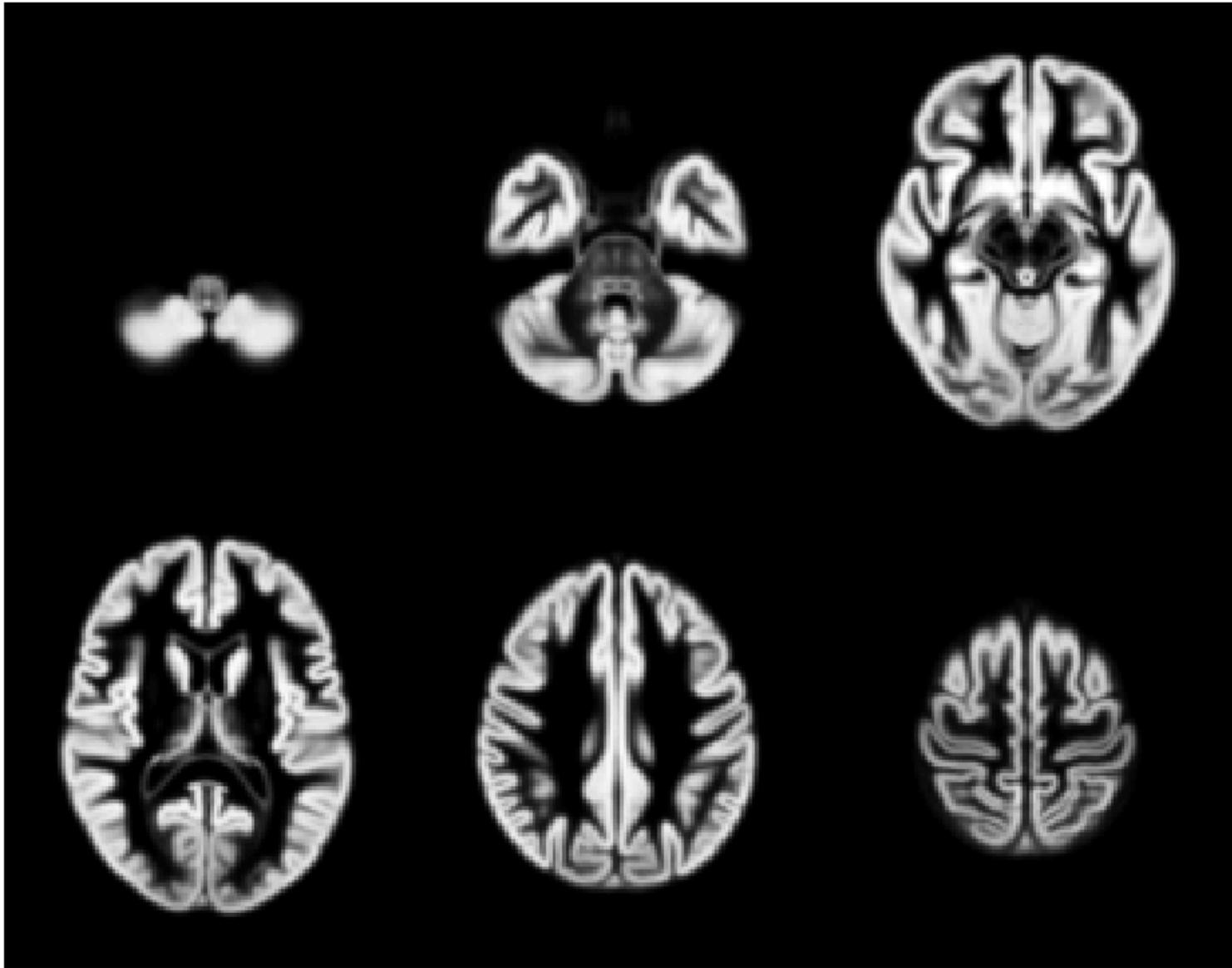


Initial  
GM images

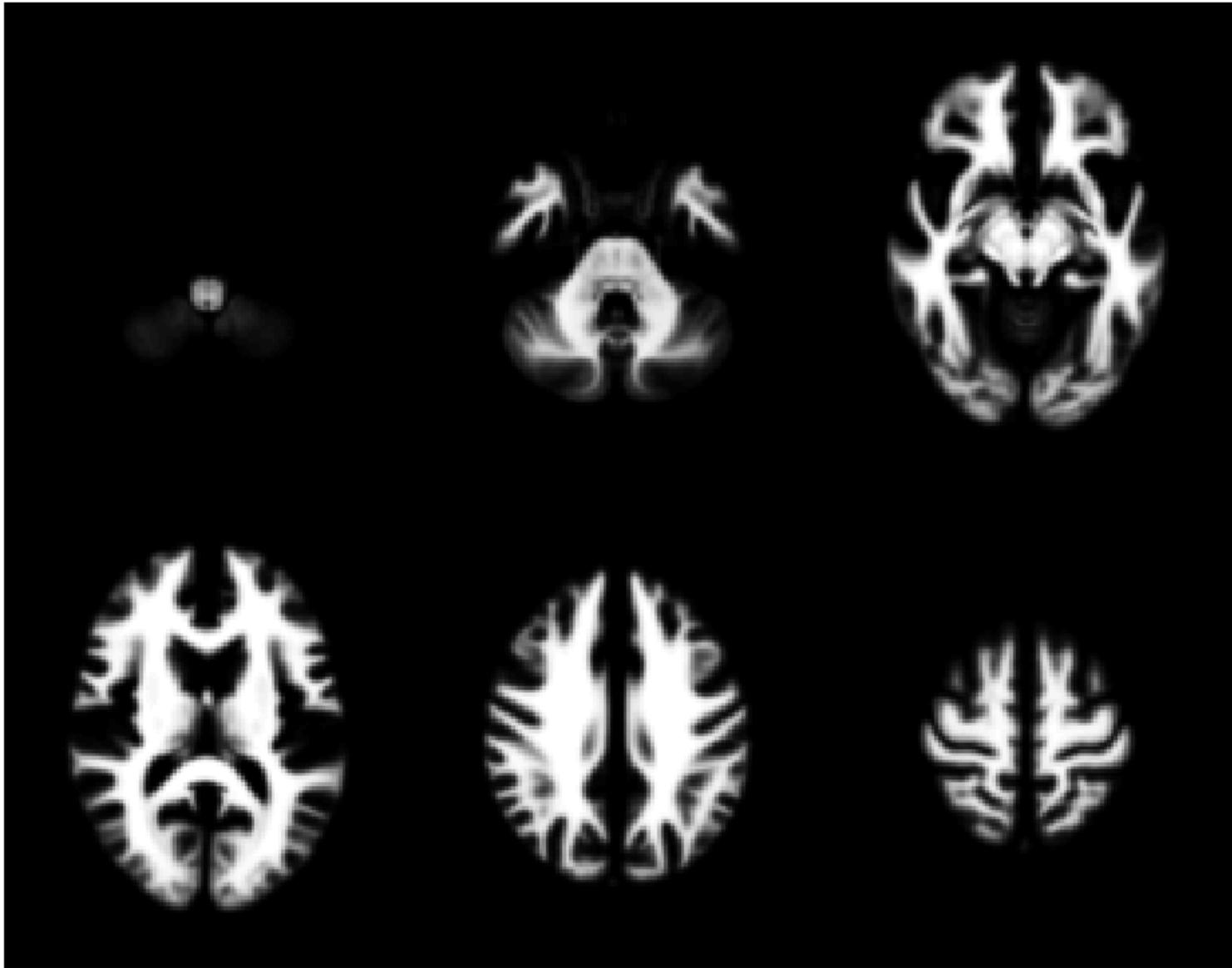


Warped  
GM images

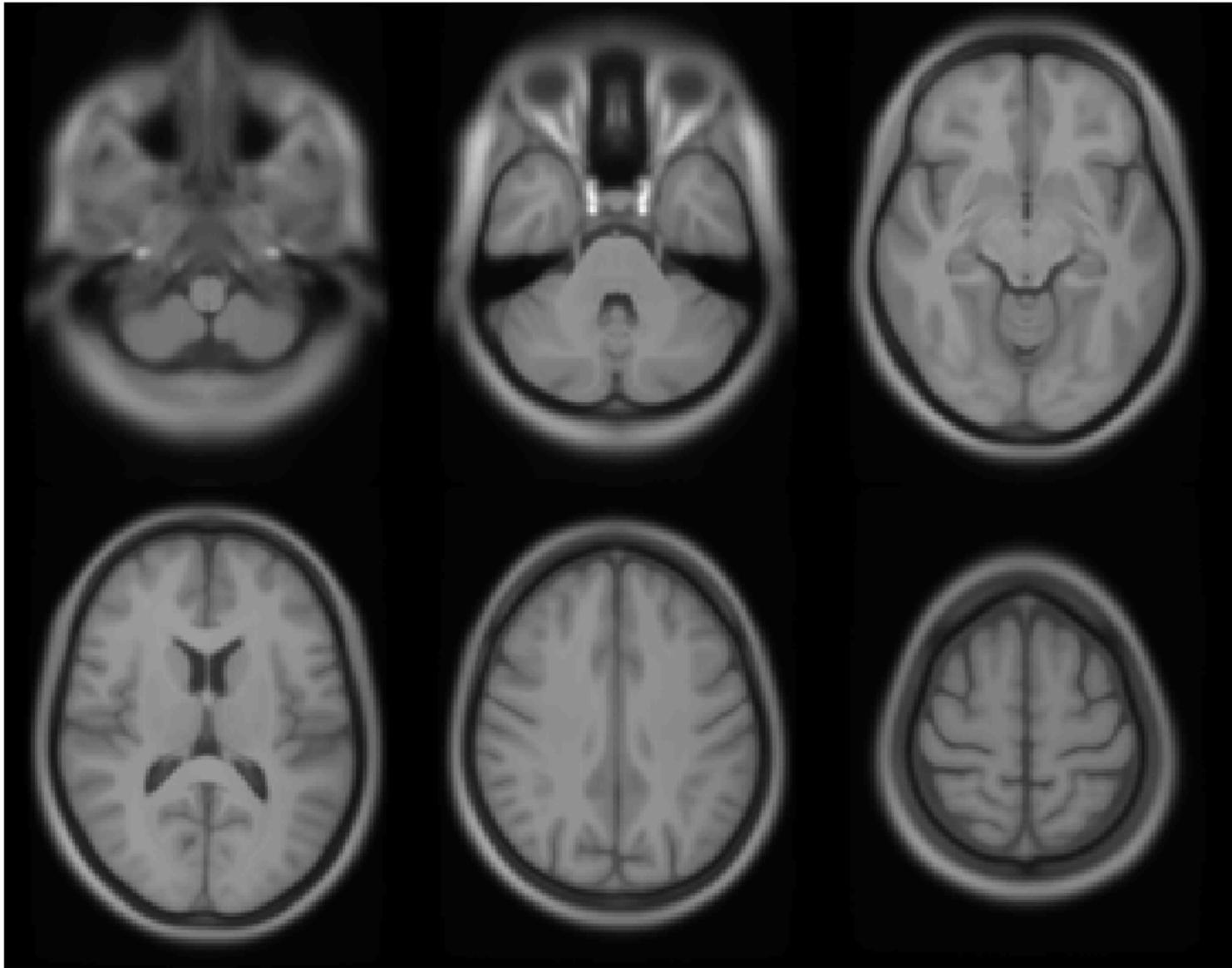




471 Subject Average



471 Subject Average

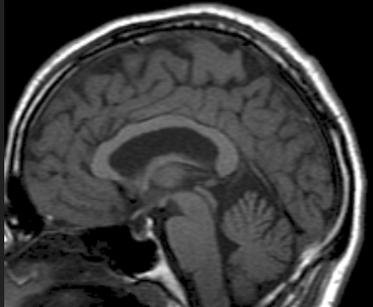


471 Subject Average

# SPM for Anatomical MRI

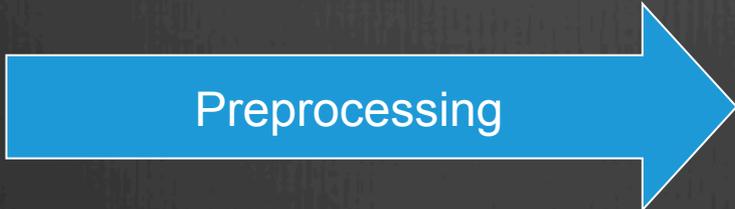
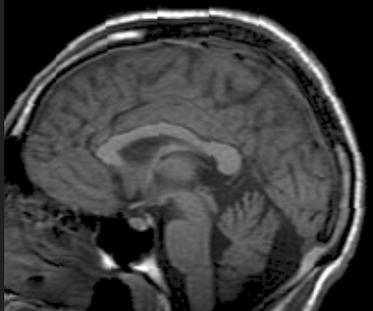
Group-wise statistics

Anatomical MRI



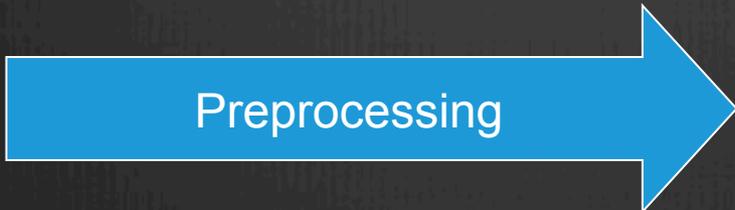
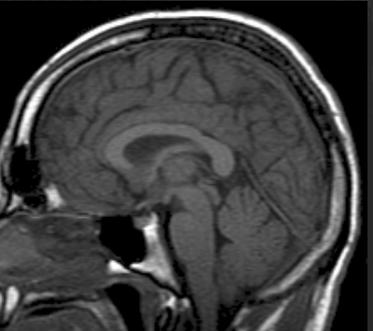
Spatially Normalised Grey Matter Image

Anatomical MRI



Spatially Normalised Grey Matter Image

Anatomical MRI



Spatially Normalised Grey Matter Image



# Some References

- ⊗ Wright, McGuire, Poline, Traverso, Murray, Frith, Frackowiak & Friston. *A voxel-based method for the statistical analysis of gray and white matter density applied to schizophrenia*. Neuroimage 2(4):244-252 (1995).
- ⊗ Ashburner & Friston. “*Voxel-based morphometry-the methods*”. Neuroimage 11(6):805-821, (2000).
- ⊗ Mechelli et al. *Voxel-based morphometry of the human brain...* Current Medical Imaging Reviews 1(2) (2005).
- ⊗ Ashburner & Friston. “*Unified Segmentation*”. NeuroImage 26:839-851, 2005.
- ⊗ Ashburner. “*A Fast Diffeomorphic Image Registration Algorithm*”. NeuroImage 38:95-113 (2007).
- ⊗ Ashburner & Friston. “*Computing Average Shaped Tissue Probability Templates*”. NeuroImage 45:333-341 (2009).
- ⊗ Klein et al. *Evaluation of 14 nonlinear deformation algorithms applied to human brain MRI registration*. NeuroImage 46(3):786-802 (2009).
- ⊗ Ashburner. “*Computational Anatomy with the SPM software*”. Magnetic Resonance Imaging 27(8):1163-1174 (2009).
- ⊗ Ashburner & Klöppel. “*Multivariate models of inter-subject anatomical variability*”. NeuroImage 56(2):422-439 (2011).

# Interpreting findings

VBM is sometimes described as  
“unbiased whole brain volumetry”

Regional  
variation in  
registration  
accuracy



Segmentation  
problems, issues  
with analysis  
mask



Intensity,  
folding, etc.



But significant blobs probably still indicate meaningful  
systematic effects!

# Summary

- ⊗ VBM performs voxel-wise statistical analysis on smoothed (modulated) normalised tissue segments
- ⊗ SPM performs segmentation and spatial normalisation in a unified generative model
  - ⊗ Based on Gaussian mixture modelling, with warped spatial prior probability maps, and multiplicative bias field

# End of the second lecture!

Next time (29 Nov. 2013):  
Basics of Diffusion tensor imaging

<http://www.fil.ion.ucl.ac.uk/spm/>