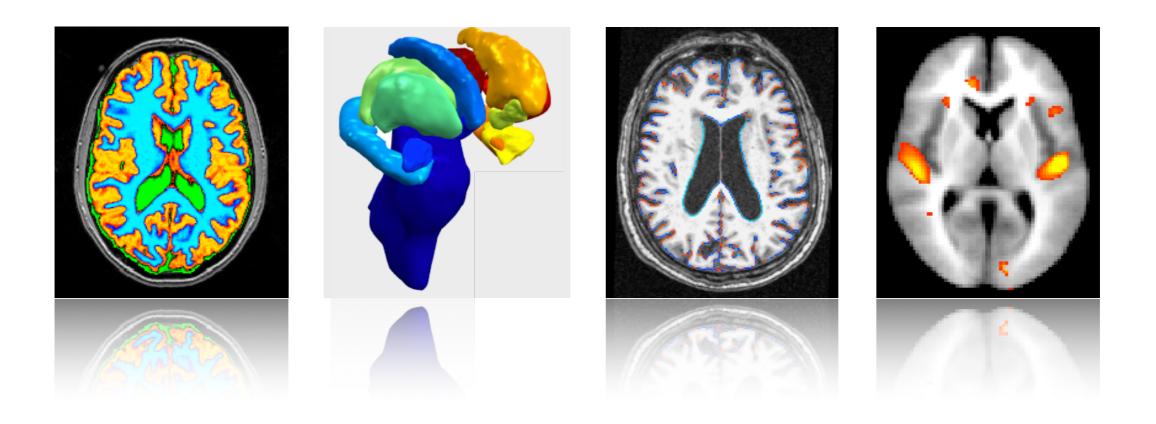


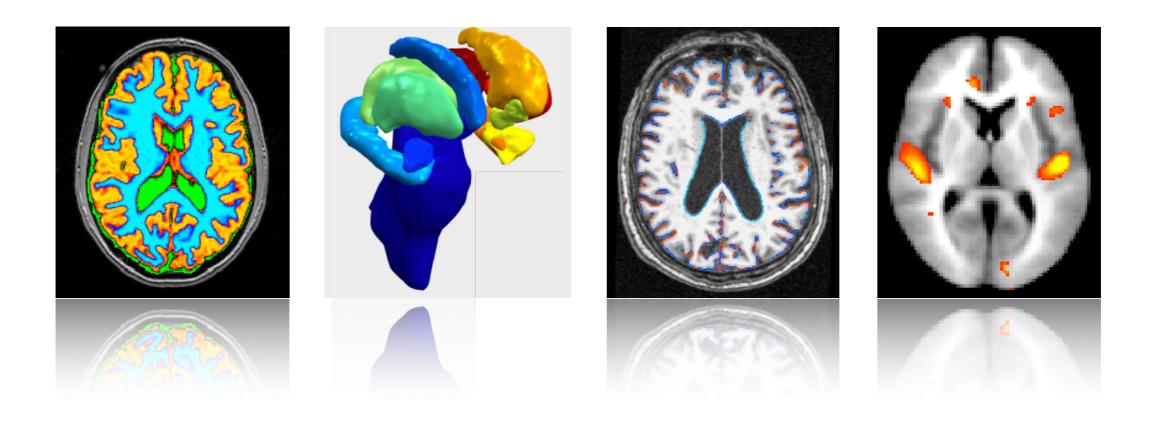
#### Structural Segmentation



- FAST tissue-type segmentation
- FIRST sub-cortical structure segmentation
- BIANCA segmentation of white matter lesions
- FSL-VBM voxelwise grey-matter density analysis
- SIENA/SIENAX global atrophy estimation



#### Structural Segmentation



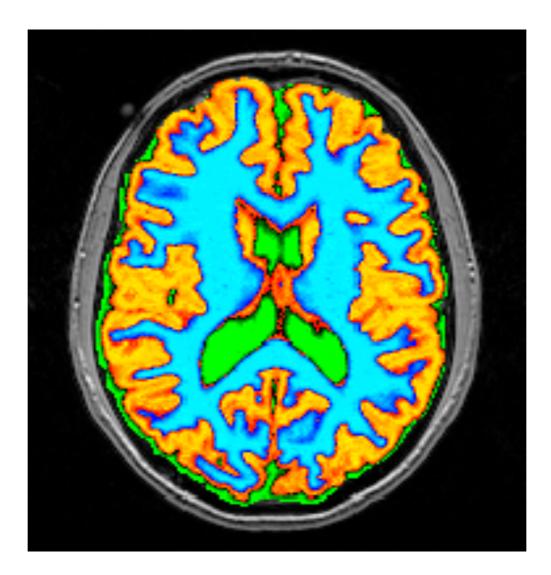
- FAST tissue-type segmentation
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- SIENA/SIENAX global atrophy estimation



#### FAST

#### FMRIB's Automated Segmentation Tool

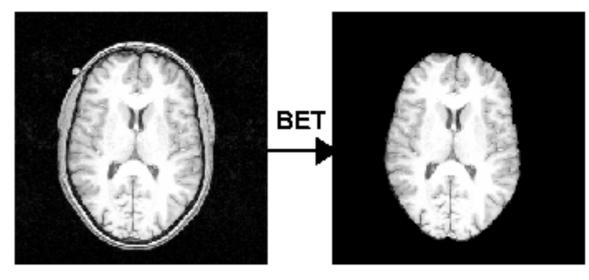
# generic tissue-type segmentation and bias field correction



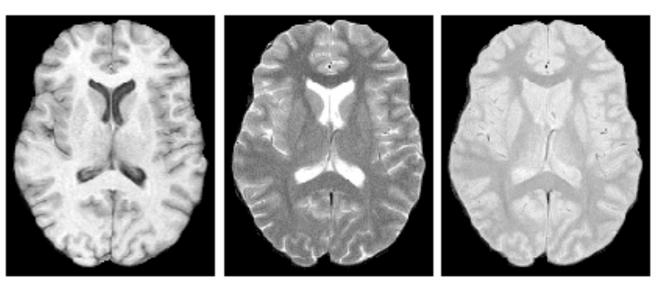


### FAST: Input

• First use BET to remove non-brain All volumetric results are highly sensitive to errors here. For bias-field correction alone the errors do not matter that much



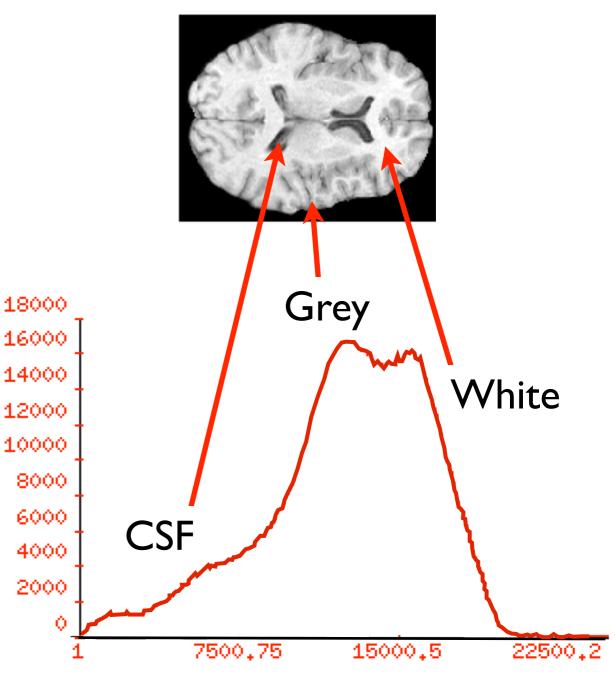
- Input is normally a single image (TI,T2, proton-density....)
- Or several inputs ("multichannel")
- For multi-channel, all must be pre-aligned (FLIRT)





# Intensity Model tissue intensity distributions

- Histogram = voxel count vs. intensity
- Model = mixture of Gaussians
- If well separated, have clear peaks; then segmentation easy
- Overlap worsened by:
  - Bias field
  - Blurring
  - Low resolution
  - Head motion
  - Noise



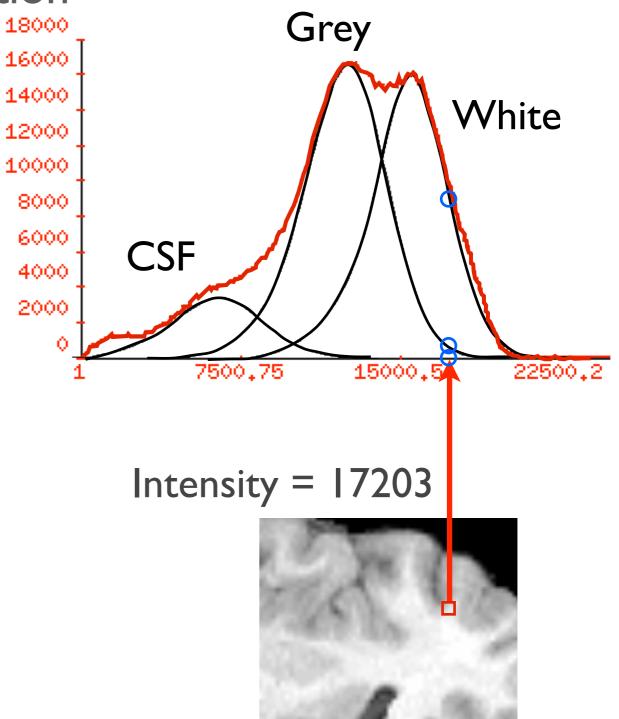


### Probability Model

- Model = mixture of Gaussians
- Probability determined for each tissue class

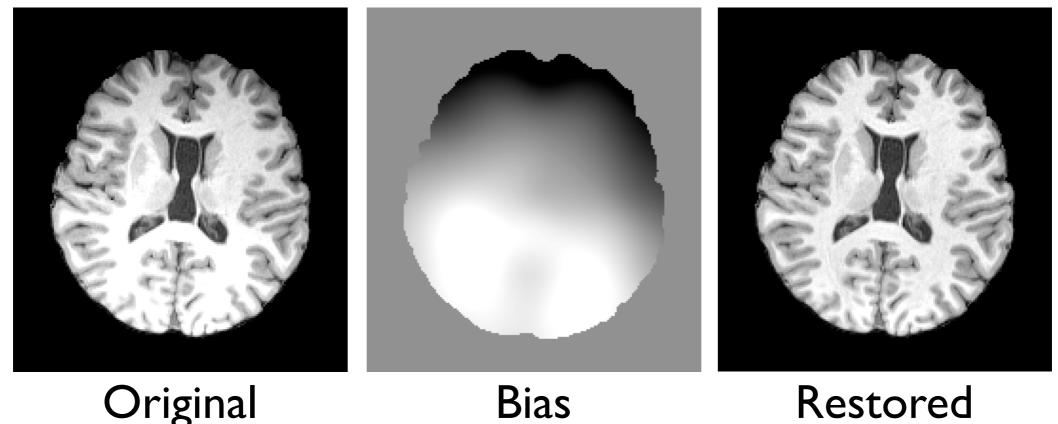
For example: Voxel near WM/GM border

P(CSF) near zero P(GM) low P(WM) moderate





#### **Bias Field Correction**

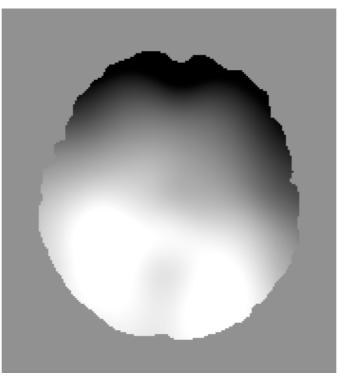


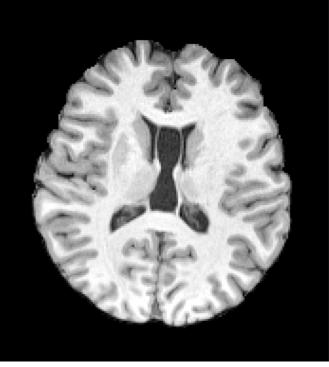
- MRI RF (radio-frequency field) inhomogeneity causes intensity variations across space
- Causes problems for segmentation
- Need to remove bias field before or during segmentation
- Becomes more common and problematic at high field



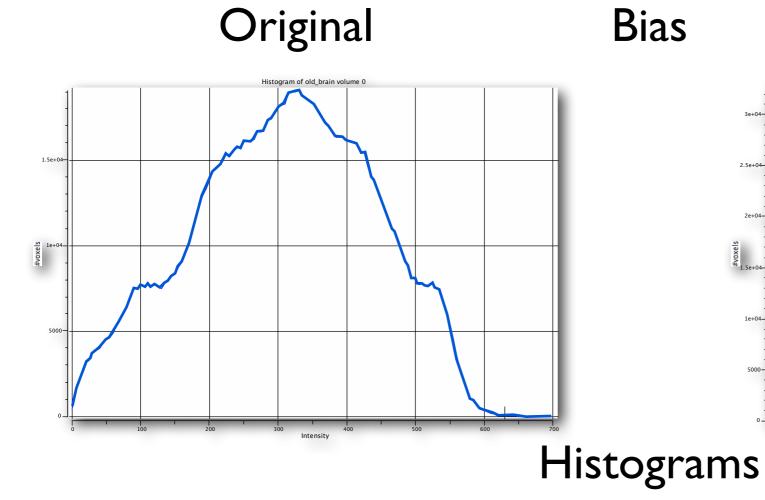
#### **Bias Field Correction**

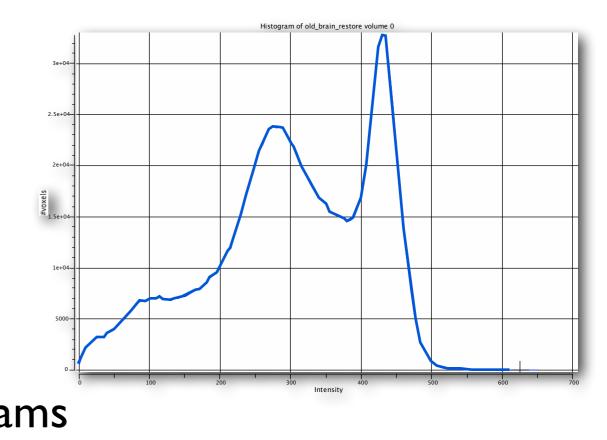






Restored

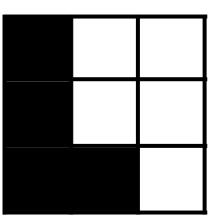




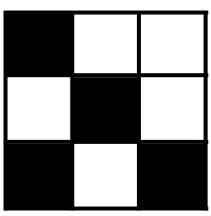


#### Use Spatial Neighbourhood Information (MRF)

- Neighbourhood information: "if my neighbours are grey matter then I probably am too"
- Simple classifiers (like K-means) do not use spatial neighbourhood information
- More robust to noise
- Need the right balance between believing neighbours or intensity



Likely configuration High probability



Unlikely configuration Low probability



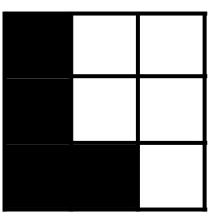
#### Use Spatial Neighbourhood Information (MRF)

Combine with probability based on Gaussian Mixture Model:

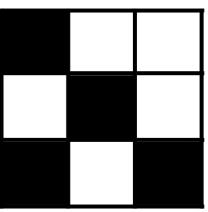
Final log probability =  $\log p(intensity) + \beta \log p(MRF)$ 

Final result depends on  $\beta$  value

This is user-adjustable



Likely configuration High probability

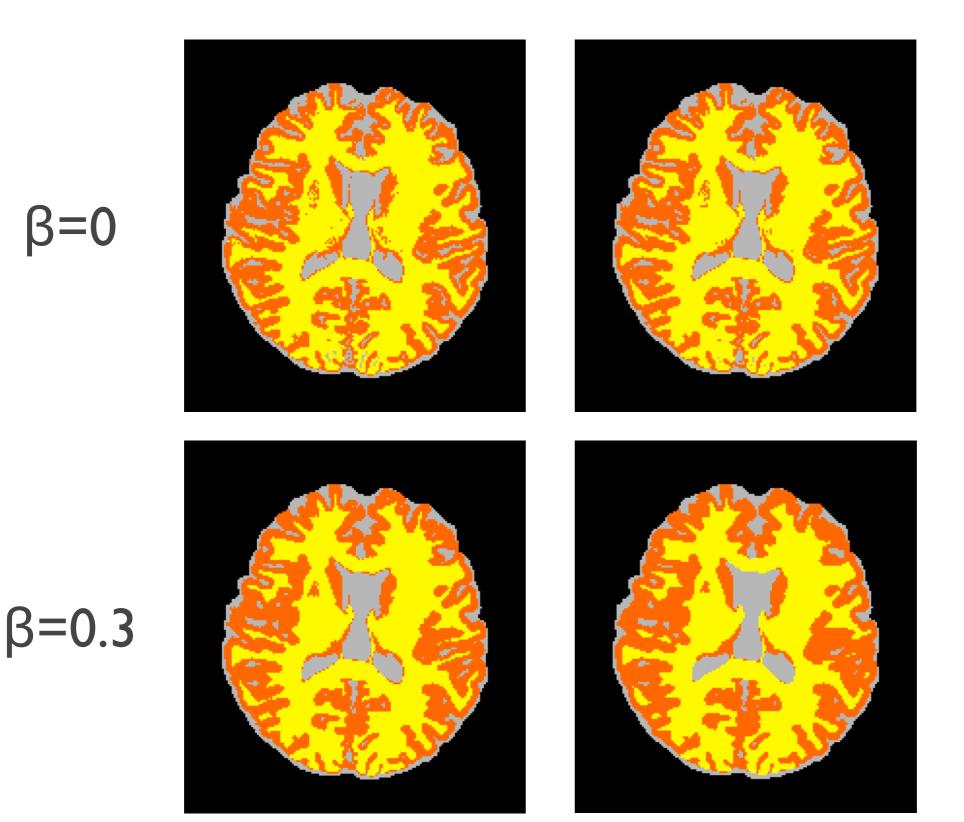


Unlikely configuration Low probability



#### Effect of MRF Weighting

β=0



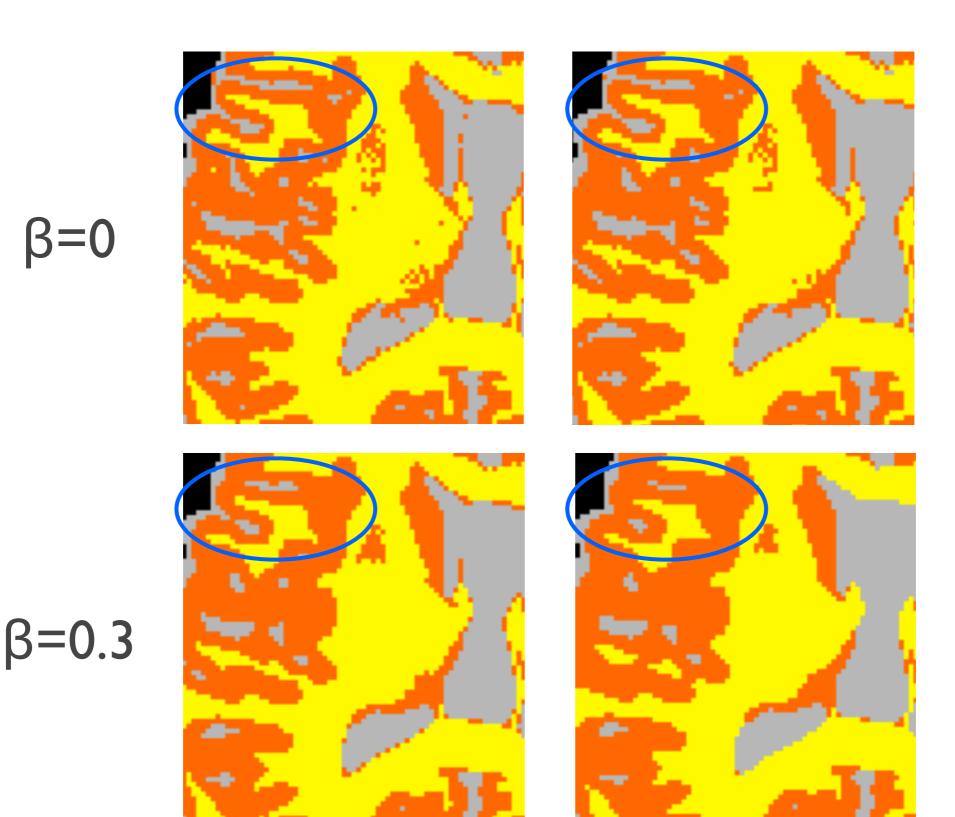
β**=**0.Ι

β=0.5



#### Effect of MRF Weighting

β=0



β=0.I

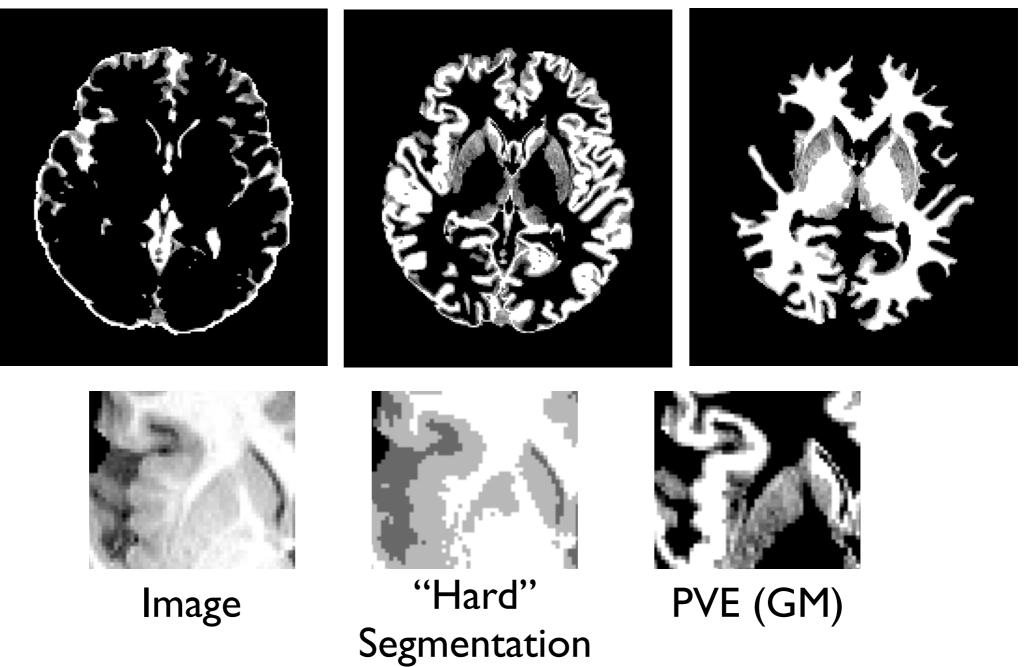
β=0.5

#### BS

### Partial Volume Modelling

- A better model is what fraction of each voxel is tissue X?
- "partial volume" = fraction of CSF, GM or WM

PVE CSF, GM, WM



• This substantially improves accuracy of volume estimation



### FAST - The Overview

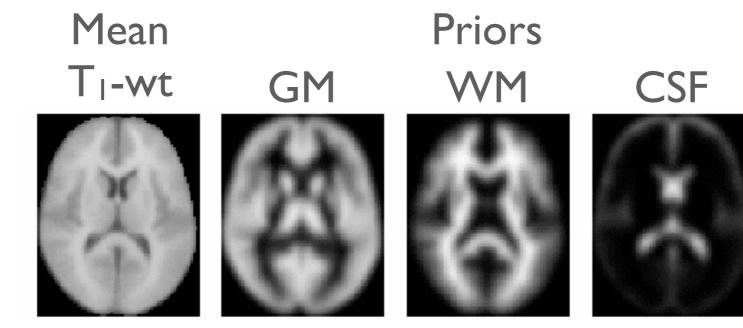
- Initial (approximate) segmentation
  - Tree-K-means
- Iterate
  - Estimate bias field
  - Estimation segmentation; iterate
    - Update segmentation (intensity + MRF)
    - Update tissue class parameters (mean and standard deviation)
- Apply partial volume model
  - MRF on mixel-type (how many tissues)
  - PV Estimation





### Optional Use of Priors (tissue probability maps)

- Segmentation priors = average of many subjects' segmentations
- Can use priors to weight segmentation, but can skew results (e.g. due to misalignment)
- FAST does not use priors by default
- If bias field is very bad, priors can be turned on to help initial segmentation (alternatively, do more iterations)
- Can also be turned on to feed into final segmentation (e.g. to aid segmentation of deep grey .... but see FIRST)





### Other Options

#### FAST:

- Bias field smoothing (-1)
  - vary spatial smoothing of the bias field
- MRF beta (-H)
  - vary spatial smoothness of the segmentation
- Iterations (-I)
  - vary number of main loop iterations

#### fsl\_anat:

- This is an alternative tool that performs brain extraction and bias field correction (along with other things) in a different way and so is worth trying out too



### FAST

#### FMRIB's Automated Segmentation Tool

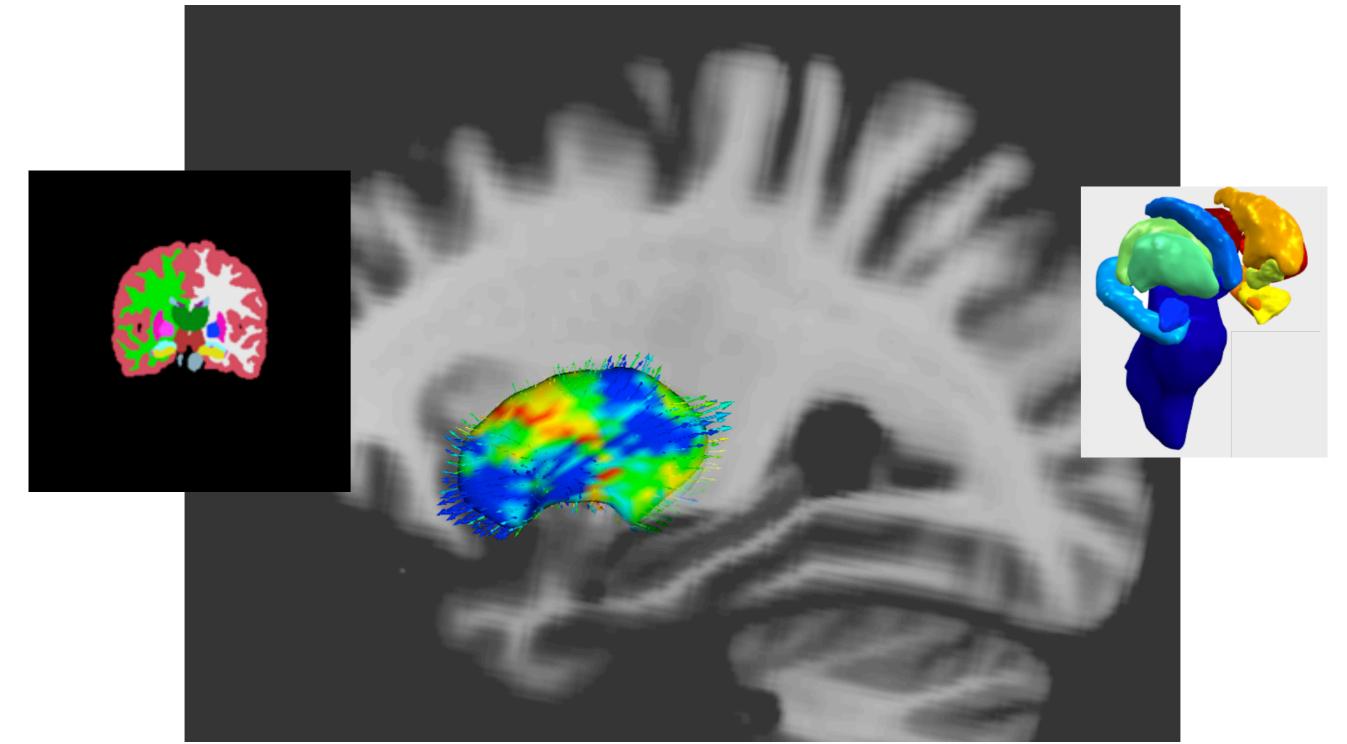
Summary

- Typically use a single T1-weighted image
- Multichannel is an option
- Segments into three main tissue-types:
  - Grey Matter, White Matter and CSF
- Models and corrects for bias field
  - Can be used just for bias field correction
- Combines intensity and neighbourhood information
- Partial Volumes Estimates (PVE) are most useful and more accurate for volume calculations
- Can use priors, but can cause bias, so not the default
- Have several adjustable parameters to optimise output



### FIRST

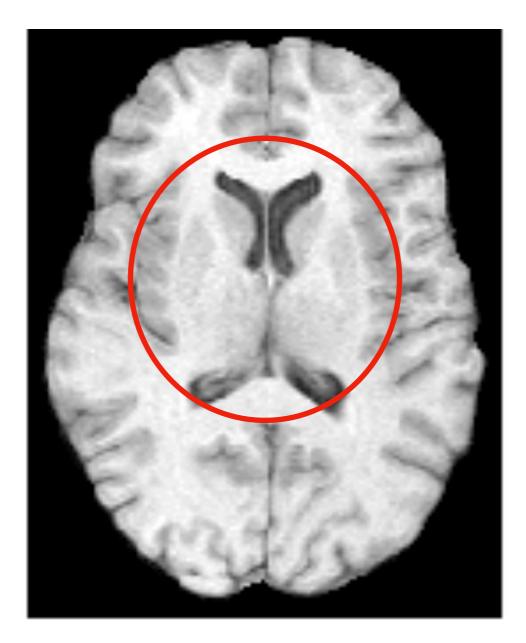
#### FMRIB's Integrated Registration & Segmentation Tool Segmentation of subcortical brain structures





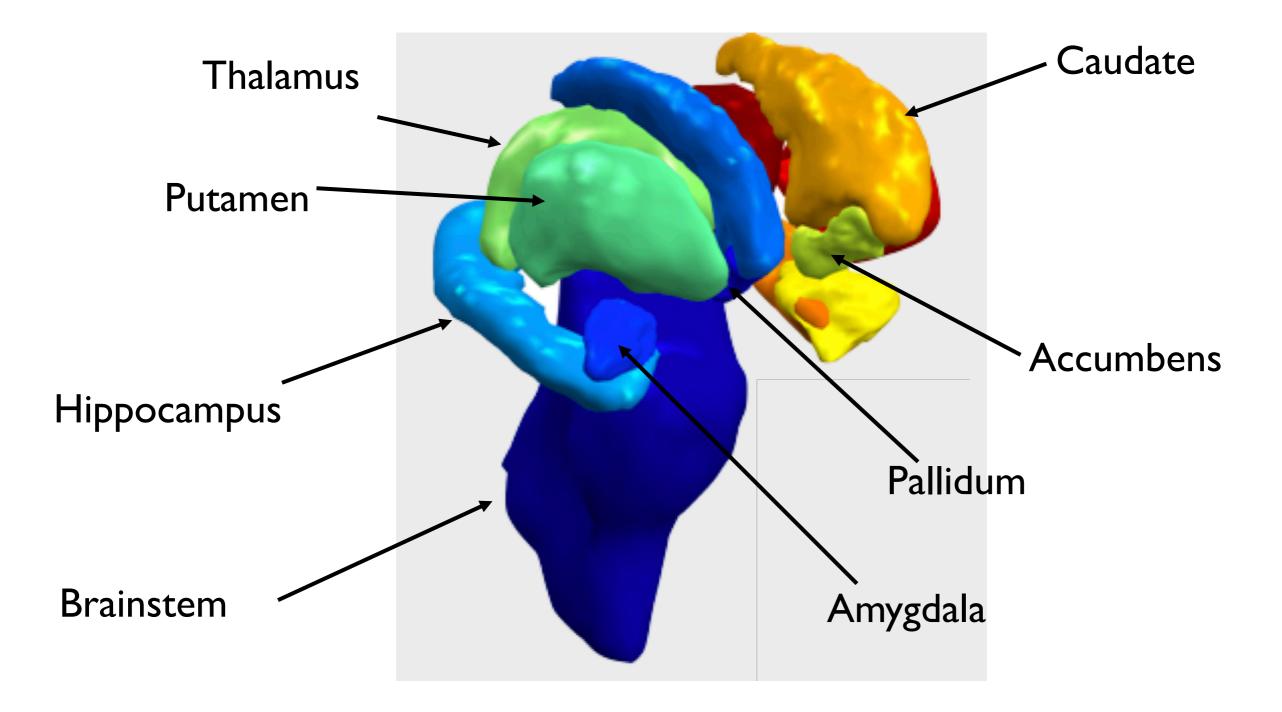
### FIRST

#### FMRIB's Integrated Registration & Segmentation Tool Segmentation of subcortical brain structures



#### Sub-Cortical Structure Models

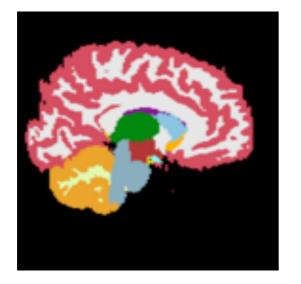
Incorporate prior anatomical information via explicit shape models Have 15 different sub-cortical structures (left/right separately)

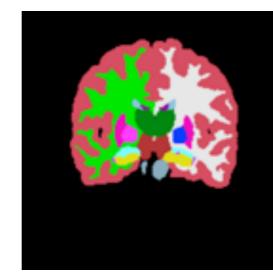




# Training Data

- Manual segmentations courtesy of David Kennedy, Center for Morphometric Analysis (CMA), Boston
- 336 complete data sets
- T<sub>1</sub>-weighted images only
- Age range 4 to 87
  - Adults: Ages 18 to 87, Normal, schizophrenia, AD
  - Children: Ages 4 to 18, Normal, ADHD, BP, prenatal cocaine exposure, schizophrenia.



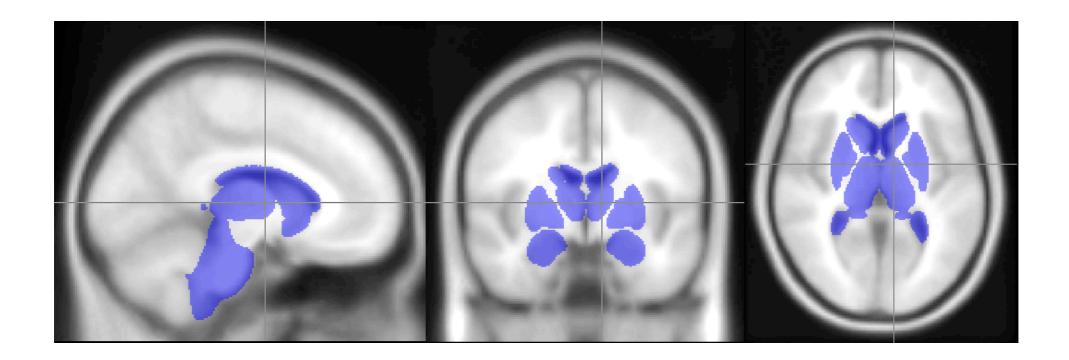






#### Model Training : Alignment to MNI152 space

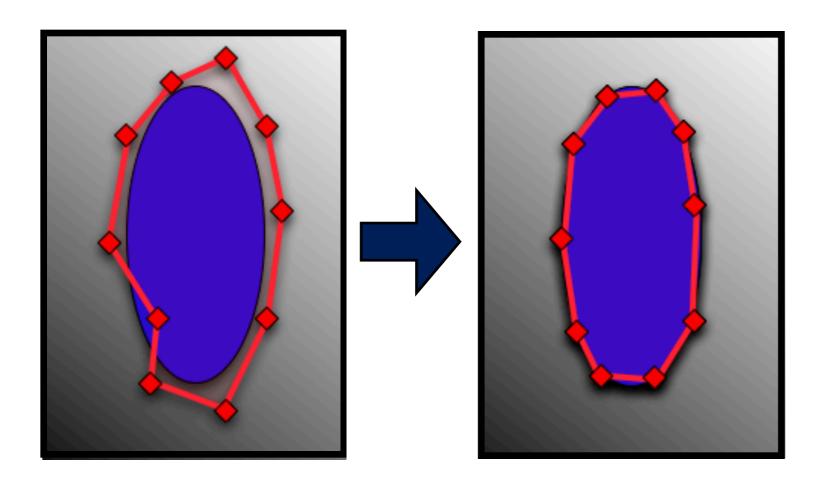
- All CMA data affine-registered to MNII52 space
  - Imm resolution, using FLIRT
- 2-stage process:
  - Whole head 12 DOF affine
  - 12 DOF affine with MNI-space sub-cortical mask

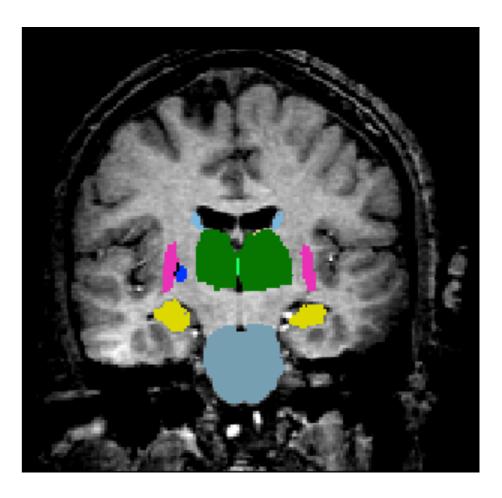




#### Deformable Models

- Model: 3D mesh
- Use anatomical info on shape & intensity (from training)
- Deformation: iterative displacement of vertices
- Maintain point (vertex) correspondence across subjects







#### The Model: Shape

- Model average shape (from vertex locations)
- Also model/learn likely variations about this mean
  - modes of variation of the population; c.f. PCA
  - also call eigenvectors
- Average shape and the modes of variation serve as prior information (known before seeing the new image that is to be segmented)
  - formally it uses a Bayesian formulation



#### The Model: Shape

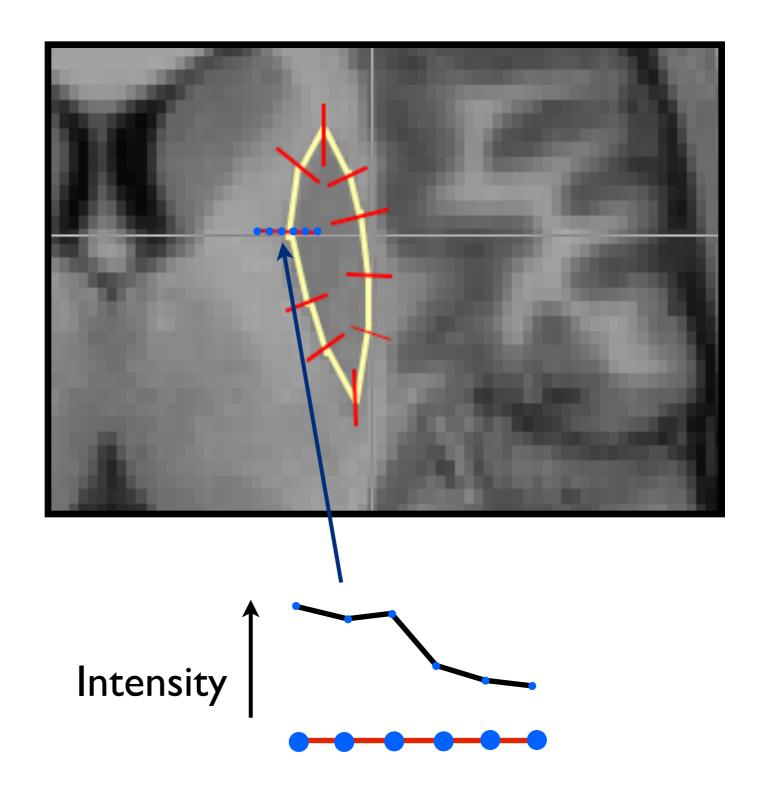
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$$X = \mu_X + UDb_X$$
  
Eigenvectors (modes) Shape parameters



### The Model: Intensity

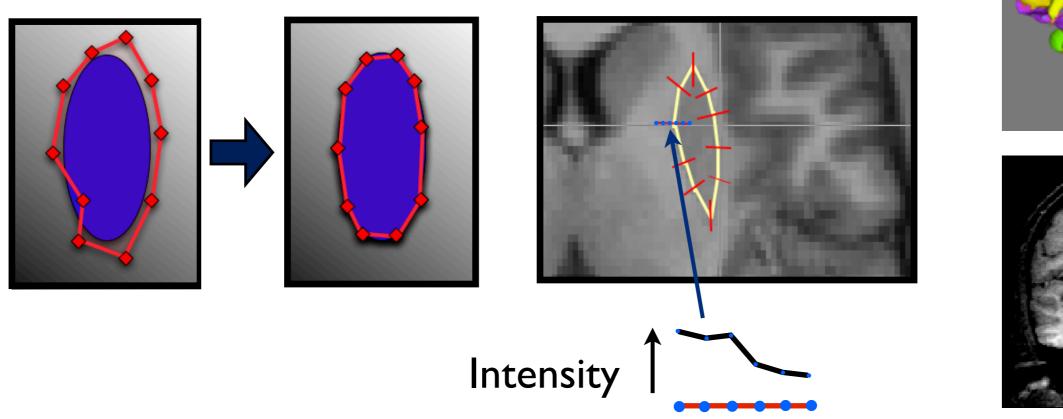
- Intensity is then sampled along the surface normal and stored
- Learn average intensity and "modes of variation"
- Aside: the intensities are re-scaled to a common range and the mode of the intensities in the structure is subtracted

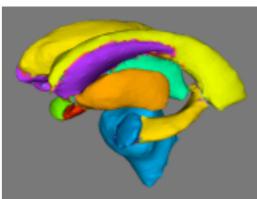


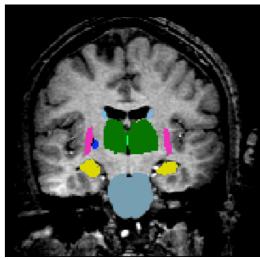


#### FIRST - Model

- <u>Model</u>: 3D mesh
- <u>Training the model</u>: learn average shape/intensity and likely variations ("modes of variation") about both
- Fitting the model: Find the "best" shape by searching along modes of variation and uses intensity match to judge fitting success



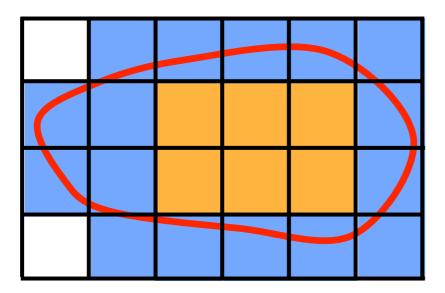


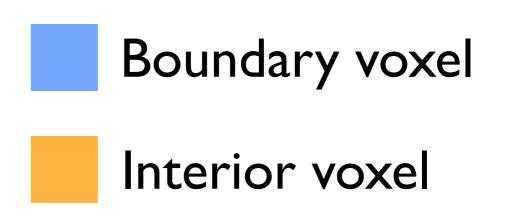




#### Boundary Correction

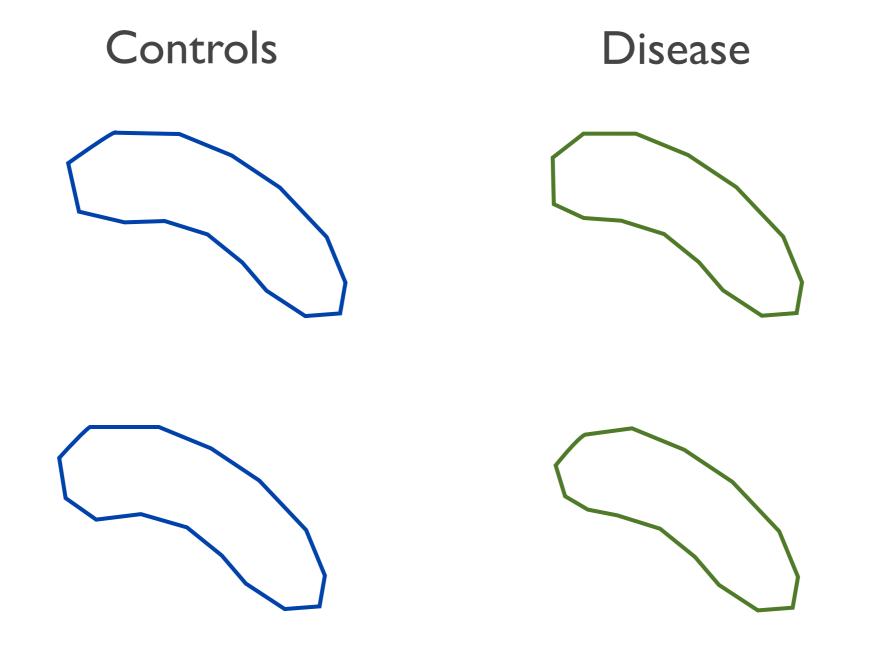
- FIRST models all structures by meshes
- Converting from meshes to images gives two types of voxels:
  - boundary voxels
  - interior voxels
- Boundary correction is necessary to decide whether the boundary voxels should belong to the structure or not
- Default correction uses FAST classification method and is run automatically (uncorrected image is also saved)
  - ensures that neighbouring structures do not overlap







• Use a univariate test at each vertex to measure difference in location (e.g. between means of two groups of subjects)

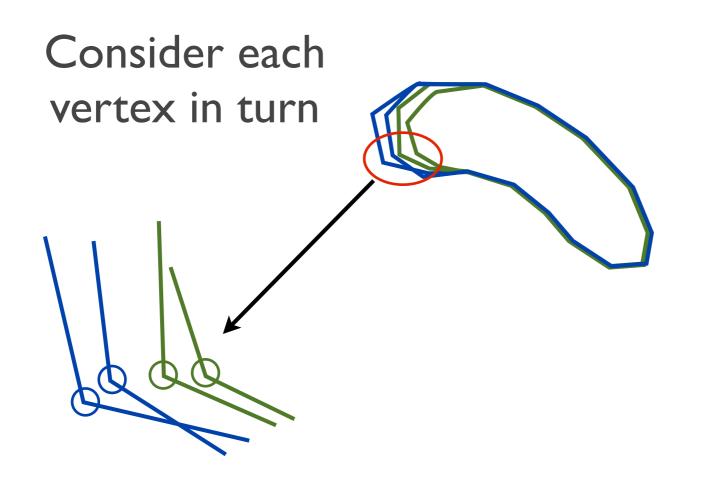




• Use a univariate test at each vertex to measure difference in location (e.g. between means of two groups of subjects)

Controls

Disease

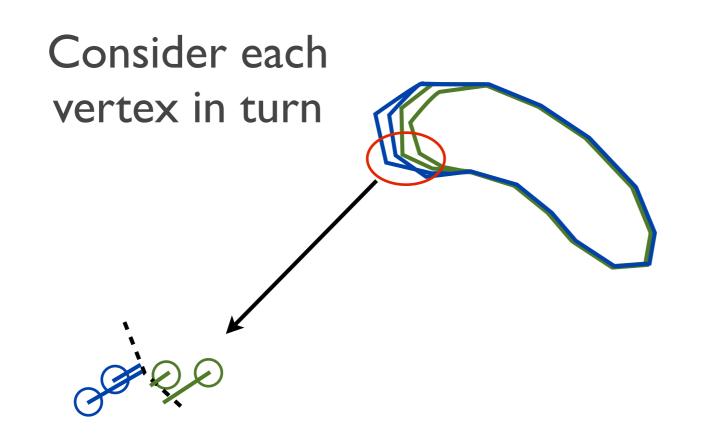




• Use a univariate test at each vertex to measure difference in location (e.g. between means of two groups of subjects)

Controls

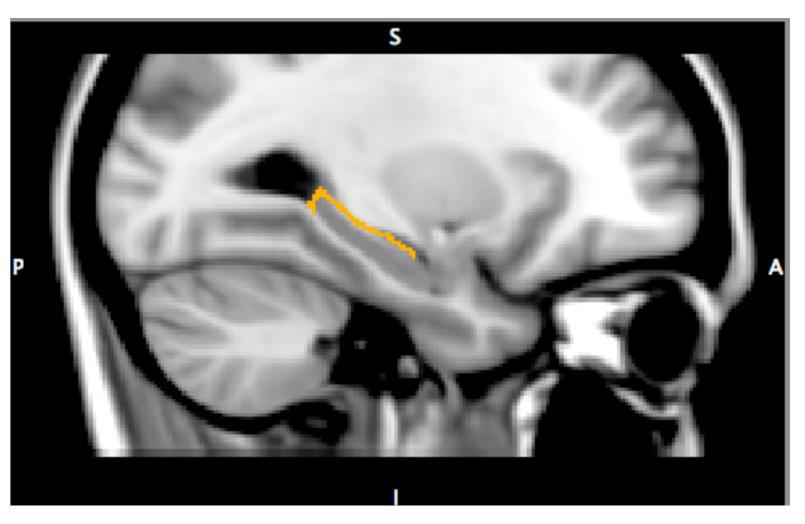
Disease



Do a test on distance of these vertices to average shape



- Use a univariate test at each vertex to measure difference in location (e.g. between means of two groups of subjects) using distance along surface normals
- Results are now given as *images* and statistics done with *randomise*
- Can do analysis in MNI space or native structural space
- MNI space analysis normalises for brain size





# Running FIRST

- Inputs:
  - T<sub>I</sub>-weighted image
  - Model (built from training data) provided with FSL
- Applying FIRST
  - A single command: run\_first\_all
    - I. registers image to MNII52 Imm template
    - 2. fits structure models (meshes) to the image
    - 3. applies boundary correction (for volumetric output)
- Analysis:
  - Use command: first\_utils
    - volumetric analysis (summary over whole structure)
    - vertex analysis (localised change in shape and/or size)
    - randomise (with multiple comparison correction)



## FIRST

FMRIB's Integrated Registration & Segmentation Tool

#### Summary

- Specific to certain deep grey structures
- Uses broad training set very general demographics
- Can only work with TI-weighted images
- Models average and variations of shape and intensity
- Represents the boundary as a set of points
- Separate boundary correction step to get voxel labels
- Can perform vertex analysis to look at changes in shape and size