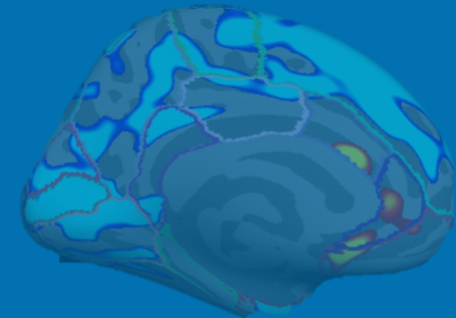
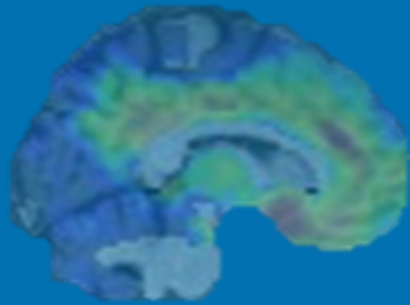




INTRODUCTION BRAIN MORPHOMETRY

Meritxell Bach Cuadra^{1,2}

¹*CIBM Center for Biomedical Imaging*

²*Radiology Department, Lausanne University Hospital and University of Lausanne*

Invited lecture, CIBM translational MR neuroimaging & spectroscopy (PHYS-760), May 2nd 2023, EPFL



C I B M . C H

OBJECTIVES

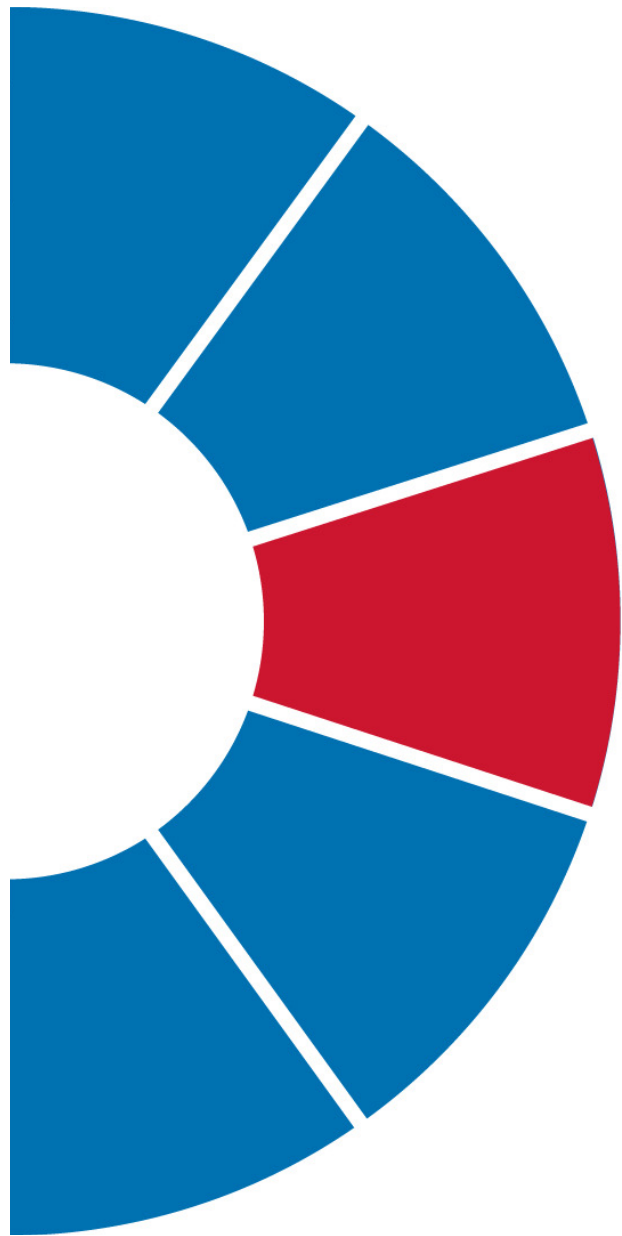


- Overview of morphometry methods for study of in-vivo brain MRI.
- Understand the **basics of voxel-wise morphometry**:
 - Pre-processing steps
- Be aware of **surface-based morphometry**:
 - and enumerate advantages and disadvantages in comparison with VBM

OUTLINE



- Definition
- Voxel-based morphometry
 - Pre-processing
- Surface-based morphometry
- Advantages and disadvantages



DEFINITION AND METHODS

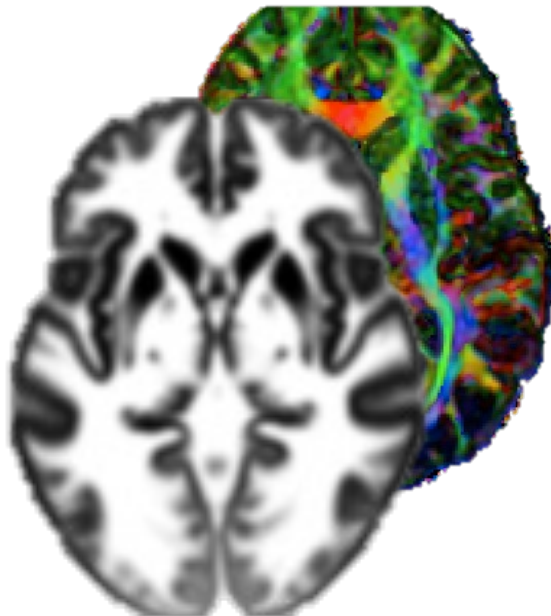
DEFINITION - IN COMPUTER VISION

Brain morphometry* measurement of brain structures and changes thereof during development, aging, learning, disease and evolution....

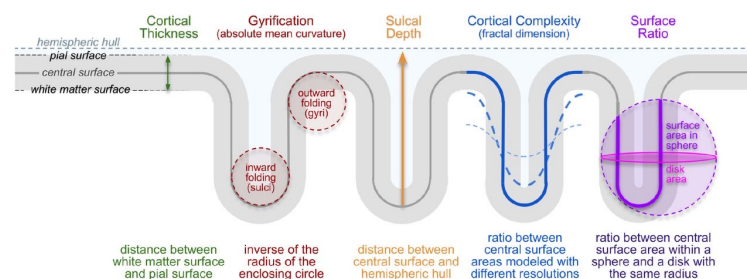
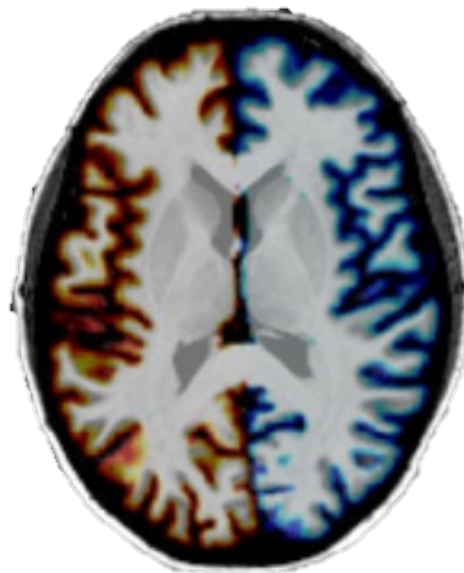


*Wikipedia

Voxel-based



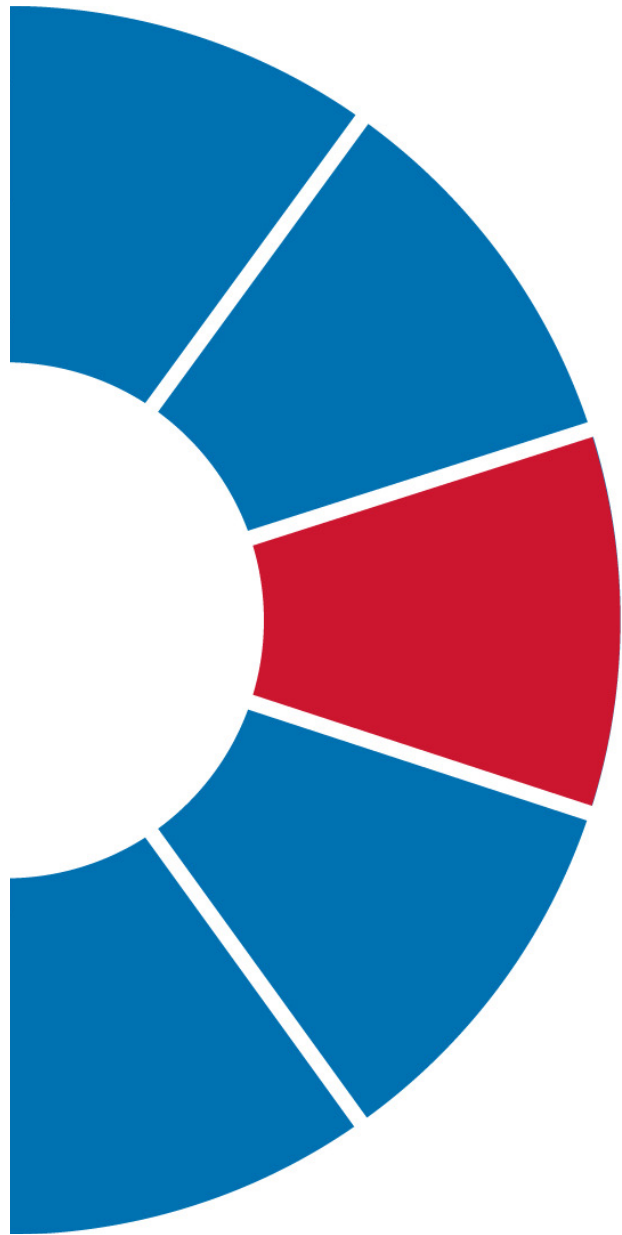
Surface-based



Deformation-based



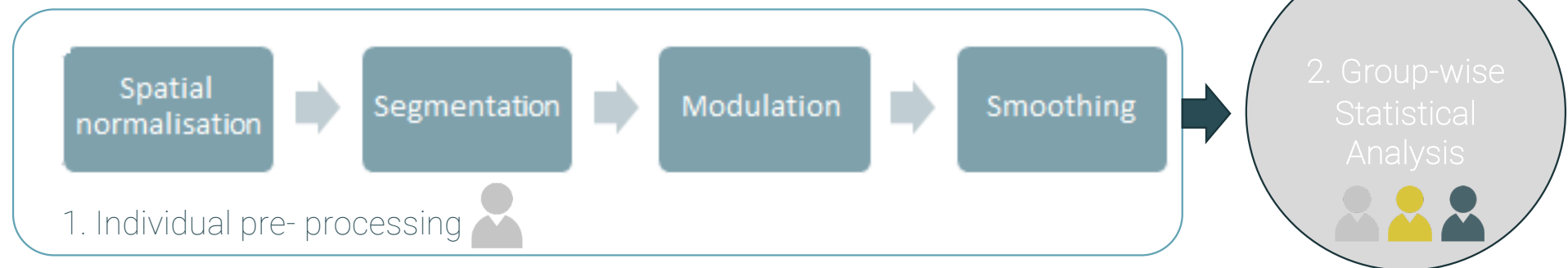
$$\mathbf{J} = \begin{bmatrix} \frac{\partial f_1}{\partial x_1} & \dots & \frac{\partial f_1}{\partial x_n} \\ \vdots & \ddots & \vdots \\ \frac{\partial f_m}{\partial x_1} & \dots & \frac{\partial f_m}{\partial x_n} \end{bmatrix}$$



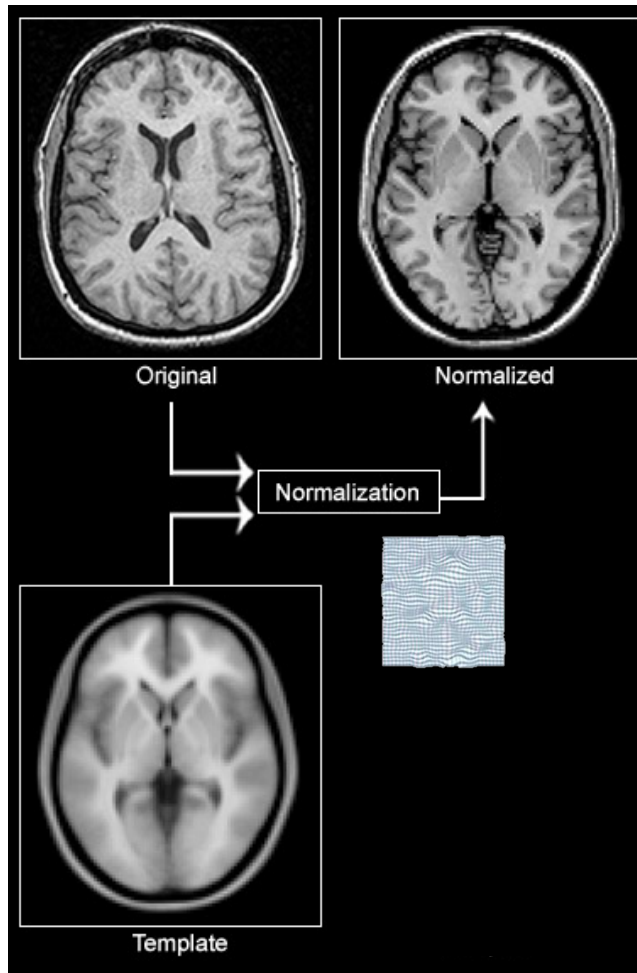
VOXEL-BASED MORPHOMETRY

VBM

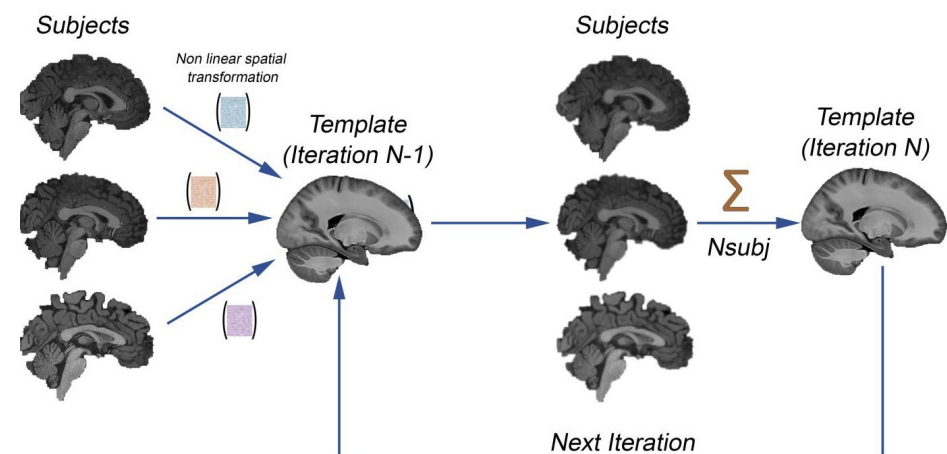
- Tissue maps : regional differences in volume
- Whole brain: no a priori on where these differences occur
- Produce a map of statistically significant differences among populations of subjects
 - compare a patient group with a control group
 - identify correlations with age, test-score etc.



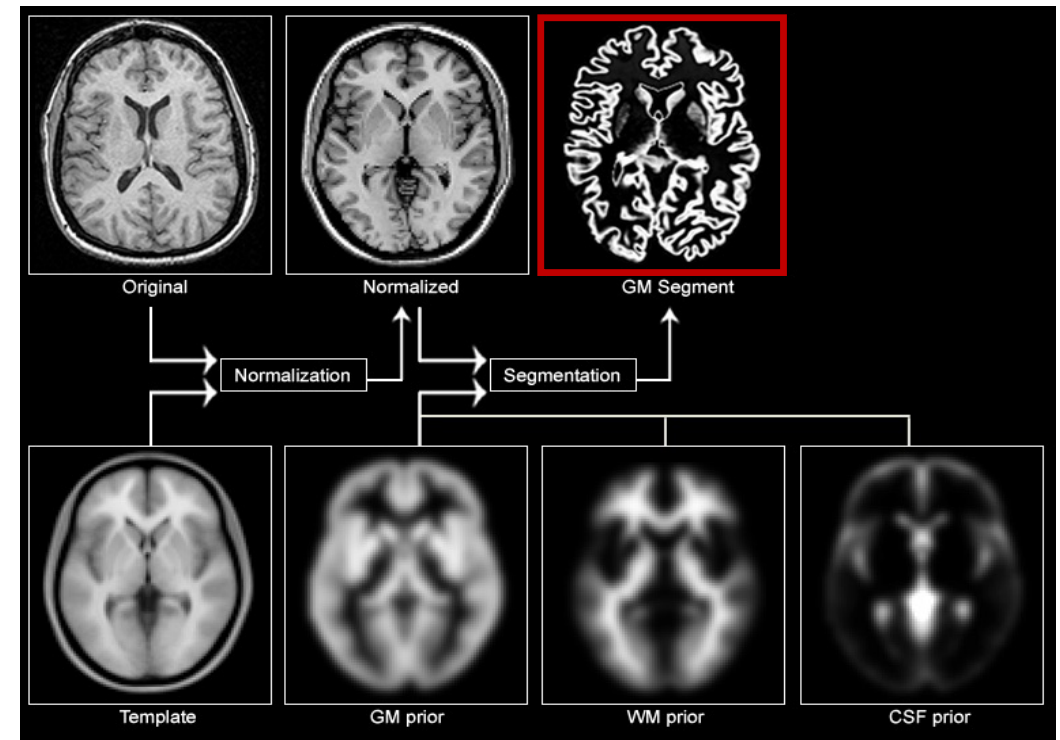
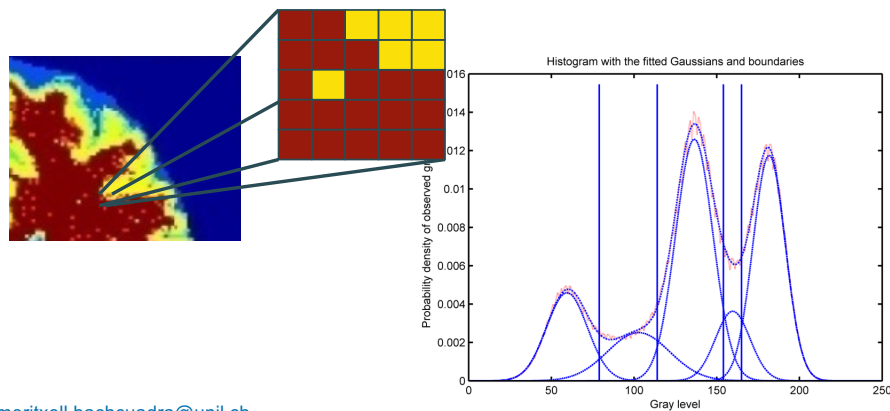
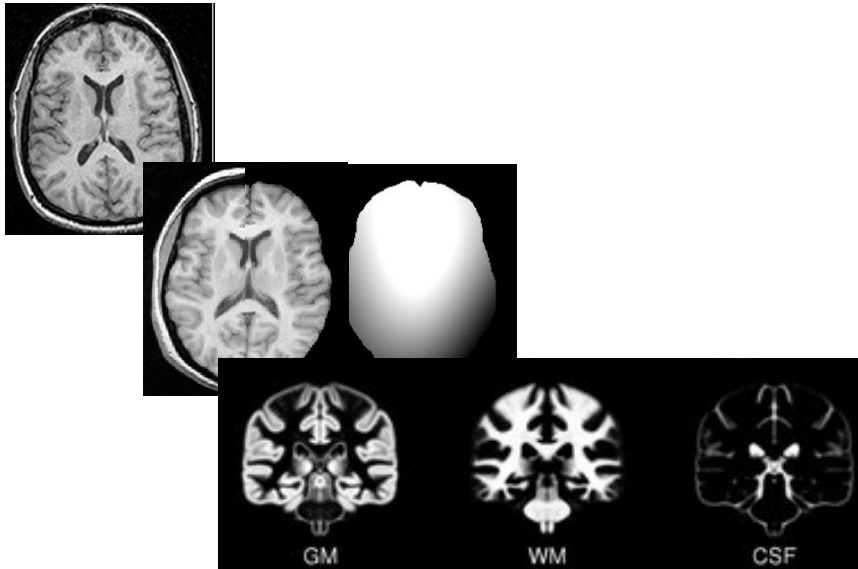
SPATIAL NORMALIZATION



- Transforming all the subject's imaging to the same reference space
- Corrects for global brain shape differences
- Choice of the template: MNI  or own template (Avants et al 2010)



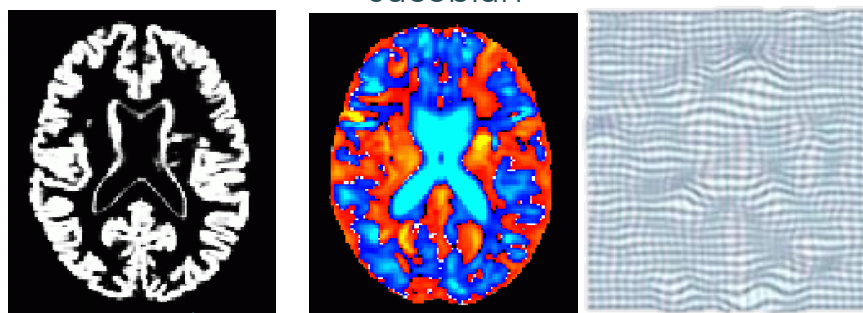
SEGMENTATION



MODULATION

Multiplying the spatially normalised scalars (GM, WM, etc) by Jacobian determinant

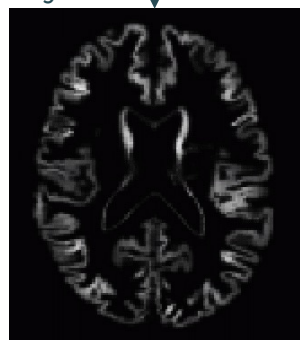
Jacobian



GM Segmentation
(Concentration)

Multiply

GM Density



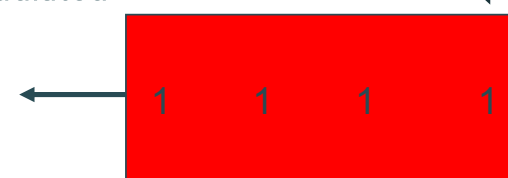
Native

intensity =
tissue probability



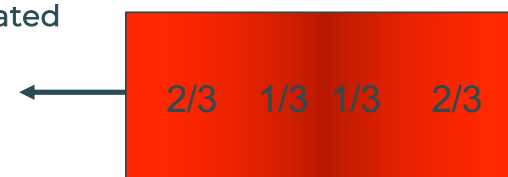
Spatial
normalization

Unmodulated



*Difficult to
interpret*

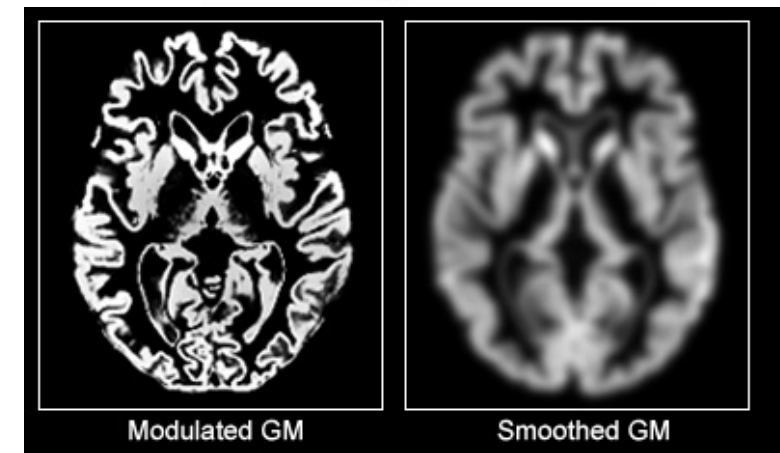
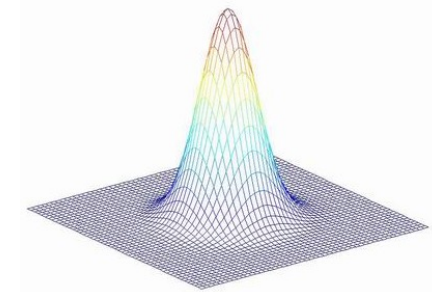
Modulated



Volume
relative to
template

SMOOTHING

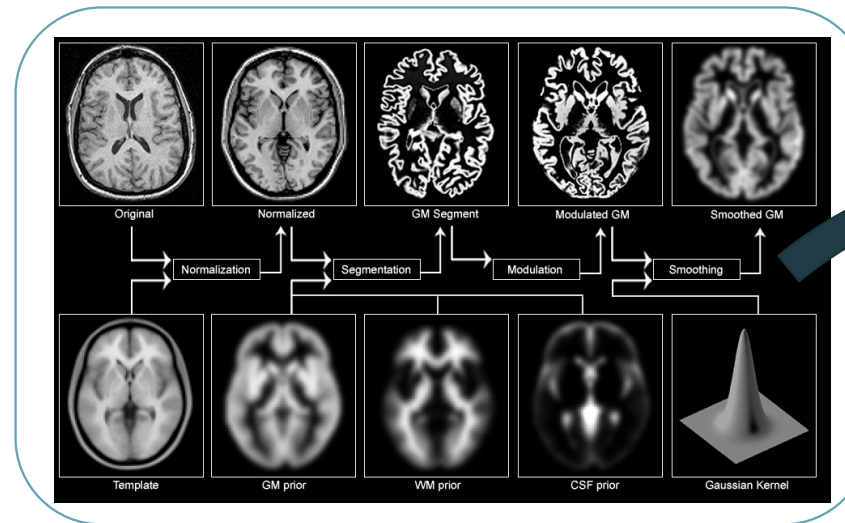
- Analysis sensitive to effects that match shape and size of the kernel
- The data will be more Gaussian and closer to a continuous random field for larger kernels
- Smoothing compensate for registration errors
- Results will be rough and noise-like if too little smoothing is used
- Usually recommend $\geq 6\text{mm}$ but $< 12\text{mm}$



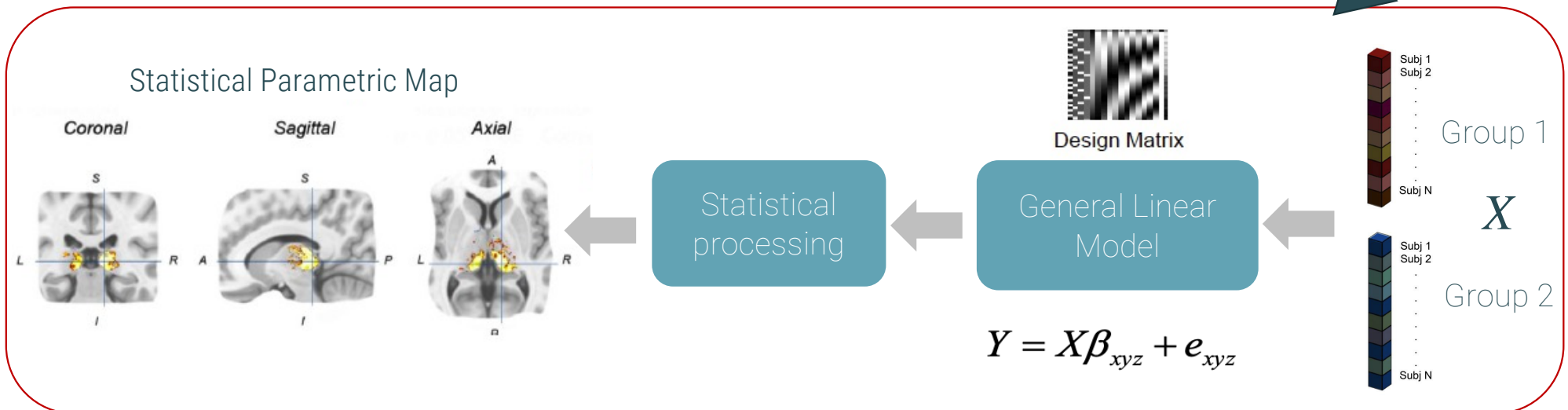
VBM

Whole pipeline

1. Pre-processing



2. Group-wise processing



VBM: SUMMARY

Advantages

- Unbiased way to localize differences
- Whole brain analysis, no knowledge on where differences are is required
- Can be adapted to be sensitive to both small and large differences
- Automated analysis: low user interaction

Disadvantages

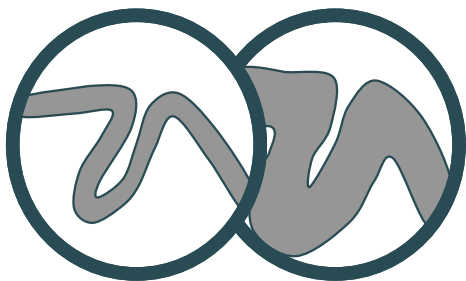
- Sensitive to:
 - Quality of image (noise and artefacts)
 - Registration errors
 - Segmentation errors
 - Priors might not be appropriate !!!
 - No model of pathology (stroke, tumor, etc)

INTERPRETATION OF FINDINGS

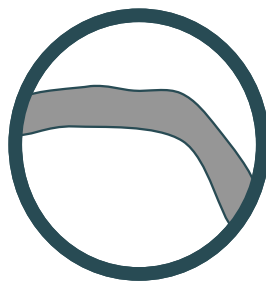
Reference



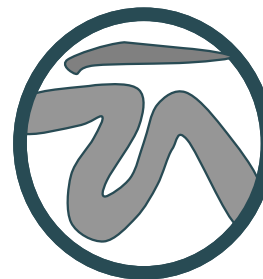
Thinner/Thicker



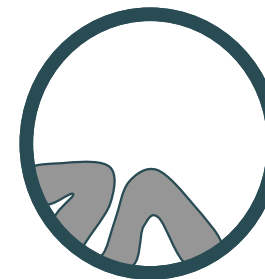
Folding



Mis-classify



Mis-register

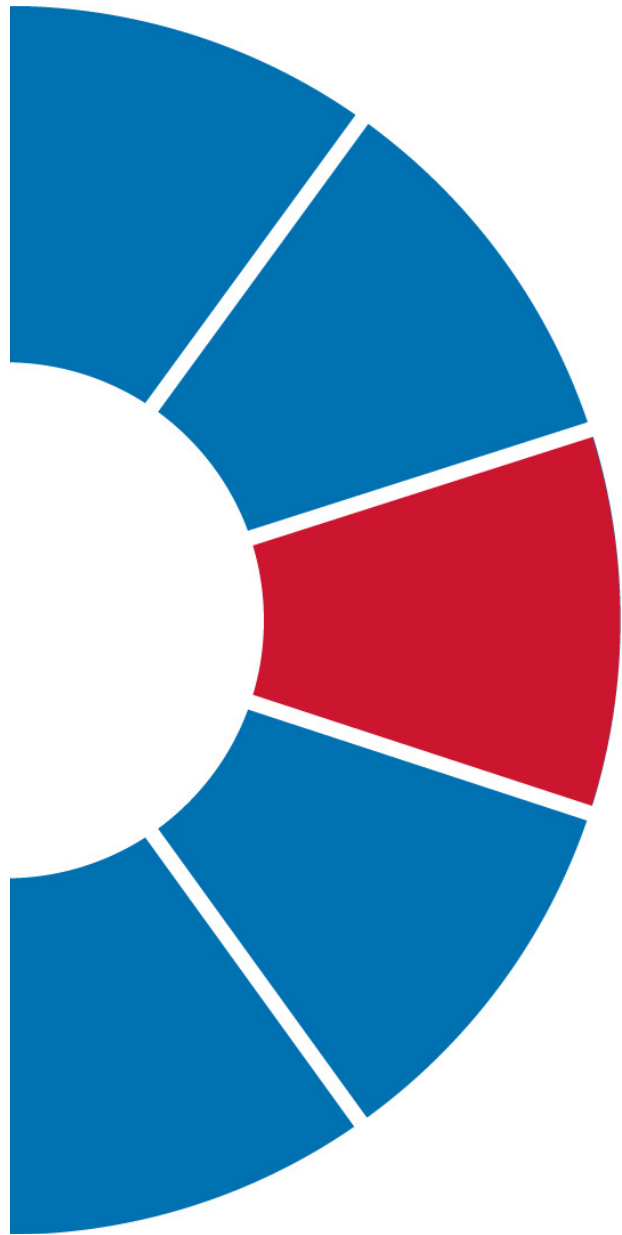


Intensity



- Statistical Parametric Mapping of regional segmented tissue **density** or **volume**

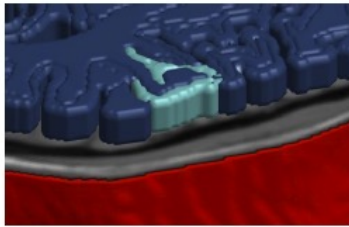
- The exact interpretation of density or volume is complicated: not interpretable as neuronal packing or other cytoarchitectonic tissue properties
- Hypothesis: changes in these microscopic properties may lead to macro-or mesoscopic VBM-detectable differences



SURFACE-BASED MORPHOMETRY

VOLUME VS SURFACE

Cortical region



Box



Pyramid

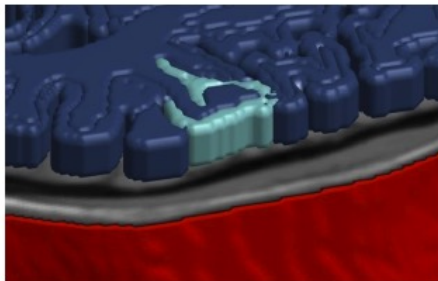


Teapot

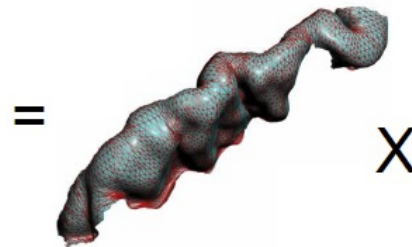


Same volume different shape

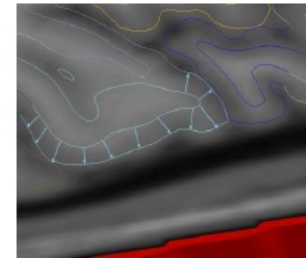
Volume



Surface Area

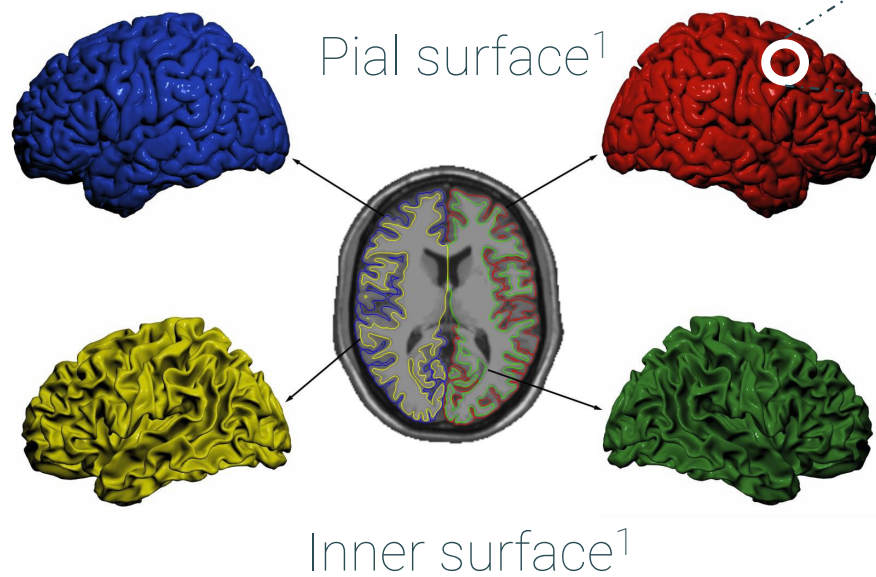


Cortical thickness

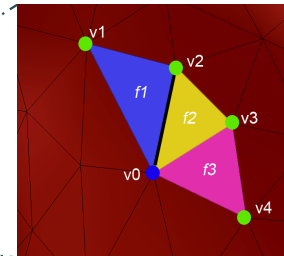


Courtesy of Dr Y. Alemán-Gómez

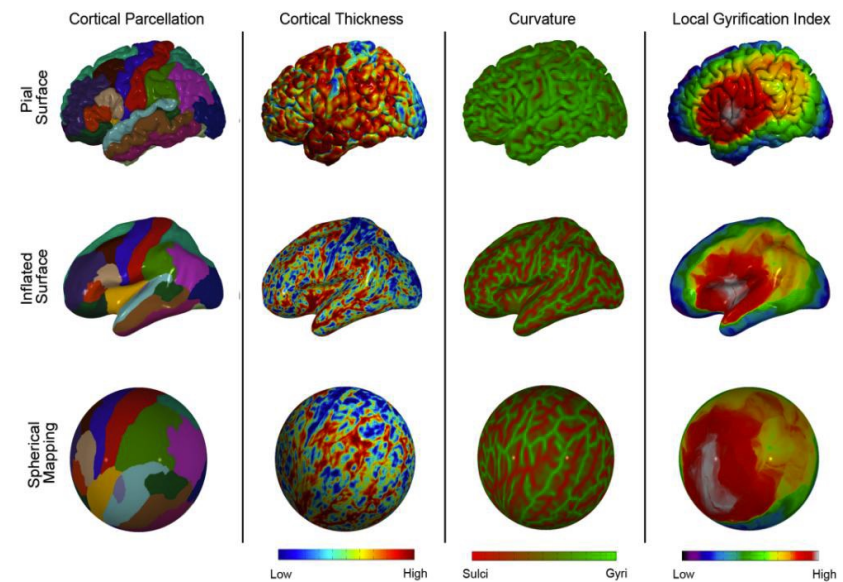
SURFACE-BASED EXTRACTION



Mesh, vertices

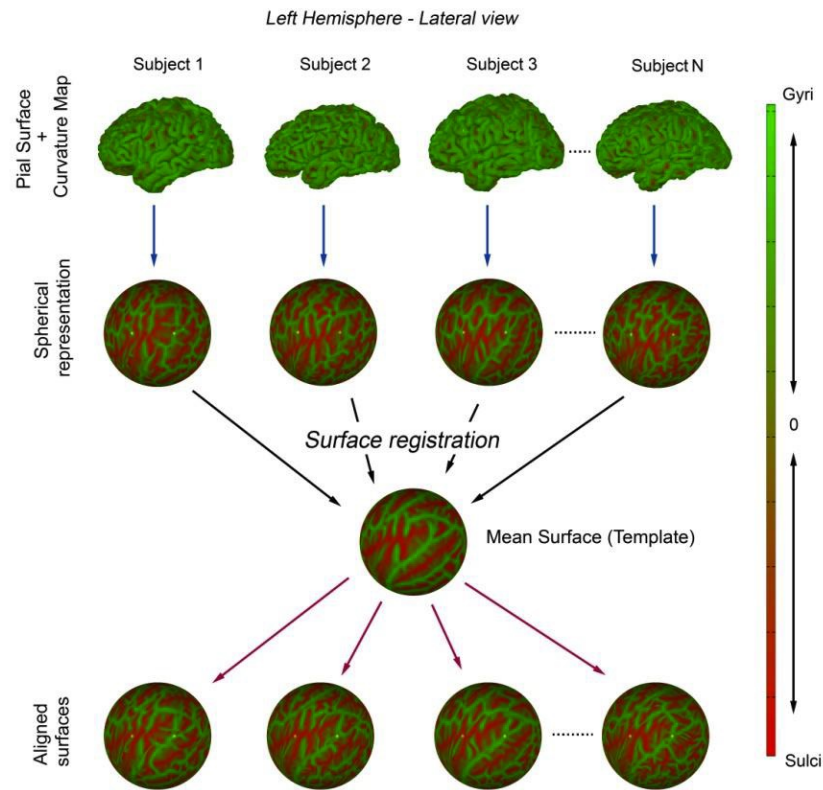


Vertex-wise metrics^{2,3,4}

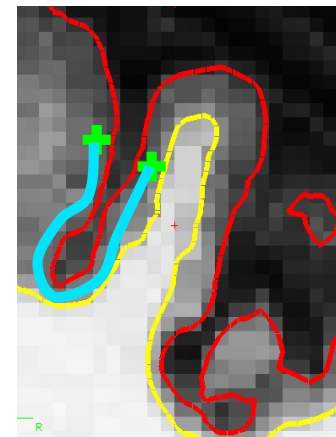


¹ Davatzikos et al. 1996. IEEE Transactions on Medical Imaging, ²Fischl et al. 2000. PNAS, 3 IEEE Trans. Med. Imag. Dale et al. 1999. Neuroimage, 4 Schaer et al. 2008.

SURFACE REGISTRATION & SMOOTHING



Fischl, et al, 1998, NI.



- 5 mm apart in 3D
- 25 mm apart on surface!

SBM: SUMMARY

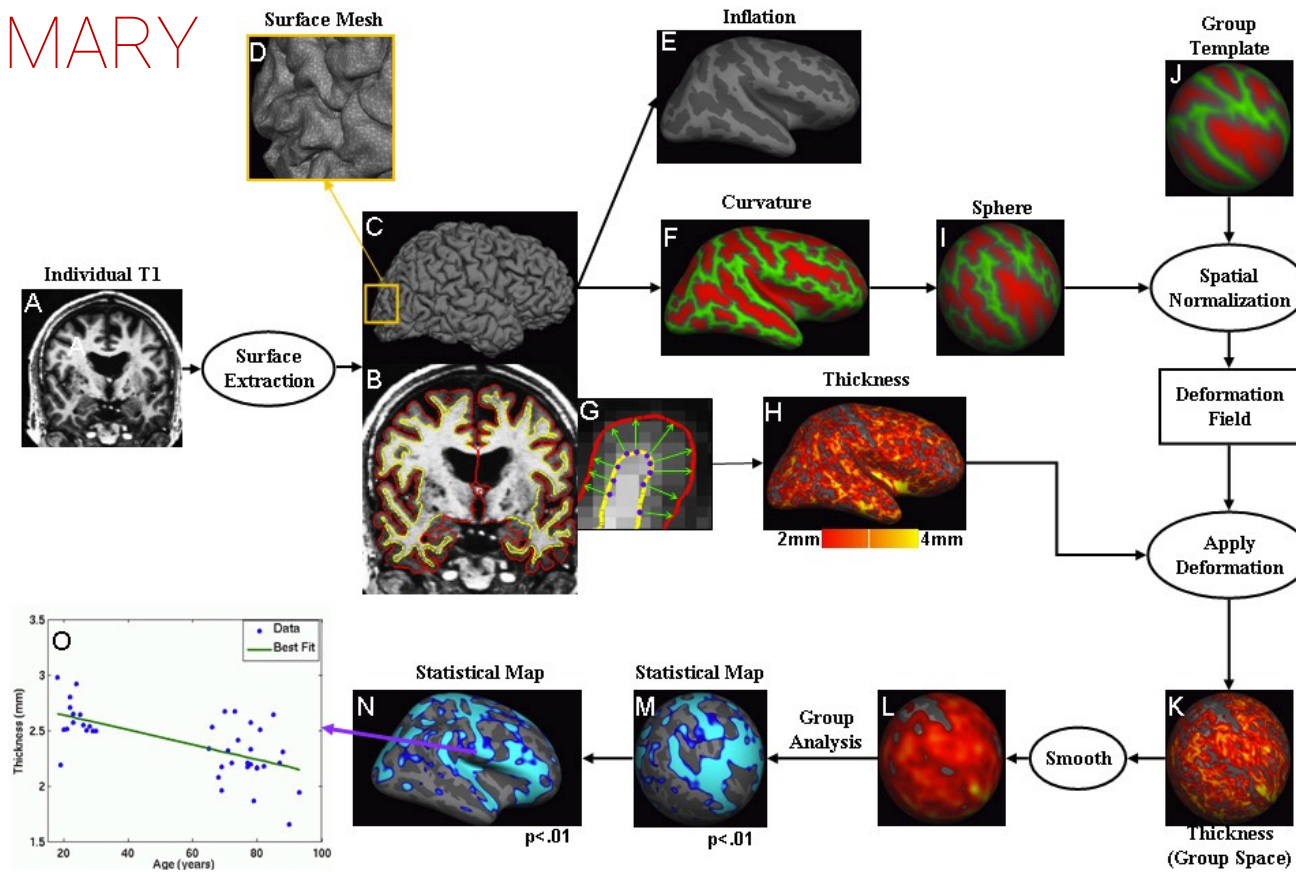
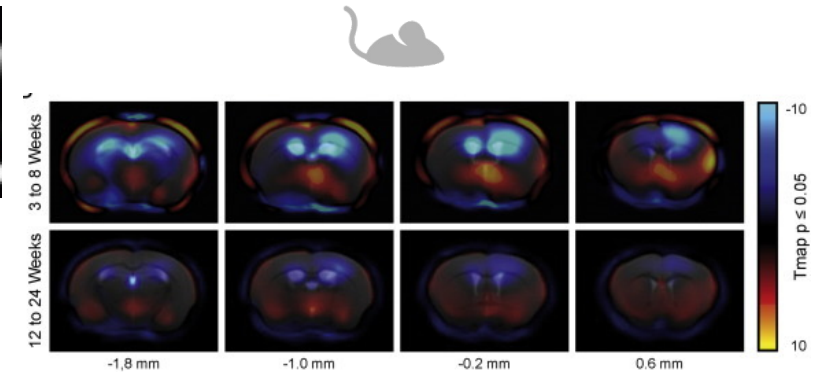
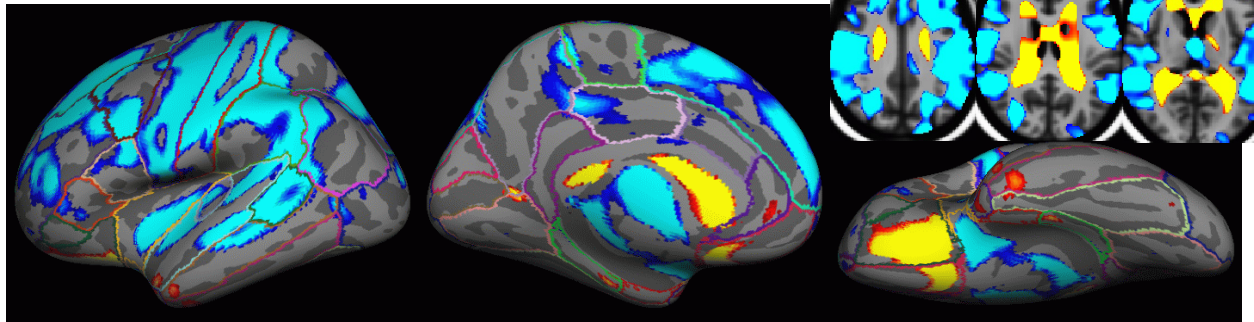


Figure from Greve et al Martinos, MGH

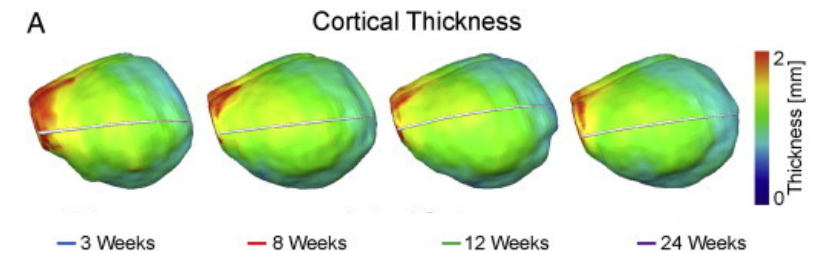
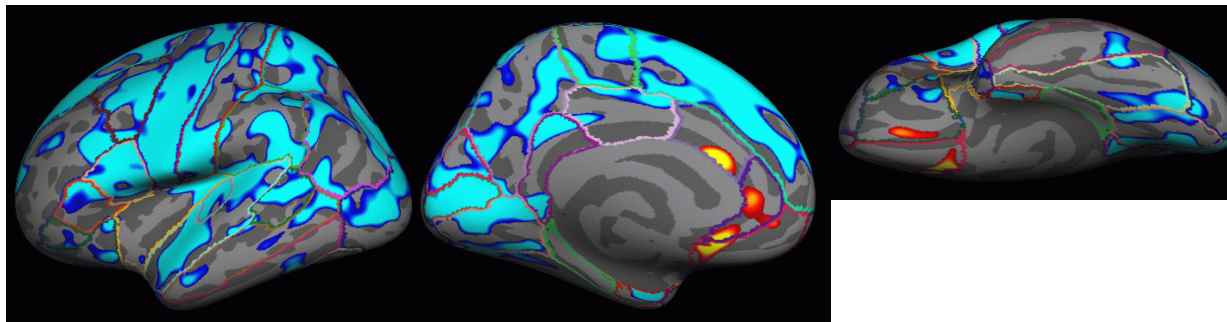
VBM VS SBM: AGE RELATED CHANGES



VBM (SPM8/VBM8)



Thickness (FreeSurfer 5.0)

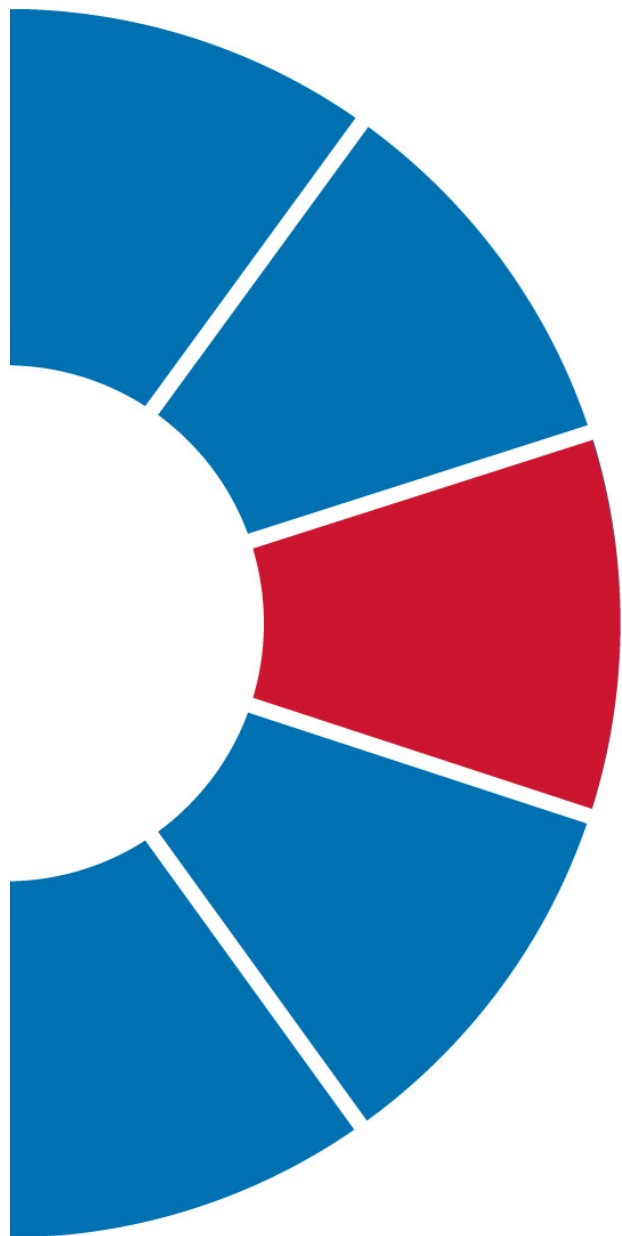


$p < .01$

Slide adapted from Martinos MGH course on VBM

Hammelrath et al, Neuroimage, 2016.

CIBM.CH



CONCLUSIONS

COMPARISON¹

VBM

- Volume = Jacobian X Segmentation
- Strengths
 - GM, WM, CSF, cortical and subcortical
 - Easy to use
- Weaknesses
 - Sensitive to registration and segmentation errors
 - Segmentation is atlas-dependent

SBM

- Thickness, gyration, etc
- Strengths
 - Surface-based Normalization
 - Surface-based Smoothing
 - Surface extraction atlas-independent
- Weaknesses
 - No subcortical, White Matter, or CSF
 - More complicated to analyze

TAKE HOME MESSAGES

Longitudinal changes

- Early development
- Aging
- Disease progression

Diagnosis / Phenotyping

- Cross-sectional group studies
- Single-subject VBM

Tools (VBM, SBM)

- SPM12
- FSL
- ANTS
- Freesurfer
- CAT
- CIVET
- SIENA

Goto et al, MR Med Sciences 2022, Zhou et al Nature Comm. Biology 2022, Antonopoulos ARXIV, 2023

FURTHER INFO

■ References & materials

- Thanks to Y. Aleman Gomez
- Support slides from Martinos MGH
- Ashburner & Friston. VBM – the Methods. Neuroimage. 2000
- Good et al. Neuroimage. 2004
- <http://www.fil.ion.ucl.ac.uk/spm/>
- <http://en.wikibooks.org/wiki/SPM>
- <http://www.mrc-cbu.cam.ac.uk/Imaging/Common/spm.shtml>
- <http://en.wikibooks.org/wiki/SPM-VBM>



THANK YOU FOR YOUR ATTENTION



C I B M . C H