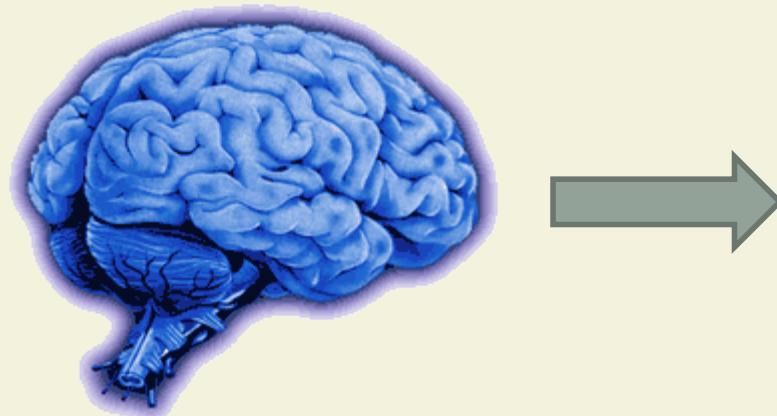
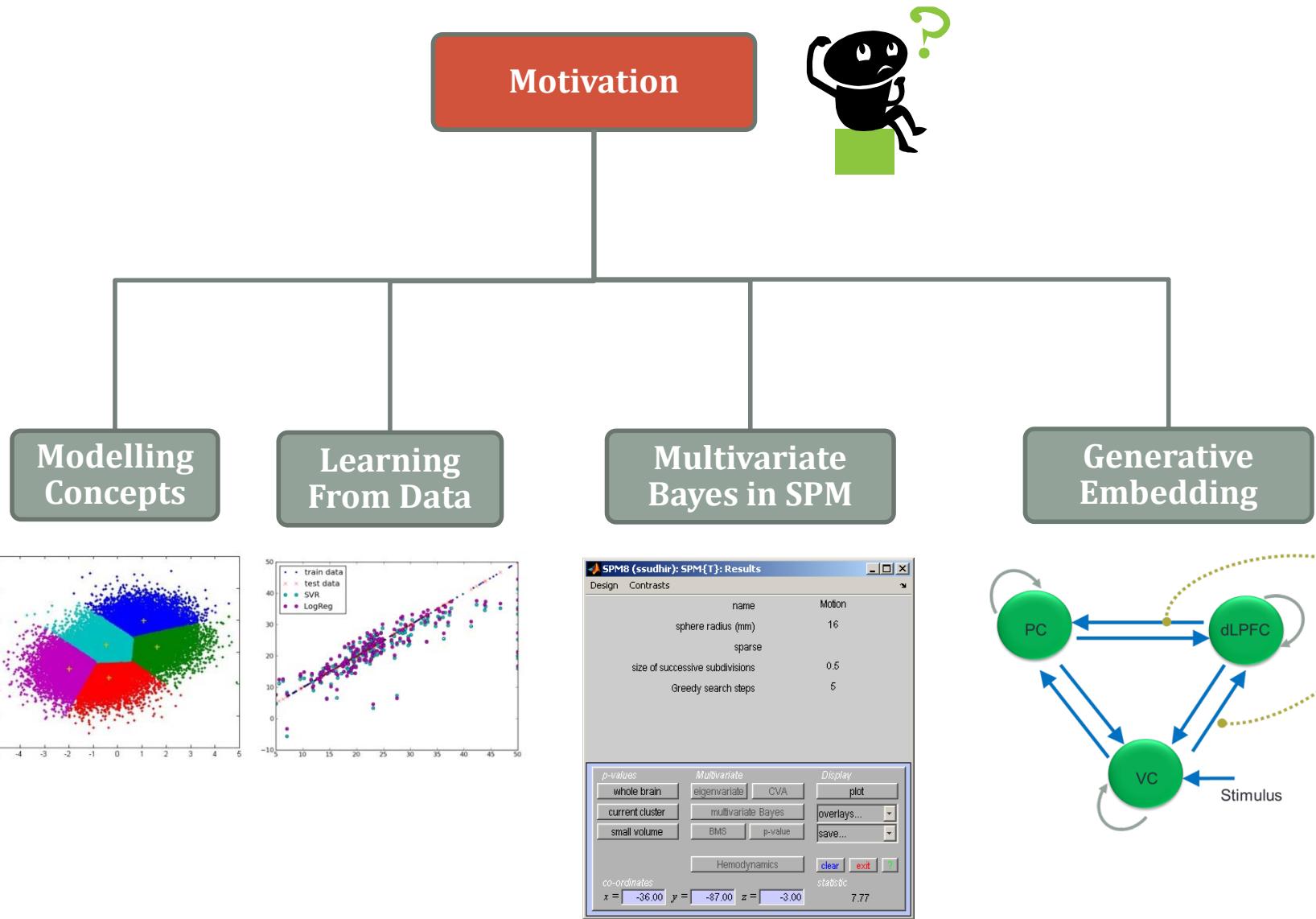


MULTIVARIATE ANALYSES WITH fMRI DATA

Sudhir Shankar Raman

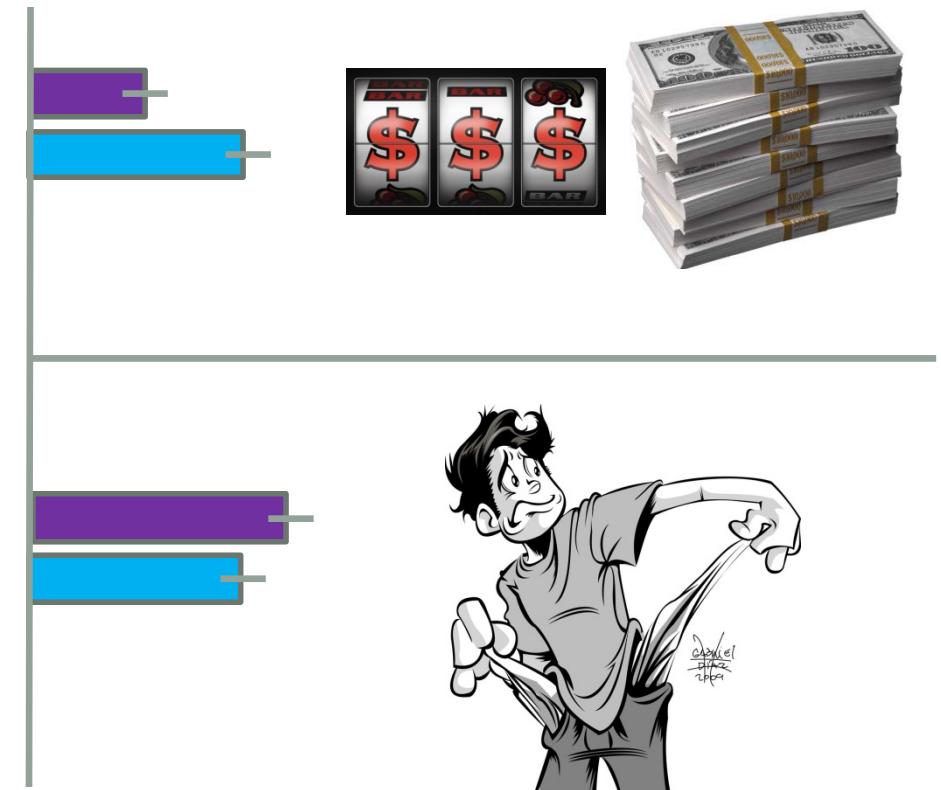
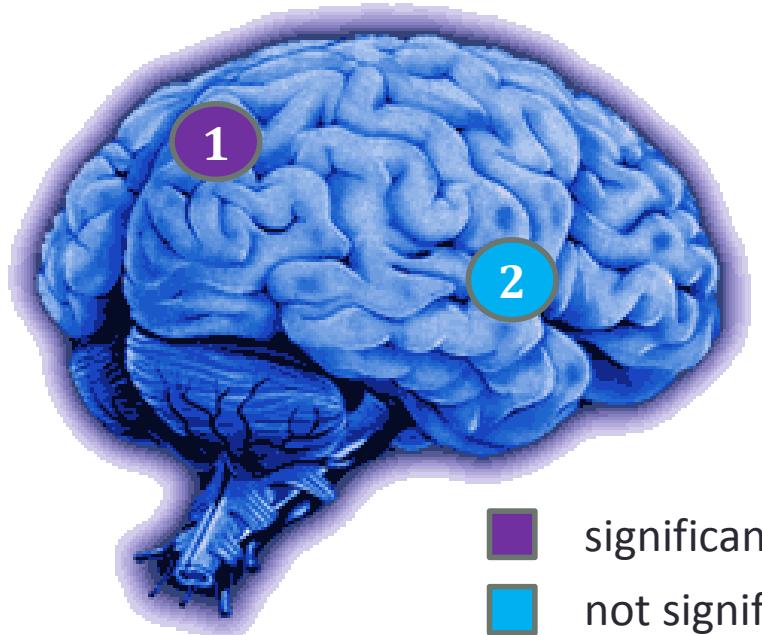
Translational Neuromodeling Unit (TNU)
Institute for Biomedical Engineering
University of Zurich & ETH Zurich



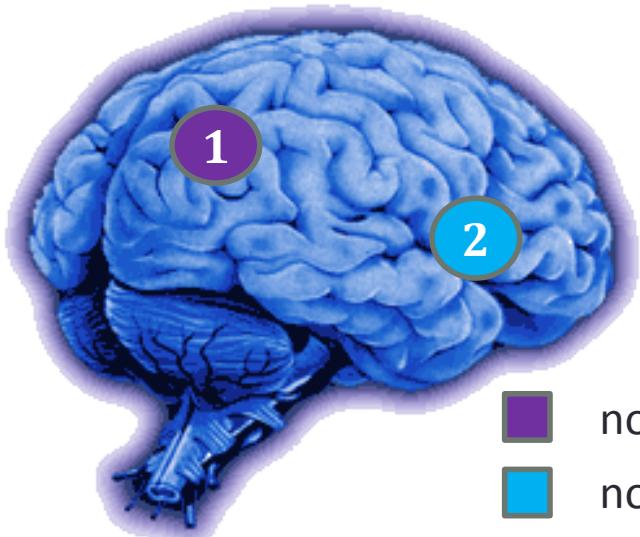


Motivation

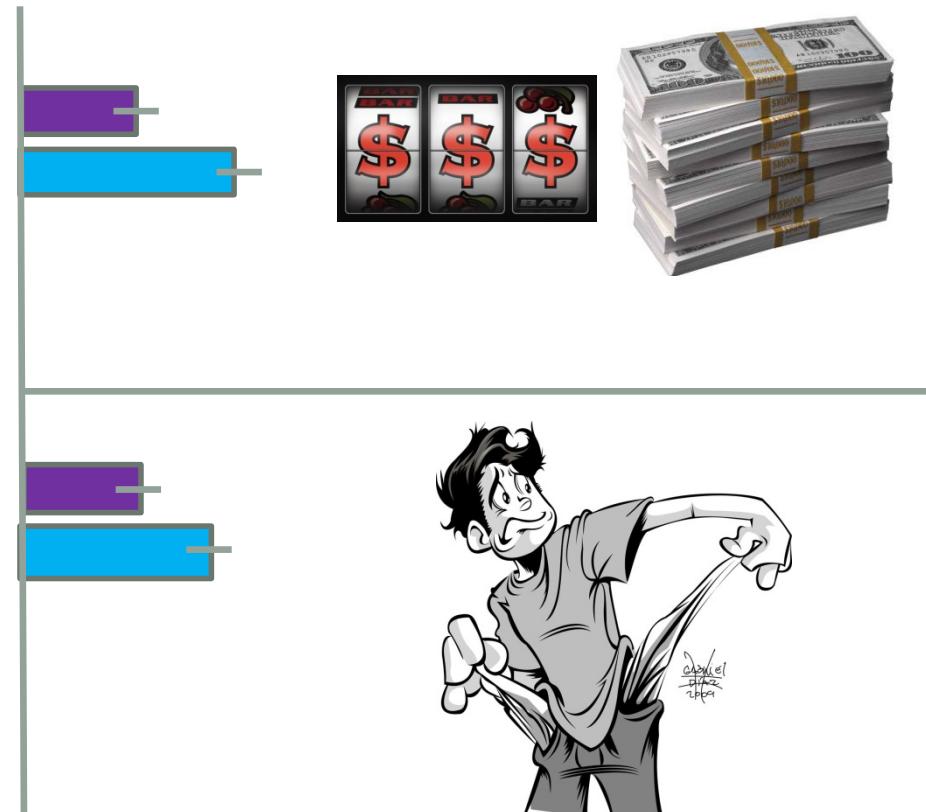
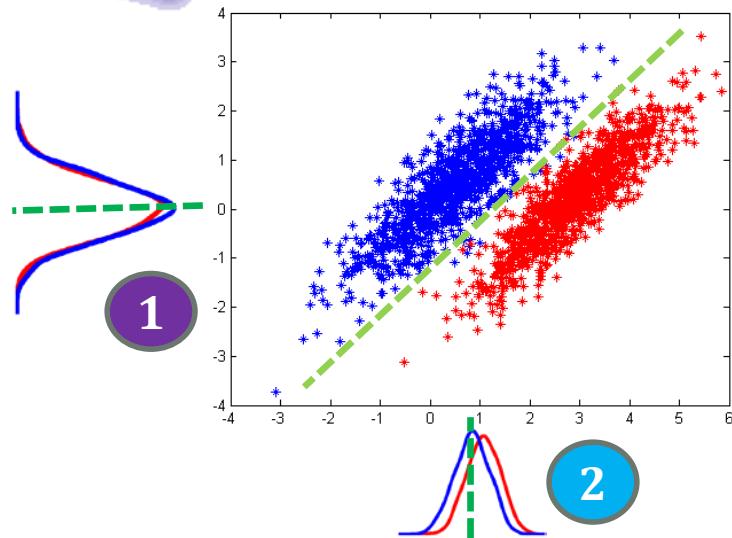
- Local activations – Univariate approach

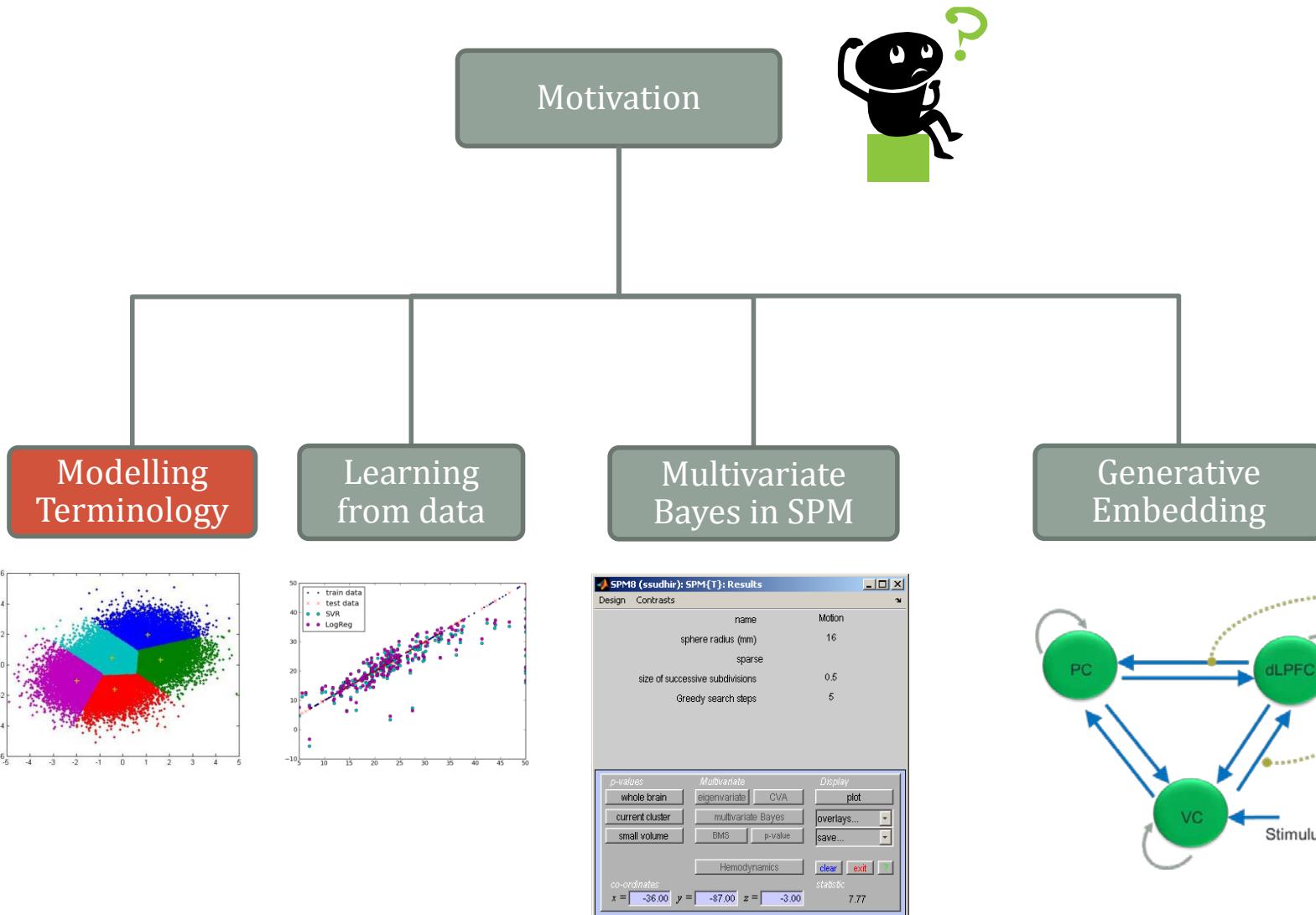


Univariate to Multivariate

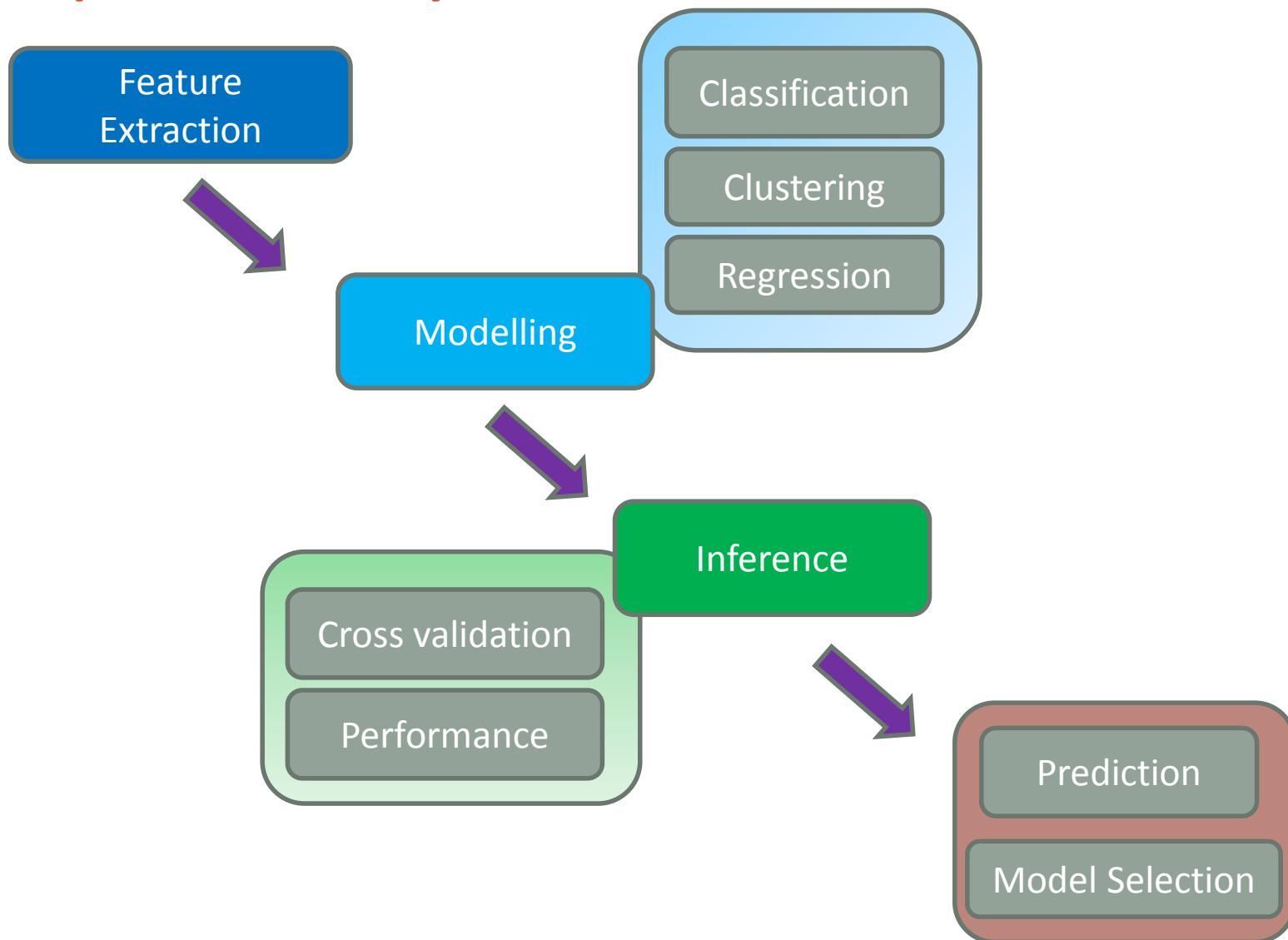


not significant
not significant



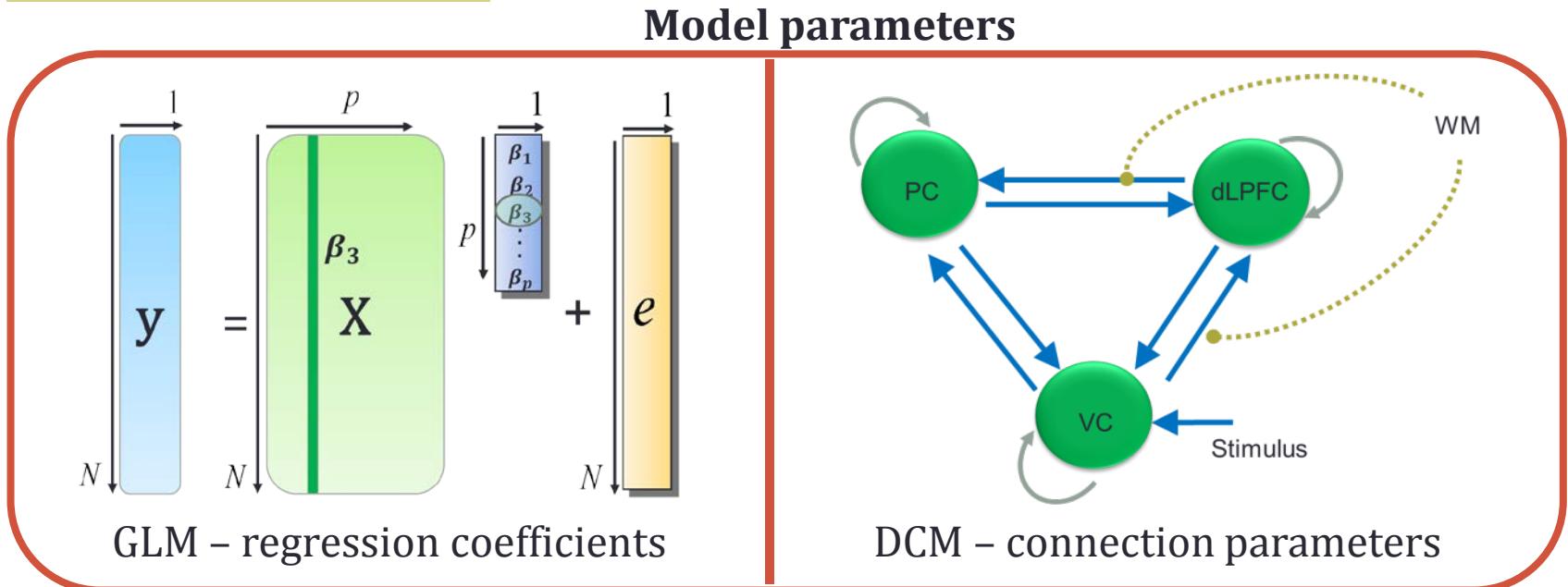
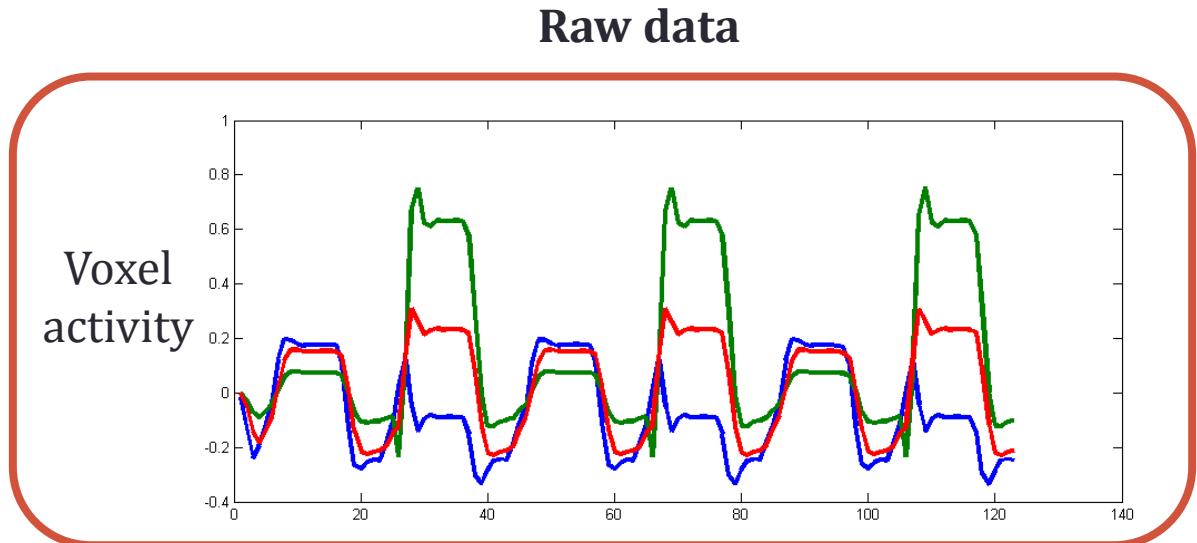


Steps for analysis

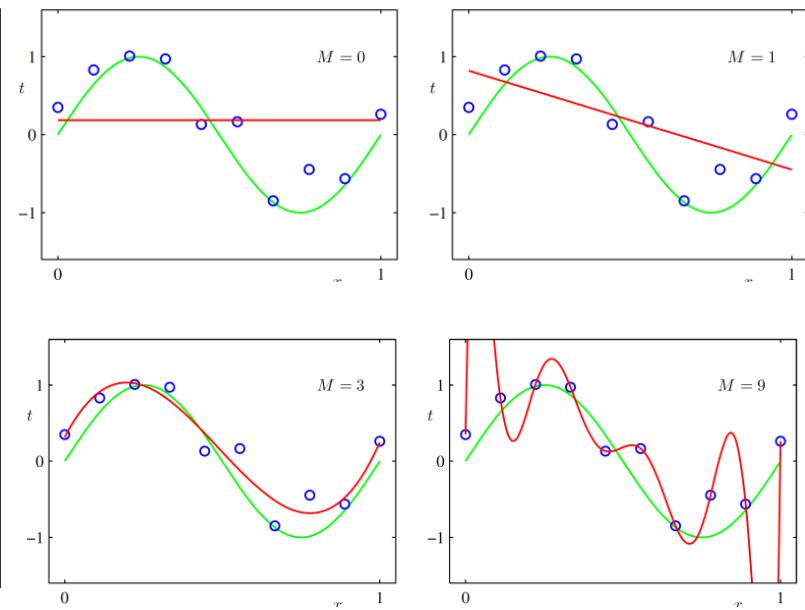
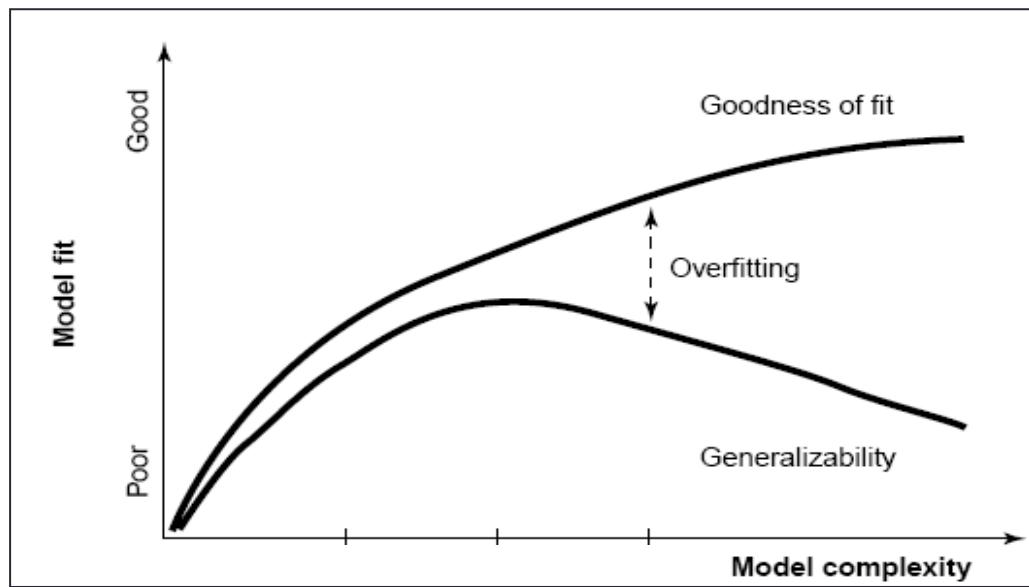


Feature Spaces

	F_1	F_2	.	.	.	F_p
S_1						
S_2	Data Point or Feature Vector					
.						
S_N	<ul style="list-style-type: none">• High dimensionality• Class/Cluster distributions• Interpretation					



Model Selection - Generalizability



Modelling goals

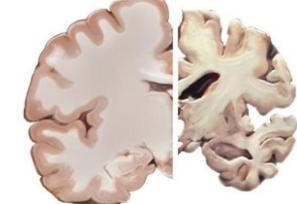
- Prediction



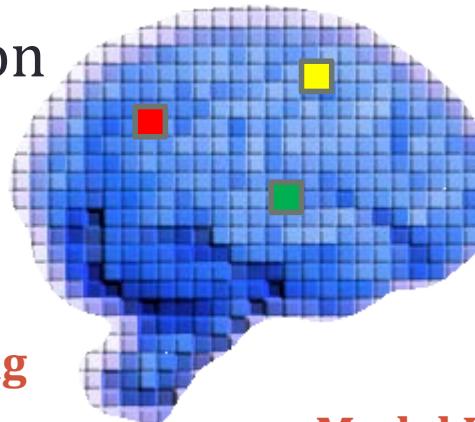
Healthy Brain Severe AD

Predictive Density

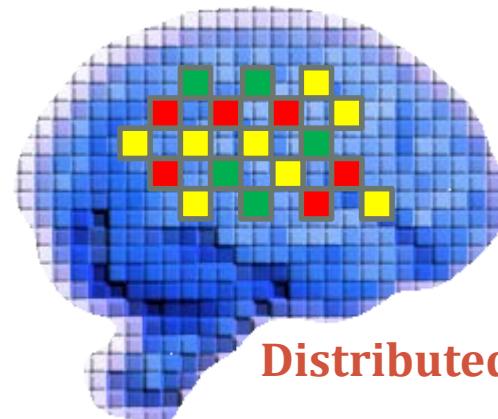
$$p(X_{\text{new}}|Y_{\text{new}}, X, Y) = \int p(X_{\text{new}}|\theta, Y_{\text{new}})q(\theta)d\theta$$



- Model Selection



vs

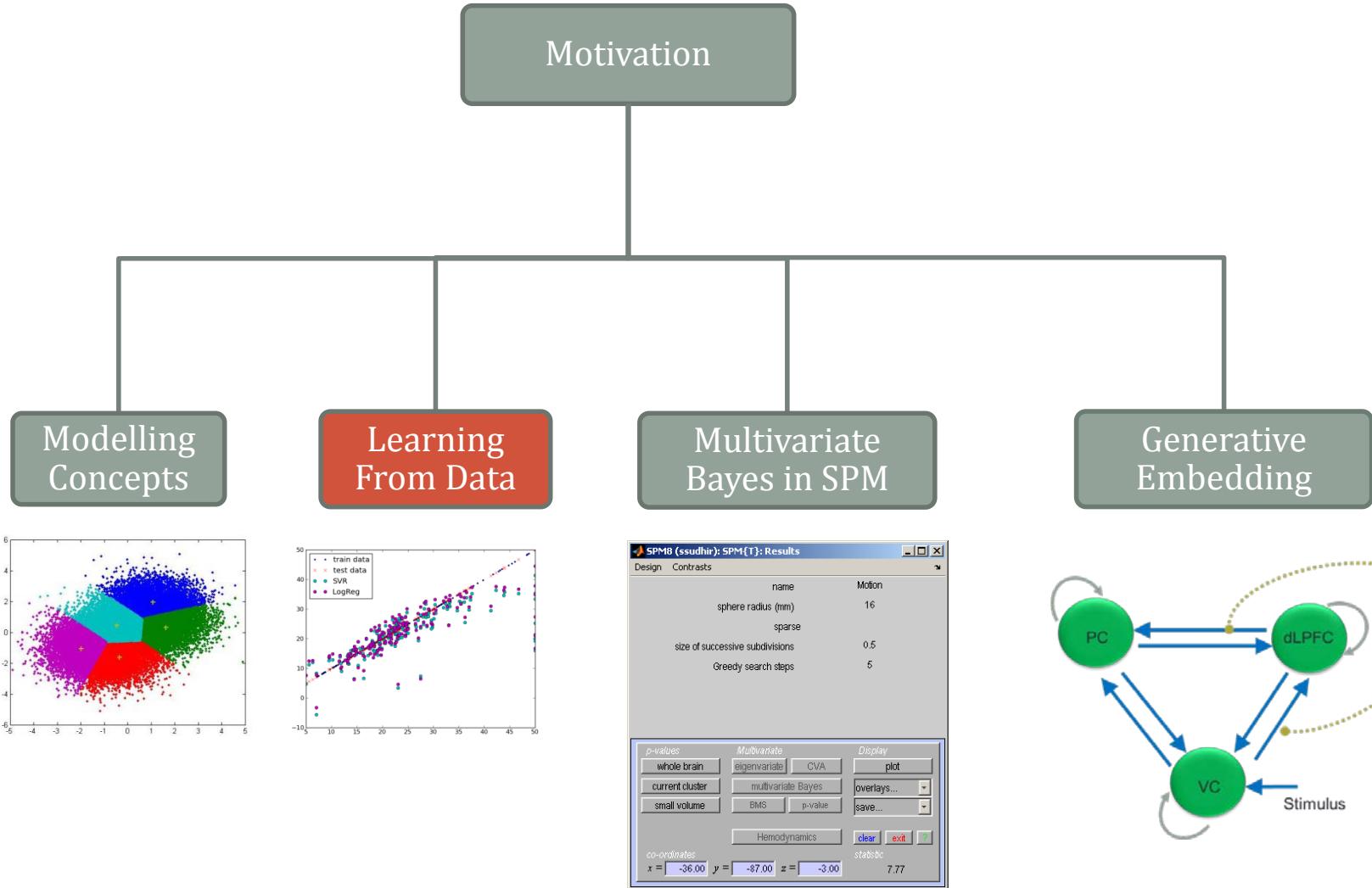


Sparse Coding

Distributed Coding

Model Evidence

$$p(Y|X) = \int p(Y, \theta|X)d\theta$$



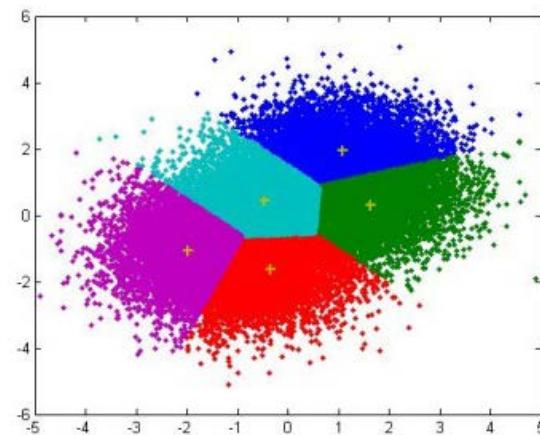
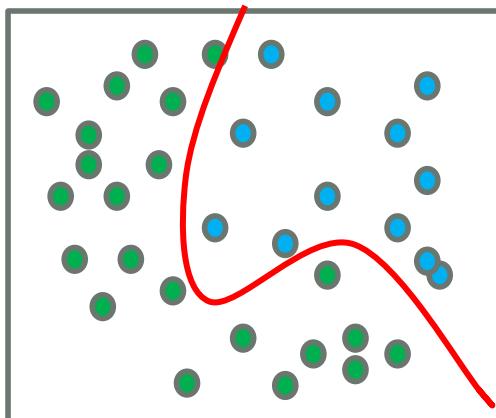
Learning from Data

Supervised Learning

Unsupervised Learning

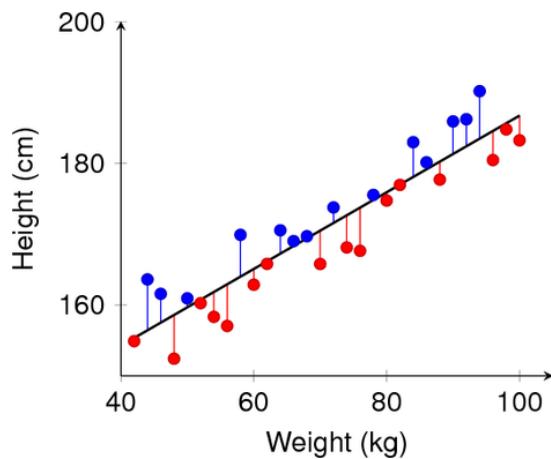
Reinforcement Learning

Semi-supervised Learning



Supervised Learning

Regression



Independent variables

X



Function - f

dependent variable
Y

Continuous

Classification

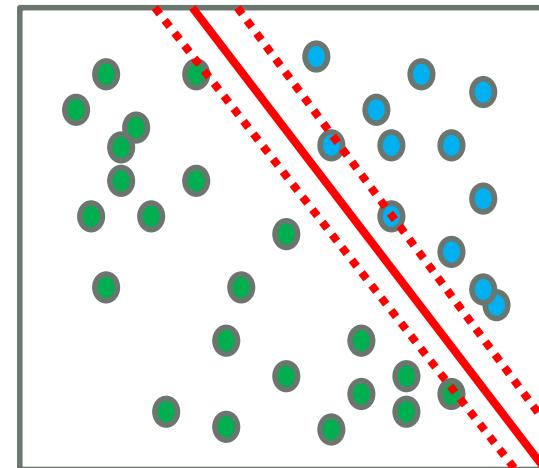
Categorical



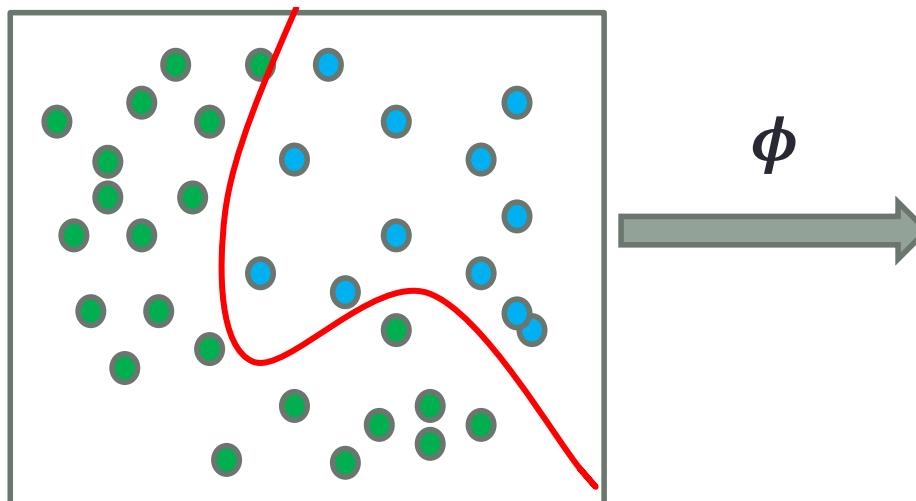
Classification



- Generative classifier
- Discriminative classifier



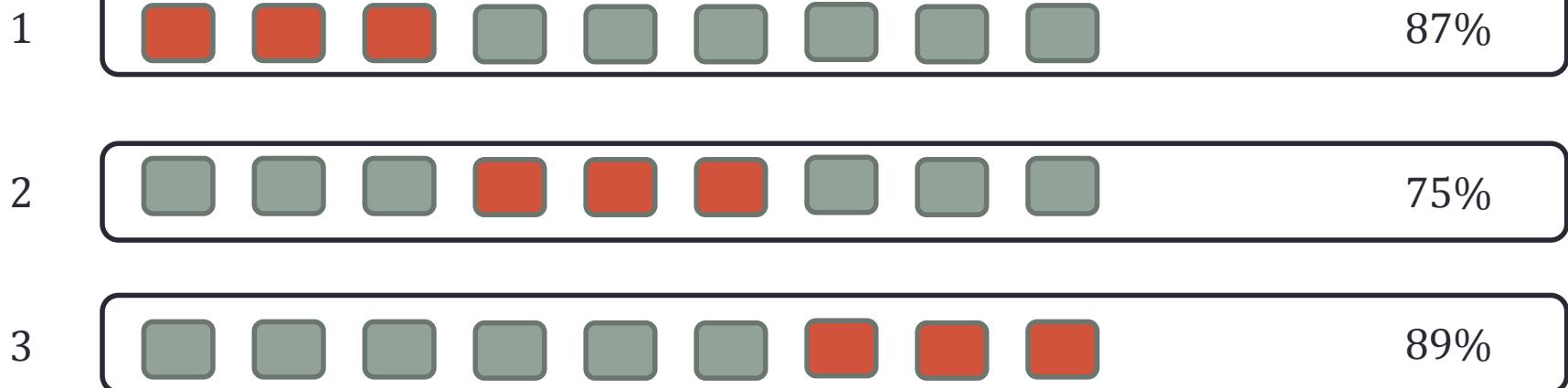
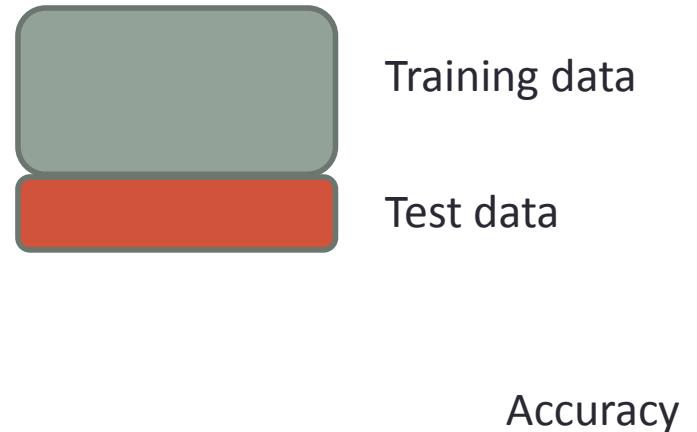
Kernel Methods



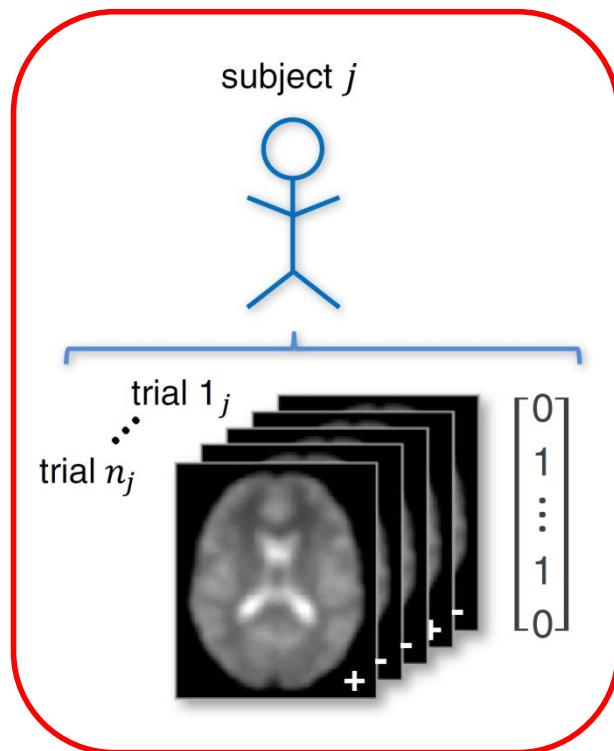
- Kernel Function – $K(x_i, x_j) = \phi(x_i) \cdot \phi(x_j)$

K-fold Cross Validation

- Model Selection
- Performance evaluation



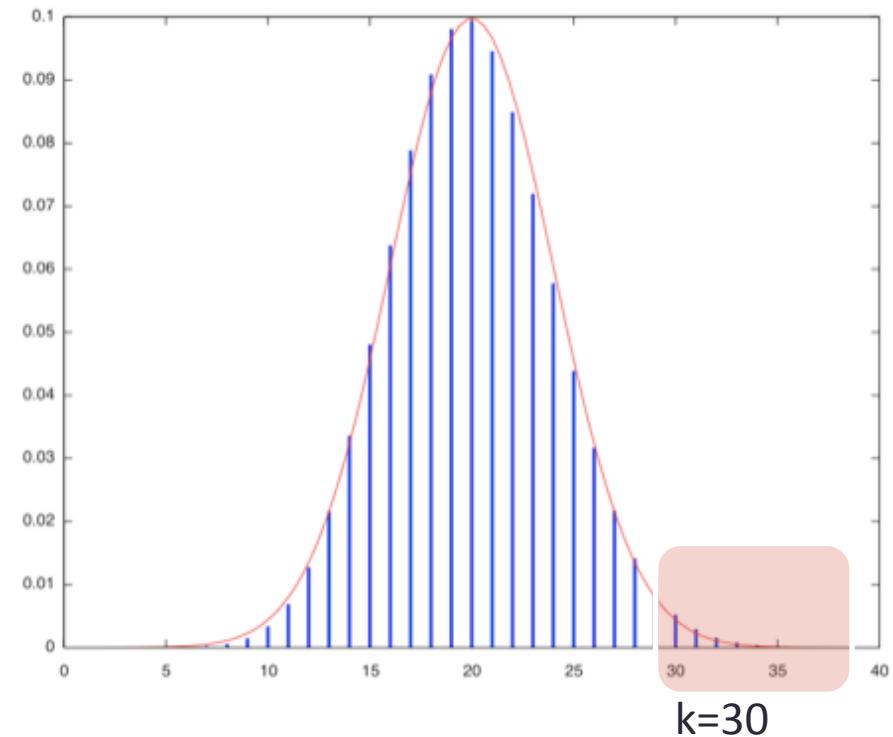
Performance – Single Subject



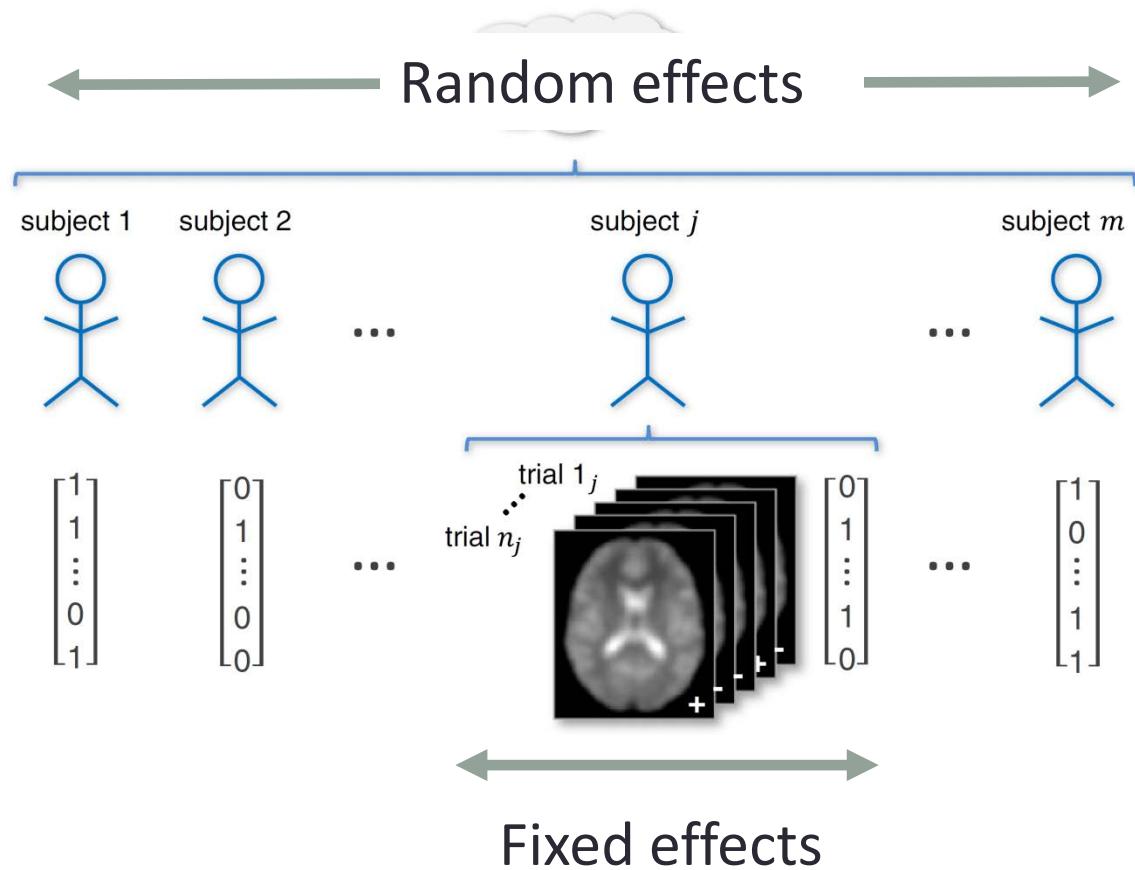
Brodersen et al. 2013, *NeuroImage*

Binomial Test

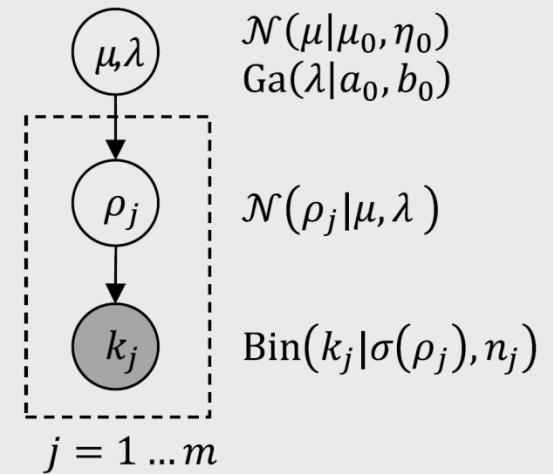
$$p = P(X \geq k | H_0) = 1 - B(k | n, \pi_0)$$



Performance – Multiple Subjects



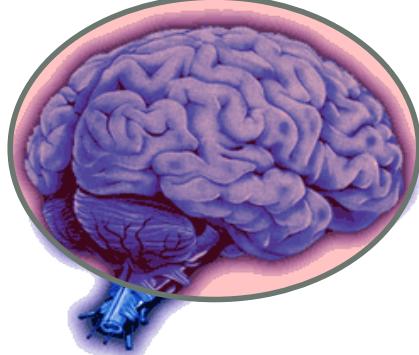
b Bayesian mixed-effects inference (univariate normal-binomial model)



<http://www.translationalneuromodeling.org/tapas/>

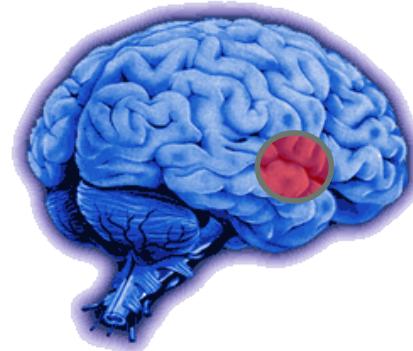
Using classification for fMRI data

Whole brain



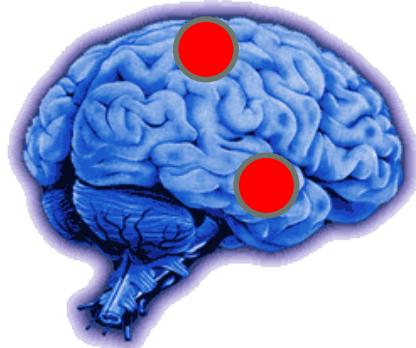
Mourao-Miranda et al. (2005) *NeuroImage*

Searchlight classifier



Nandy & Cordes (2003) *MRM*, Kriegeskorte et al. (2006) *PNAS*

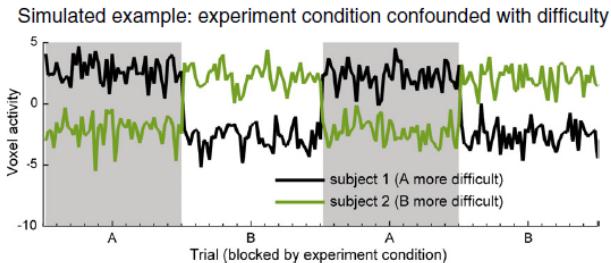
Pattern localization



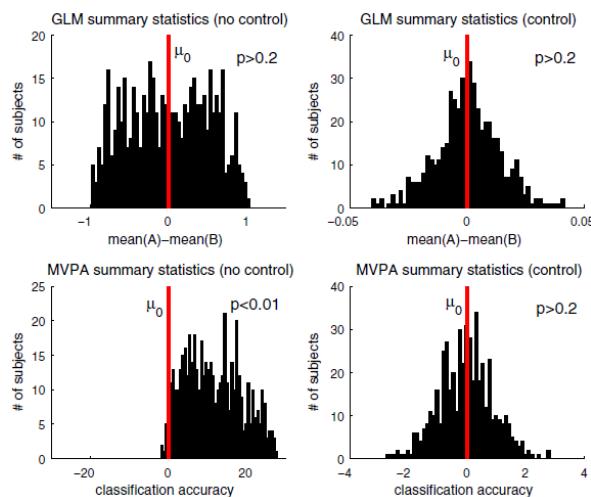
Word Category
Food
Building
People

Pereira et al. (2009) *NeuroImage*, Mitchell et al. (2004) *Machine Learning*

Confounds – GLM vs MVPA

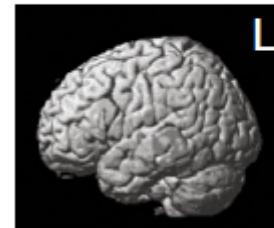


Individual-Subject Summary Statistics		
Subject	Experimental Effect (GLM)	Discrimination Success (MVPA)
Subject 1	mean(A)-mean(B) = +4.75	classification accuracy = +13.15, within-minus-across = +3.826
Subject 2	mean(A)-mean(B) = -5.56	classification accuracy = +13.44, within-minus-across = +3.848
Group Test Statistics (two-tailed <i>t</i> -test)		
Experimental Effect (GLM)		
	mean(A)-mean(B): $t_1=-0.0780, p=0.9504, \text{n.s.}$	classification accuracy: $t_1=94.0, p<0.01, \text{sig.}$

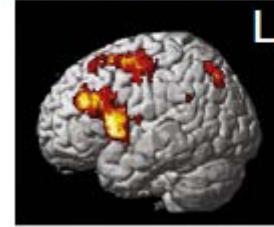


Task Rule (A vs. B)

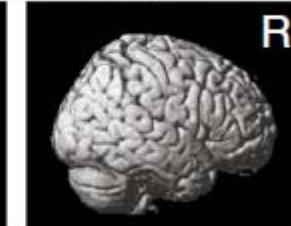
GLM



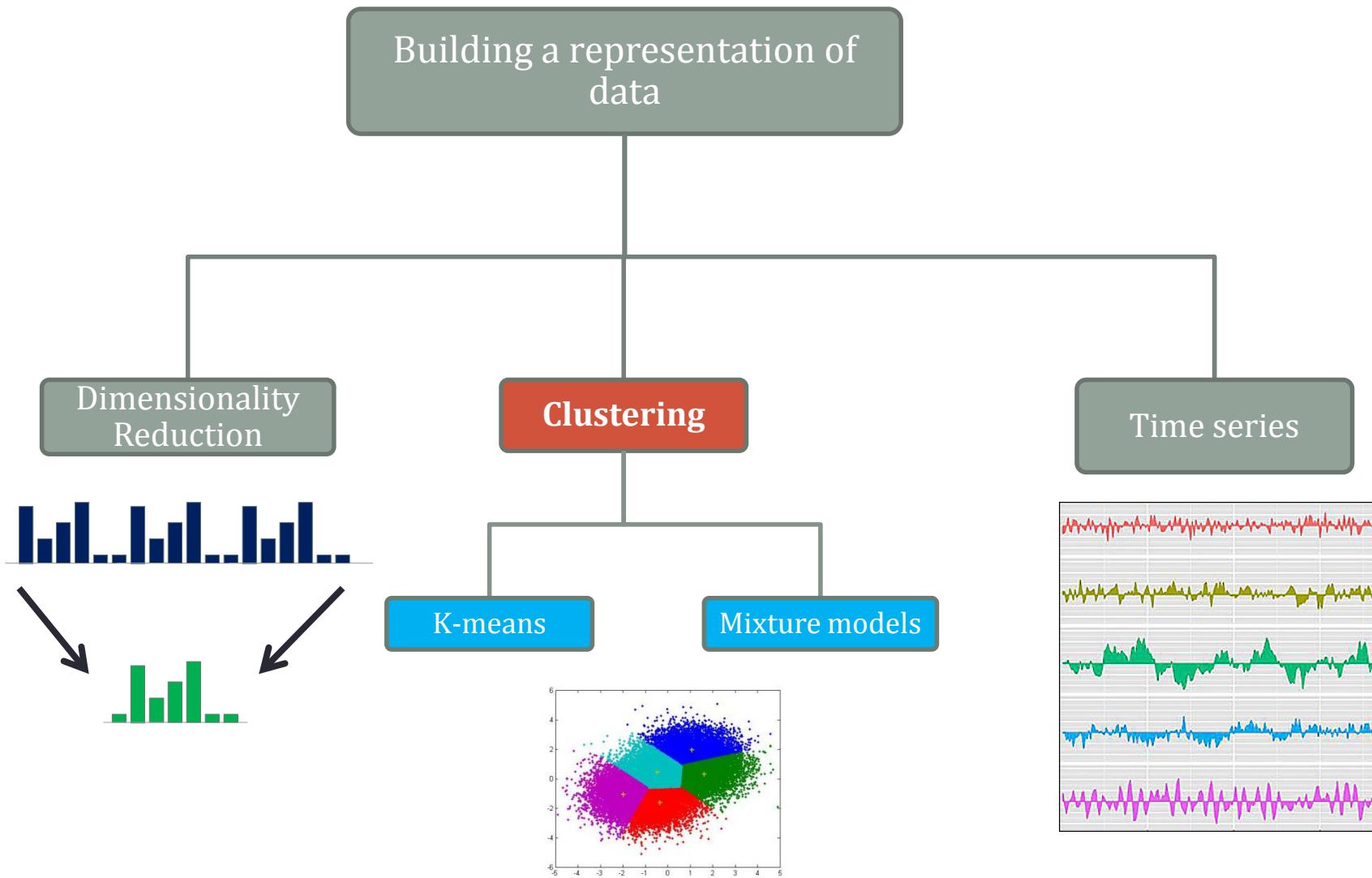
MVPA (no control)



MVPA (after RT regression)



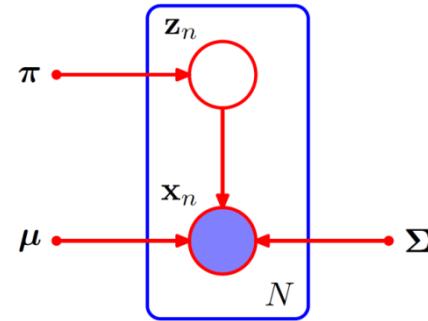
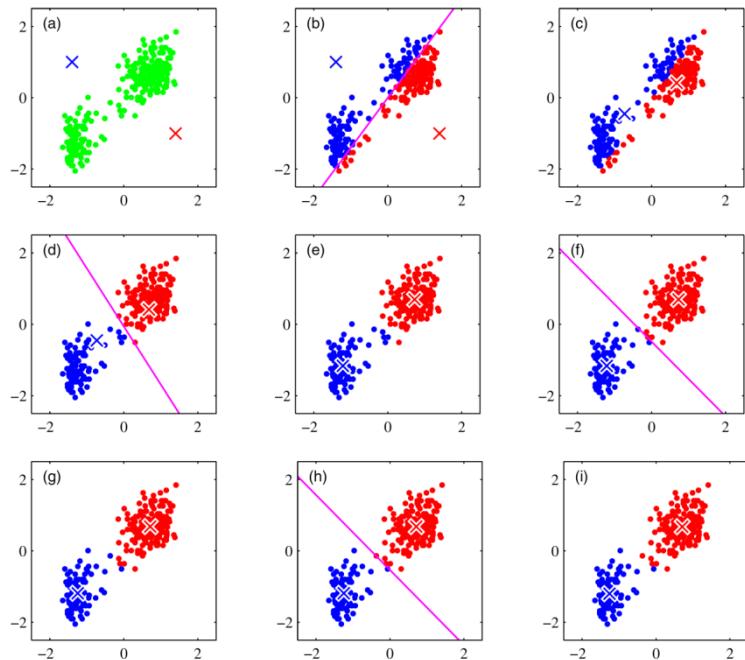
Unsupervised Learning



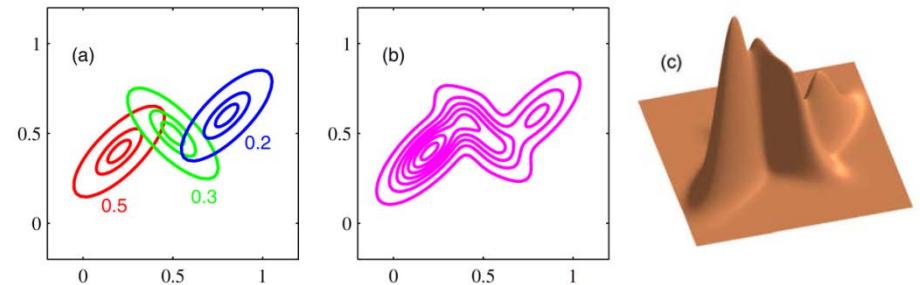
K-means

Mixture of Gaussians

$$\tilde{J} = \sum_{n=1}^N \sum_{k=1}^K r_{nk} \mathcal{V}(\mathbf{x}_n, \boldsymbol{\mu}_k)$$

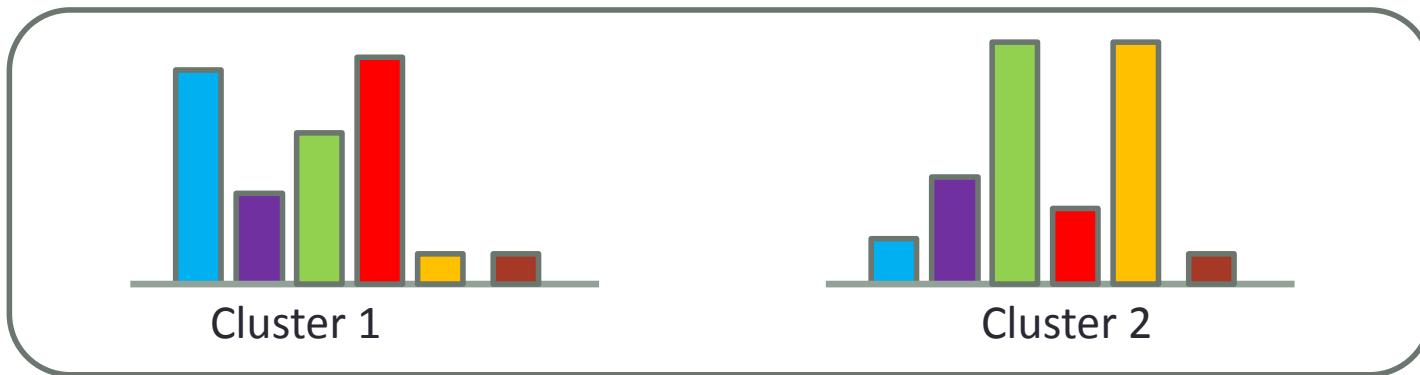


$$p(\mathbf{x}) = \sum_{k=1}^K \pi_k \mathcal{N}(\mathbf{x} | \boldsymbol{\mu}_k, \boldsymbol{\Sigma}_k)$$

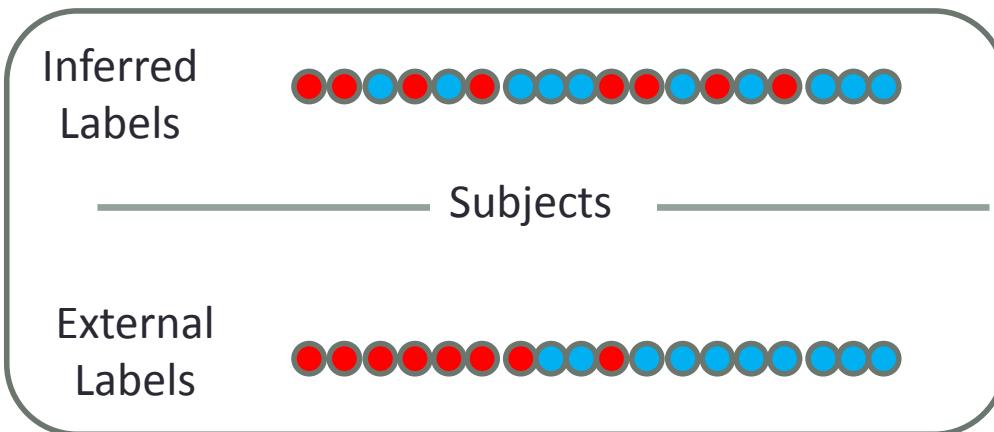


Interpretation

- Cluster parameters



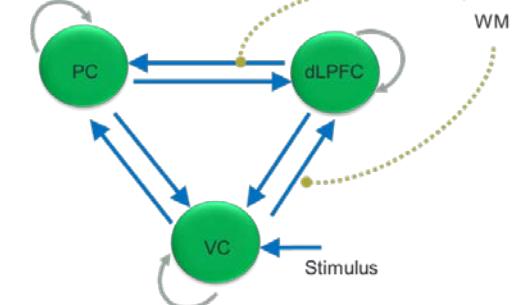
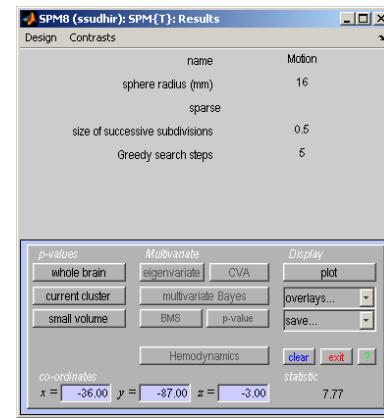
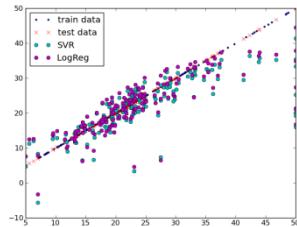
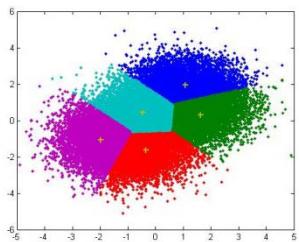
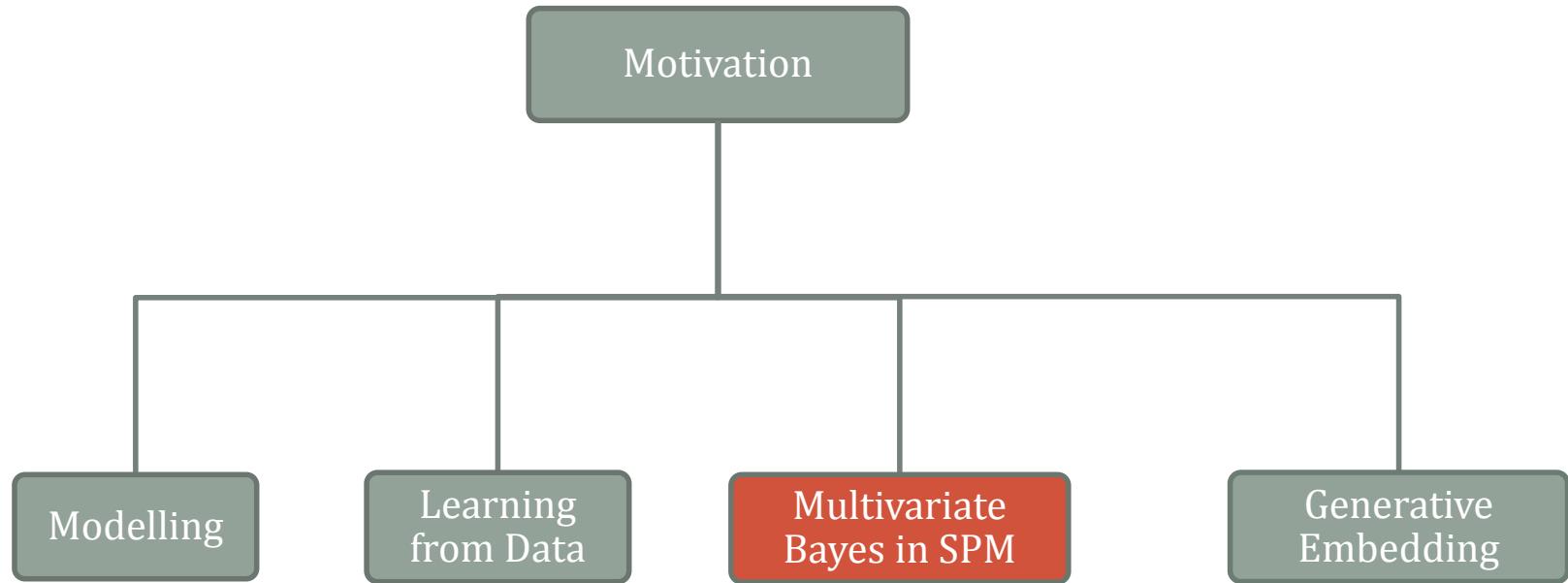
- Purity



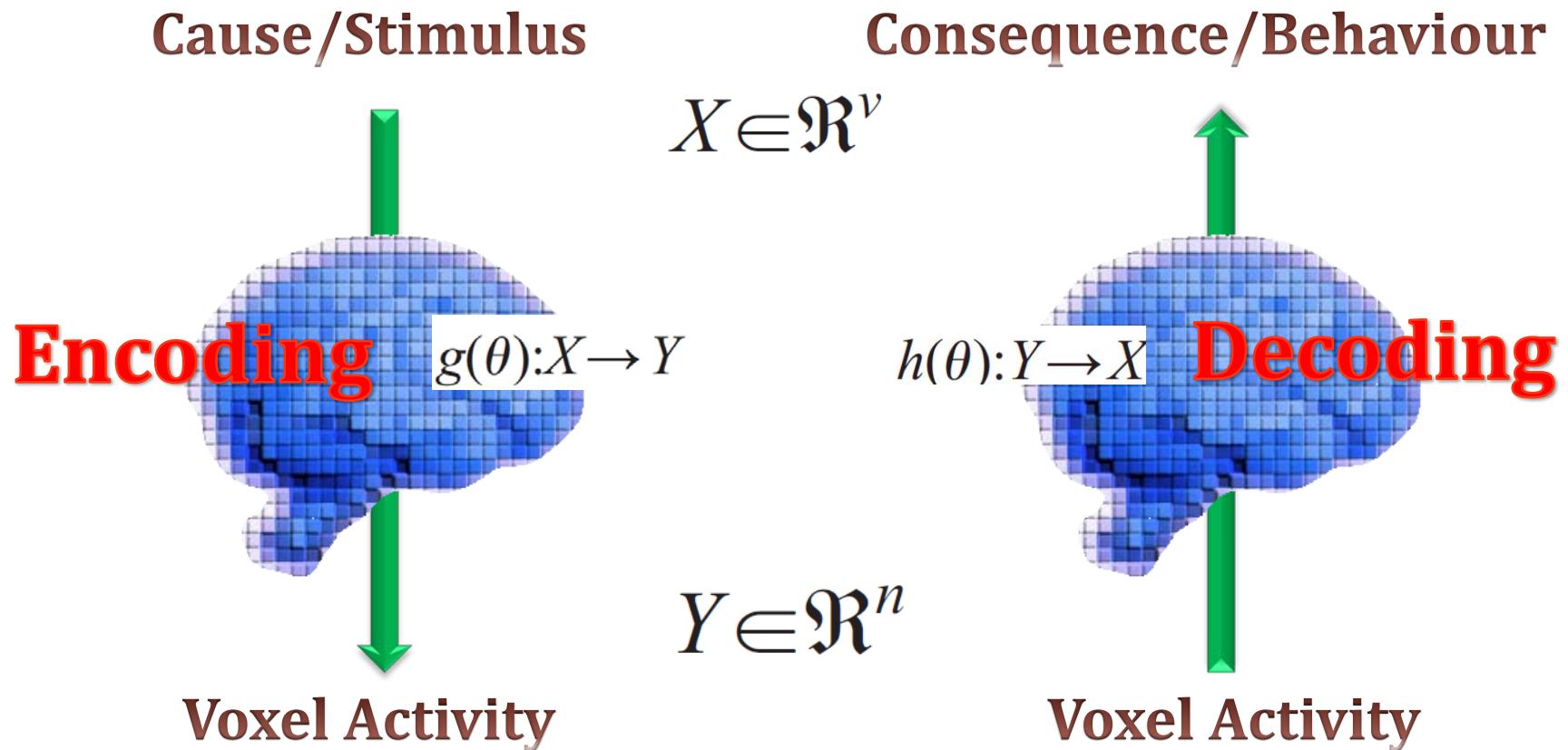
Normalized Mutual Information (NMI)

Balanced purity

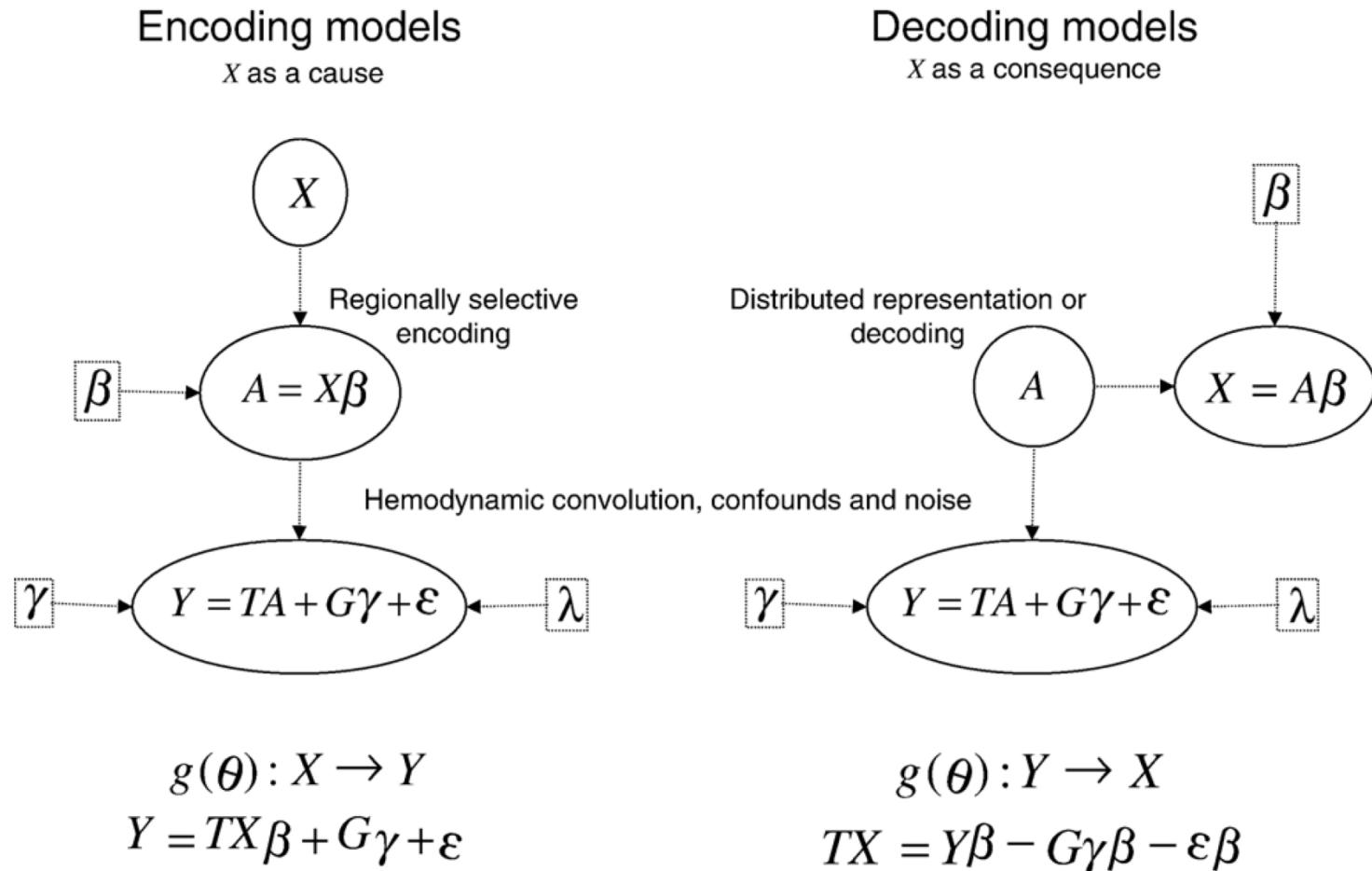
Rand Index



Encoding Vs Decoding Models

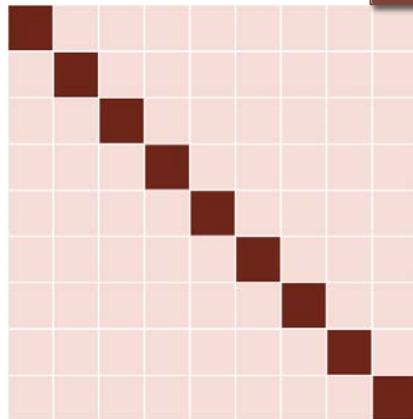


Encoding Vs Decoding

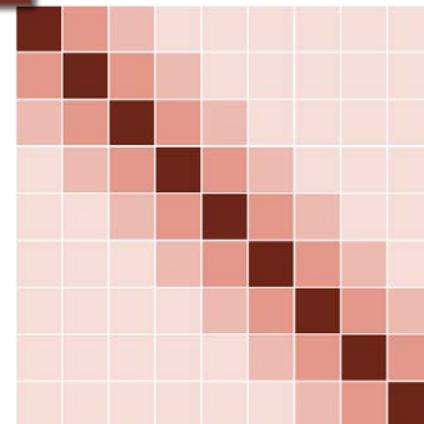


Coding hypotheses

Sparse vectors

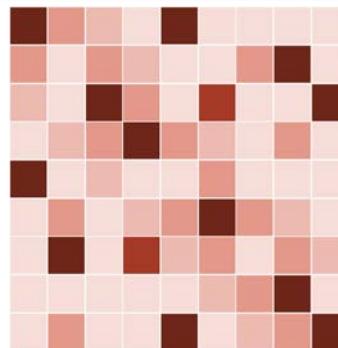


Spatial vectors



Smooth vectors

Distributed vectors



Singular vectors
of data

$$UDV^T = RY^T$$

Support vectors

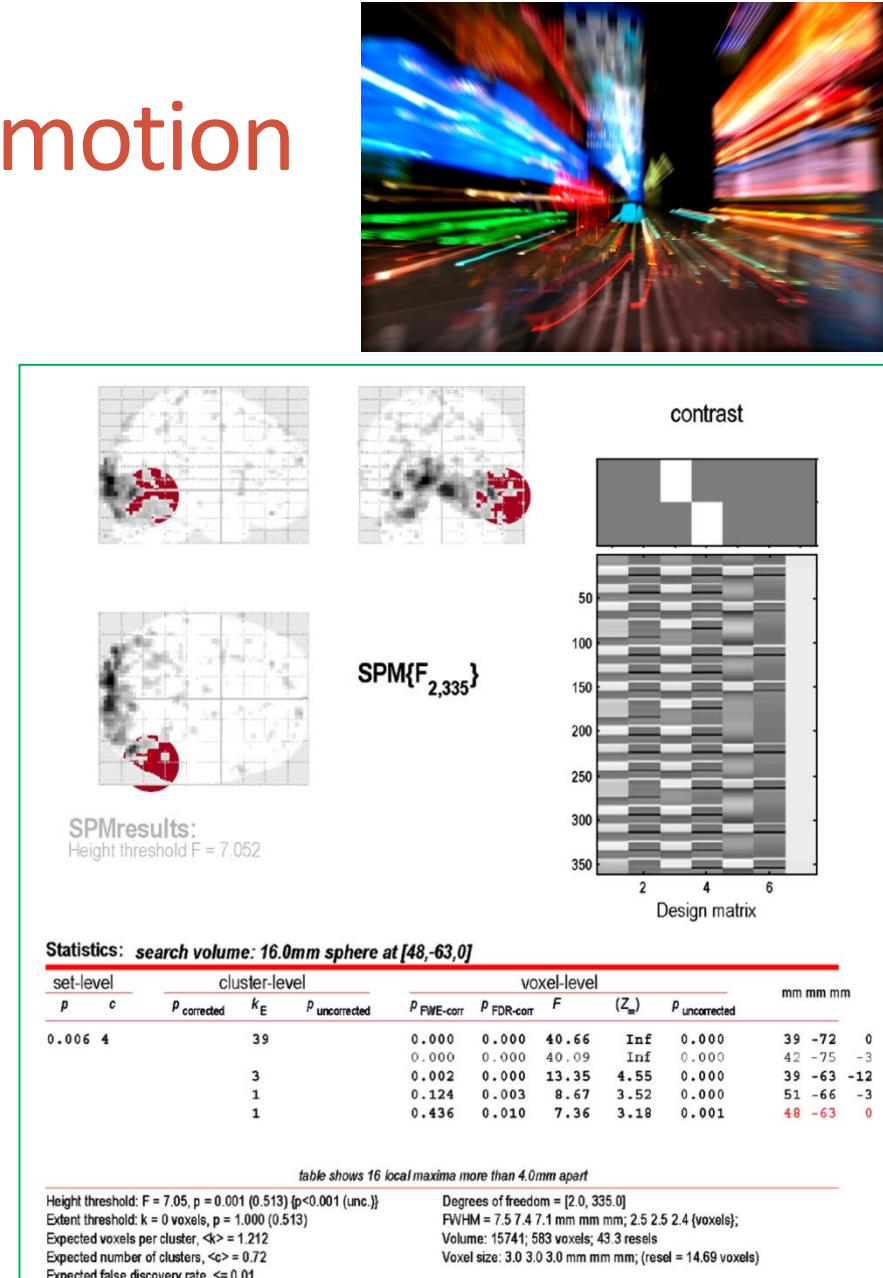
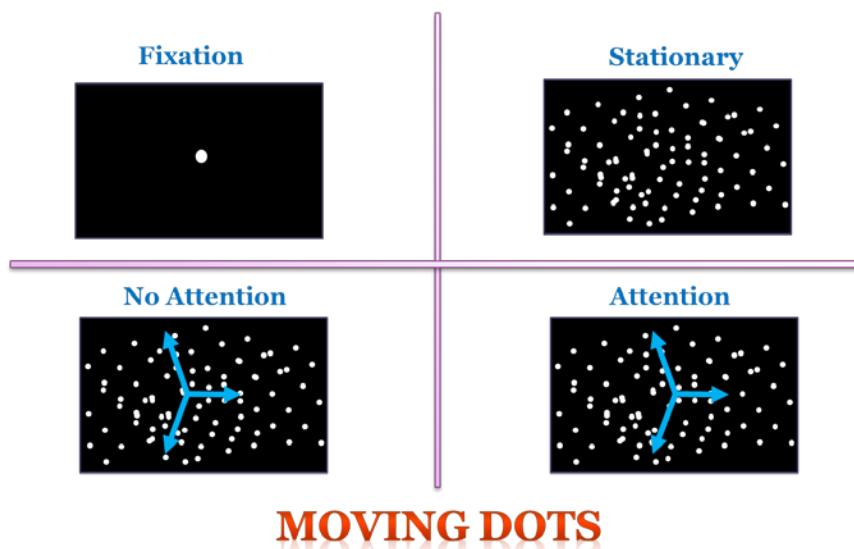
$$U = RY^T$$

Bayesian decoding of motion

Attention to motion dataset - Büchel &
Friston 1999 Cerebral Cortex

Experimental factors:

1. Photic
2. Motion
3. Attention



Multivariate Bayes in SPM

SPM12 (6225): Menu

Scans Previews

- Realign (Es...)
- Slice timing
- Smooth
- Coregister (...)
- Normalise (...)
- Segment

Inference

- Specify 1st-level
- Review
- Specify 2nd-level
- Estimate
- Results (circled in red)
- Dynamic Causal Modelling

SPM for functional MRI

- Display
- Check Reg
- Render... FMRI
- Toolbox:
- Help

SPM8 (ssudhir): SPM{T}: Results

Design Contrasts

		Motion
sphere radius (mm)		16
sparseness		
size of successive subdivisions		0.5
Greedy search steps		5

p-values Multivariate Display

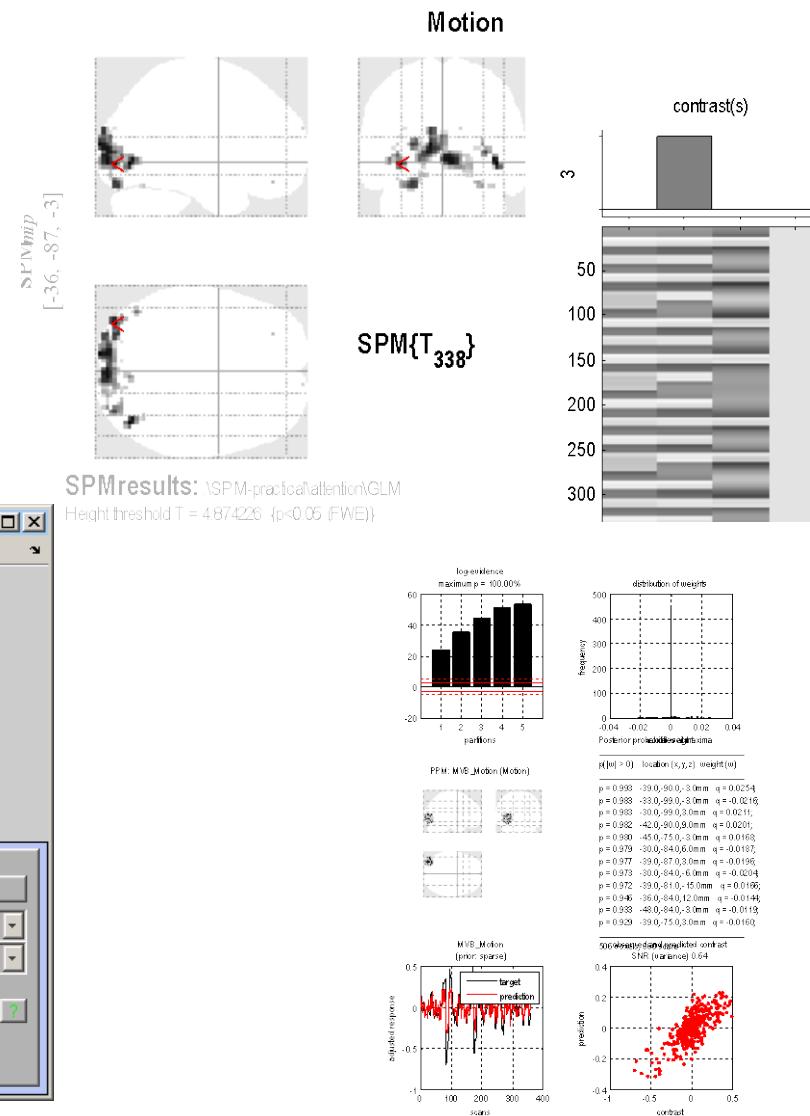
- whole brain
- eigenvalue
- CVA
- plot
- current cluster
- multivariate Bayes
- BMS
- p-value
- overlays...
- save...

co-ordinates

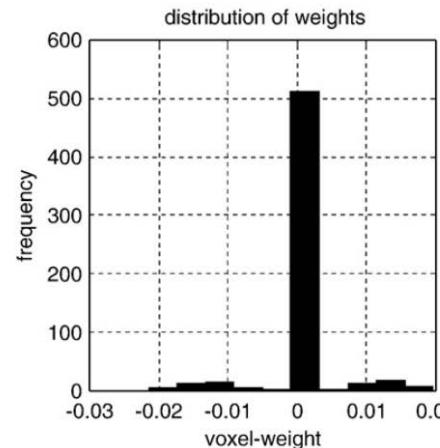
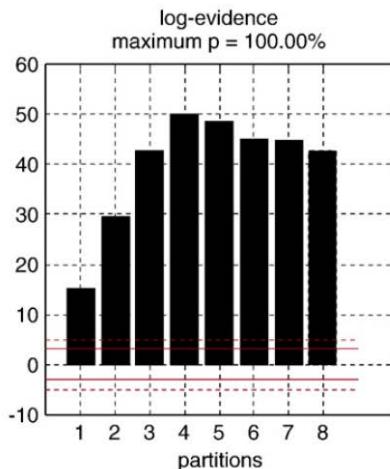
x = -36.00 y = -87.00 z = -3.00

statistic 7.77

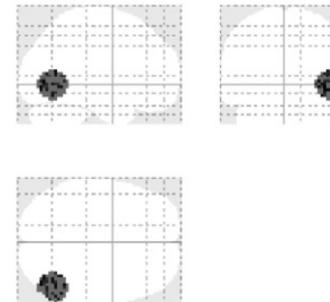
Hemodynamics clear exit ?



Results



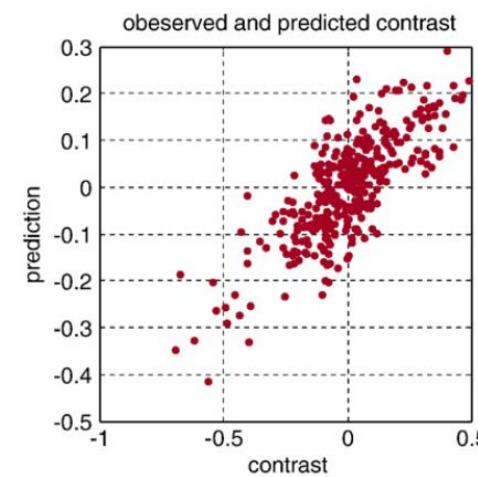
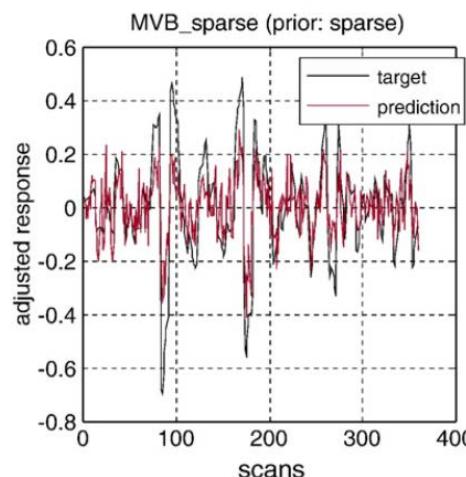
MVB_sparse (motion)



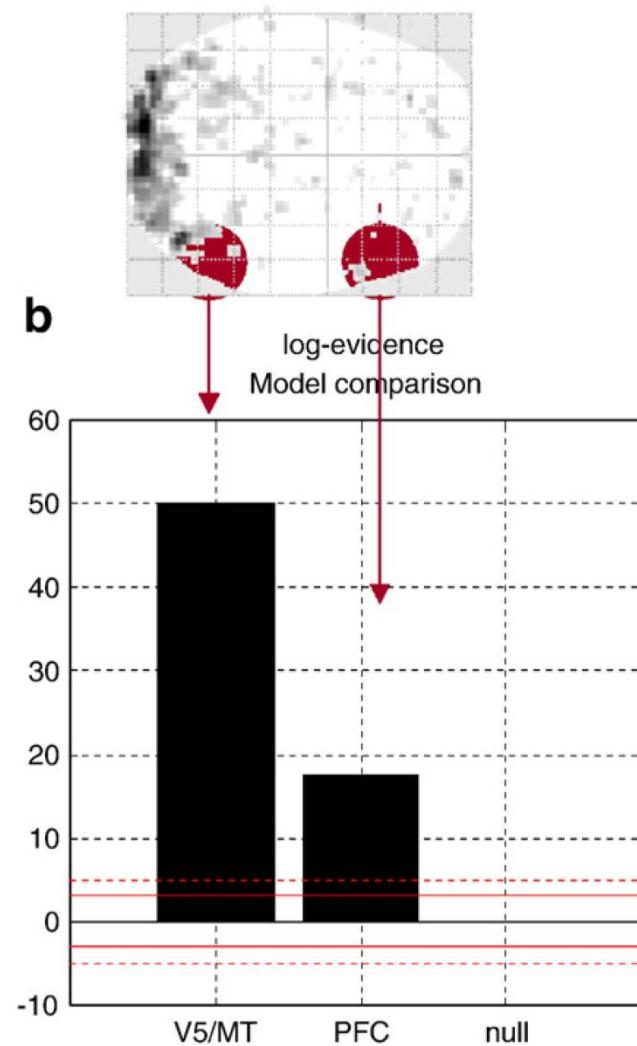
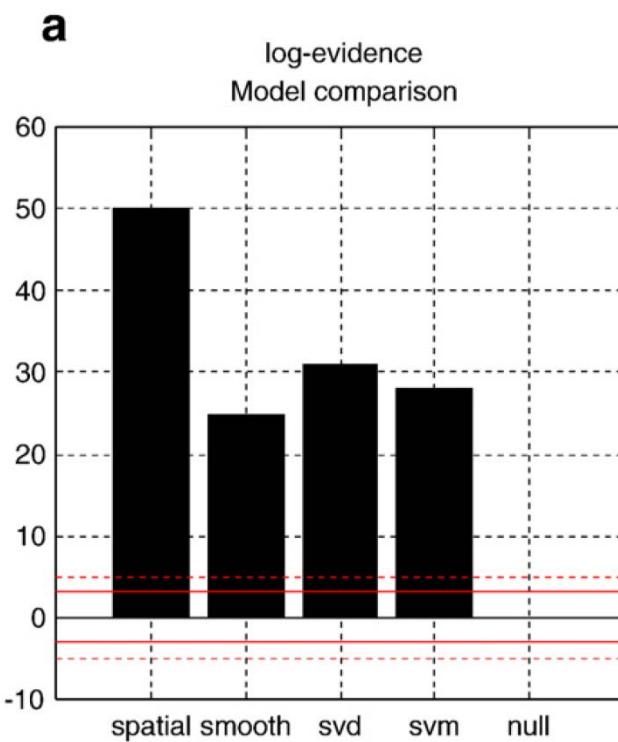
Posterior probabilities at maxima

p(iwl > 0)	location (x,y,z)	weight (w)
p = 0.991	48,-78,0mm	q = -0.0208;
p = 0.977	48,-72,-3mm	q = -0.0215;
p = 0.973	36,-72,3mm	q = 0.0185;
p = 0.972	45,-51,9mm	q = 0.0188;
p = 0.968	39,-66,-6mm	q = -0.0180;
p = 0.966	42,-54,-3mm	q = -0.0168;
p = 0.963	45,-75,-6mm	q = 0.0196;
p = 0.954	54,-54,9mm	q = 0.0154;
p = 0.947	63,-60,3mm	q = -0.0161;
p = 0.945	42,-63,0mm	q = 0.0150;
p = 0.942	60,-60,-9mm	q = -0.0136;
p = 0.942	36,-57,6mm	q = -0.0167;

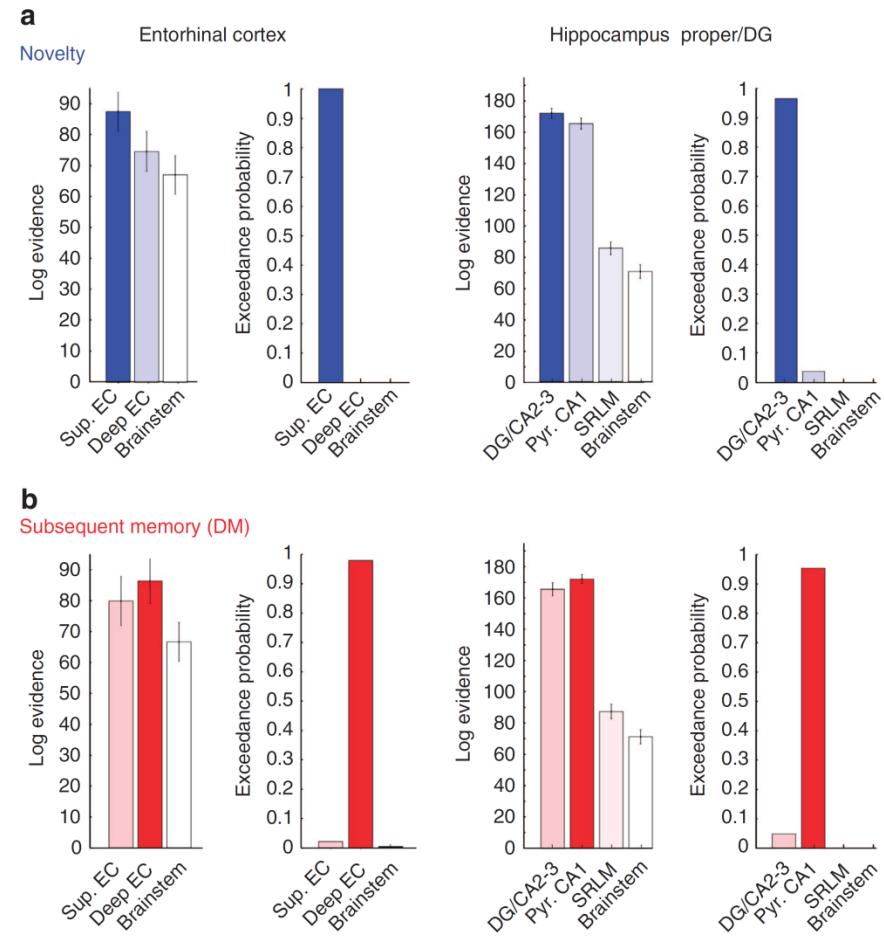
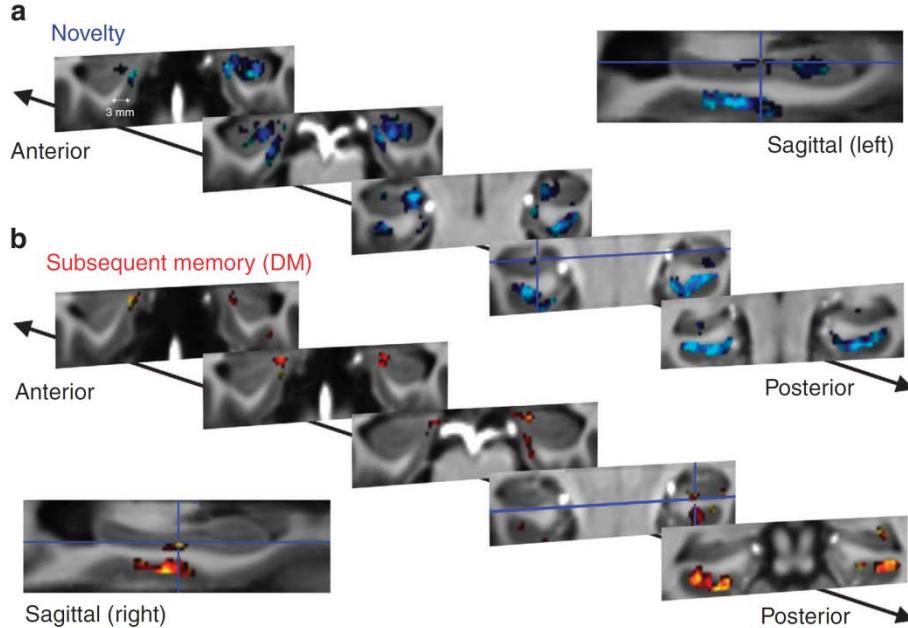
583 voxels; 360 scans

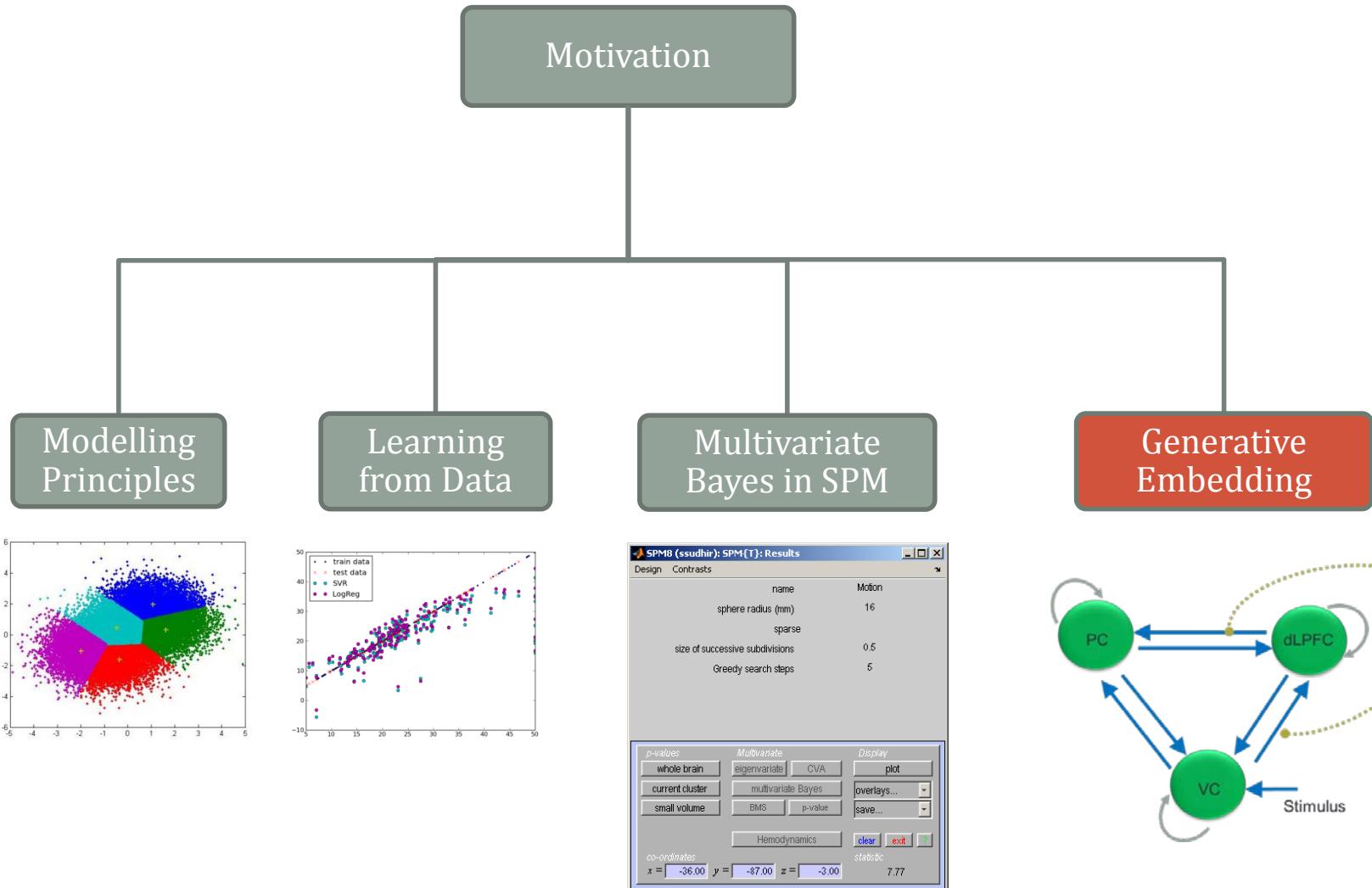


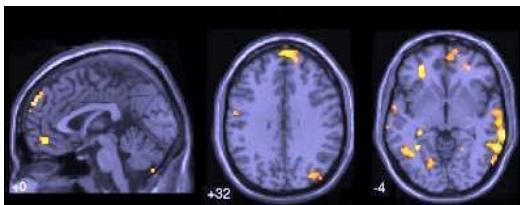
Results



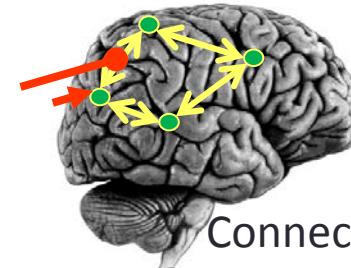
Laminar activity related to novelty and episodic encoding



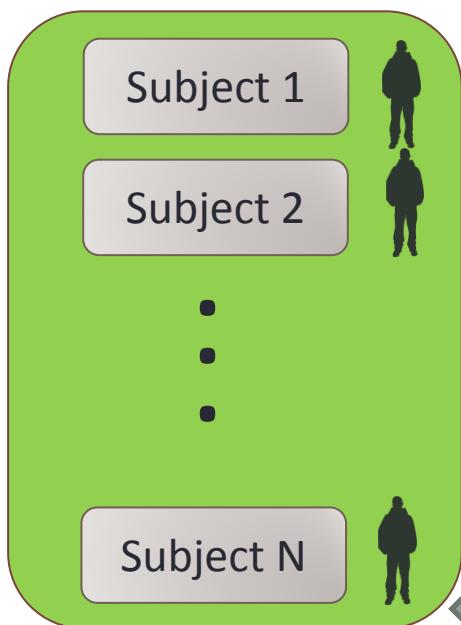




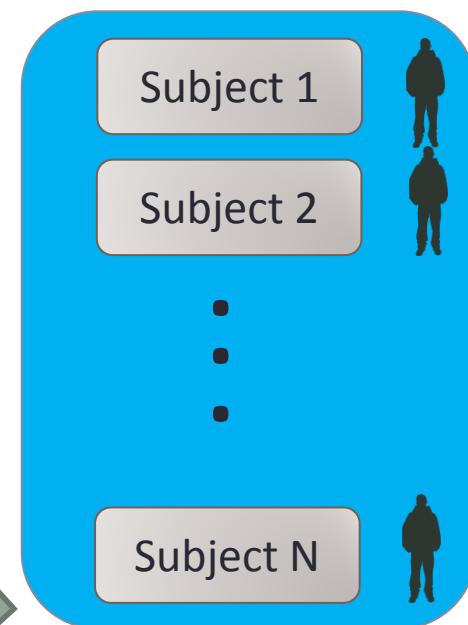
Voxel activity



Connectivity



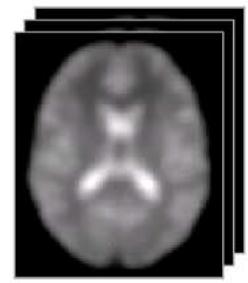
Dynamic causal
model (DCM)



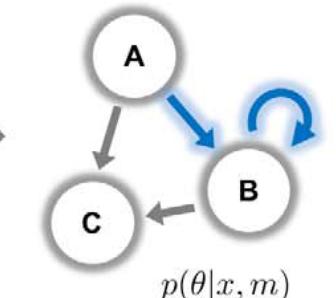
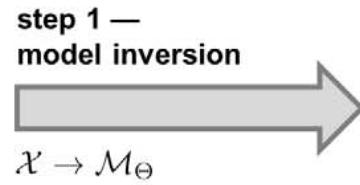
Classification
Clustering



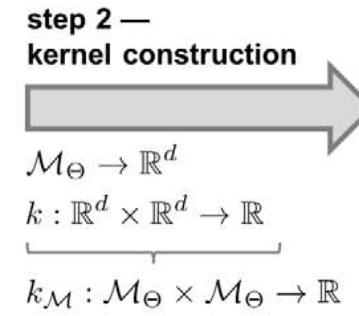
Generative Embedding - Classification



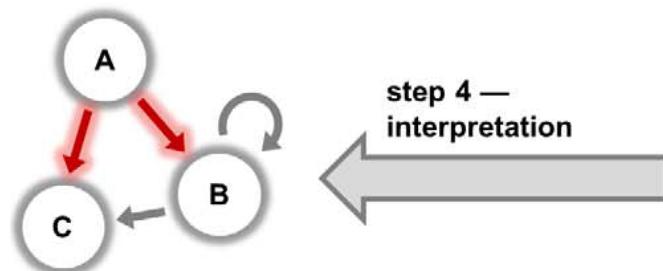
measurements from
an individual subject



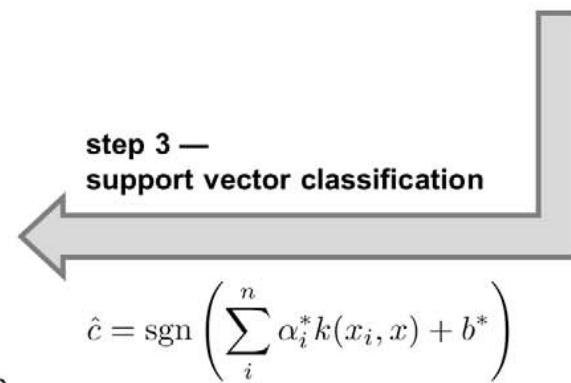
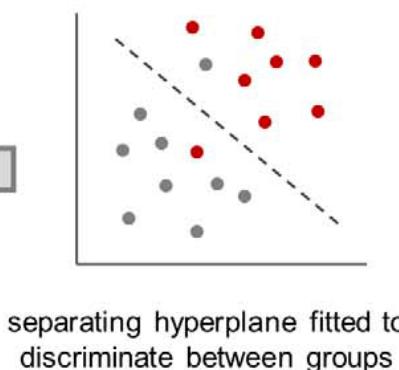
subject-specific
inverted generative model



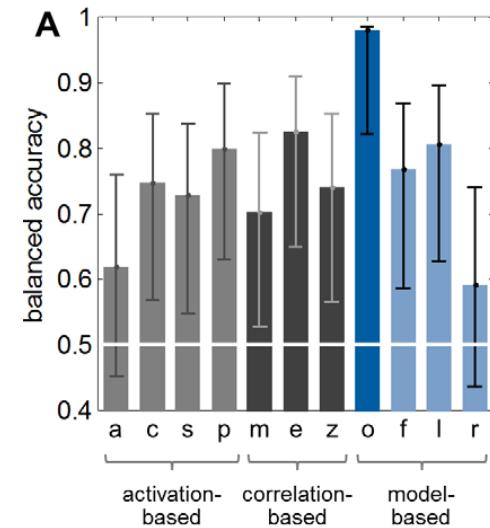
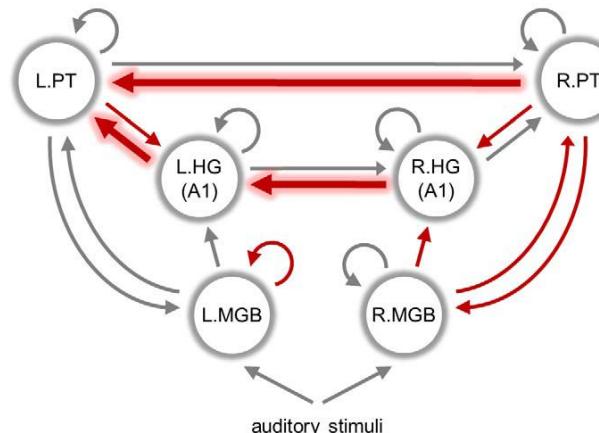
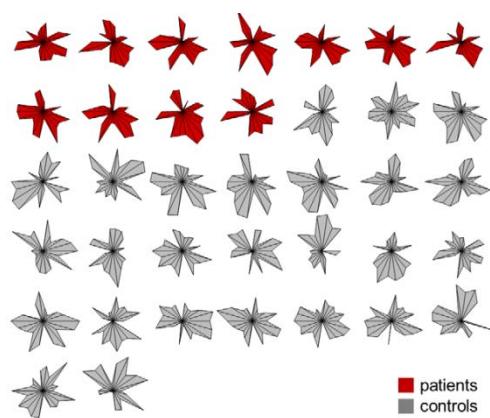
subject representation in the
generative score space



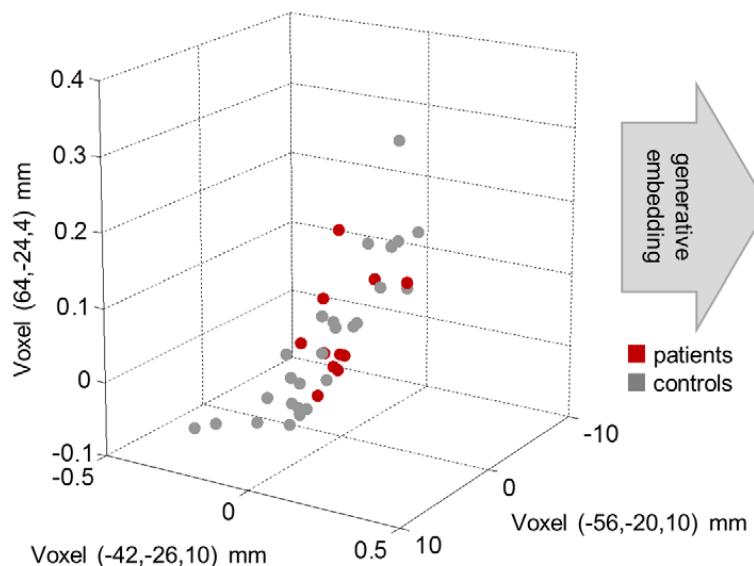
jointly discriminative
connection strengths



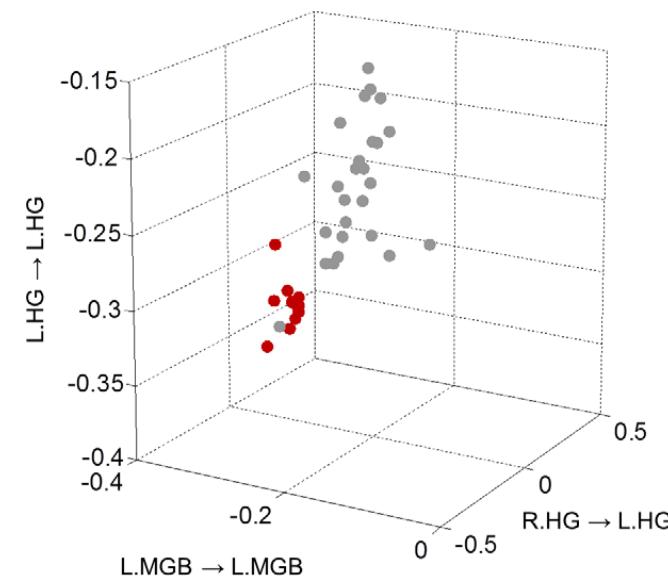
DCM - Speech processing



Voxel-based feature space

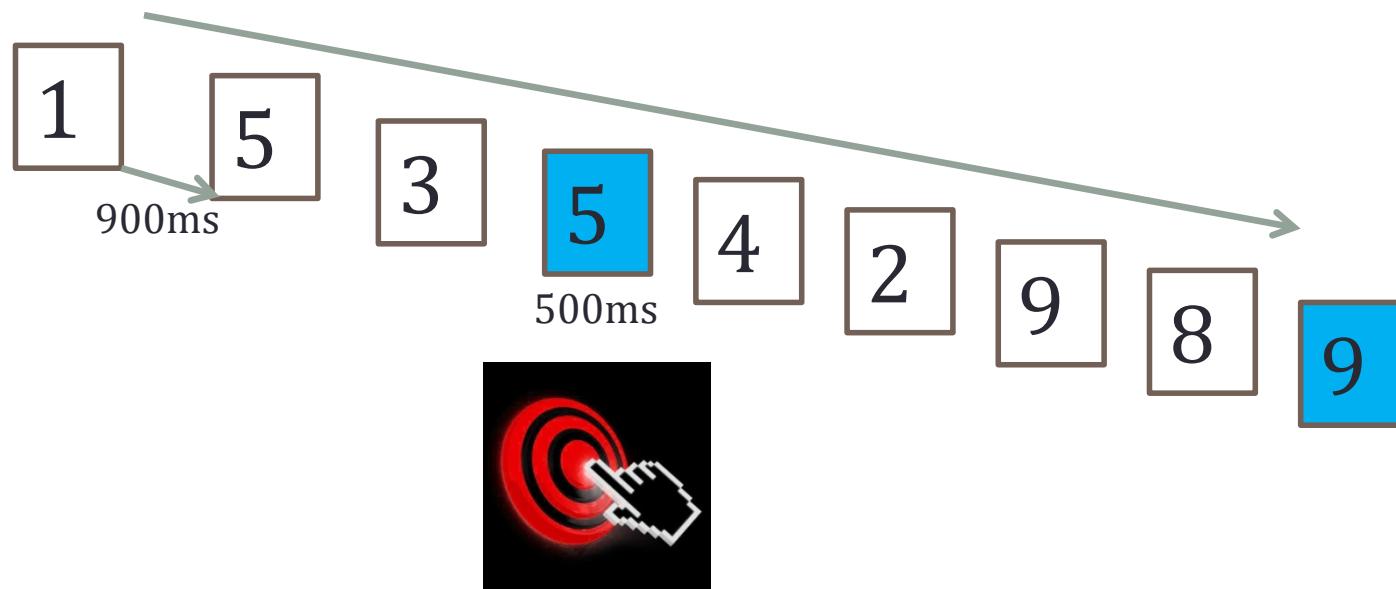


Generative score space

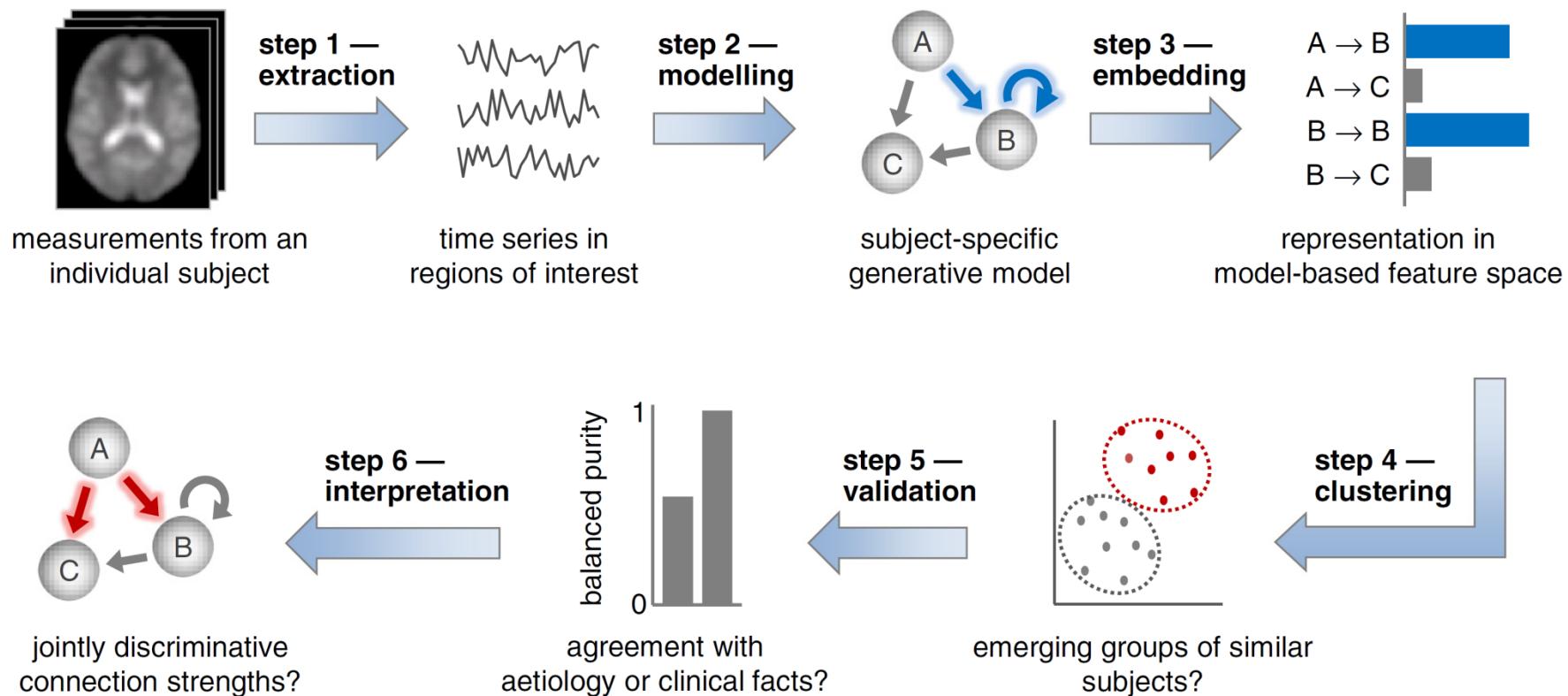


Working memory - fMRI

- 41 Schizophrenia patients (DCM IV, ICD 10), 42 controls
- Visual numeric n-back working memory task

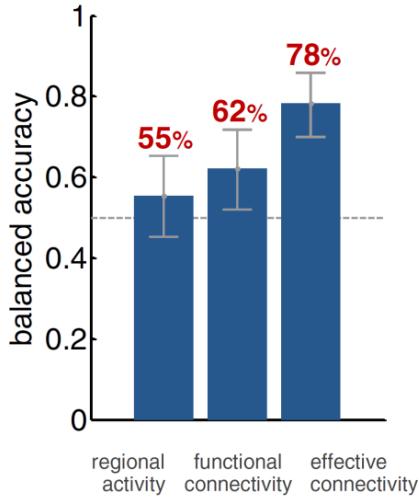


Model based clustering

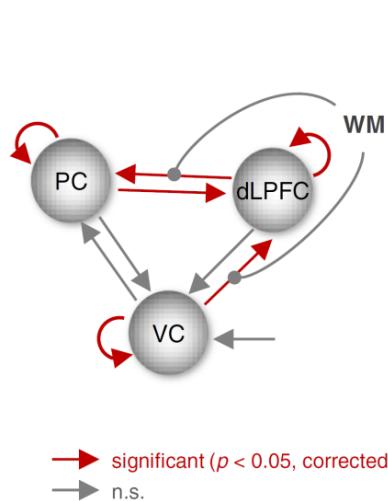


Results – Healthy vs Schizophrenic

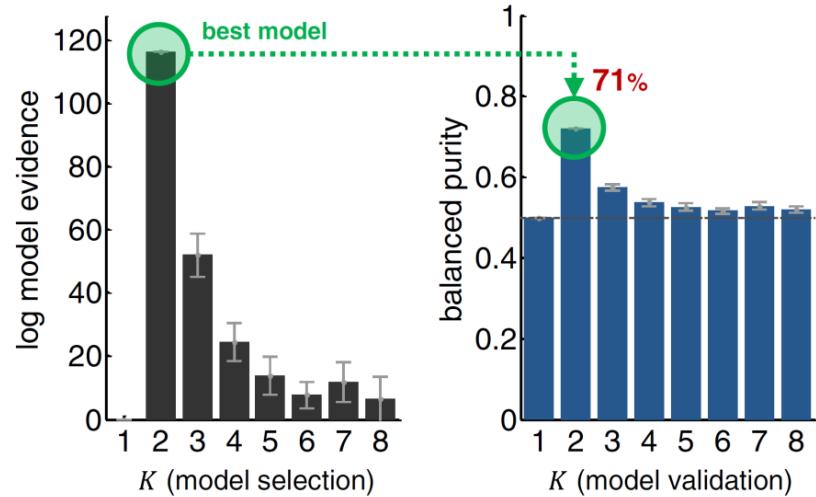
a supervised learning:
SVM classification



b discriminative
model parameters

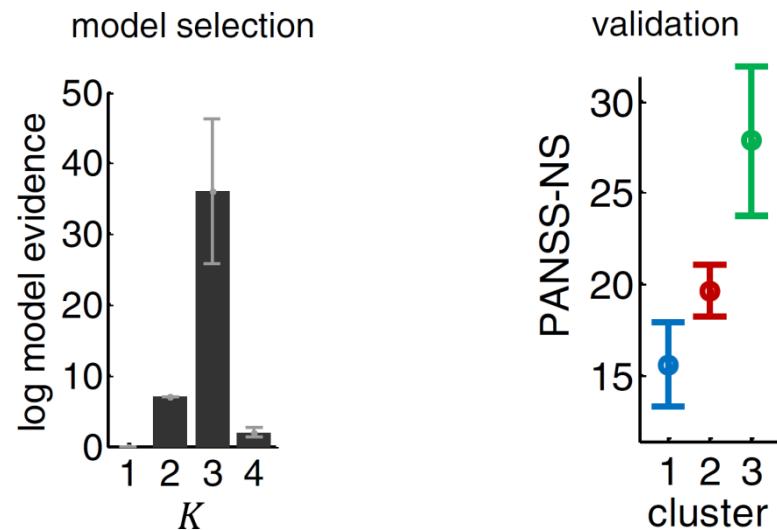
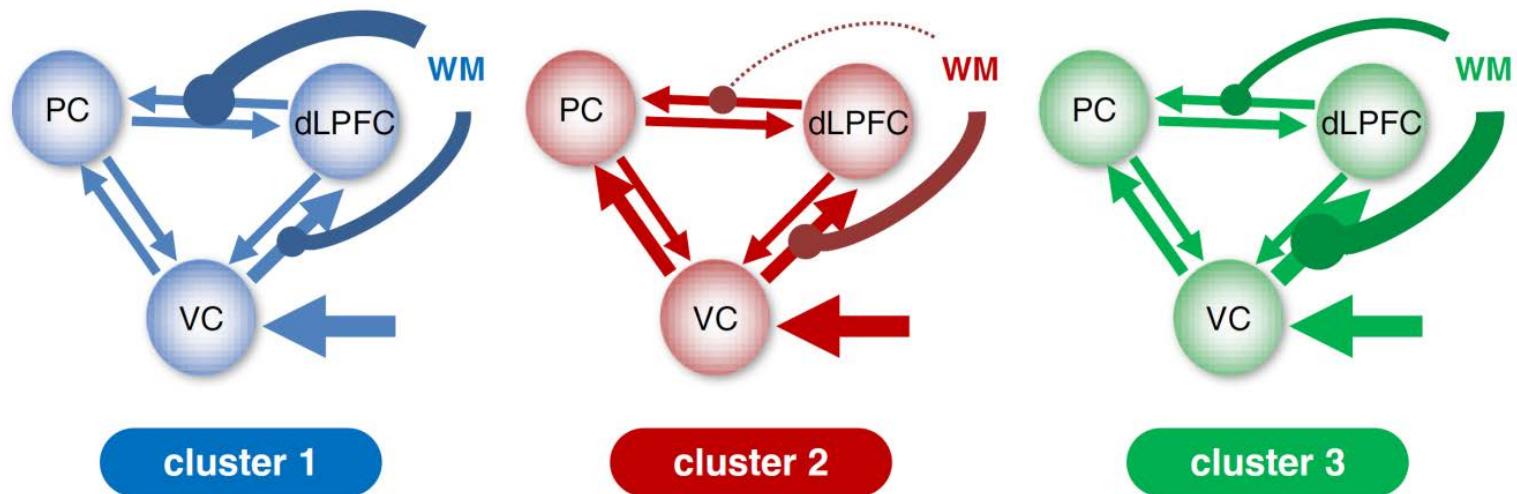


c unsupervised learning:
variational GMM clustering (using effective connectivity)



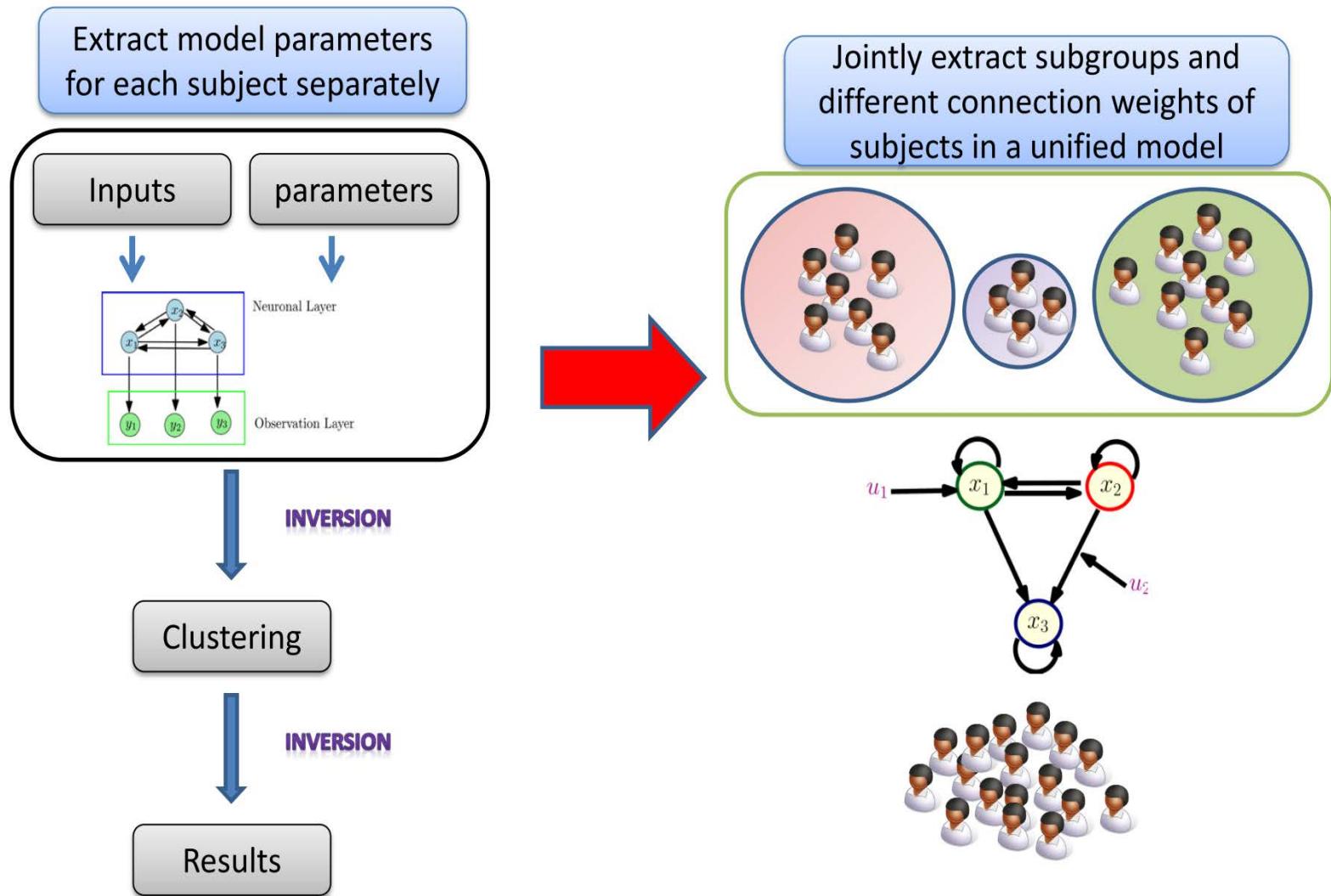
Brodersen et al 2014 Neuroimage

Results – Schizophrenia patients



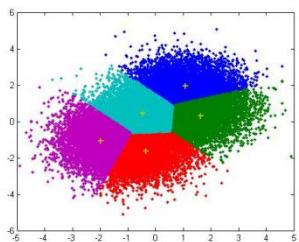
**COMING
SOON**

Unified model for identifying subgroups

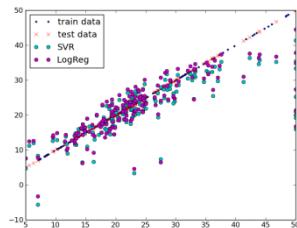


Summary

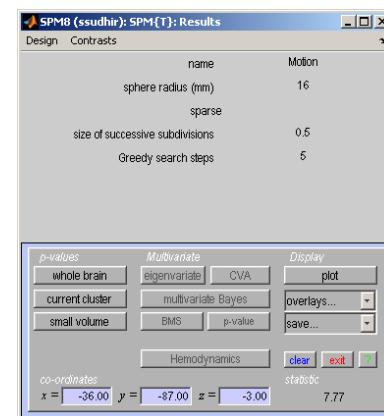
Modelling Principles



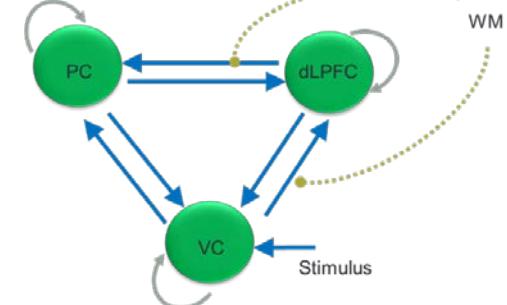
Learning from Data



Multivariate Bayes in SPM

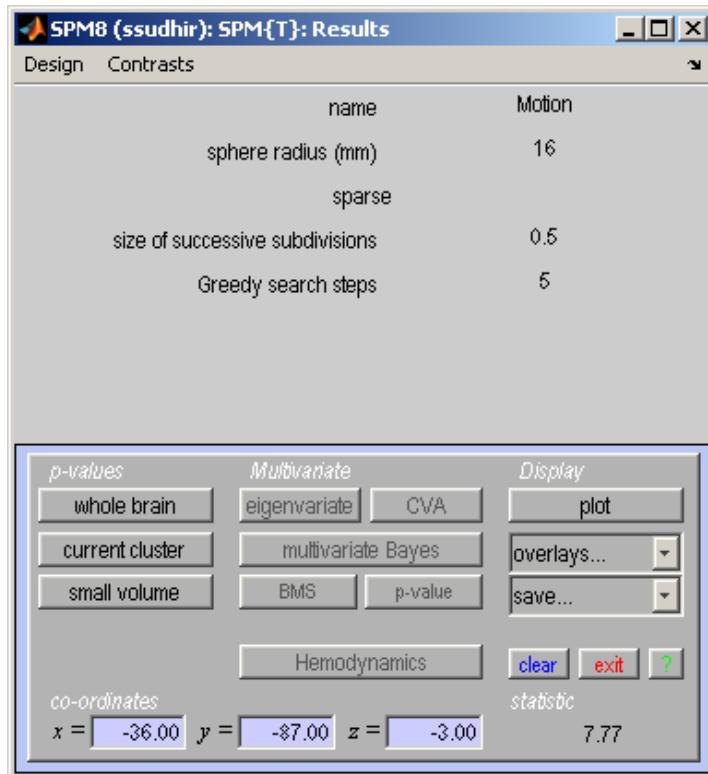


Generative Embedding

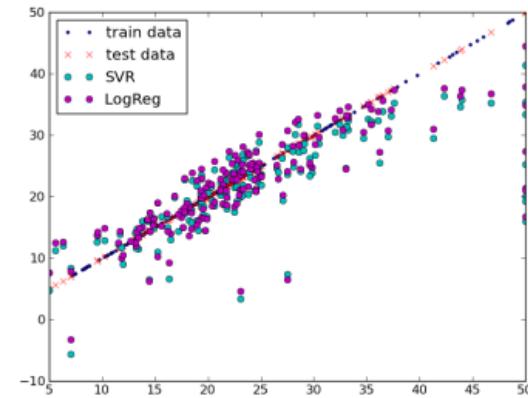


Practicals

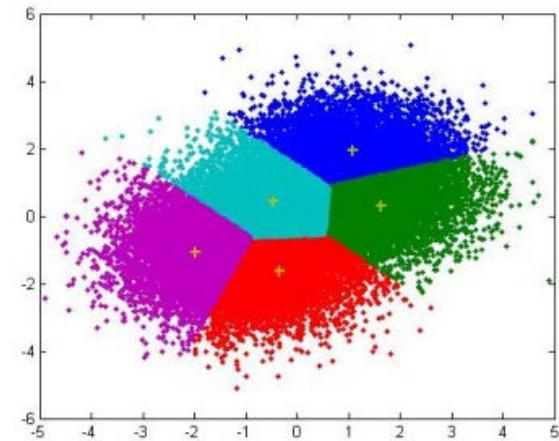
- Multivariate Bayes in SPM



- Classification



- Clustering



Thank you