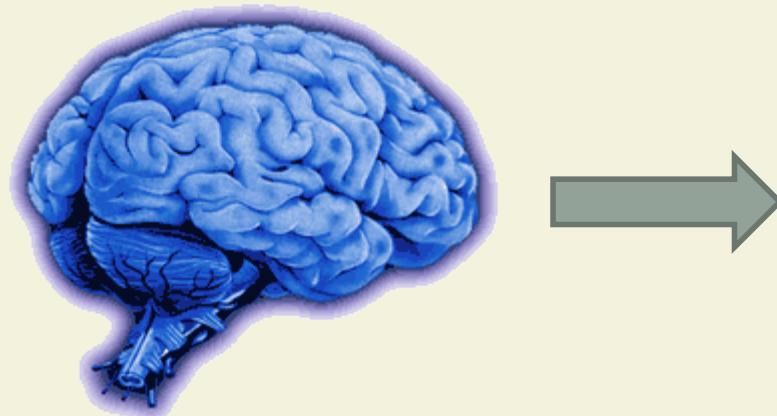


MULTIVARIATE ANALYSES WITH fMRI DATA

Sudhir Shankar Raman

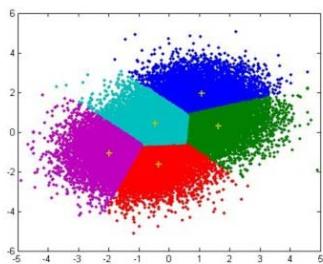
Translational Neuromodeling Unit (TNU)
Institute for Biomedical Engineering
University of Zurich & ETH Zurich



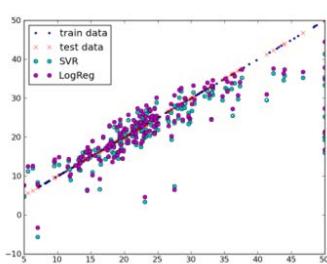
Motivation



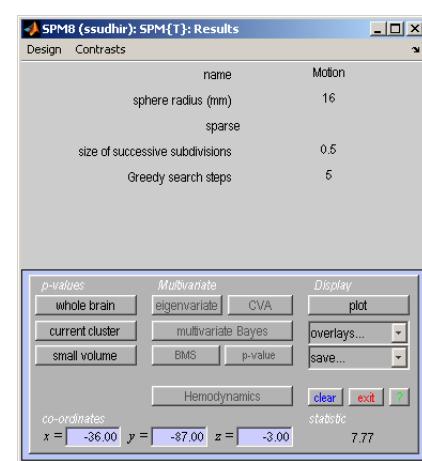
Modelling Concepts



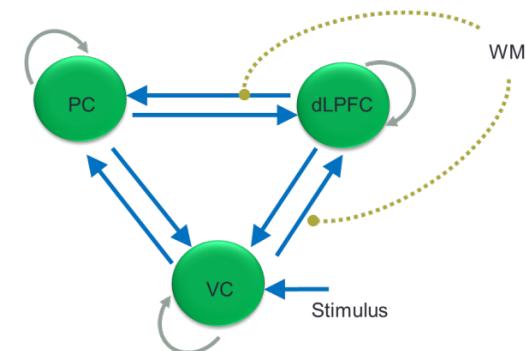
Learning From Data



Multivariate Bayes in SPM

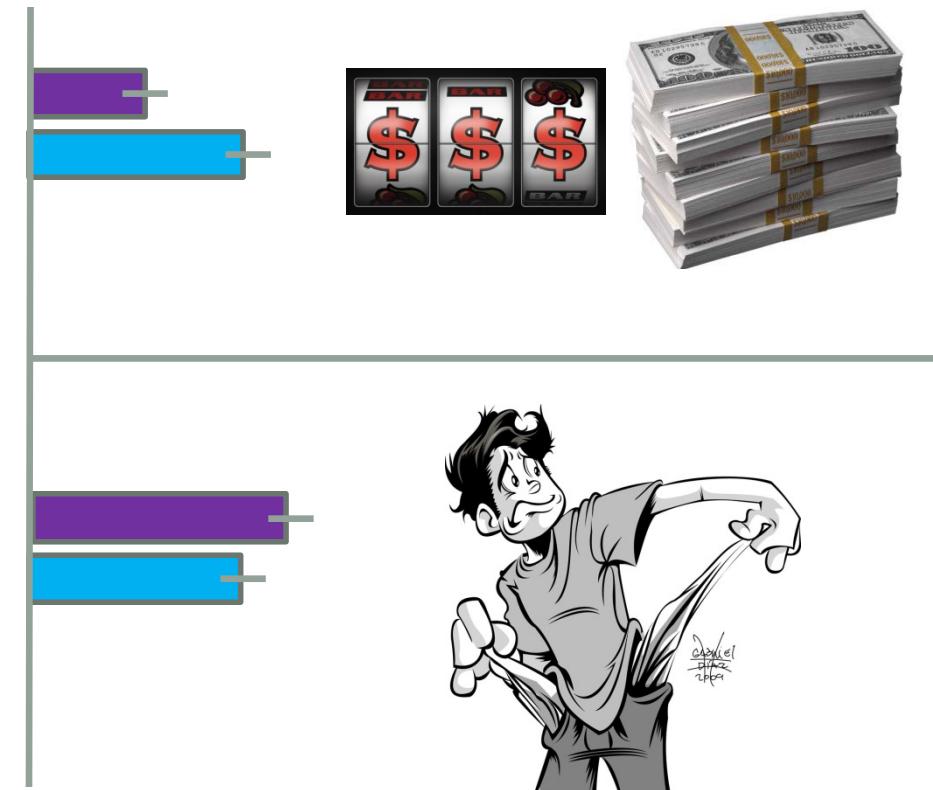
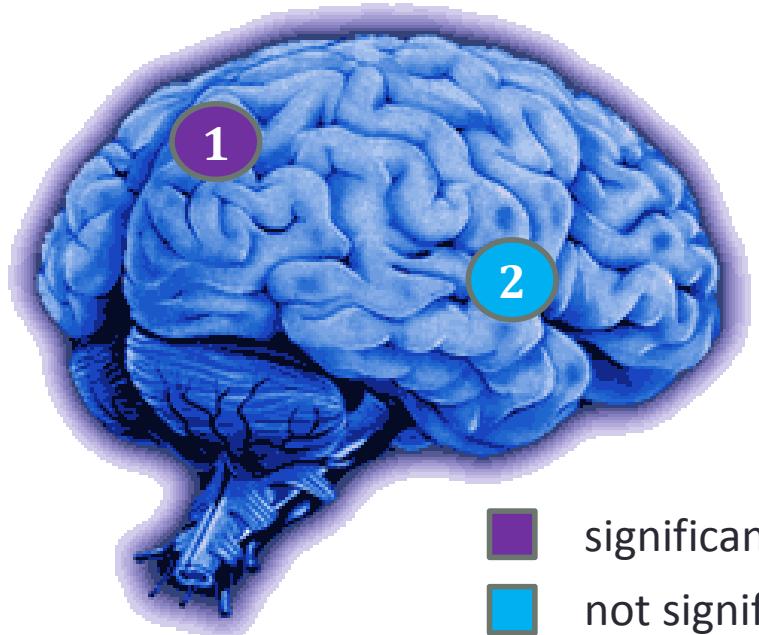


Generative Embedding

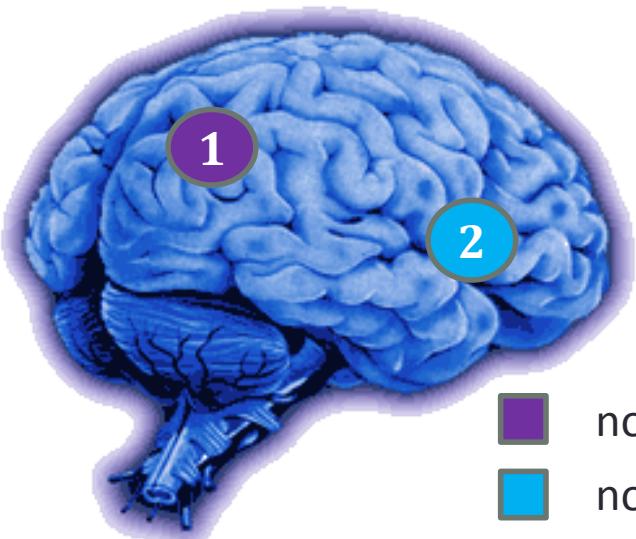


Motivation

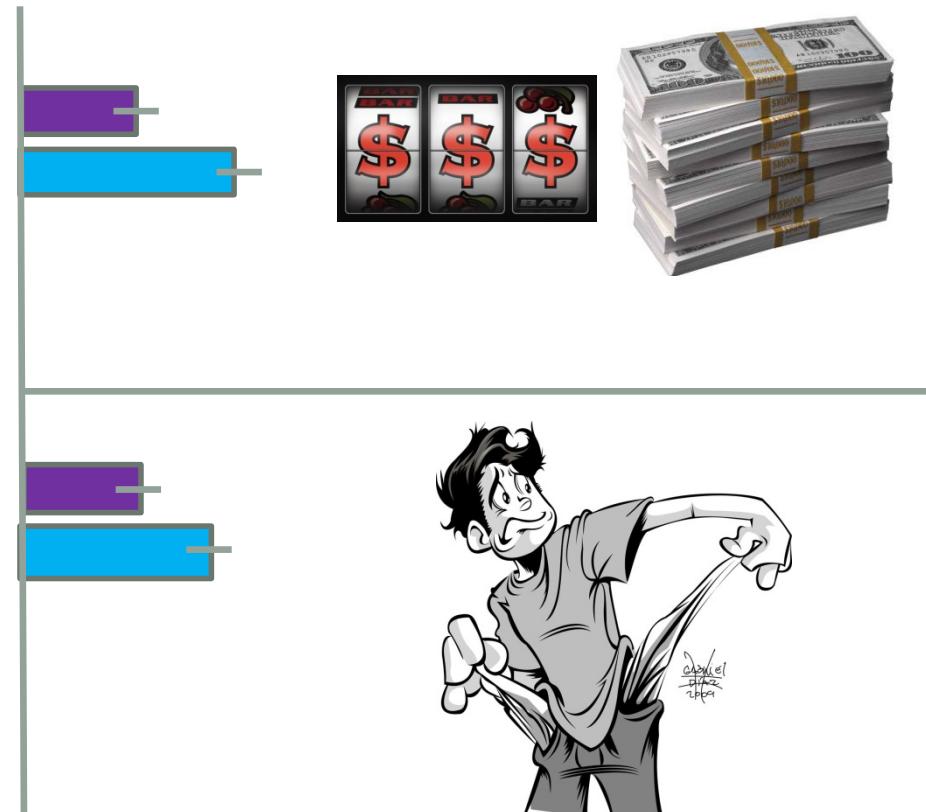
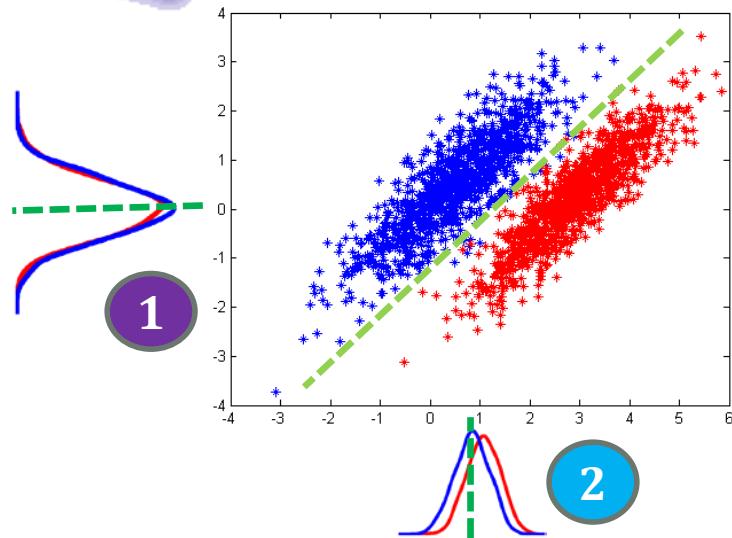
- Local activations – Univariate approach

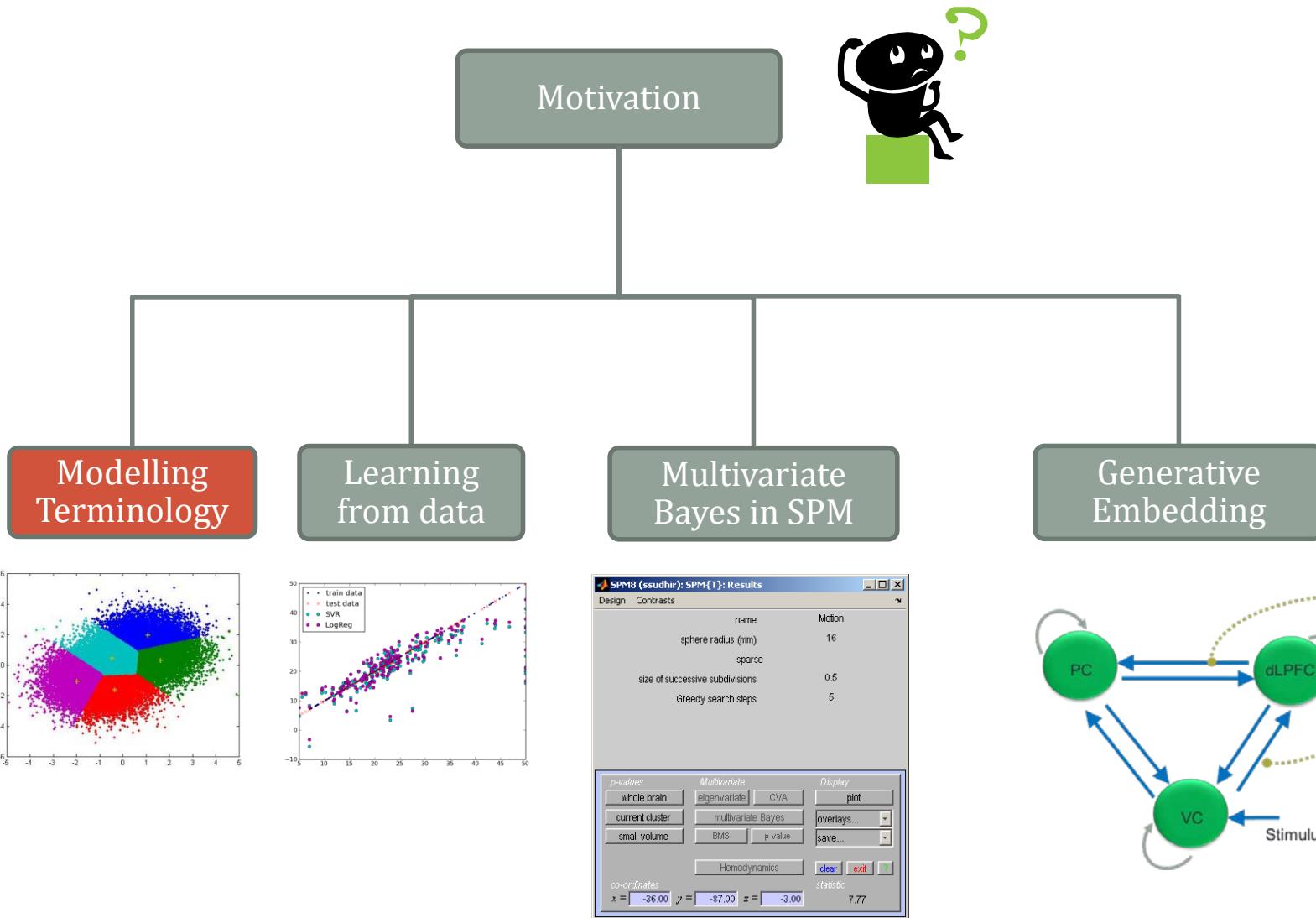


Univariate to Multivariate

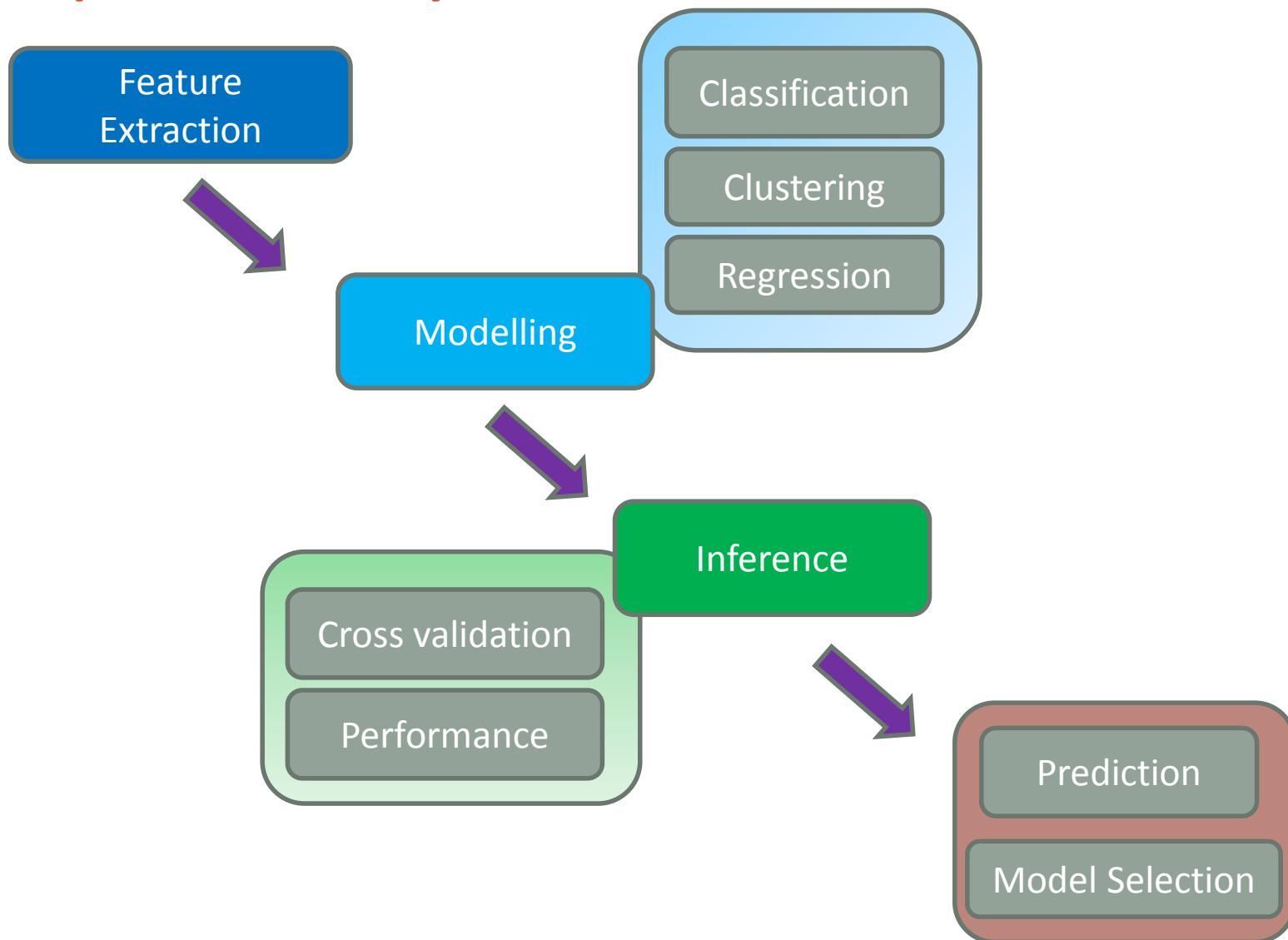


not significant
not significant





Steps for analysis



Feature Space

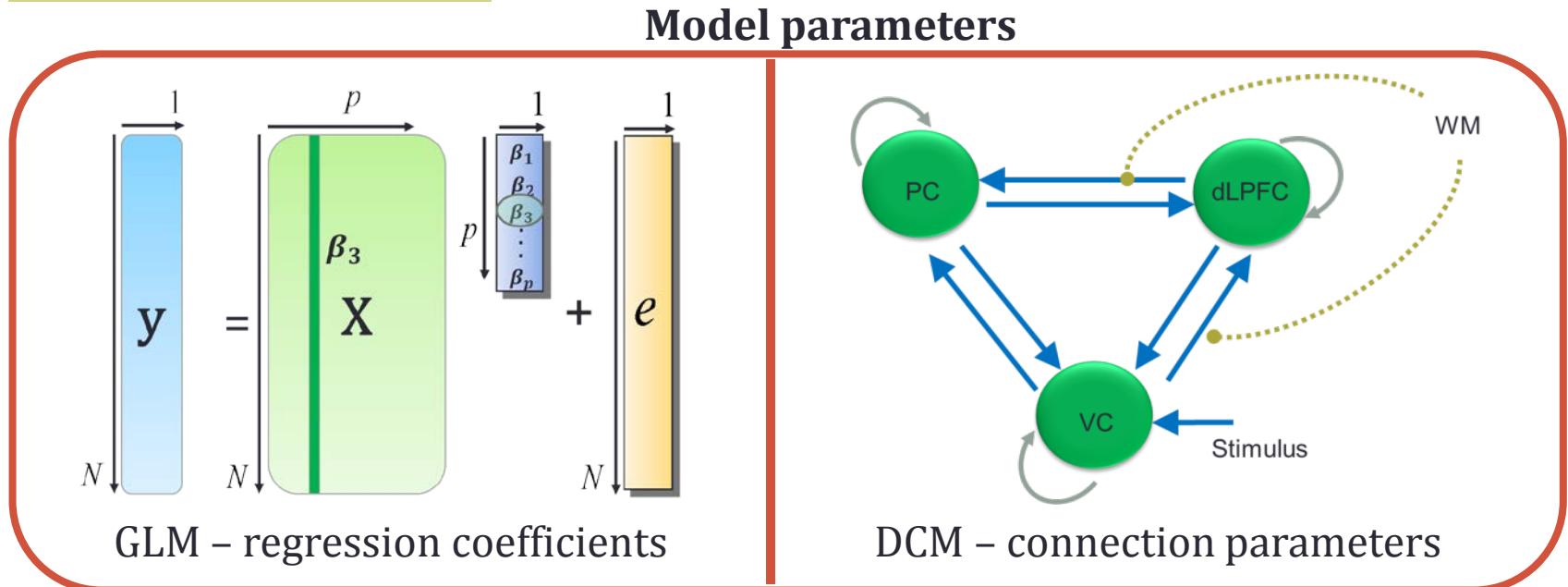
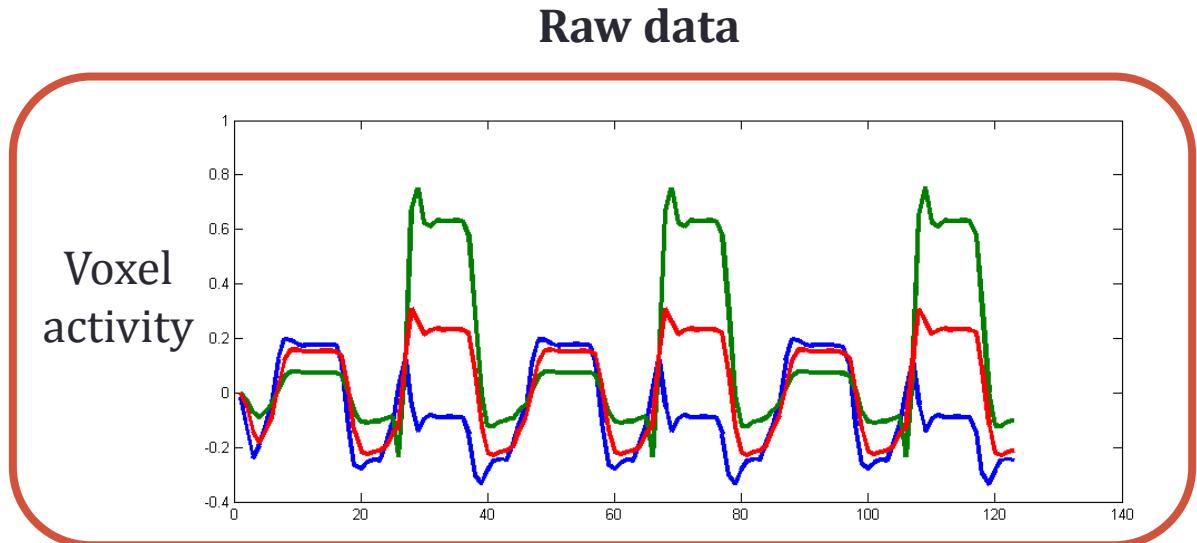
Features

	F_1	F_2	.	.	.	F_P
Data Points	S_1	1	0.5			
	S_2	0	5.7			
	.	1	4			
	.	1	5.3			
	S_N	1	6.6			

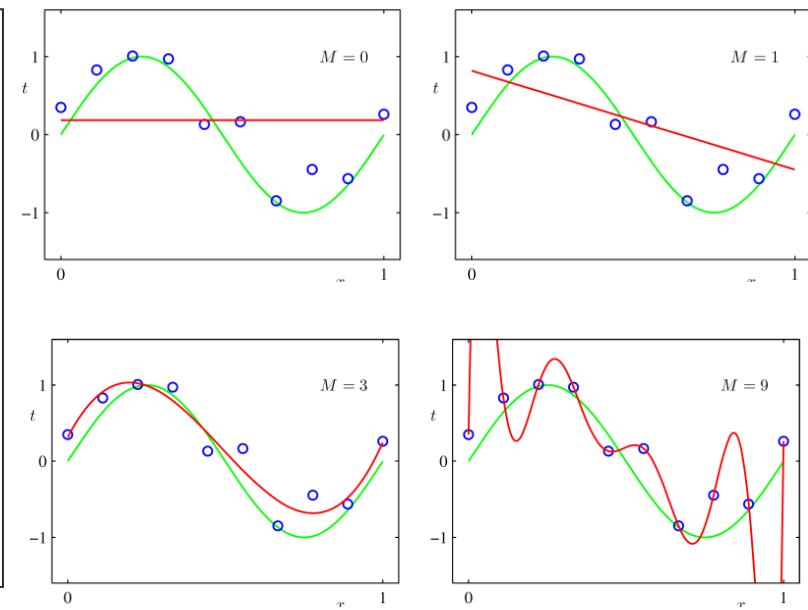
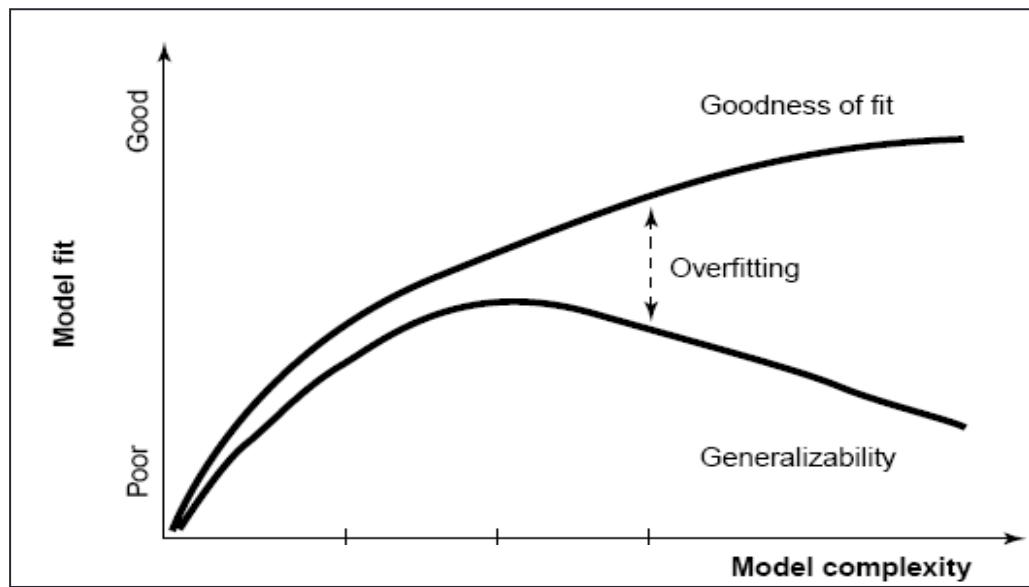
- Discrete
- Continuous

Feature Space Examples

	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



Predictive Density

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

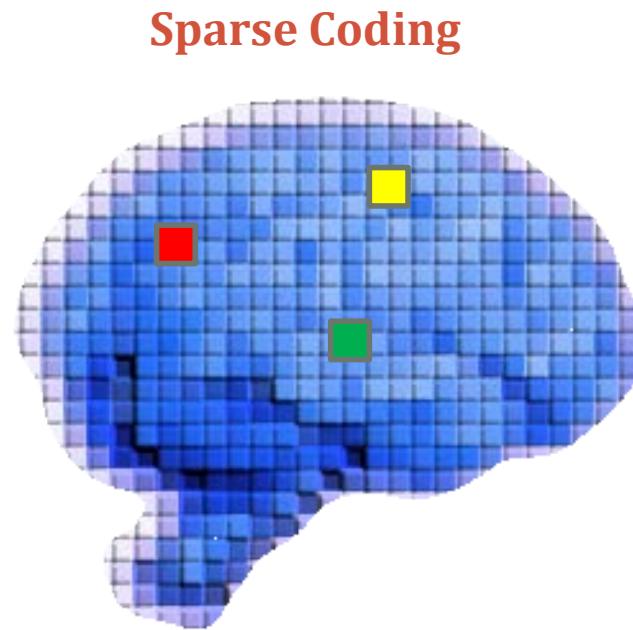


Healthy Brain Severe AD

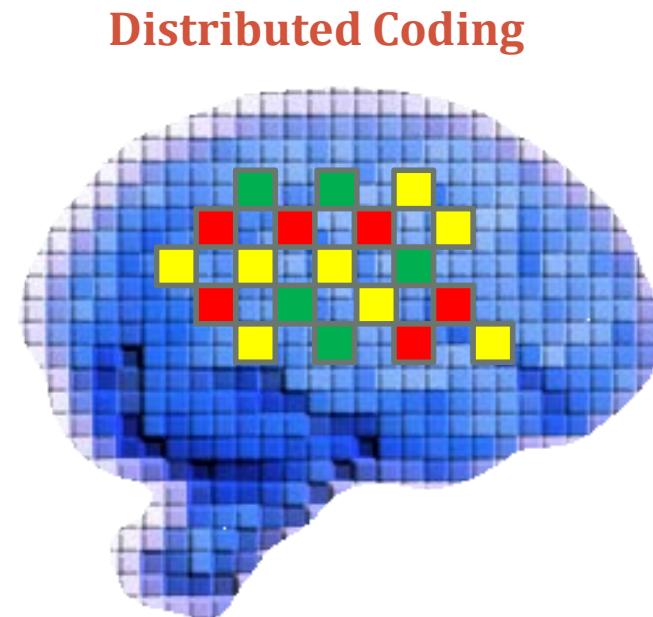


Modelling goals

- Model Selection

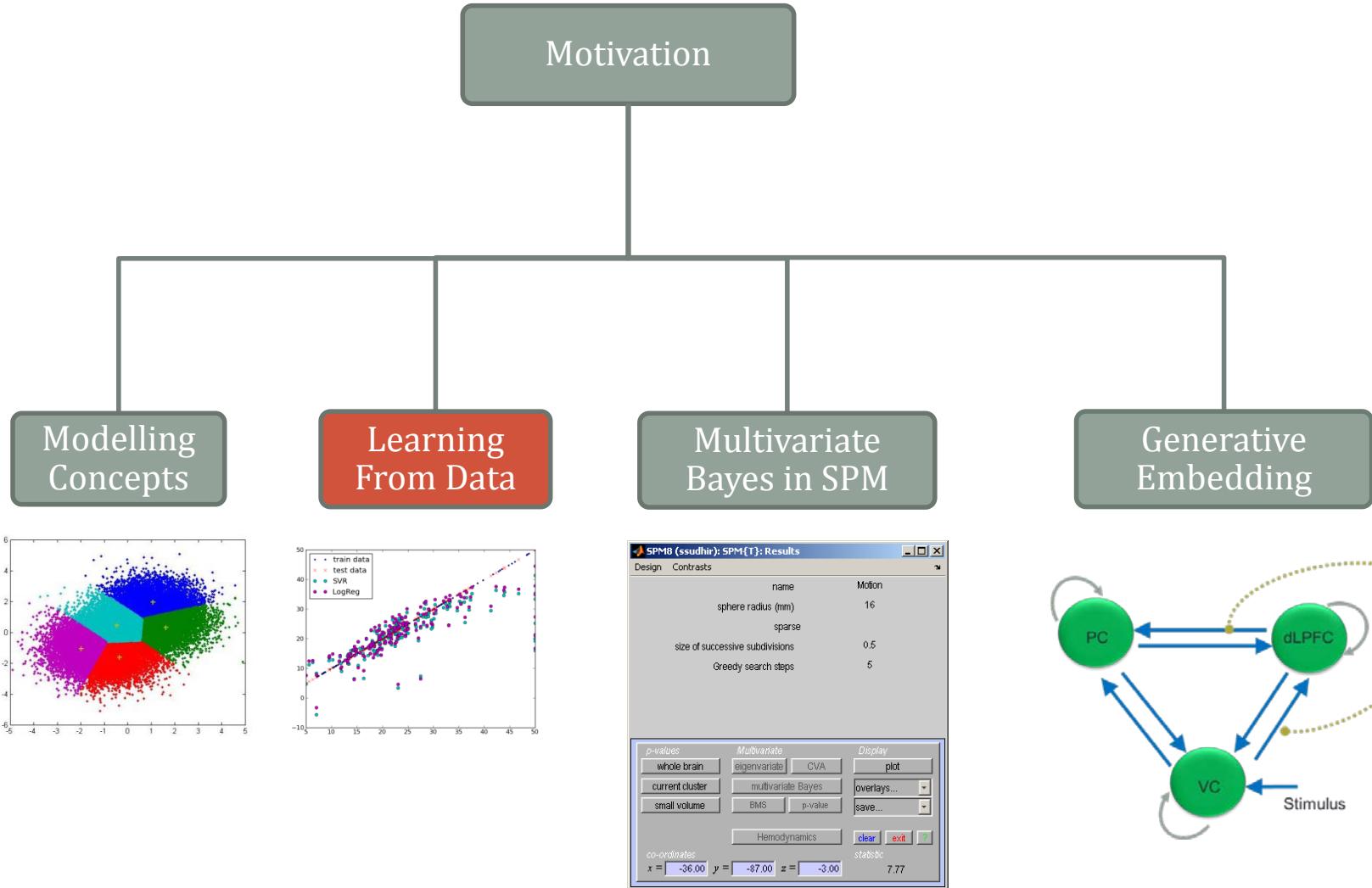


VS



Model Evidence

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



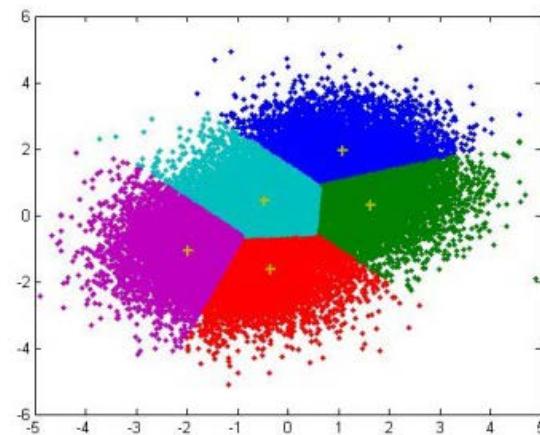
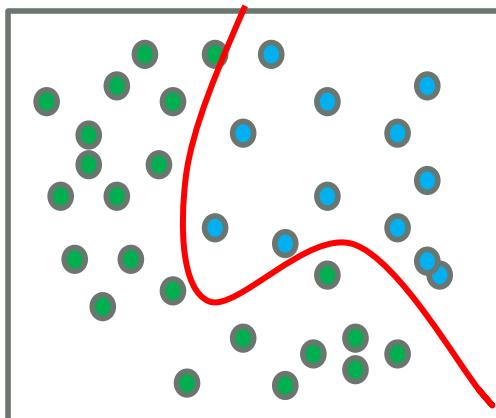
Learning from Data

Supervised Learning

Unsupervised Learning

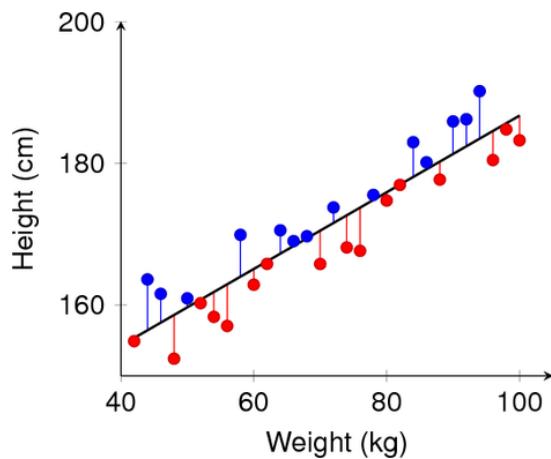
Reinforcement Learning

Semi-supervised Learning



Supervised Learning

Regression



Independent variables

X



Function - f

dependent variable
Y

Continuous

Classification

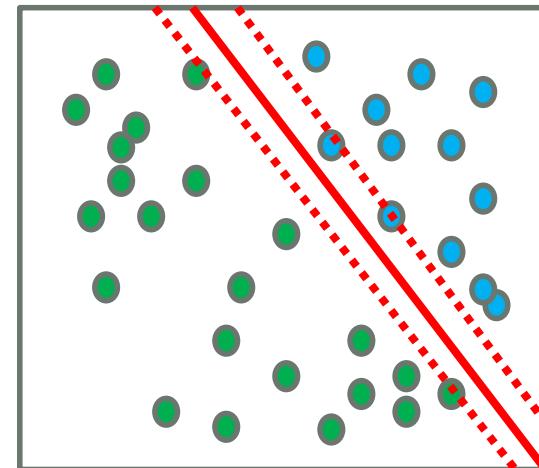
Categorical



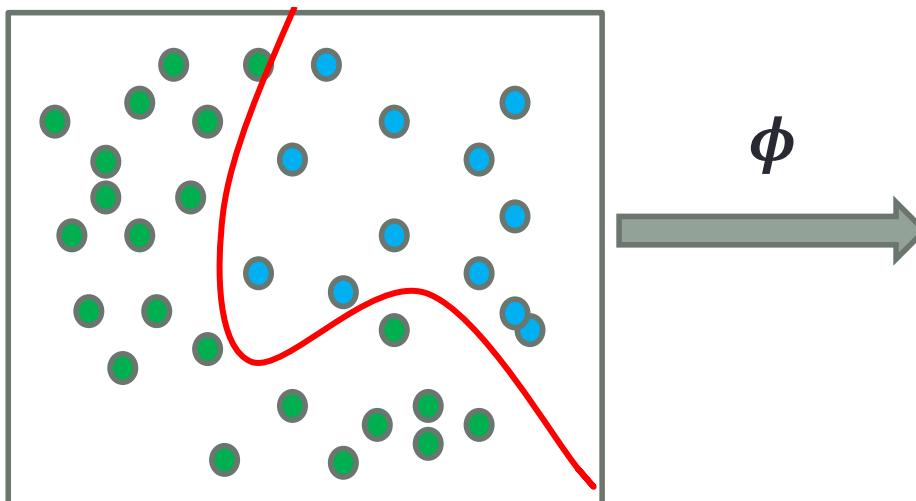
Classification



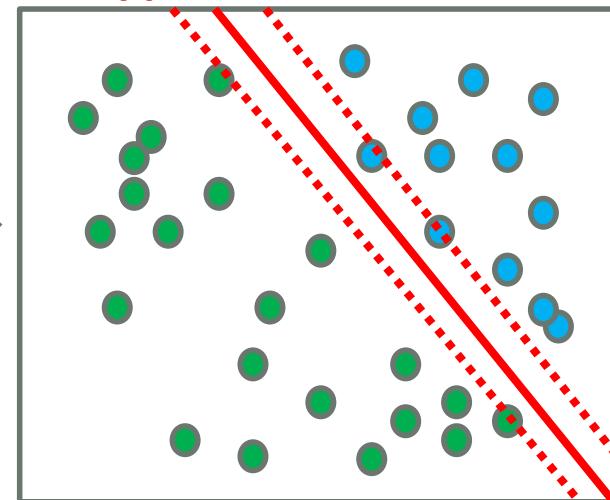
- Generative classifier
- Discriminative classifier



Kernel Methods



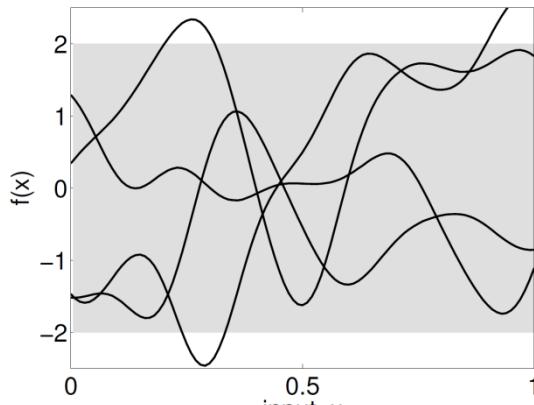
Support Vector Machines



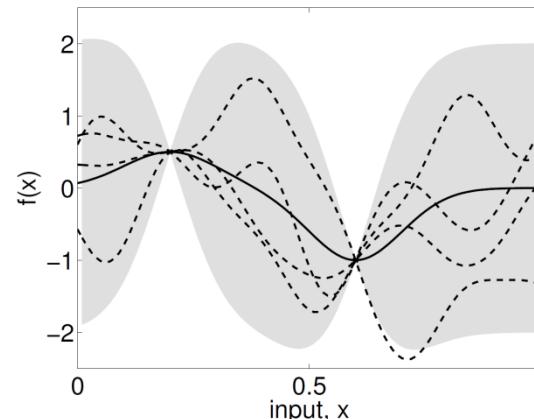
- Kernel Function – $K(x_i, x_j) = \phi(x_i) \cdot \phi(x_j)$

Other popular classifiers

- Gaussian Processes



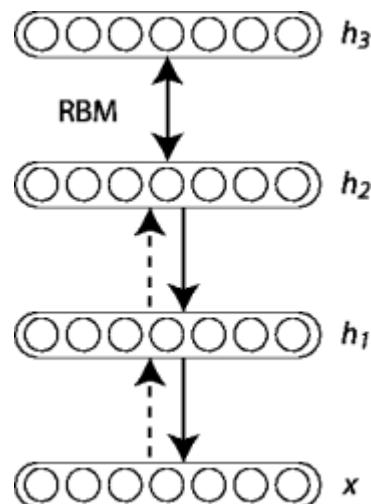
(a), prior



(b), posterior

C. E. Rasmussen & C. K. I. Williams, Gaussian Processes for Machine Learning, the MIT Press, 2006,

- Deep Belief networks



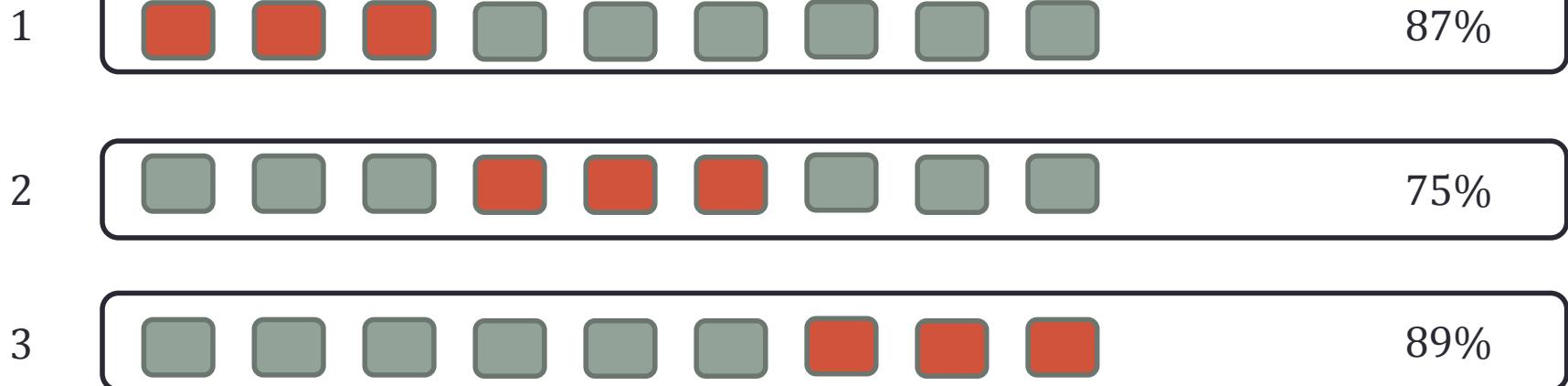
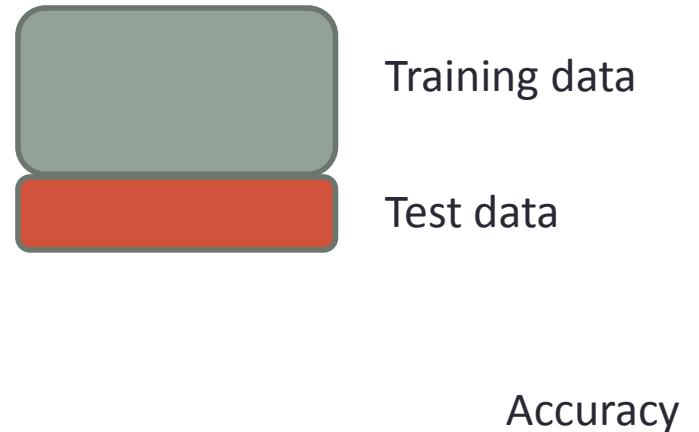
$$P(x, h^1, \dots, h^\ell) = \left(\prod_{k=0}^{\ell-2} P(h^k | h^{k+1}) \right) P(h^{\ell-1}, h^\ell)$$

<http://deeplearning.net/tutorial/DBN.html>

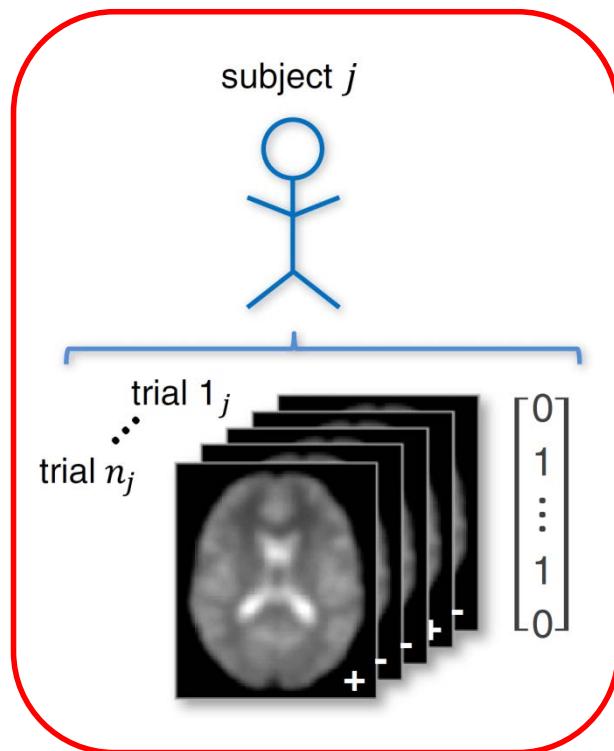
G.E. Hinton, S. Osindero, and Y. Teh, "A fast learning algorithm for deep belief nets", Neural Computation, vol 18, 2006

K-fold Cross Validation

- Model Selection
- Performance evaluation
 - Balanced Accuracy
 - F1 Score



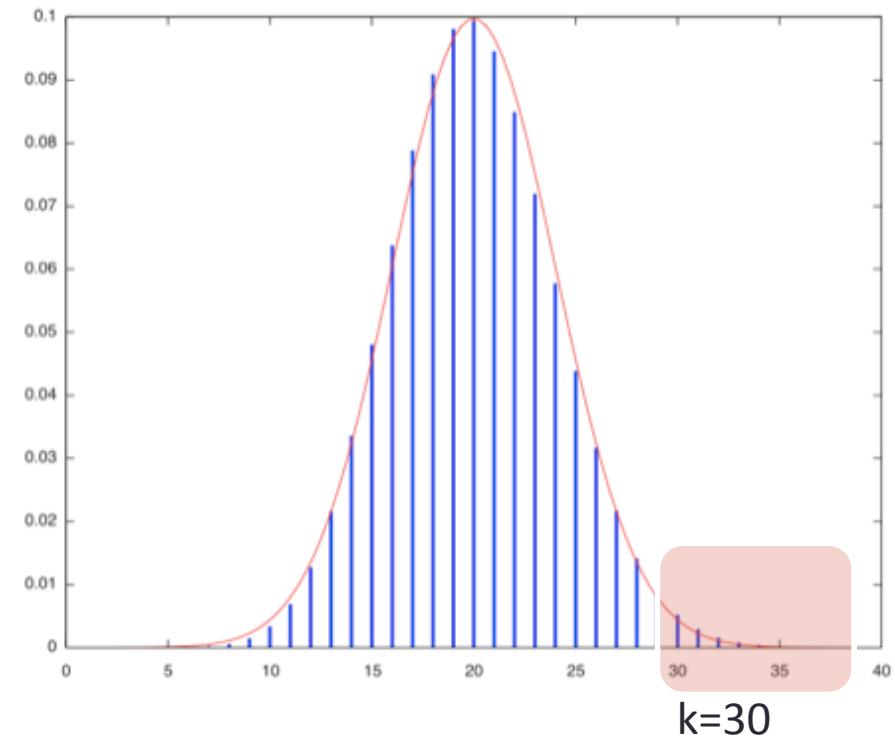
Performance – Single Subject



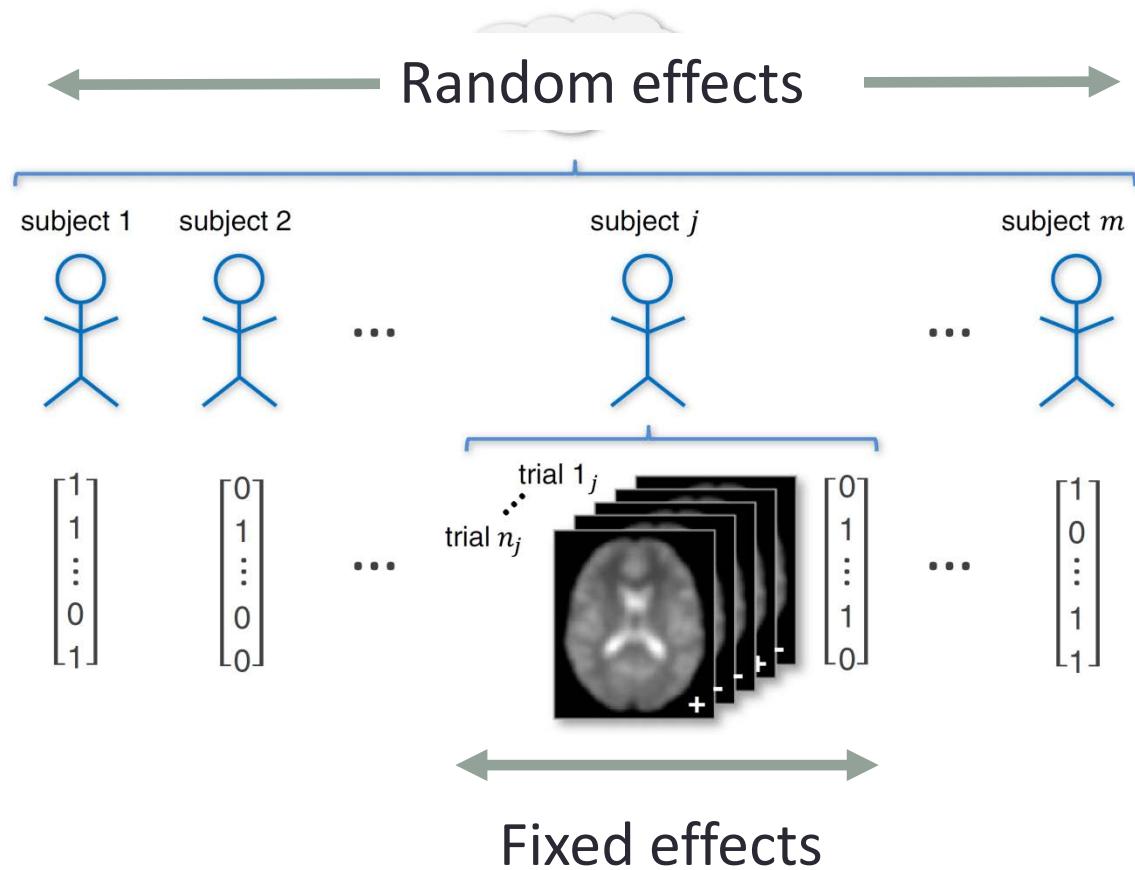
Brodersen et al. 2013, *NeuroImage*

Binomial Test

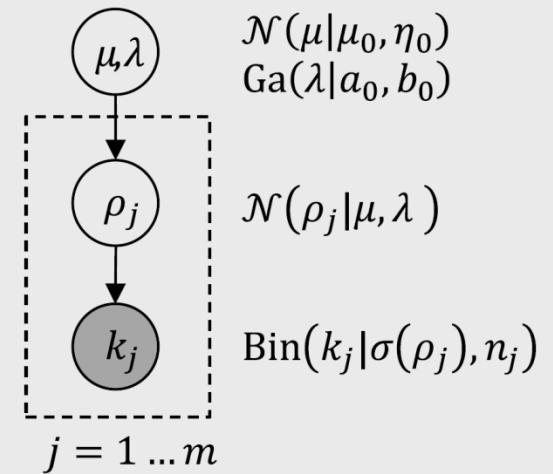
$$p = P(X \geq k | H_0) = 1 - B(k | n, \pi_0)$$



Performance – Multiple Subjects



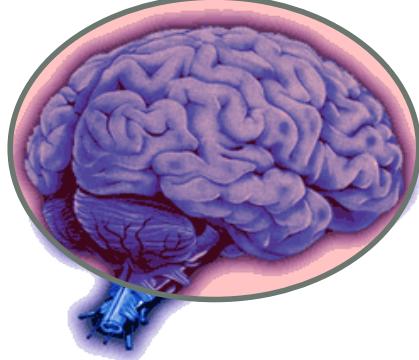
b Bayesian mixed-effects inference (univariate normal-binomial model)



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

Using classification for fMRI data

Whole brain



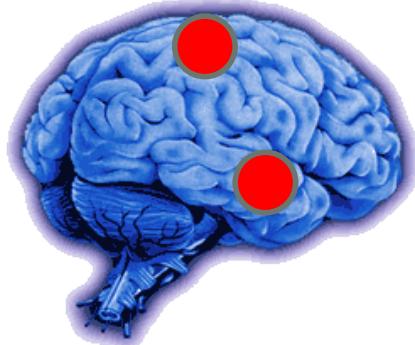
Mourao-Miranda et al. (2005) *NeuroImage*,
Marquand et al. (2010) *NeuroImage*

Searchlight classifier



Kriegeskorte et al. (2006) *PNAS*

Pattern localization

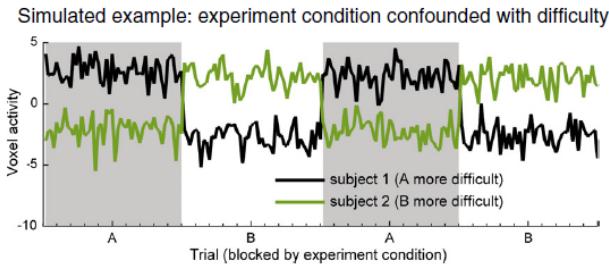


Word Category

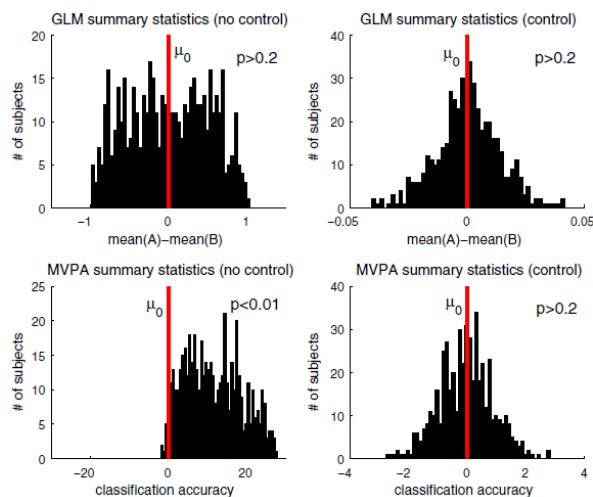
Food
Building
People

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

Confounds – GLM vs MVPA

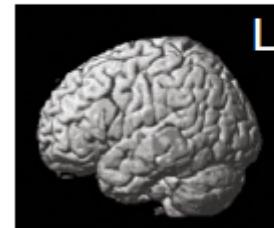


Individual-Subject Summary Statistics		
Subject	Experimental Effect (GLM)	Discrimination Success (MVPA)
Subject 1	mean(A)-mean(B) = +4.75	classification accuracy = +13.15, within-minus-across = +3.826
Subject 2	mean(A)-mean(B) = -5.56	classification accuracy = +13.44, within-minus-across = +3.848
Group 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: $t_1=94.0, p<0.01, \text{sig.}$

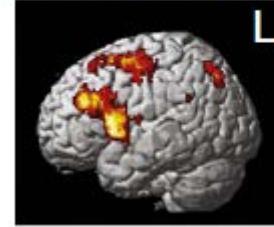


Task Rule (A vs. B)

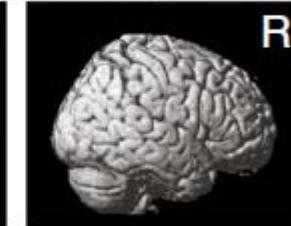
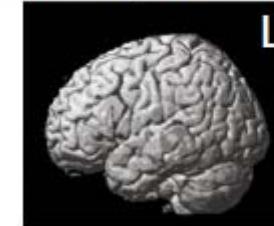
GLM



MVPA (no control)

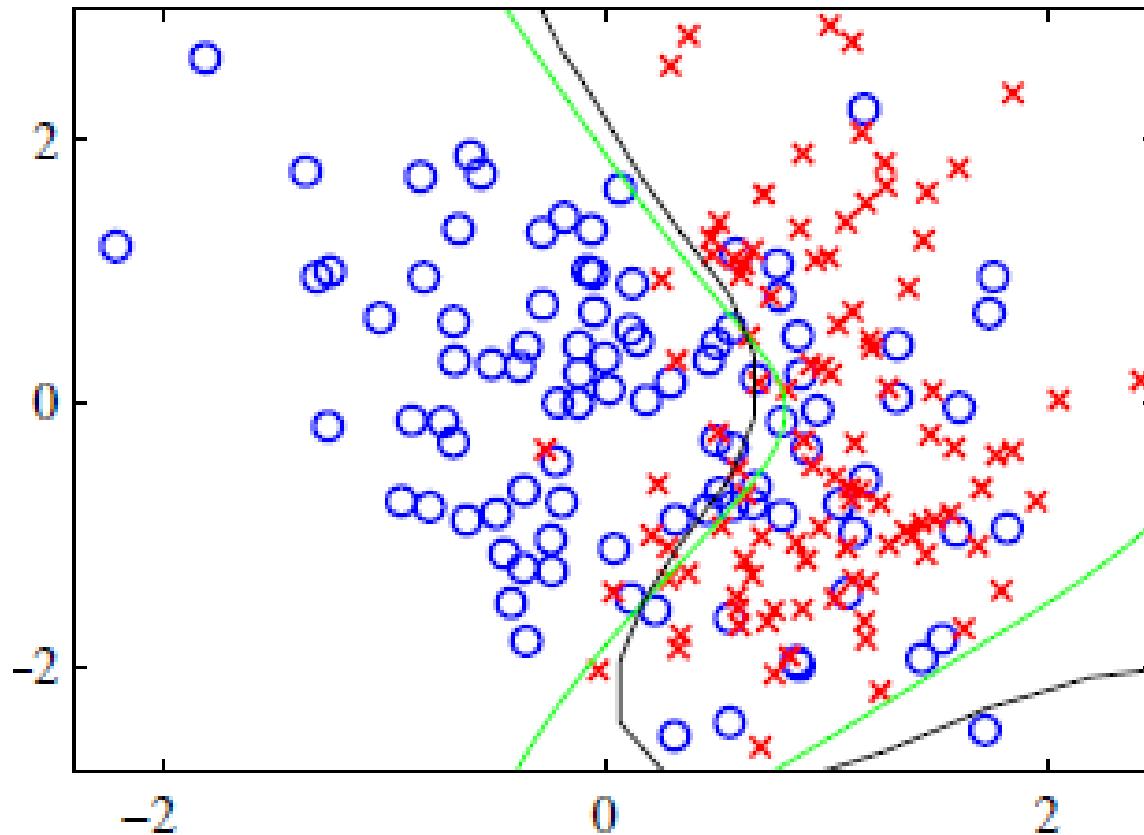


MVPA (after RT regression)

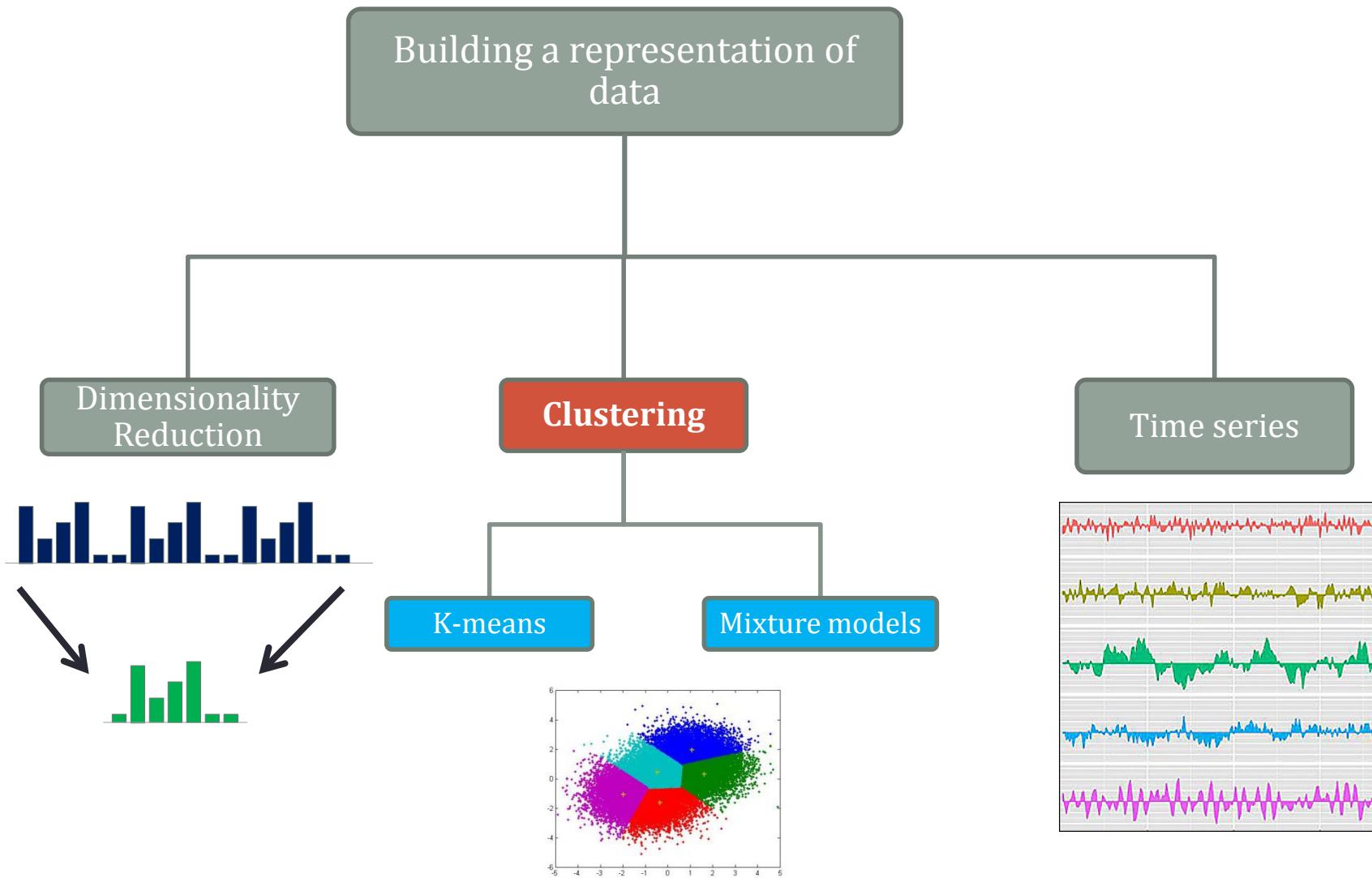


Practicals

- Classification using SVM



Unsupervised Learning

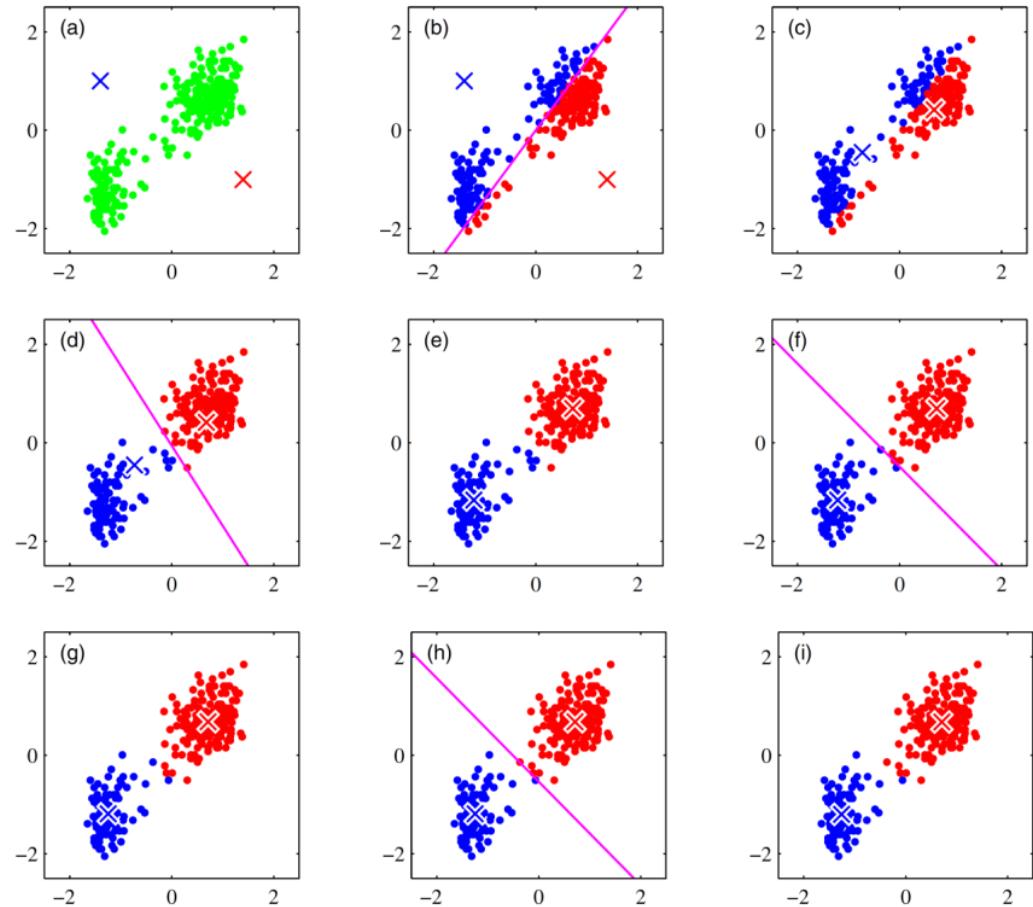


Clustering using K-means

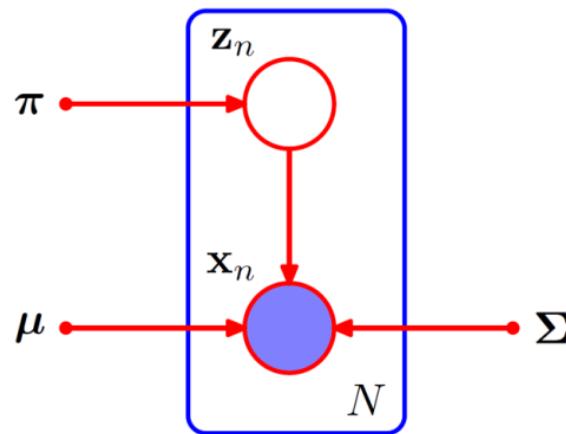
- Cost function

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

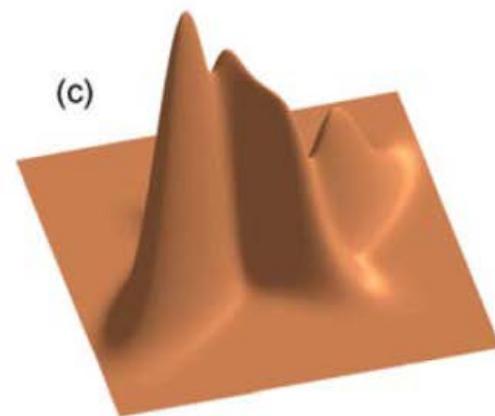
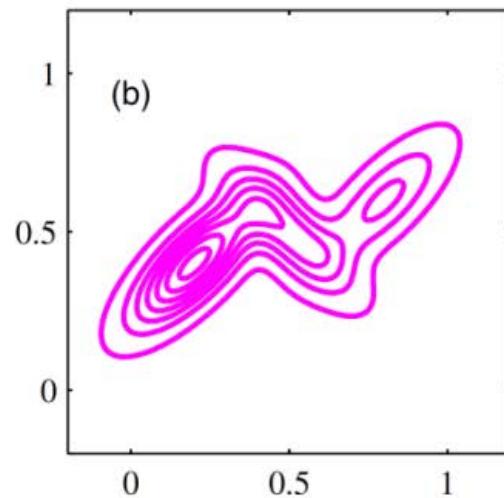
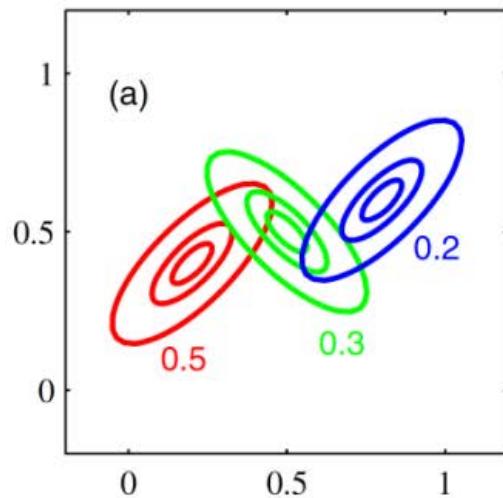
- Algorithm
 1. Initialize
 2. Estimate assignments
 3. Estimate cluster centroids
 4. Repeat 2,3 until convergence



Clustering – Mixture of Gaussians

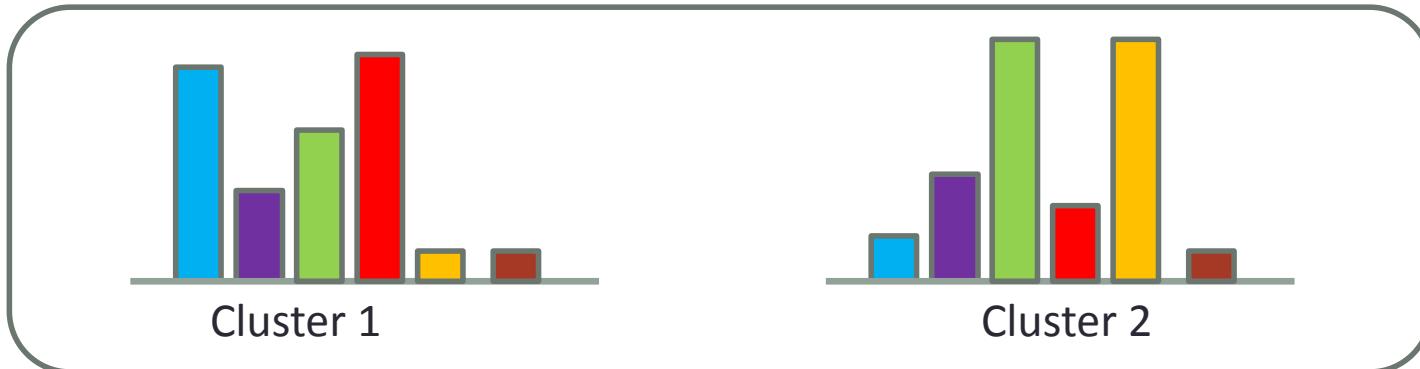


$$p(\mathbf{x}) = \sum_{k=1}^K \pi_k \mathcal{N}(\mathbf{x} | \boldsymbol{\mu}_k, \boldsymbol{\Sigma}_k)$$

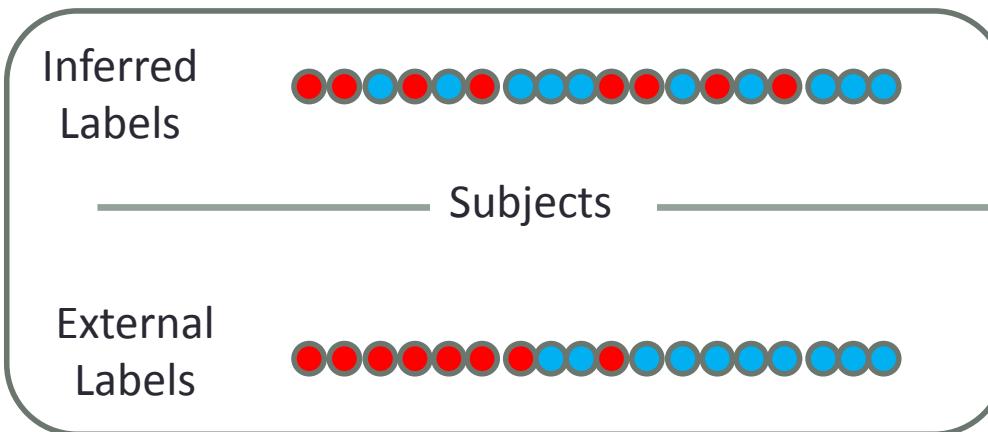


Interpretation

- Cluster parameters



- Internal Criterion – Model Evidence
- External Criterion - Purity



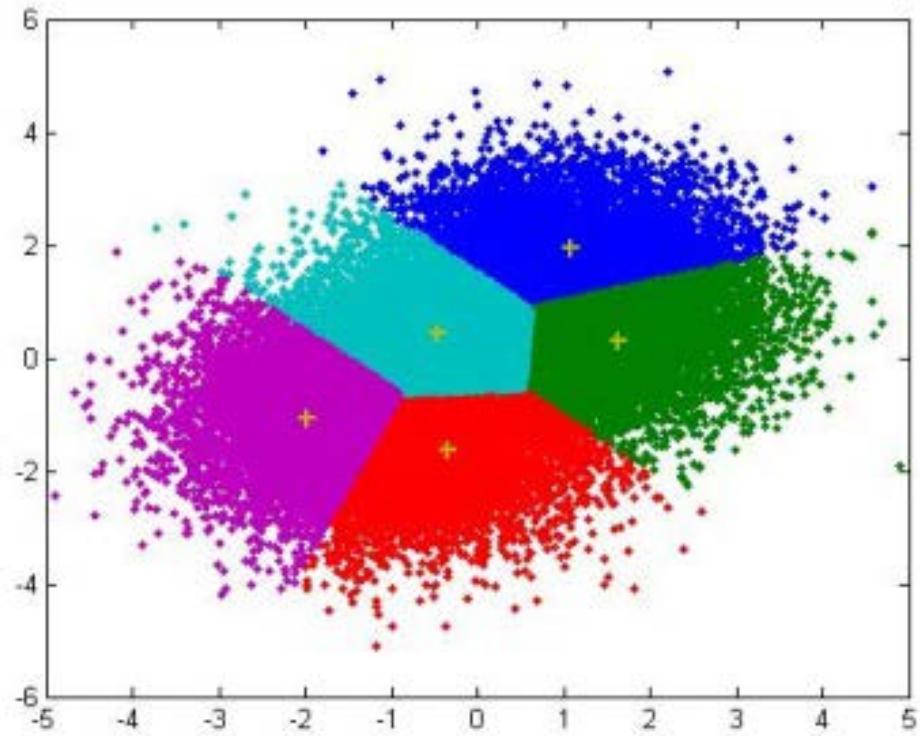
Normalized Mutual
Information (NMI)

Balanced purity

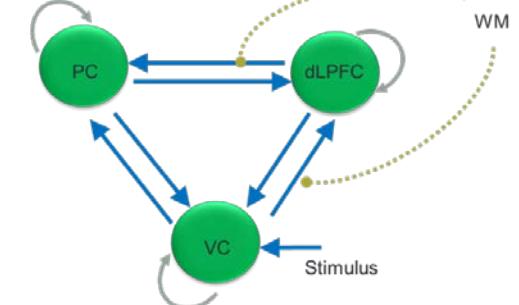
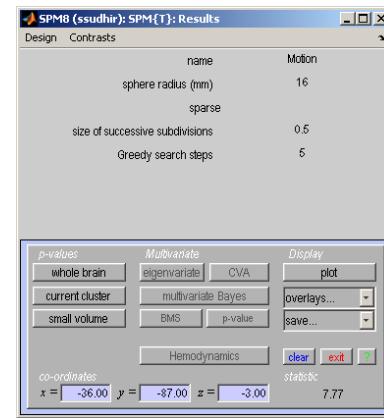
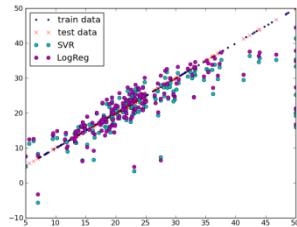
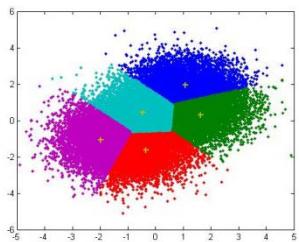
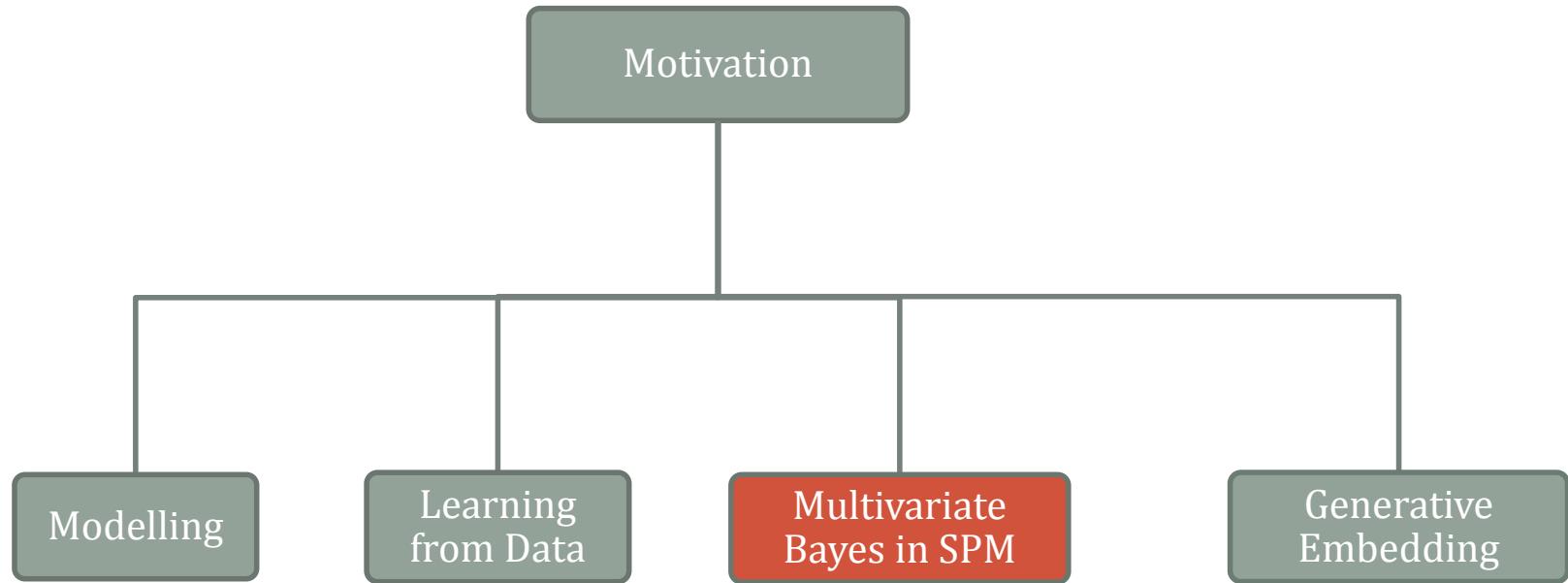
Rand Index

Practicals

- Clustering
 - K-Means
 - Finite Gaussian mixture model



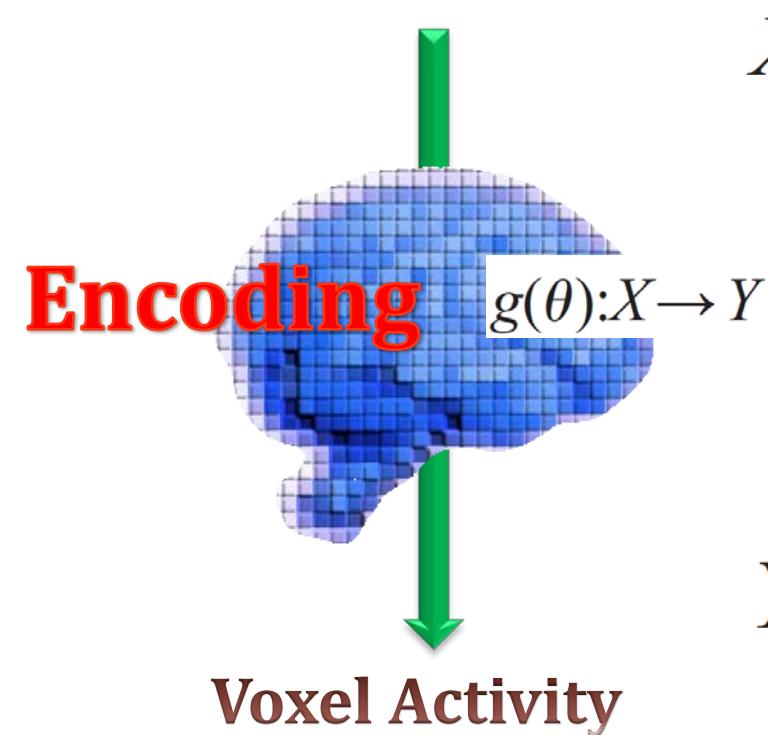
Bishop (2006) PRML



Encoding Vs Decoding Models

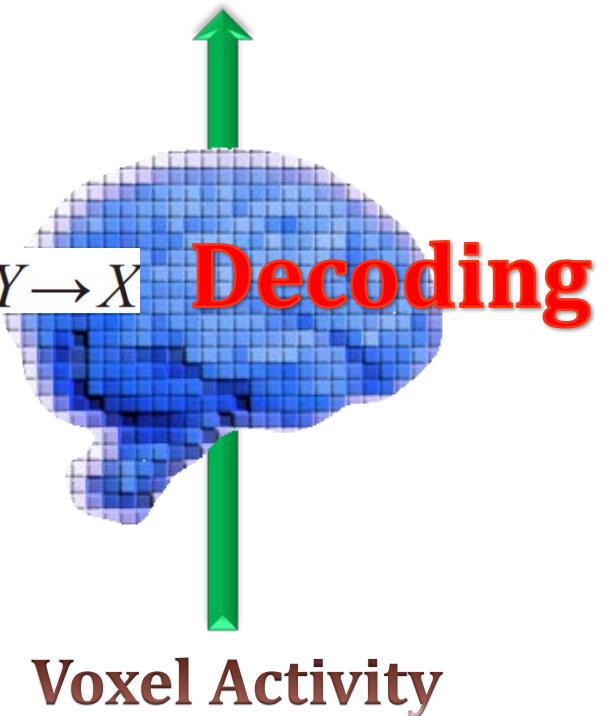
Cause/Stimulus

Consequence/Behaviour



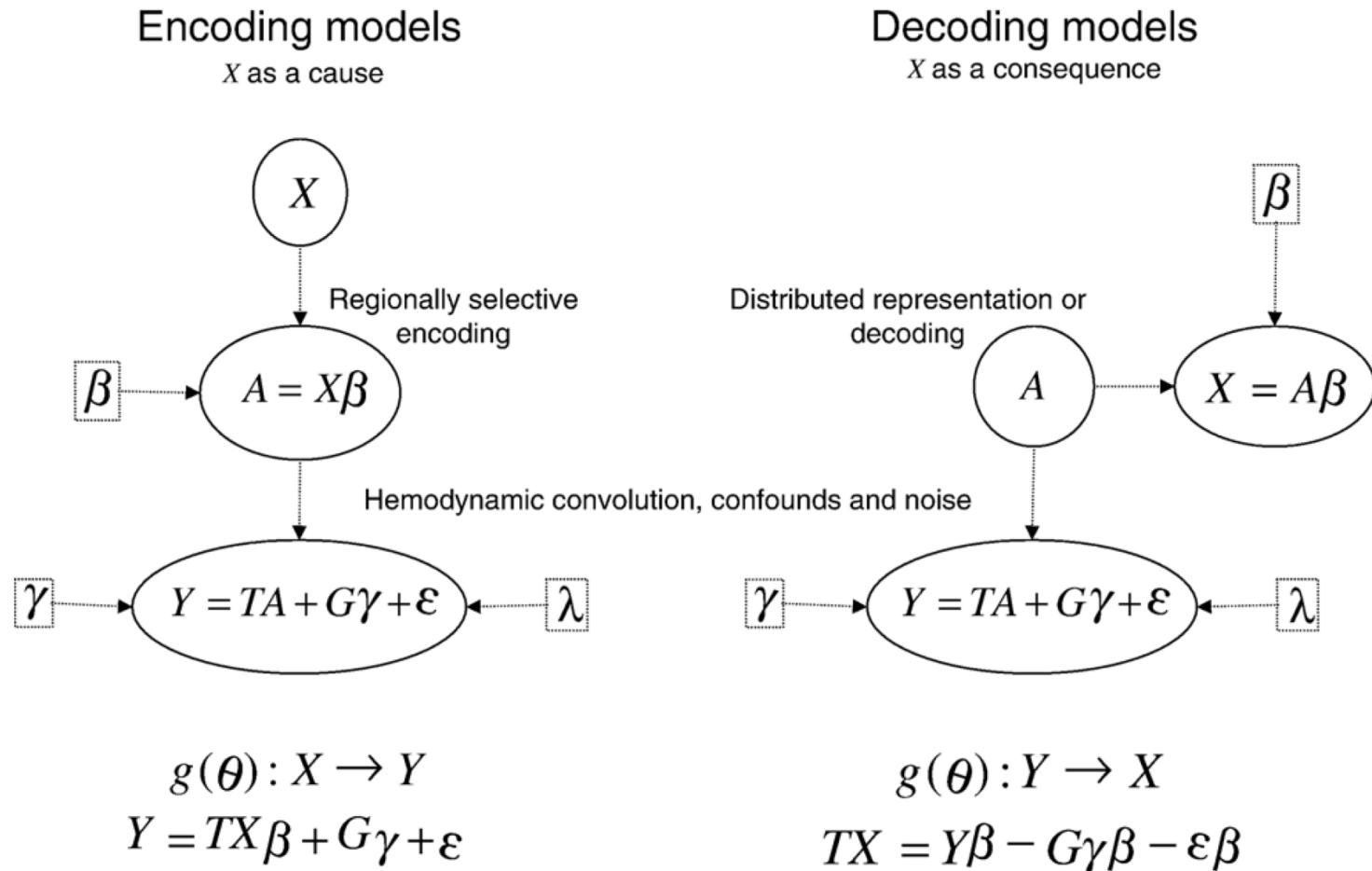
$$X \in \Re^v$$

$$h(\theta):Y \rightarrow X$$



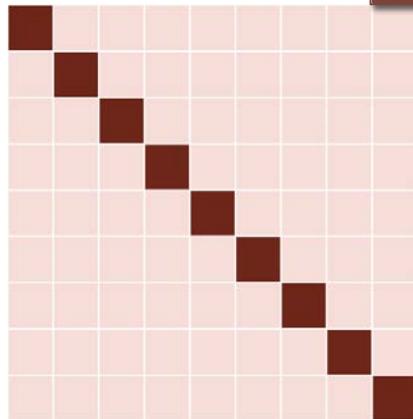
$$Y \in \Re^n$$

Encoding Vs Decoding

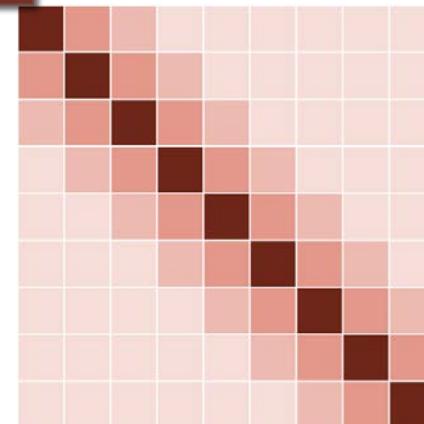


Coding hypotheses

Sparse vectors

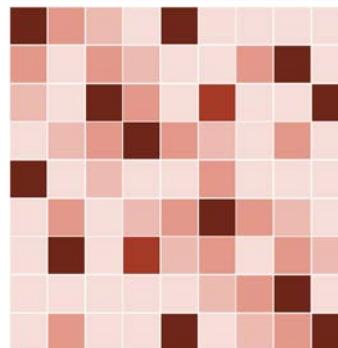


Spatial vectors



Smooth vectors

Distributed vectors



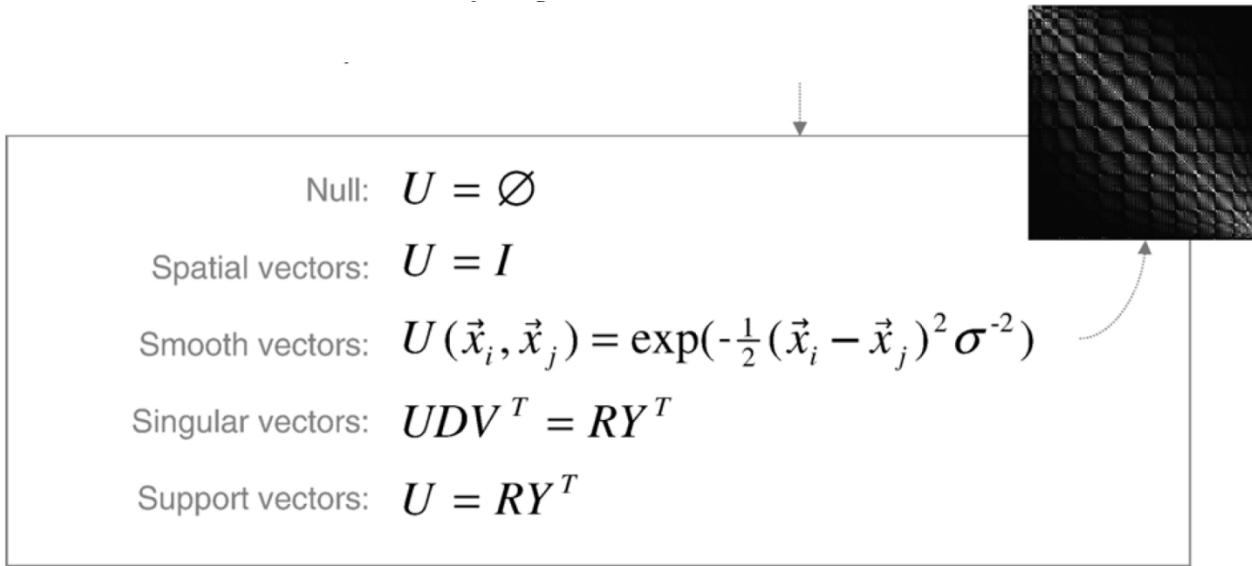
Singular vectors
of data

$$UDV^T = RY^T$$

Support vectors

$$U = RY^T$$

Coding hypotheses



$$WX = RY\beta + \varsigma$$

$$\beta = U\eta$$

$$\text{cov}(\varsigma) = \Sigma^\varsigma(\lambda) = \exp(\lambda^\varsigma)RVR^T$$

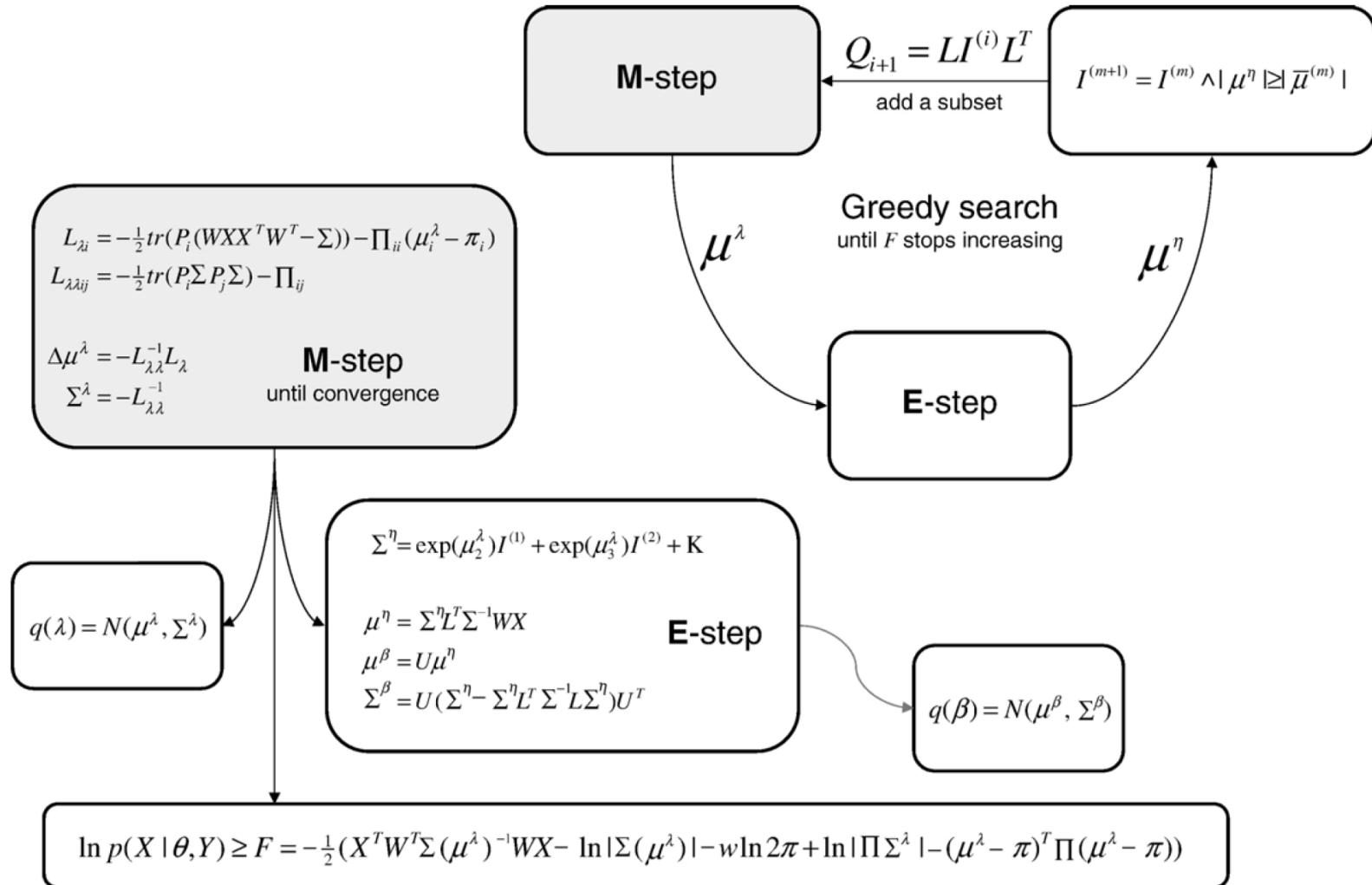
$$\text{cov}(\eta) = \Sigma^\eta(\lambda) = \exp(\lambda_1^\eta)I^{(1)} + \dots + \exp(\lambda_m^\eta)I^{(m)}$$

$$p(\beta) = N(0, U\Sigma^\eta U^T)$$

$$W = RT$$

$$R = \text{orth}(I - GG^-)^T$$

VB Inference

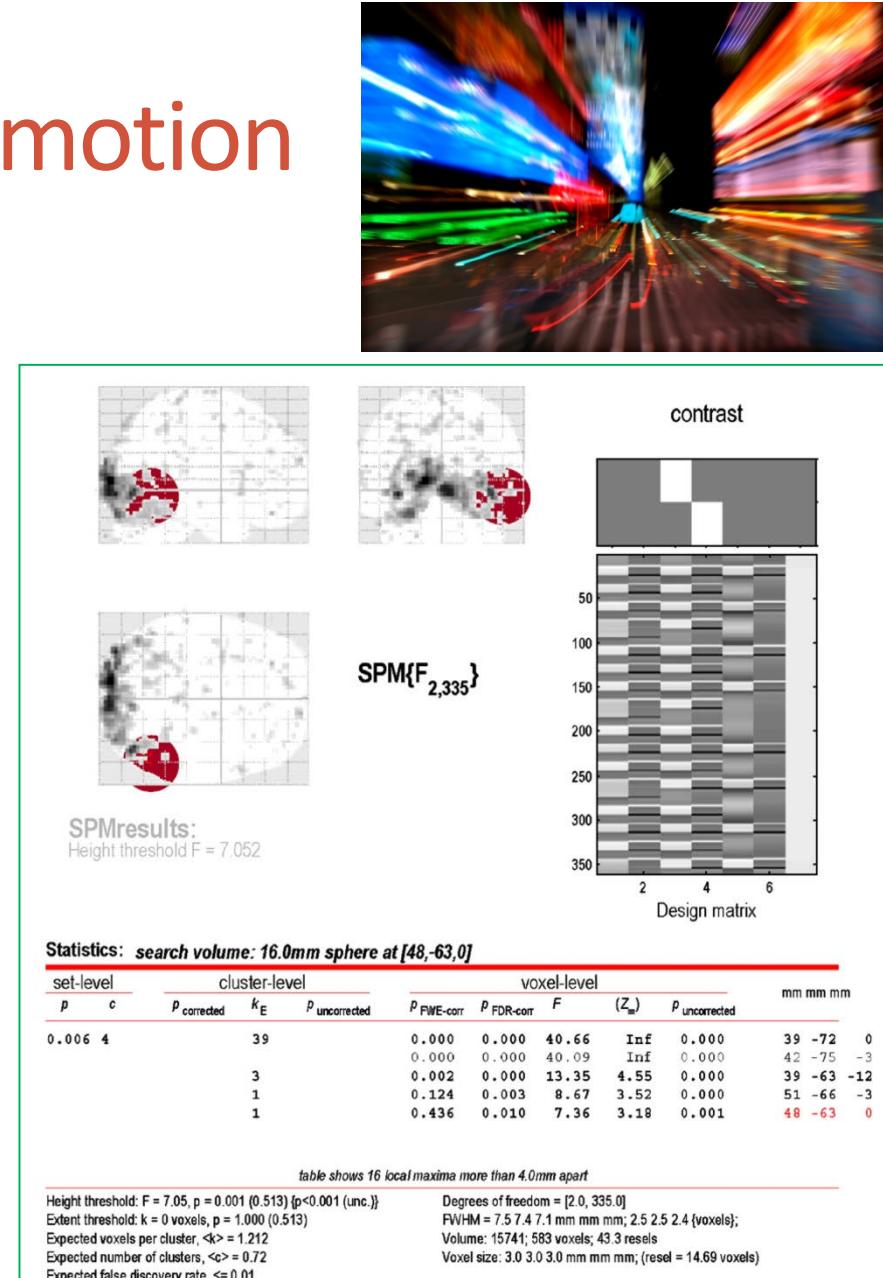
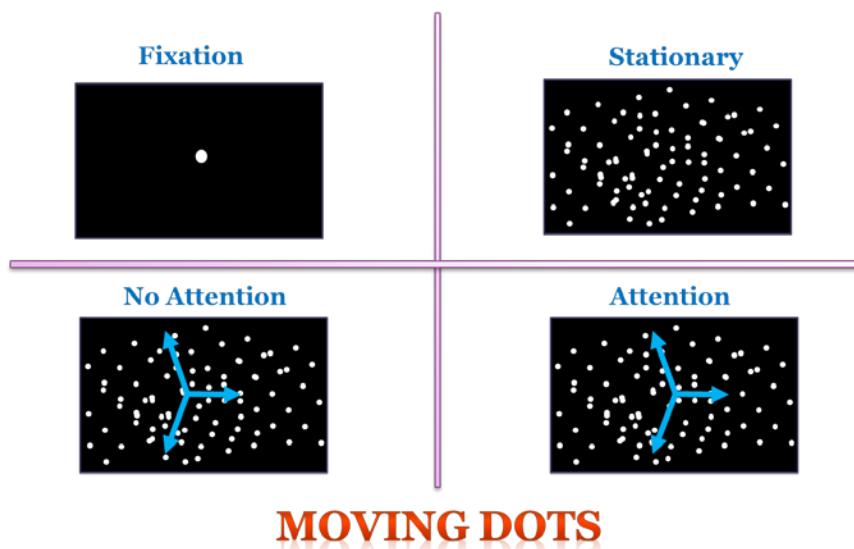


Bayesian decoding of motion

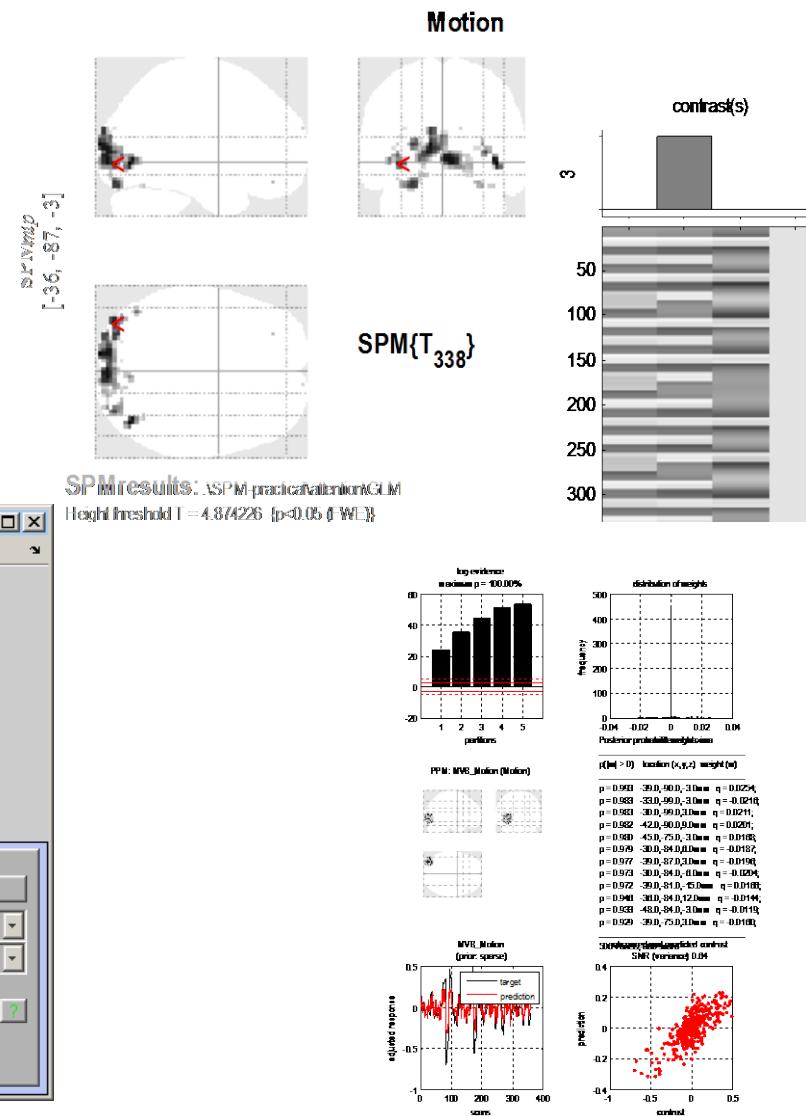
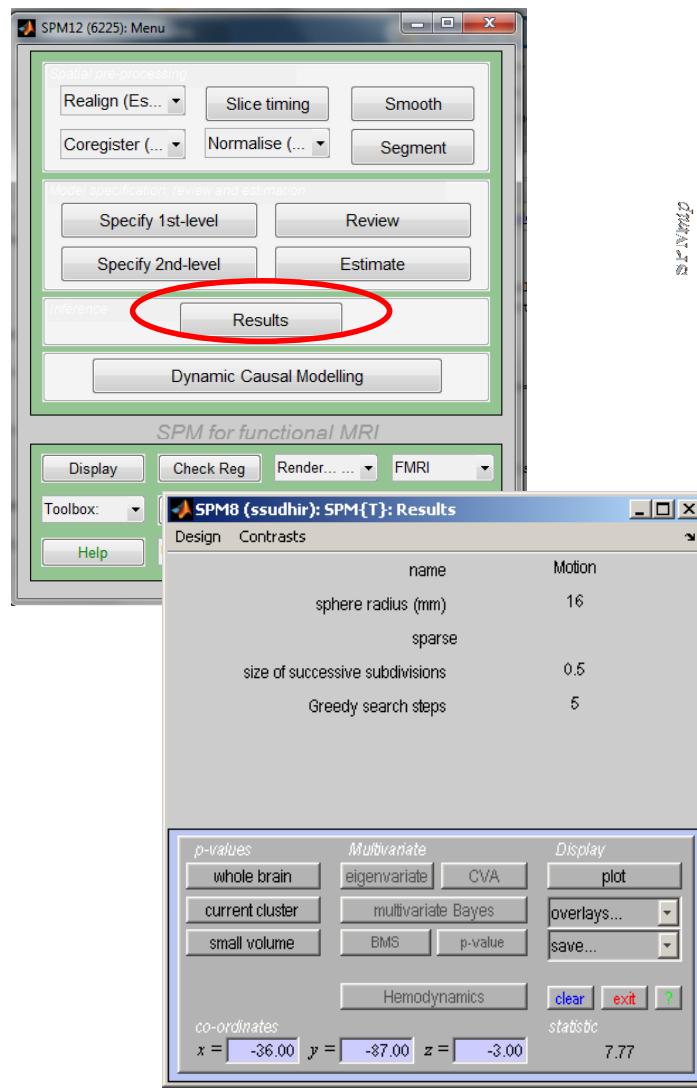
Attention to motion dataset - Büchel &
Friston 1999 Cerebral Cortex

Experimental factors:

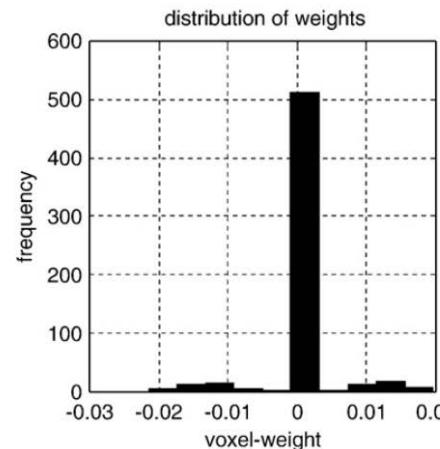
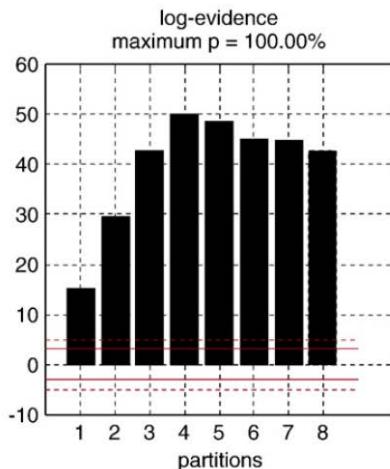
1. Photic
2. Motion
3. Attention



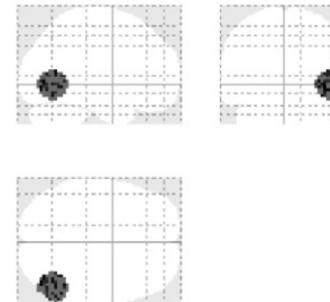
Multivariate Bayes in SPM



Results



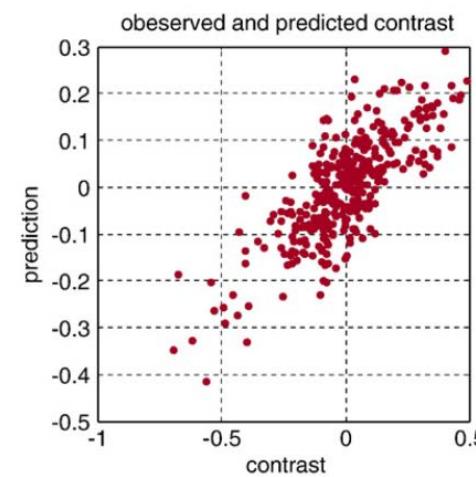
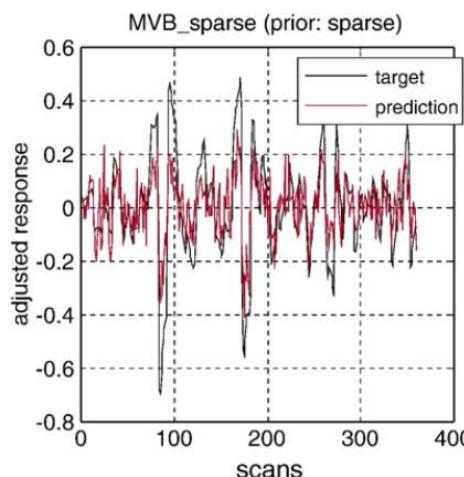
MVB_sparse (motion)



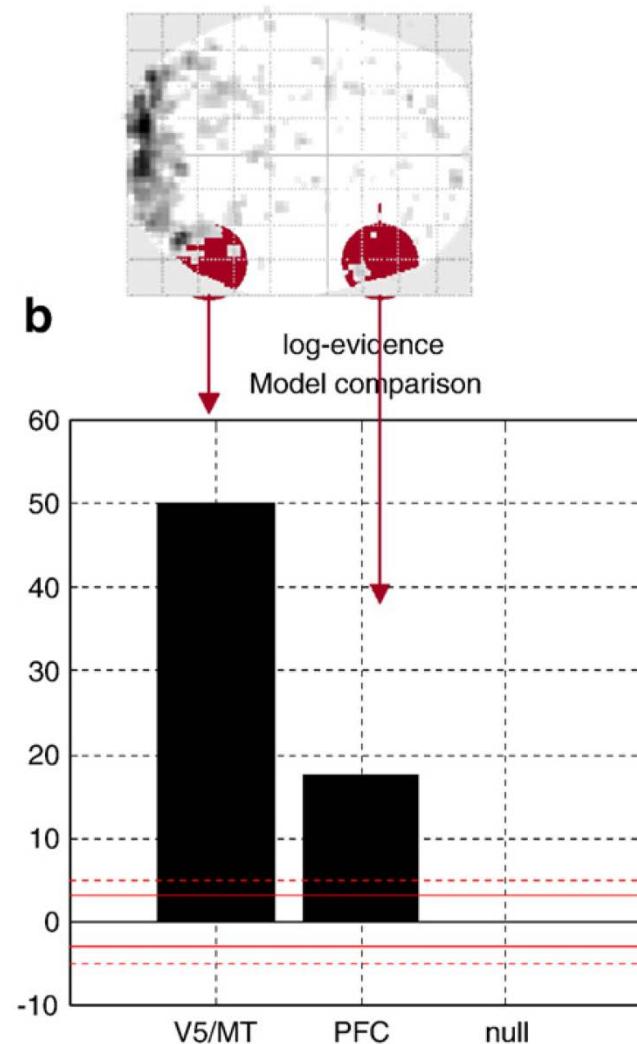
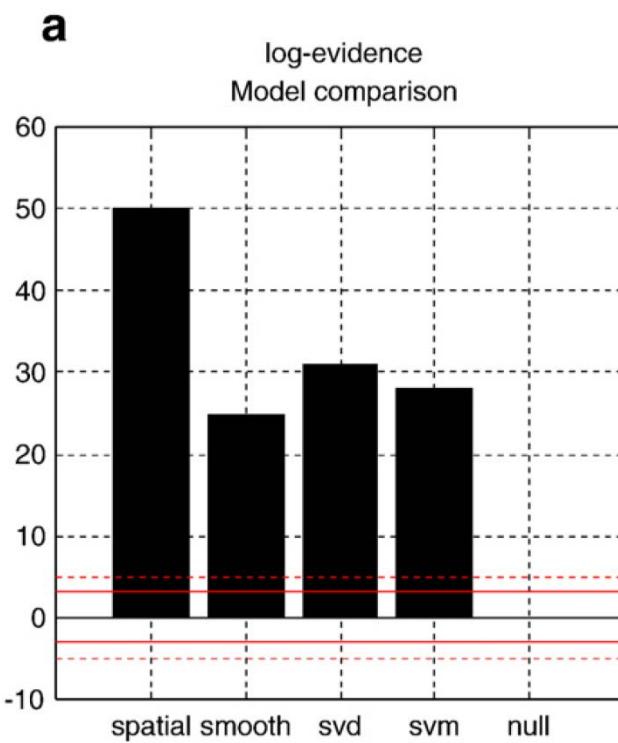
Posterior probabilities at maxima

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;

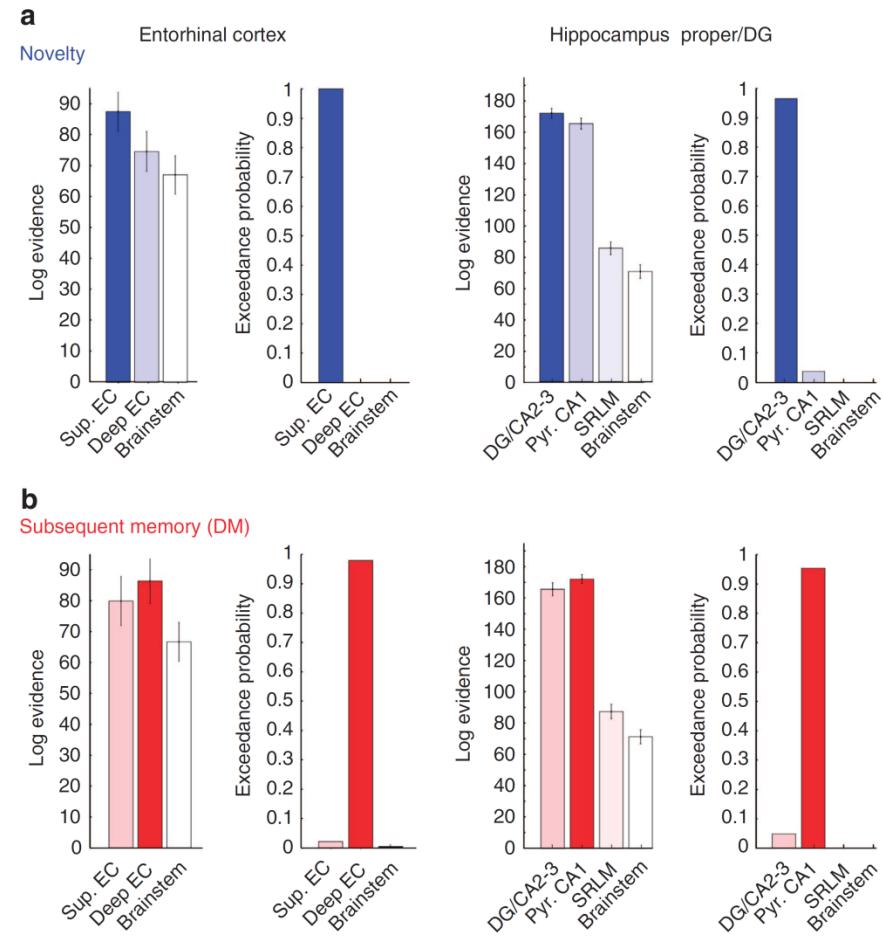
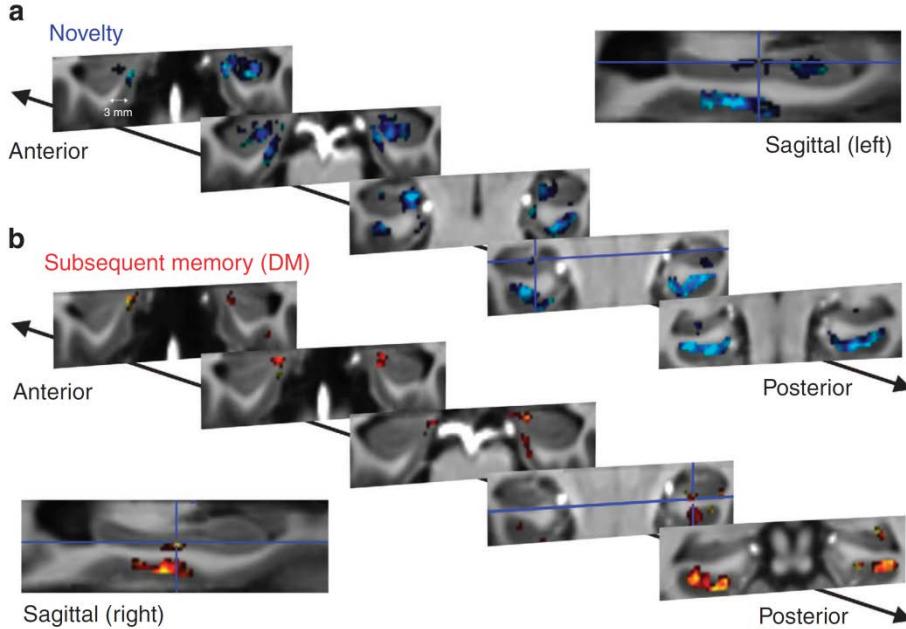
583 voxels; 360 scans



Results

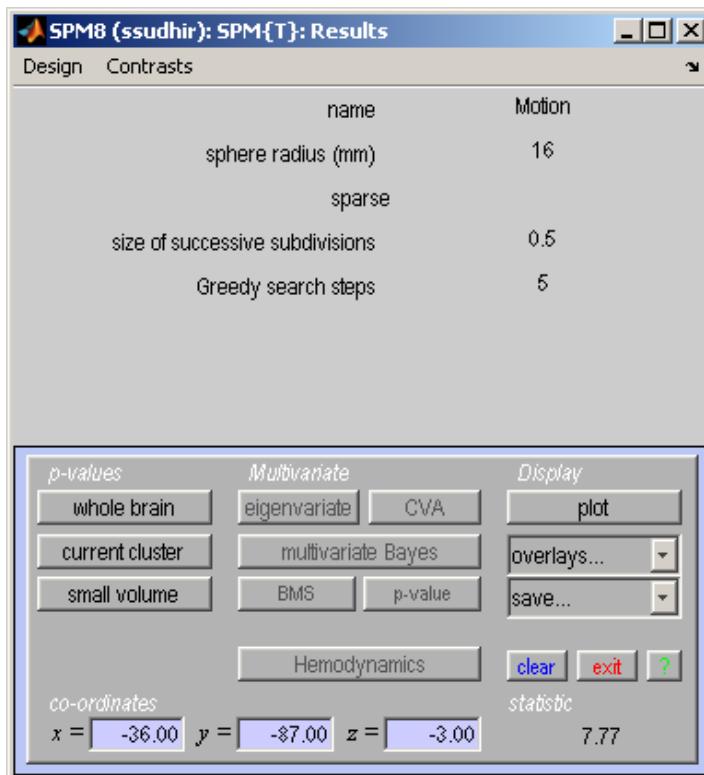


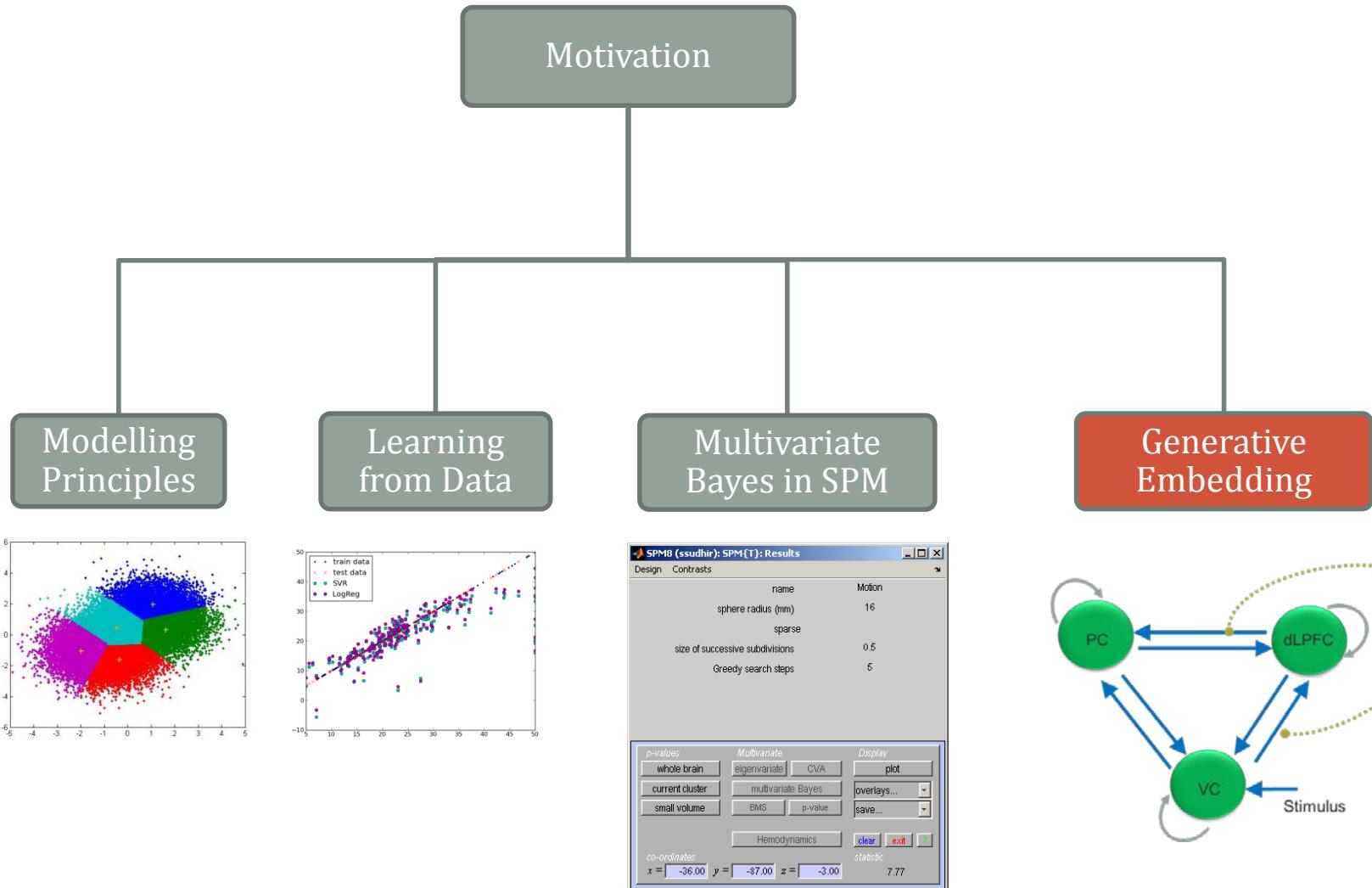
Laminar activity related to novelty and episodic encoding

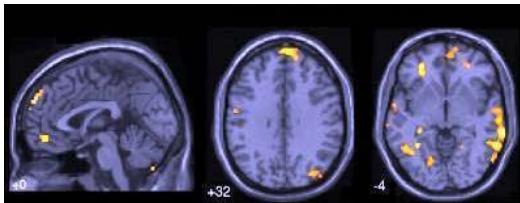


Demo

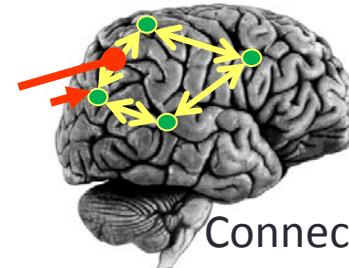
- Bayesian decoding of motion in SPM



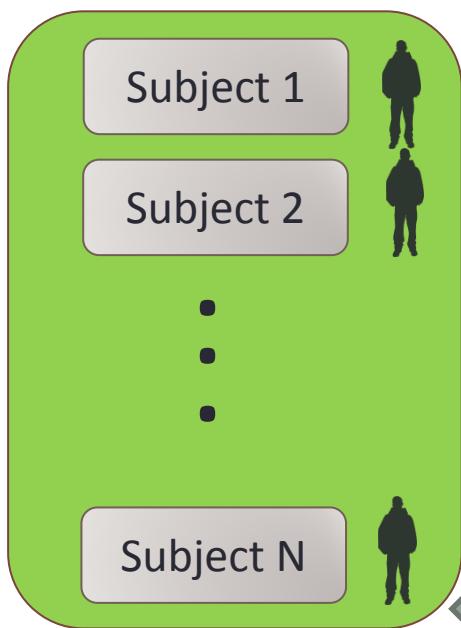




Voxel activity

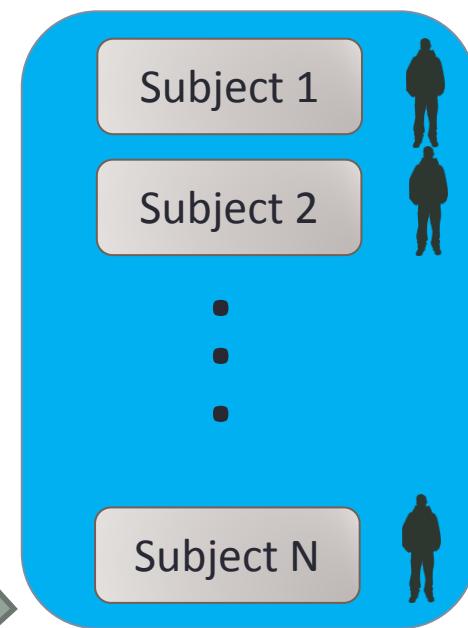


Connectivity



Dynamic causal
model (DCM)

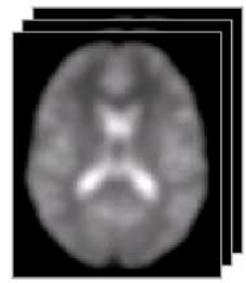
- High dimensionality
- Unusual cluster distributions
- Lack of interpretation



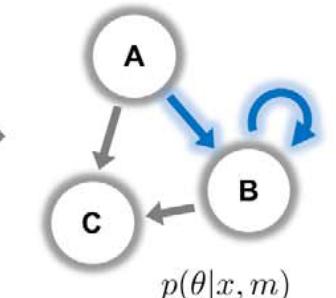
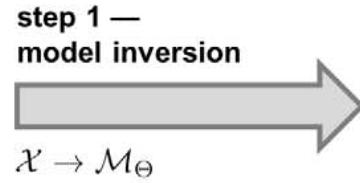
Classification
Clustering



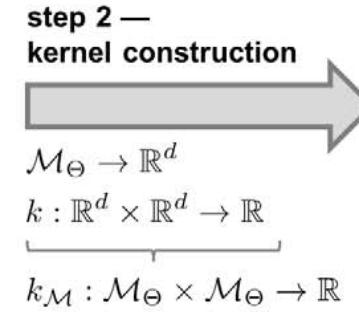
Generative Embedding - Classification



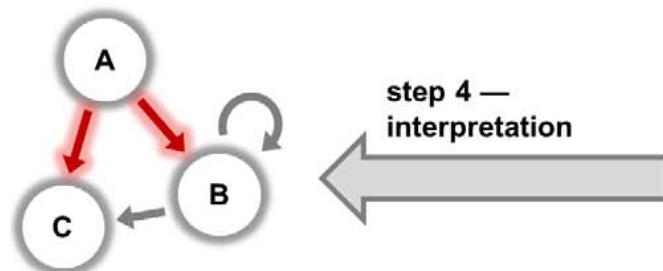
measurements from
an individual subject



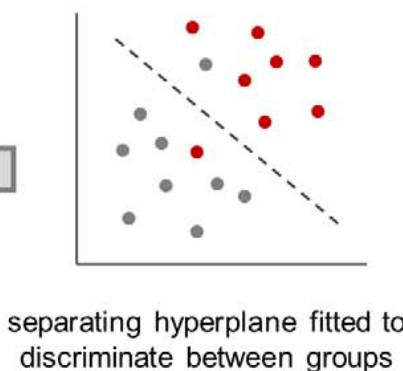
subject-specific
inverted generative model



subject representation in the
generative score space



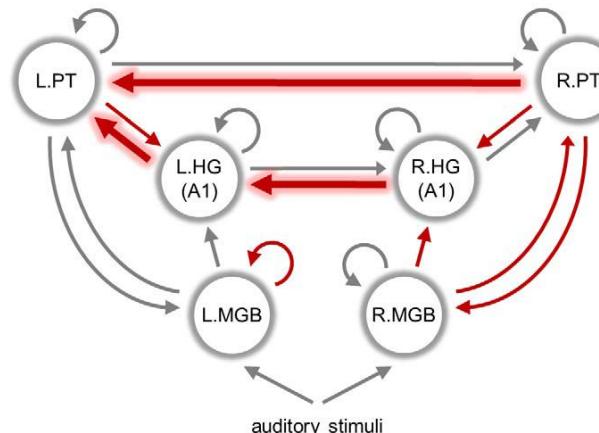
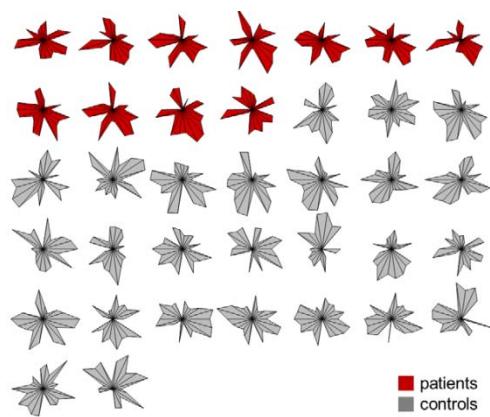
jointly discriminative
connection strengths



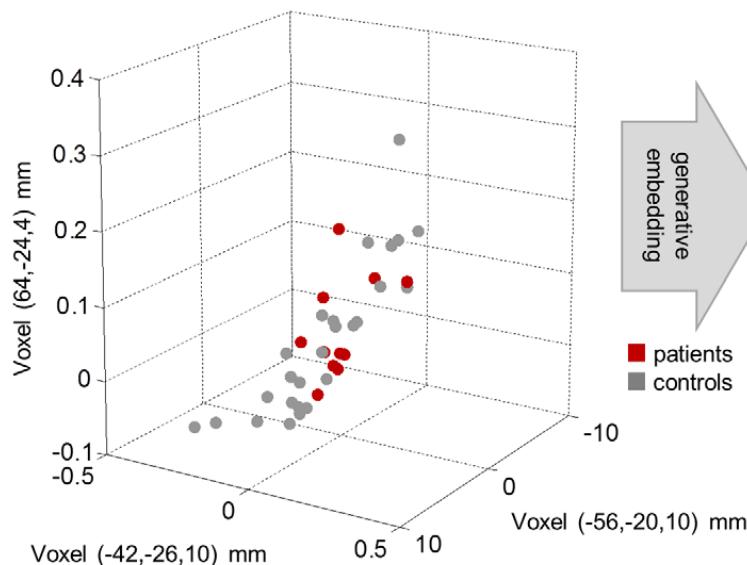
step 3 —
support vector classification

$$\hat{c} = \text{sgn} \left(\sum_i^n \alpha_i^* k(x_i, x) + b^* \right)$$

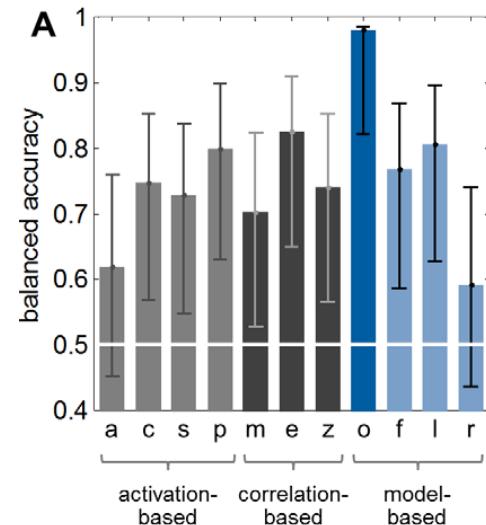
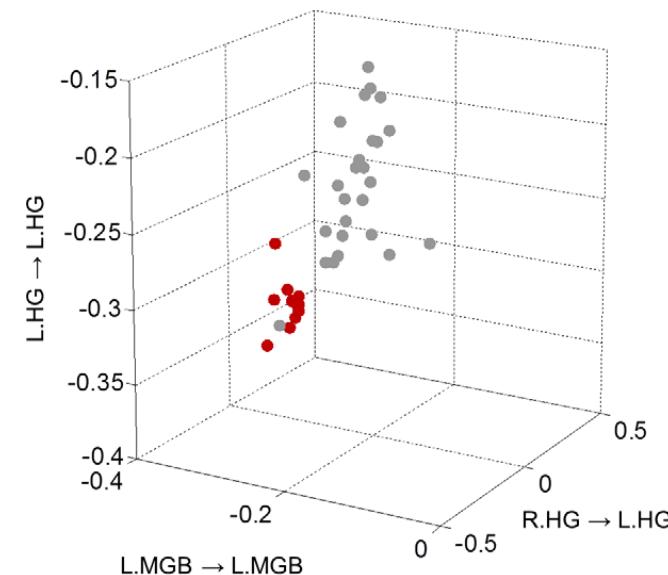
DCM - Speech processing



Voxel-based feature space

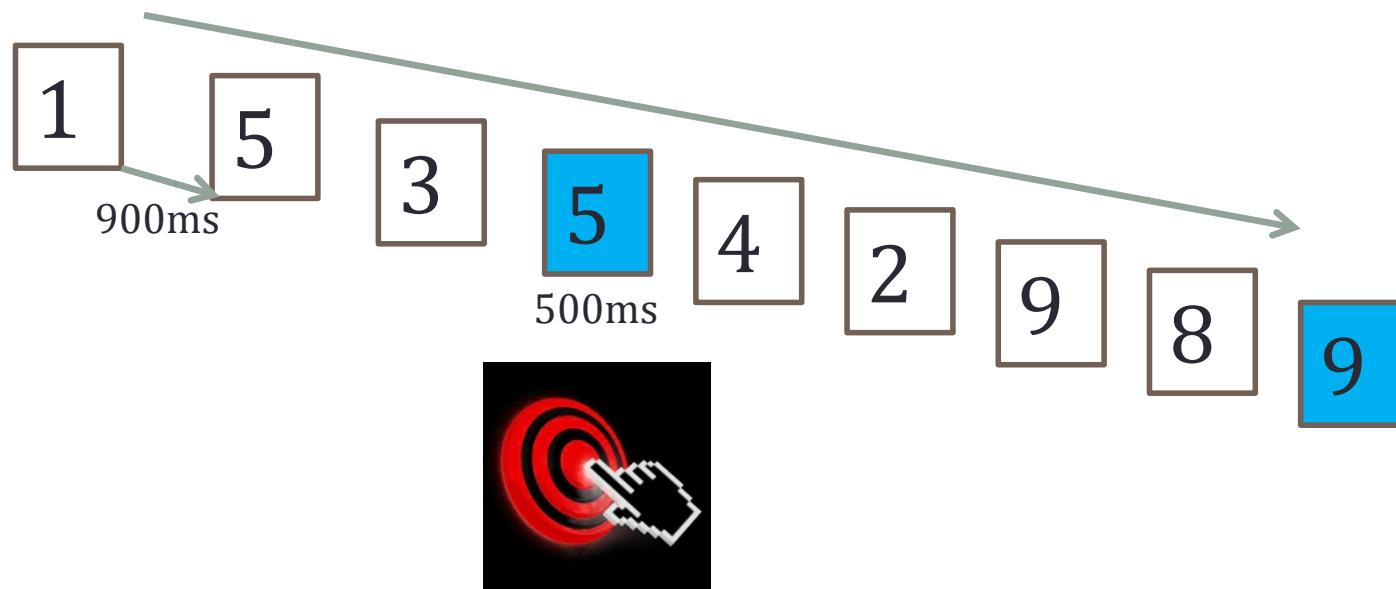


Generative score space

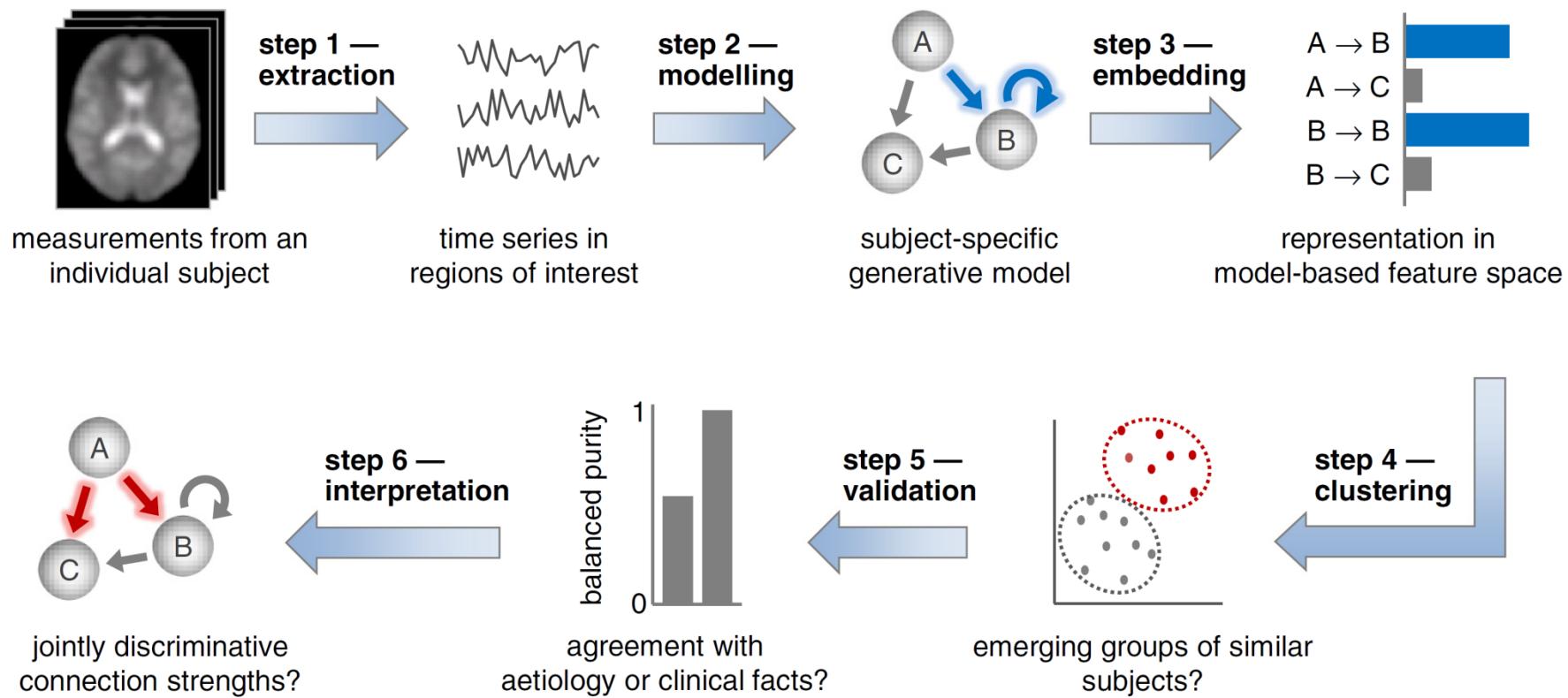


Working memory - fMRI

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

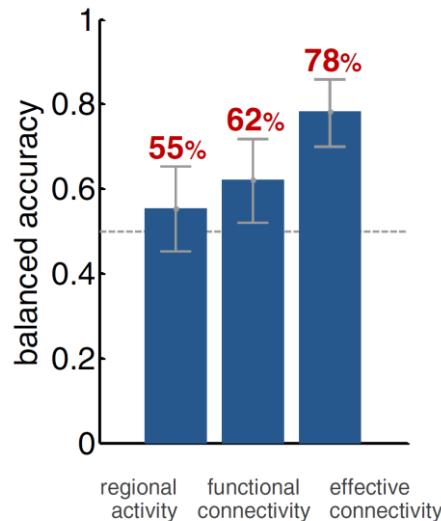


Model based clustering

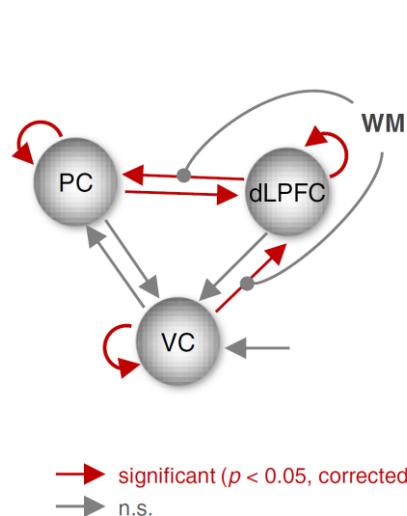


Results – Healthy vs Schizophrenic

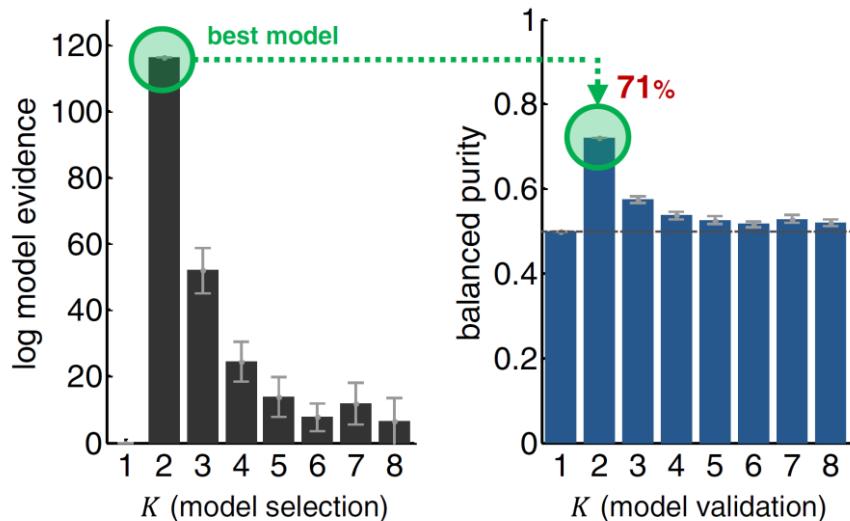
a supervised learning:
SVM classification



b discriminative
model parameters

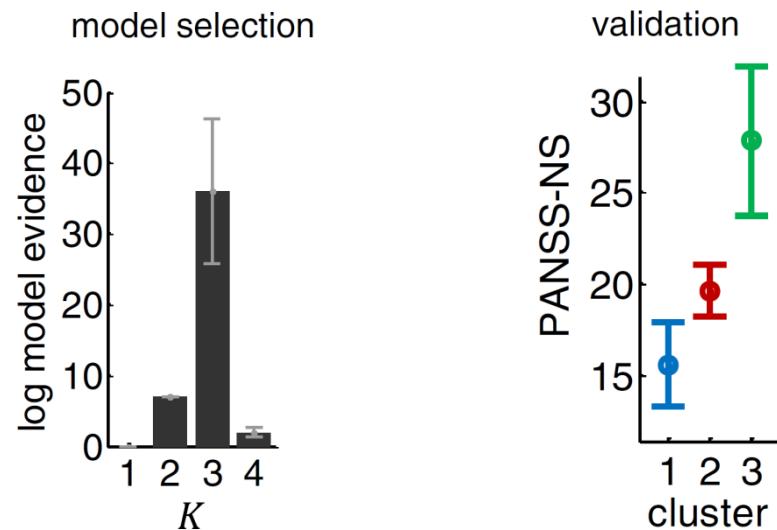
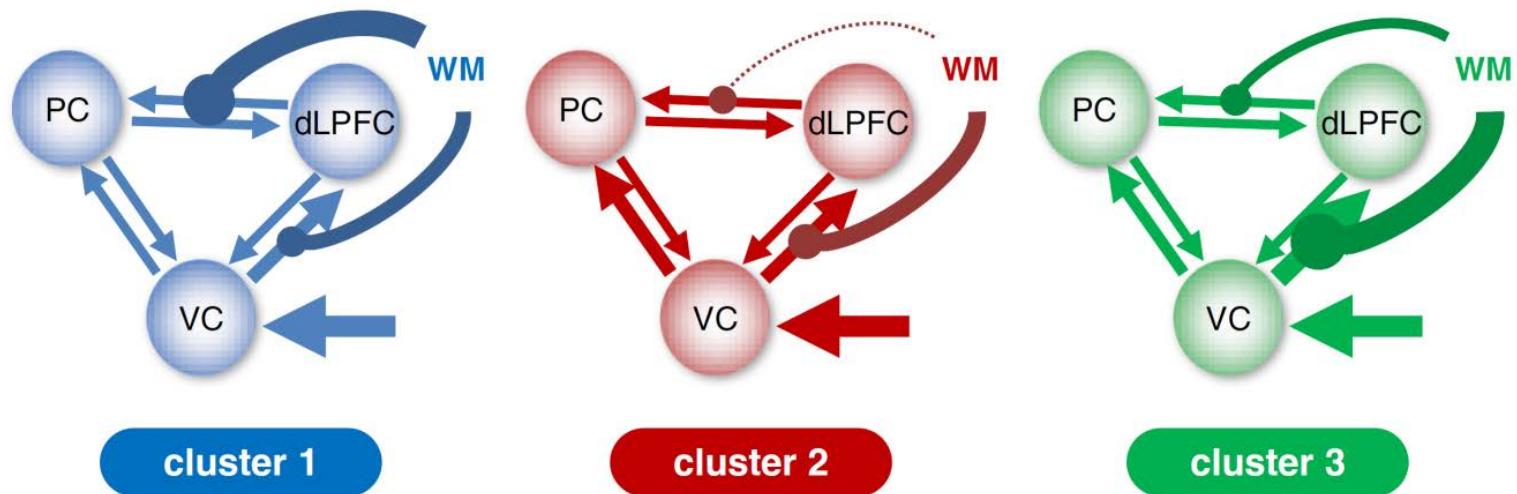


c unsupervised learning:
variational GMM clustering (using effective connectivity)

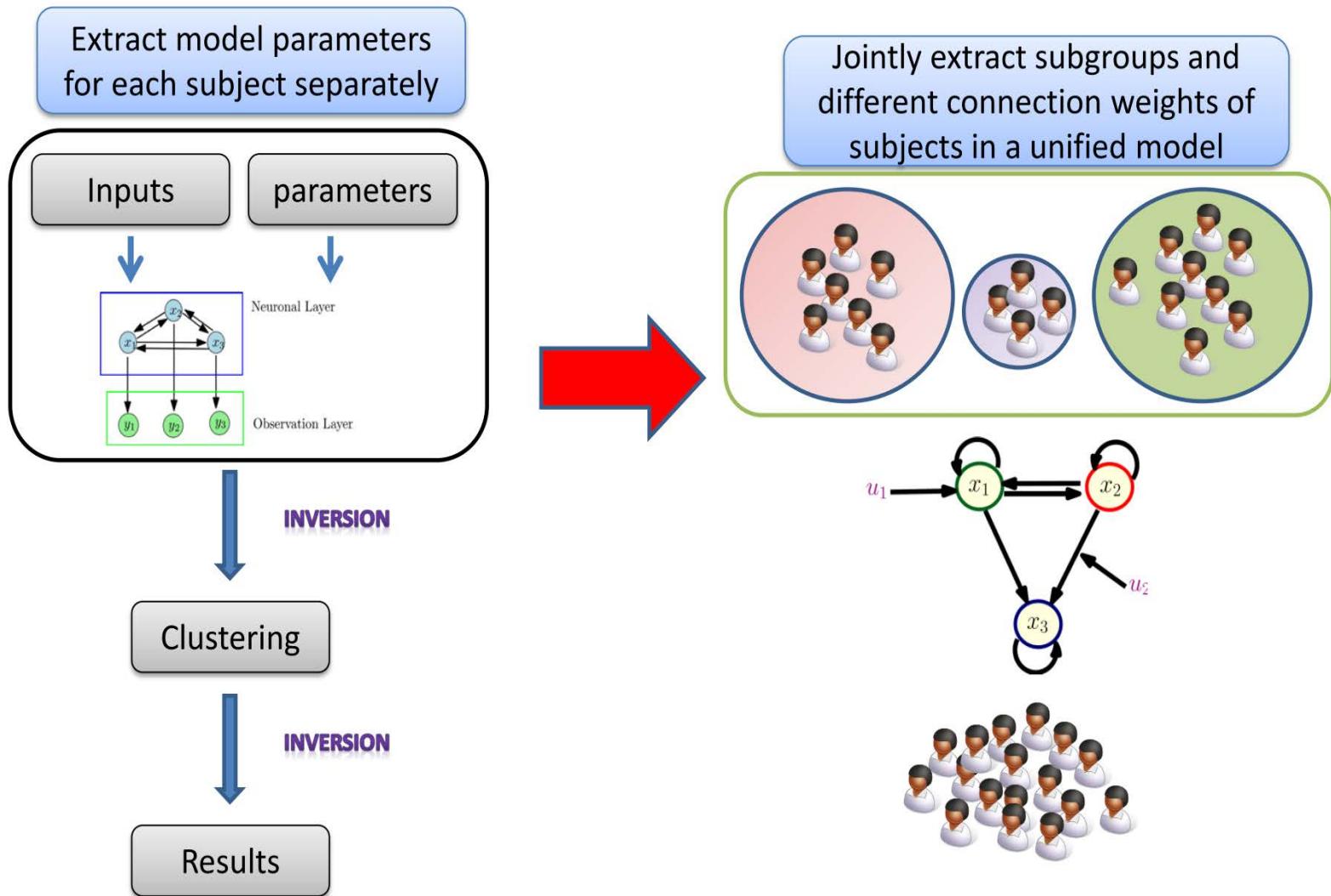


Brodersen et al 2014 Neuroimage

Results – Schizophrenia patients

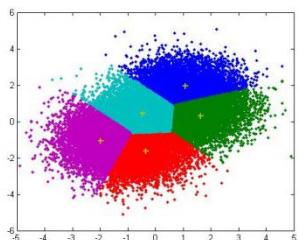


Unified model for identifying subgroups

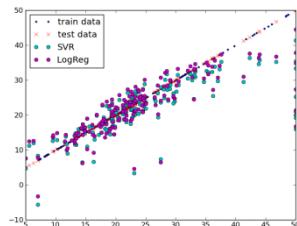


Summary

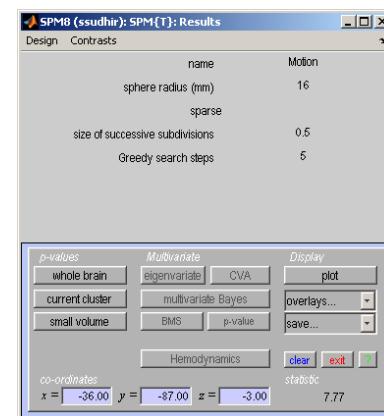
Modelling Principles



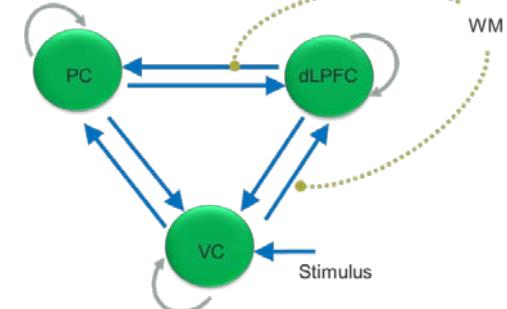
Learning from Data



Multivariate Bayes in SPM



Generative Embedding



Thank you