



Donders Institute  
for Brain, Cognition and Behaviour

**Brain Reading (MKI43)**

# **Lecture 1: Introduction**

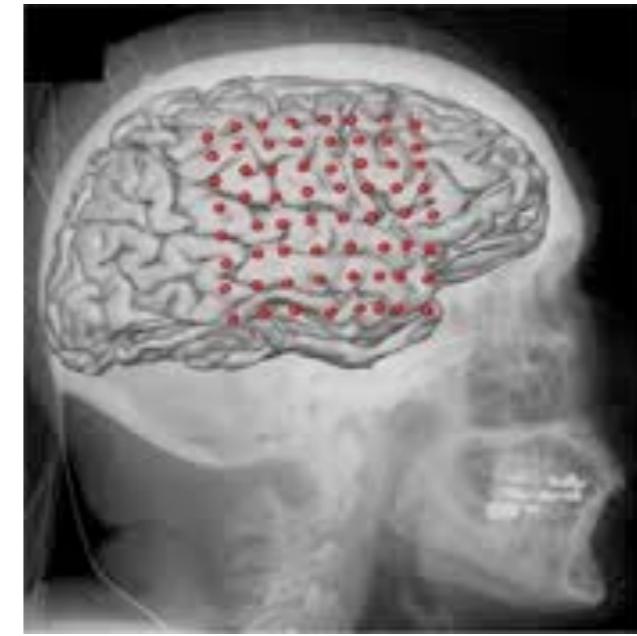
**Marcel van Gerven**  
**Assistant Professor**  
**Distributed Representations Group**  
**Donders Centre for Cognition**

**Radboud University Nijmegen**





- Marcel van Gerven
- Assistant professor in Artificial Intelligence
- Principal investigator at the Donders Institute
- Research group: [www.distrep.org](http://www.distrep.org)
- Understanding brain function using machine learning techniques:
  - Decoding cognitive states from neural data
  - Inferring connectivity patterns from neural data





- Course outline
- Introduction to brain reading:

Working definition:

Understanding cognitive processing based on  
brain activity as analysed using sophisticated  
statistical machine learning techniques



## Some motivating examples





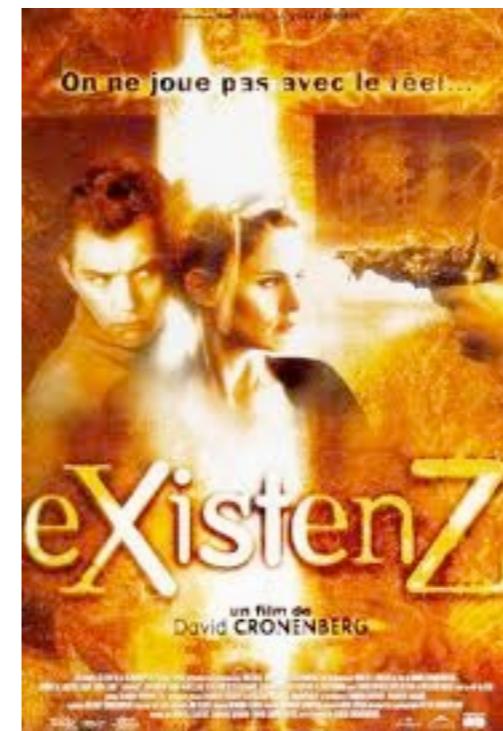
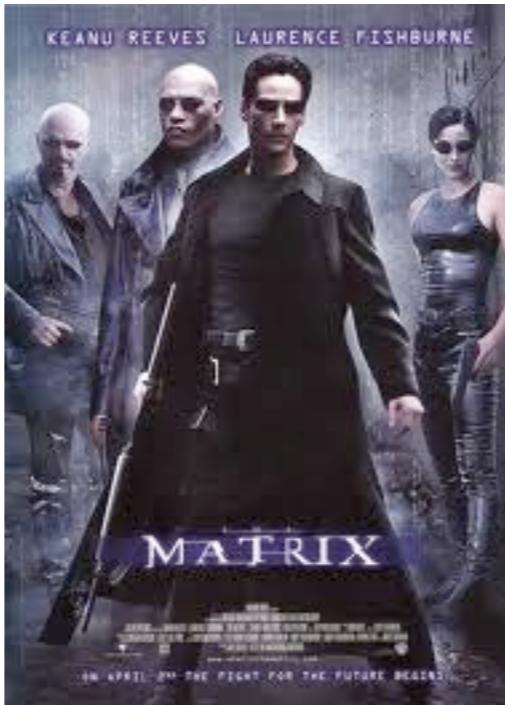
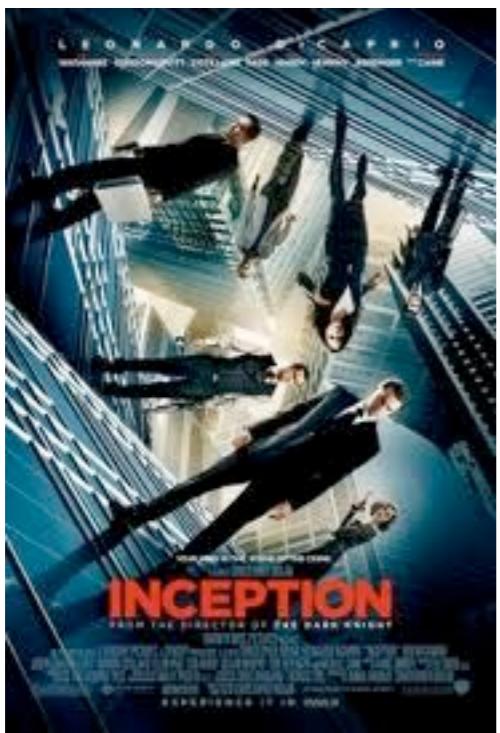
## Presented clip



## Clip reconstructed from brain activity



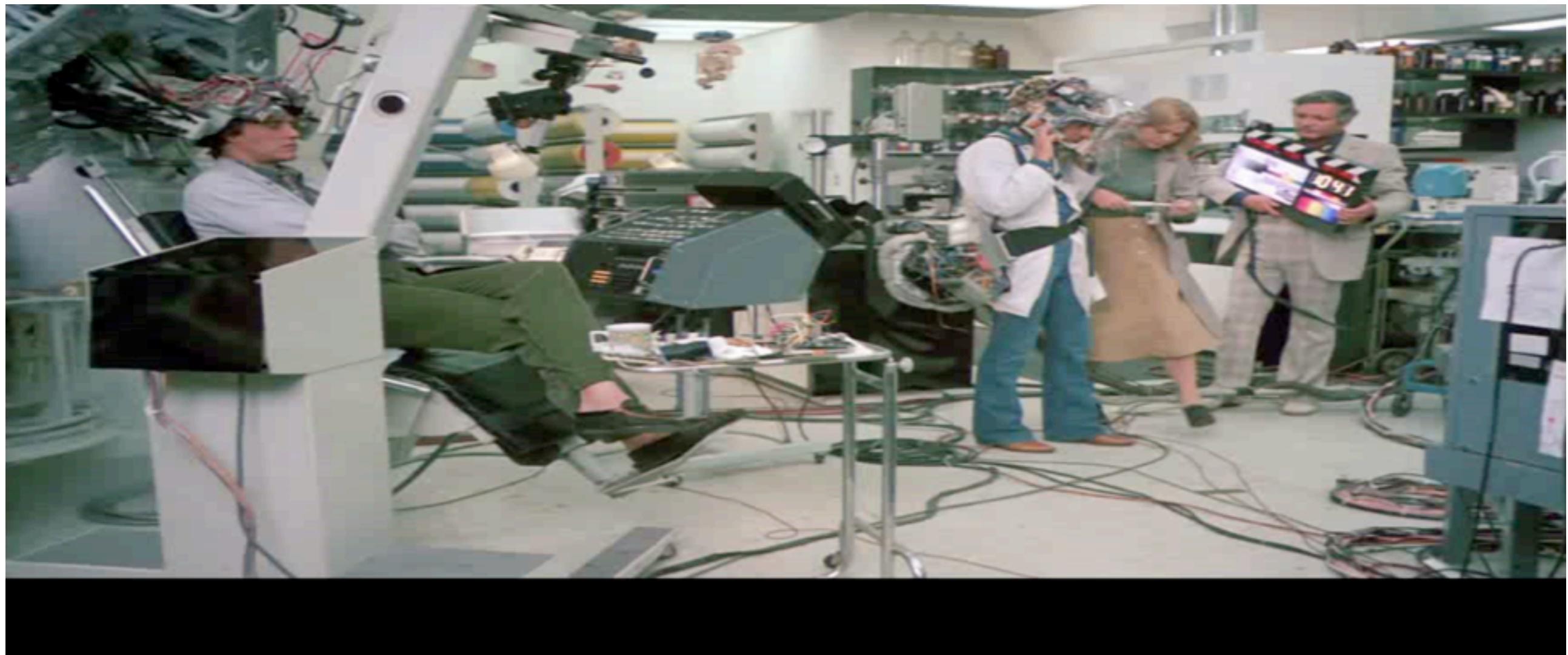
Decoding speaks to the imagination



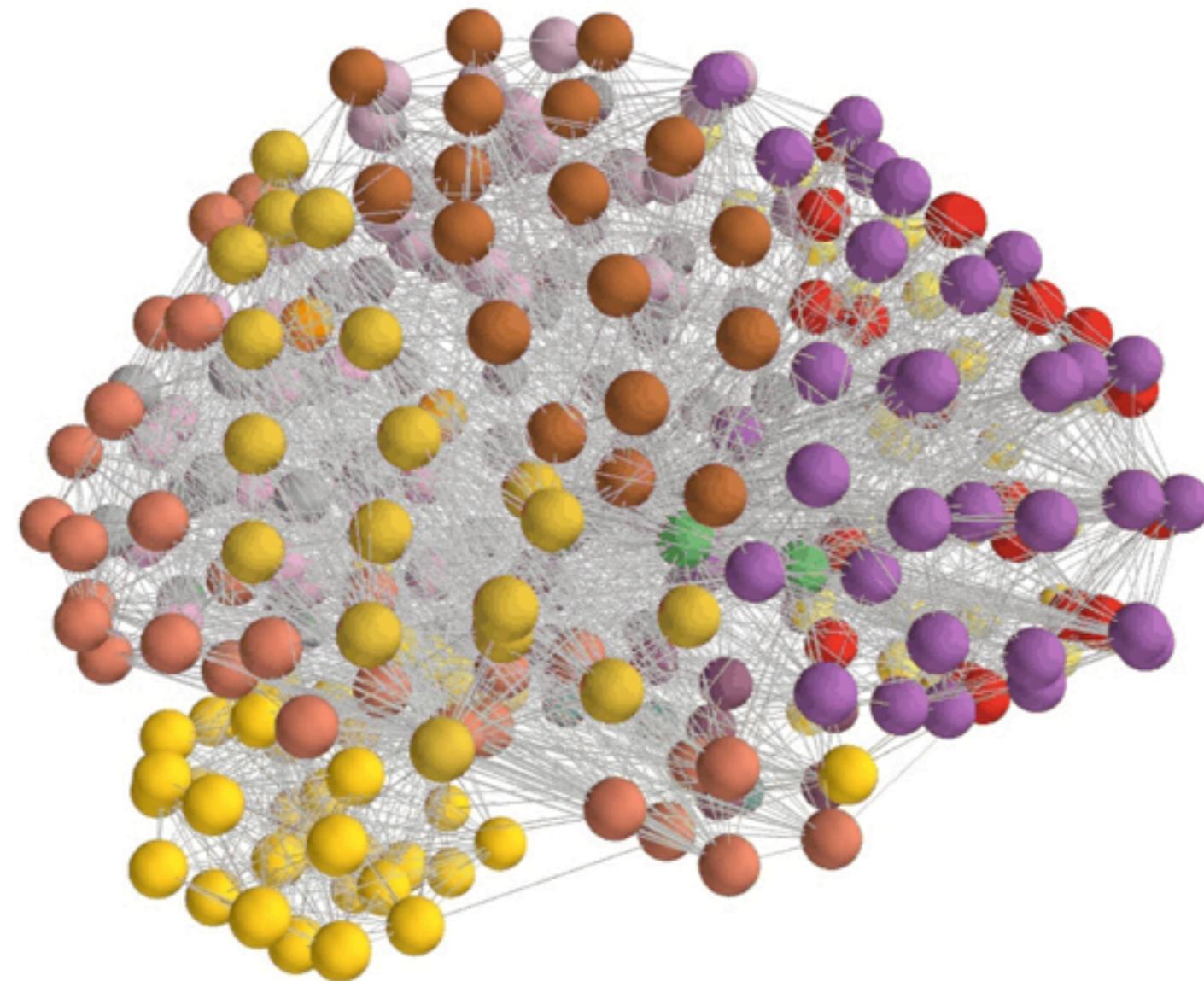
# Brainstorm (1983)



# Brainstorm (1983)



# Connectivity



# Dynamic changes







## Study goals:





## Study goals:

- Understand how to decode cognitive states from neuroimaging data





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- Understand how to decode cognitive states from neuroimaging data
- Get acquainted with the mathematics underlying multivariate analysis





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- Analyse (preprocessed) fMRI data using multivariate analysis methods





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- Understand how to decode cognitive states from neuroimaging data
- Get acquainted with the mathematics underlying multivariate analysis
- Analyse (preprocessed) fMRI data using multivariate analysis methods
- Read and understand the literature on this topic





## Study goals:

- Understand how to decode cognitive states from neuroimaging data
- Get acquainted with the mathematics underlying multivariate analysis
- Analyse (preprocessed) fMRI data using multivariate analysis methods
- Read and understand the literature on this topic
- Being able to place this research in a broader context





The course consists of the following components:

1. Lecture
2. Practical sessions
3. Essay
4. Final exam





For each lecture, required reading (journal papers or book chapter) and slide handouts will be made available in Blackboard.



A second component of the course is the practical. In the practical sessions, you will learn how to perform multivariate analysis of fMRI data. The practical assignments must be handed in before the exam date. Consult the practical manual for further details.



A third component of the course is writing an essay of about five to seven pages (2000-3000 words) on multivariate decoding.

The essay should address a topic that is of relevance in the context of multivariate decoding and will be evaluated on originality, scientific relevance and scientific soundness.



The final mark will be based on three parts:

- i. Practical assignments (20%)
- ii. Essay on multivariate decoding (30%)
- iii. Final written, closed book exam (50%)

The exam consist of a series of open questions.

Examinable is:

- ♣ All that has been discussed during lectures
- ♣ All required background reading material

Prerequisites:

- Hand in of practical assignments
- Hand in of essay

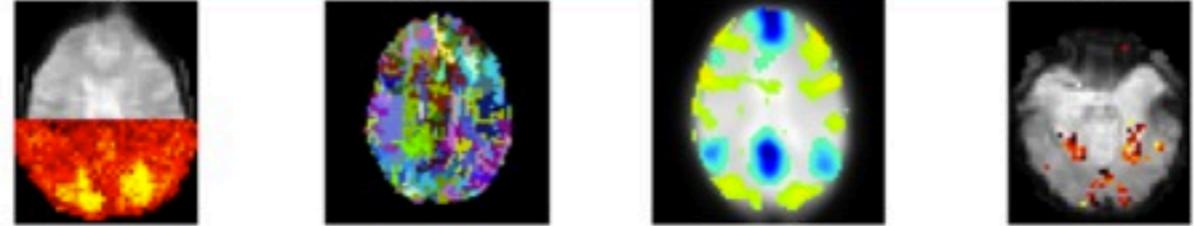


<http://nisl.github.com>

**scikit-learn**

Data Manipulation Supervised Learning Unsupervised Learning Examples Google Custom Search Search ×

## Machine learning for Neuroimaging in Python



**News**  
Gaël Varoquaux will present a tutorial on neuroimaging with scikit-learn at PRNI, in London July 2nd 2012.

**Licensing**  
All material Open source: BSD license (3 clause).

**About**  
Authors

**Giving credit**  
Please consider citing the scikit-learn if you use it.

**Learn machine learning for fMRI**  
This document compiles examples and tutorials to learn to apply machine learning to fMRI using Python and the scikit-learn. It requires nibabel and the scikit-learn.

**Download**

- PDF, 2 pages per side
- HTML and example files
- Source code (github)

### 1. Introduction

- 1.1. Machine Learning in Neuroimaging: what and why
- 1.2. Python and the scikit-learn: a primer

### 2. Basic dataset manipulation: loading and visualisation

- 2.1. Downloading the tutorial data
- 2.2. Loading Nifti or analyze files
- 2.3. Visualizing brain images
- 2.4. Masking the data

### 3. Supervised learning

- 3.1. Decoding on simulated data
- 3.2. fMRI decoding: predicting which objects a subject is viewing
- 3.3. Searchlight : finding voxels containing maximum information

### 4. Unsupervised learning

- 4.1. fMRI clustering
- 4.2. ICA of resting-state fMRI datasets

### 5. Code examples





also see: <https://github.com/distrep/DMLT>

The screenshot shows the GitHub repository page for 'distrep / DMLT'. The repository has 10 issues and 84 commits. The README.md file contains an 'About' section describing the DMLT toolbox as a Machine Learning toolbox written in Matlab and C, developed at the Donders Institute for Brain, Cognition and Behaviour. It provides a general interface for statistical machine learning methods and allows complex methods to be built from simple building blocks. The code requires Matlab distribution 7.6.0.324 (R2008a). Functions are licensed under the GNU General Public License (GPL).

Donders Machine Learning Toolbox — Read more

Clone in Mac ZIP HTTP SSH Git Read-Only https://github.com/distrep/DMLT.git Read+Write access

branch: master Files Commits Branches 1 Tags Downloads

DMLT / 84 commits

whitener

File	Author	Date	Message
+dml	distrep	a month ago	variational garrote added [distrep]
external	distrep	a month ago	added linux support for repop and tprod [distrep]
html	distrep	a month ago	added linux support for repop and tprod [distrep]
README.md	distrep	a month ago	whitener [distrep]
info.xml	distrep	7 months ago	MSG [distrep]

README.md

## About

DMLT is a Machine Learning toolbox written in Matlab and C. This toolbox is developed at the Donders Institute for Brain, Cognition and Behaviour and provides a general interface to support the integration of new statistical machine learning methods by writing high level wrappers. It allows complex methods to be built from simple building blocks and makes the use of cross-validation and permutation testing as easy as writing one line of Matlab code. The code requires at least Matlab distribution 7.6.0.324 (R2008a).

Most functions in this toolbox are licensed under the GNU General Public License (GPL), see <http://www.gnu.org> for details. Unauthorised copying and distribution of functions that are not explicitly covered by the GPL is not allowed. This code comes without warranty of any kind.





- Check Blackboard for announcements/deadlines/course material
- Check out the Reader and Practical Manual
- Study background literature belonging to each lecture (exam material)
- Interactive lectures (don't be afraid to ask questions or give input!)



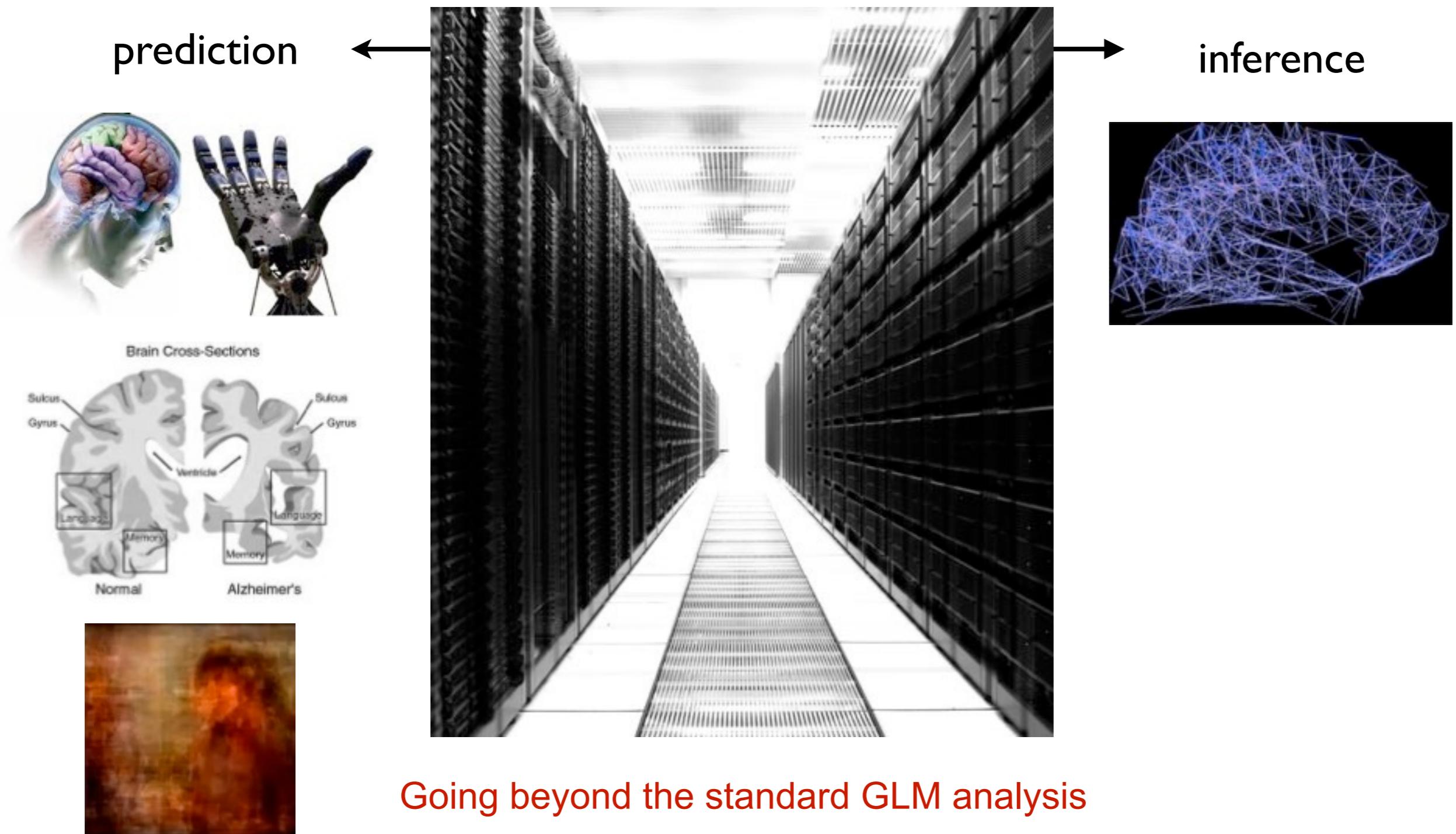


# Introduction to brain reading





## statistical machine learning



# Why Statistical machine learning





# Probabilistic models as a lingua franca





## Probabilistic models as a lingua franca

Statistics as inverse probability theory:

$$p(\theta | D) = \frac{p(D | \theta)p(\theta)}{P(D)}$$

“Getting from the data to the models”





## Probabilistic models as a lingua franca

Statistics as inverse probability theory:

$$p(\theta | D) = \frac{p(D | \theta)p(\theta)}{P(D)}$$

“Getting from the data to the models”

Bayesian statistics as the theoretically optimal way to

- analyze neural data
- think about brain function

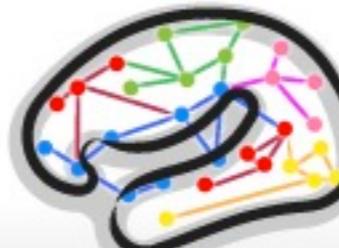




The fMRI Data Center  
**fMRI DC**



ALZHEIMER's DISEASE NEUROIMAGING INITIATIVE



HUMAN  
Connectome  
PROJECT



ALLEN INSTITUTE  
for BRAIN SCIENCE  
*Fueling Discovery*



**NEUROTYCHO** *beta*

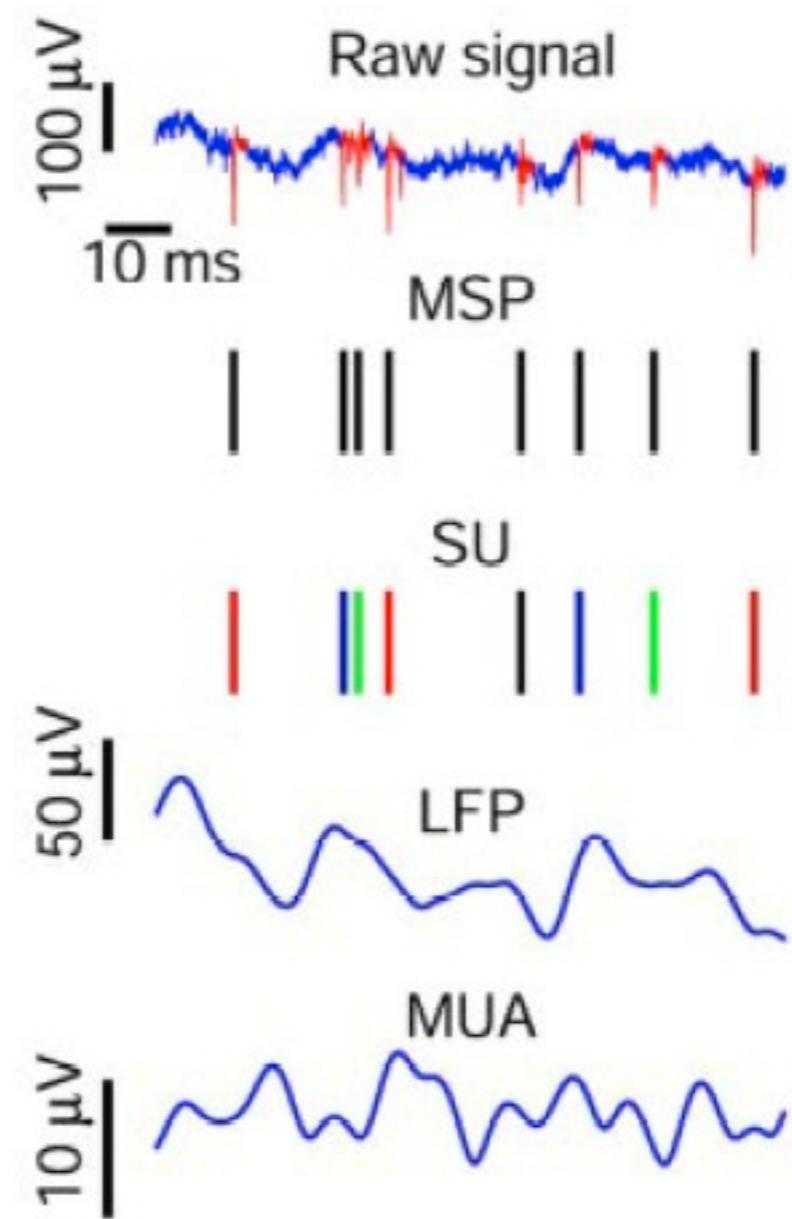
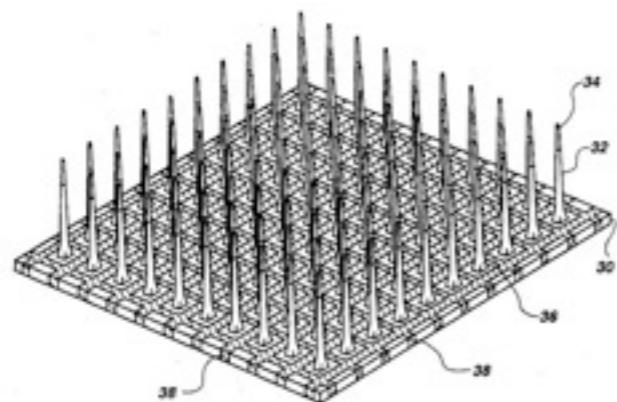
## Challenges and Opportunities in Mining Neuroscience Data

Huda Akil,<sup>1\*</sup> Maryann E. Martone,<sup>2</sup> David C. Van Essen<sup>3</sup>

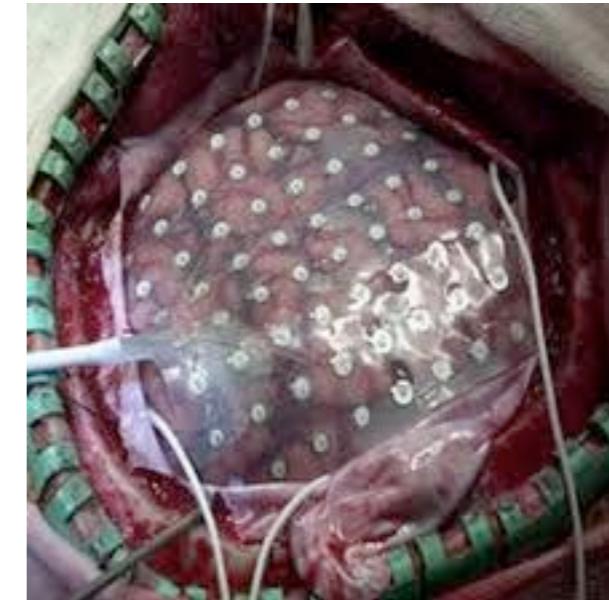
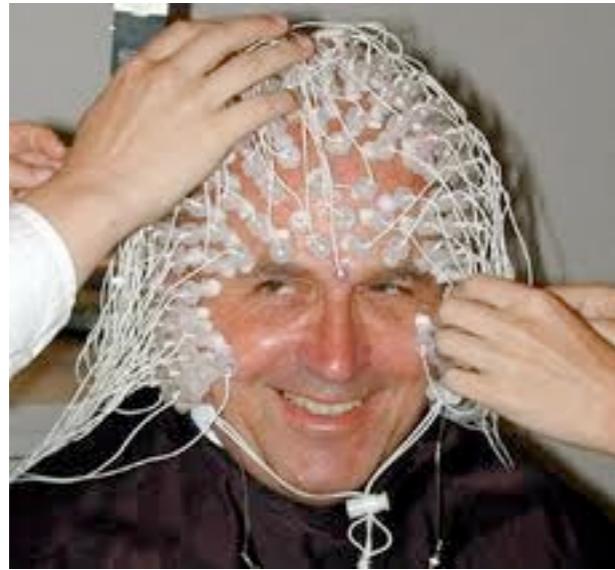
Science, 2011

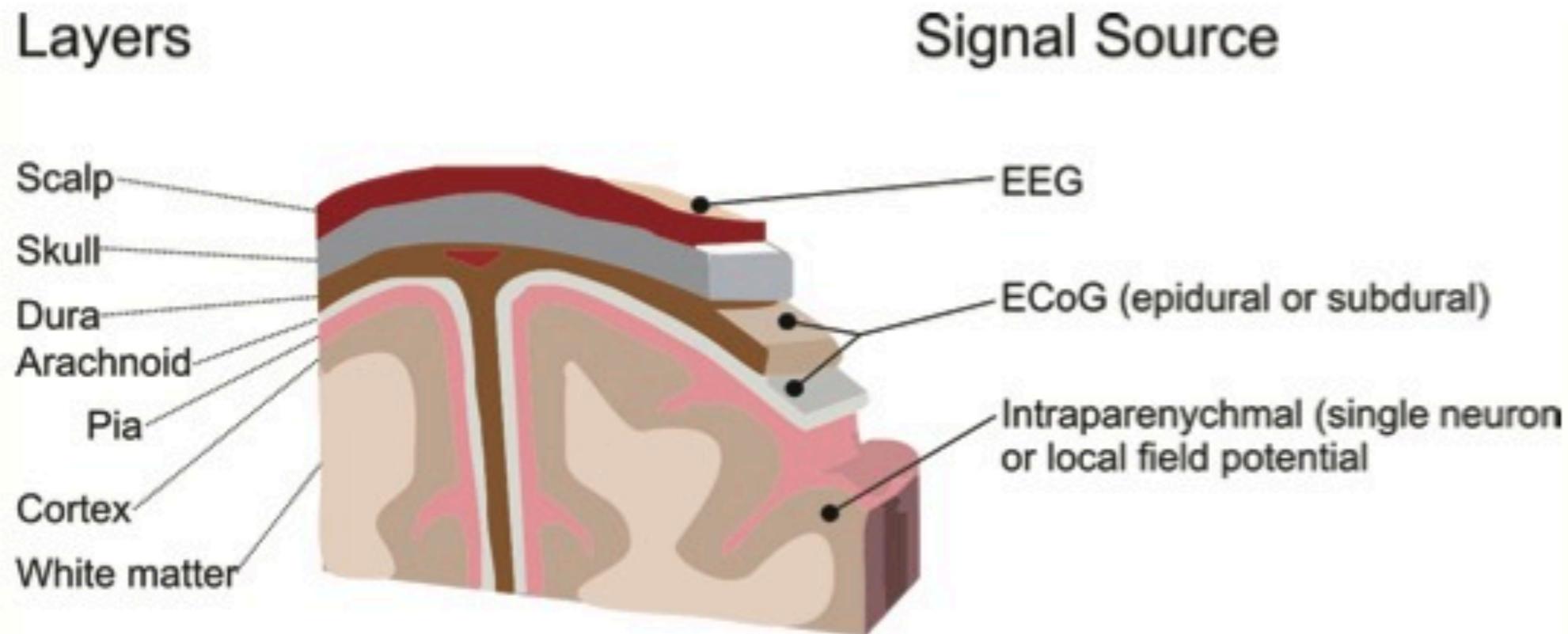


# Neuroimaging data: electrophysiology

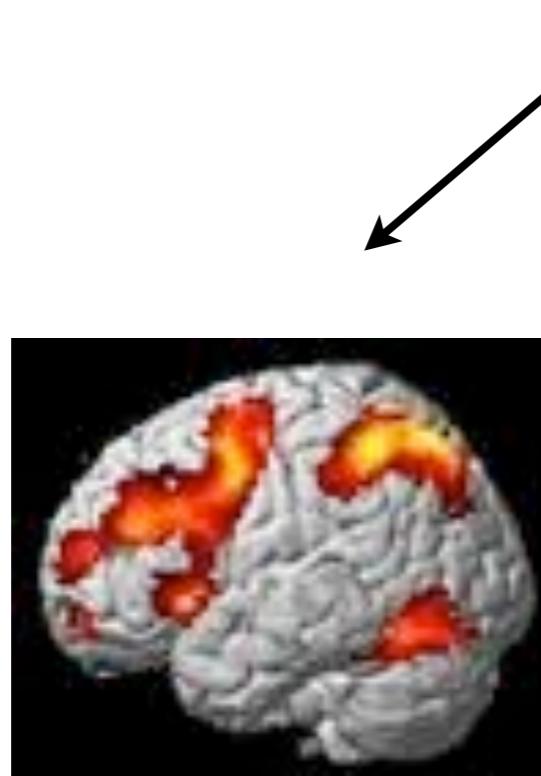


# Neuroimaging data: electrophysiology

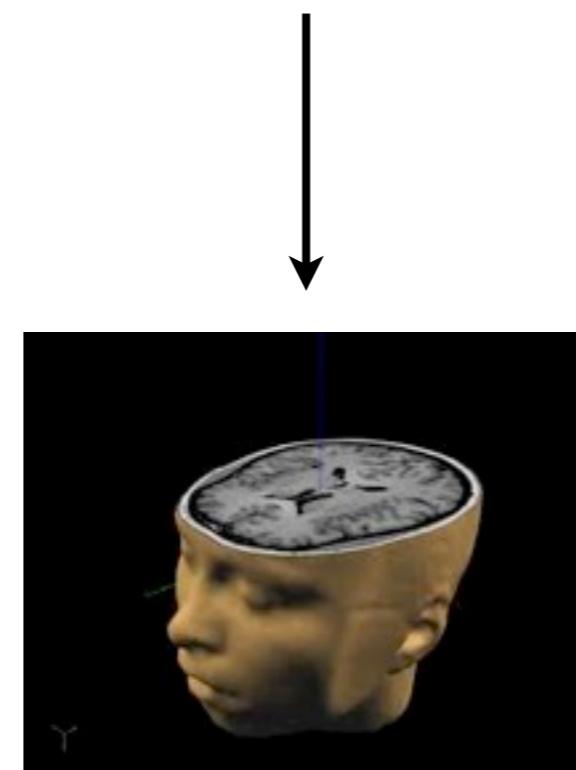




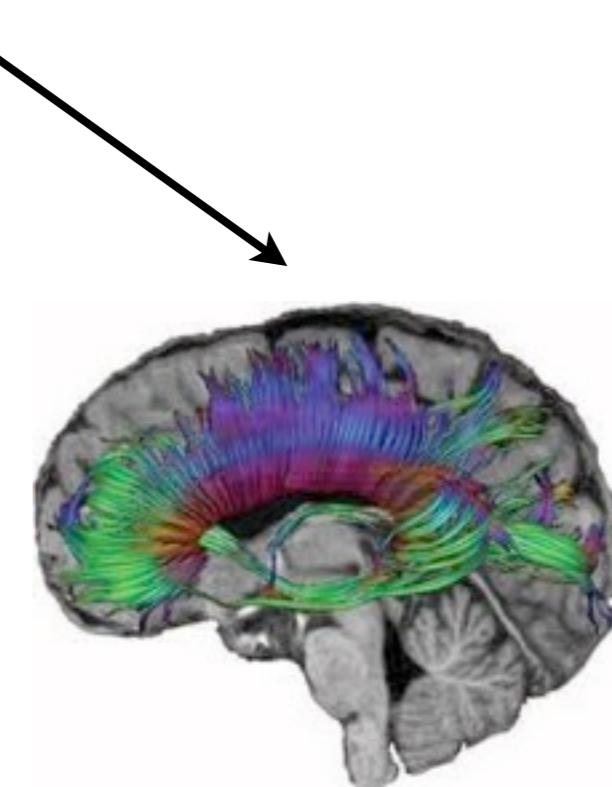
Leuthardt, E. C., Schalk, G., Roland, J., Rouse, A., & Moran, D. W. (2009). Evolution of brain-computer interfaces: going beyond classic motor physiology Neurosurgical focus, 27(1)



functional



structural



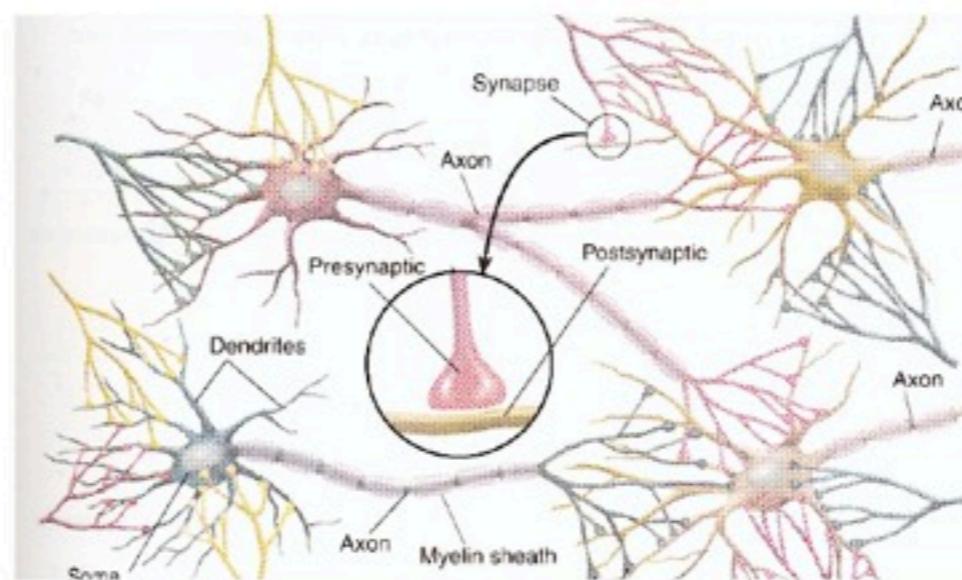
diffusion



## THE BOLD CONTRAST

BOLD (Blood Oxygenation Level Dependent) contrast = measures inhomogeneities in the magnetic field due to changes in the level of O<sub>2</sub> in the blood

**CURRENT CONCLUSION: BOLD SIGNAL  
SEEMS TO BE MORE STRONGLY CORRELATED  
TO POSTSYNAPTIC ACTIVITY**

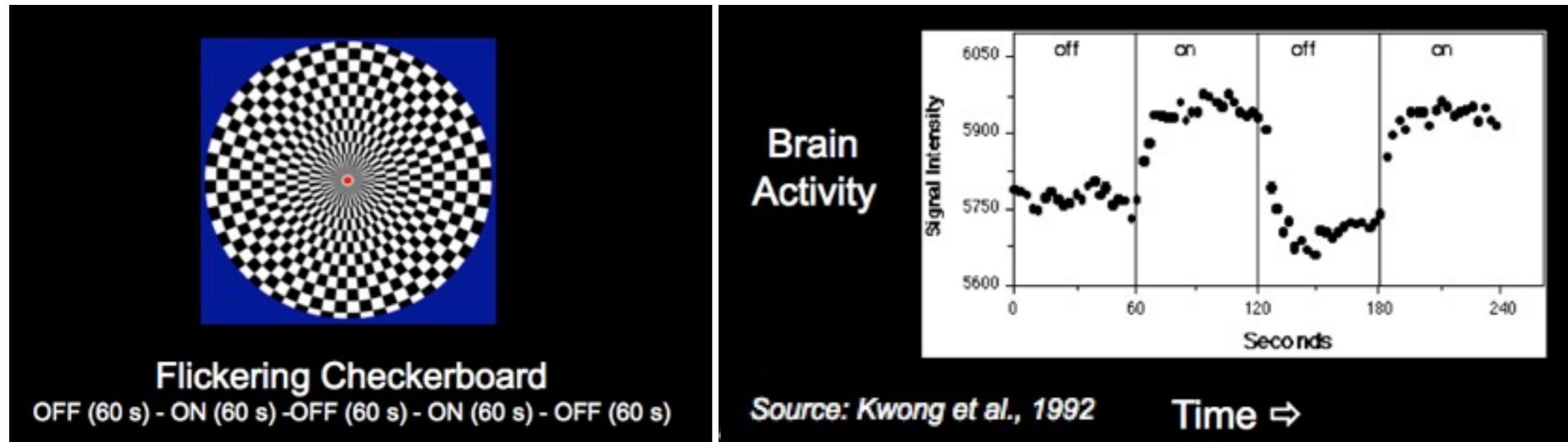


BOLD seems to reflect the input to a neuronal population as well as its intrinsic processing.

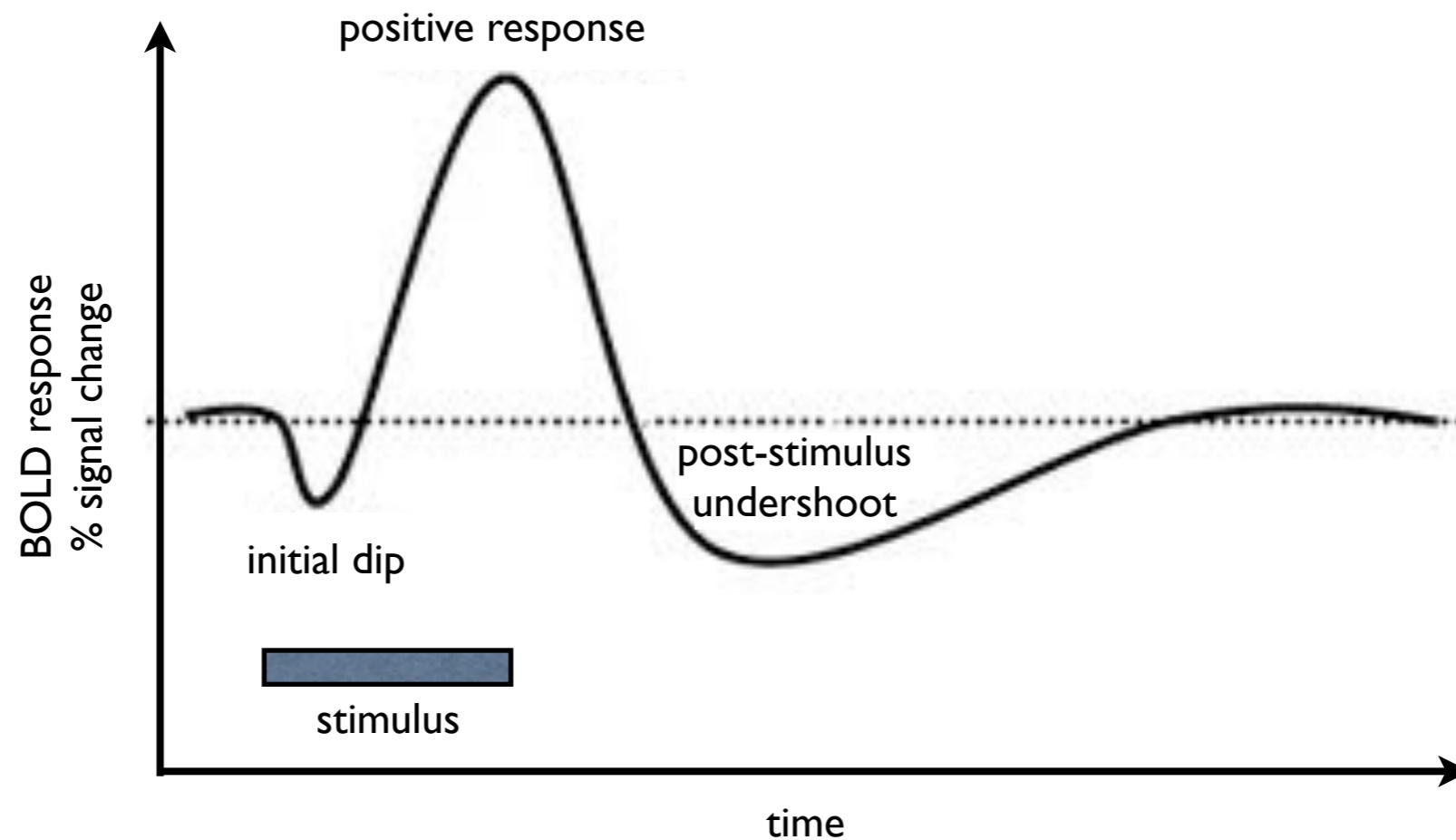
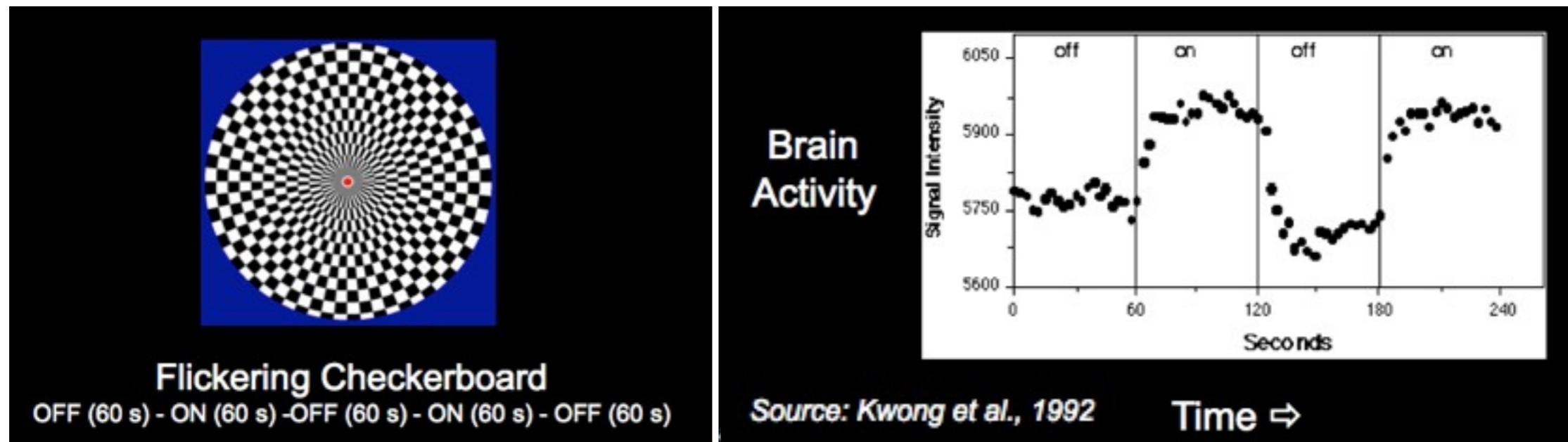
Lauritzen 2005, *Nat. Neurosci. Rev.*



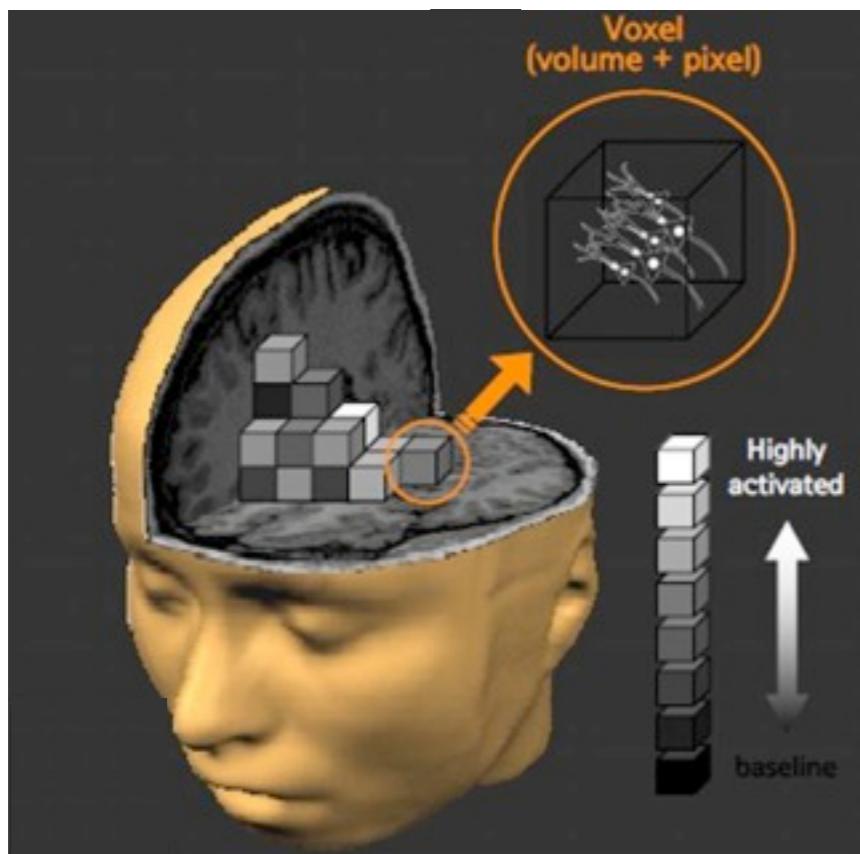
## Hemodynamic response



## Hemodynamic response

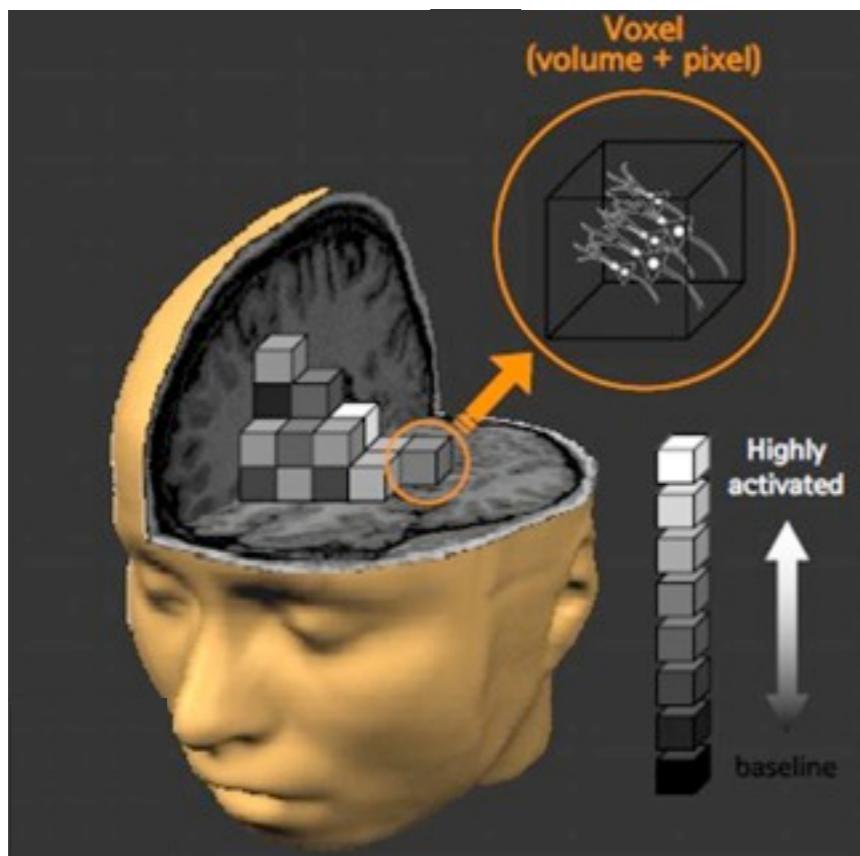


## Acquired data

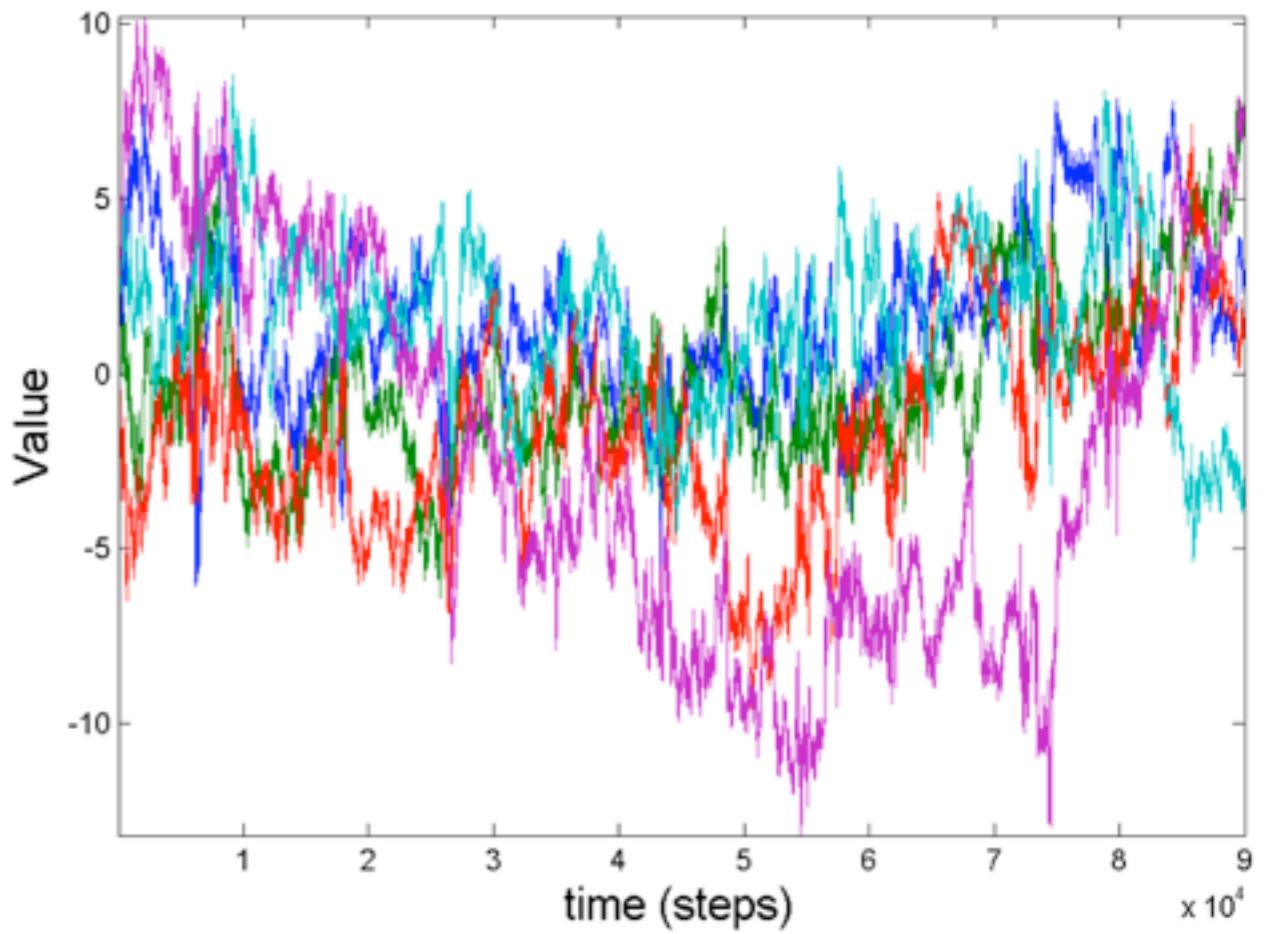


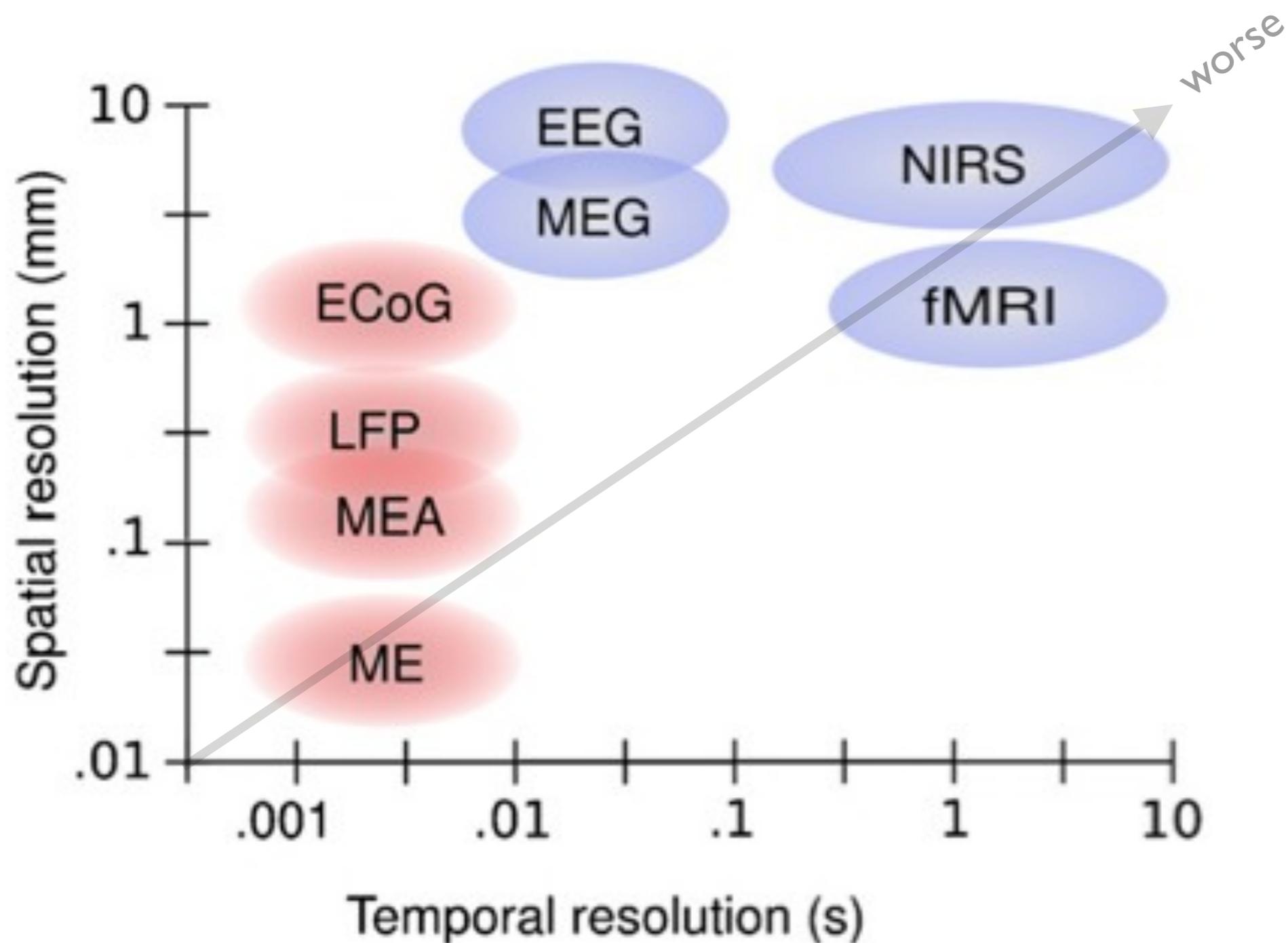
Acquisition of about 50000 2x2x2 mm voxels every 2 seconds

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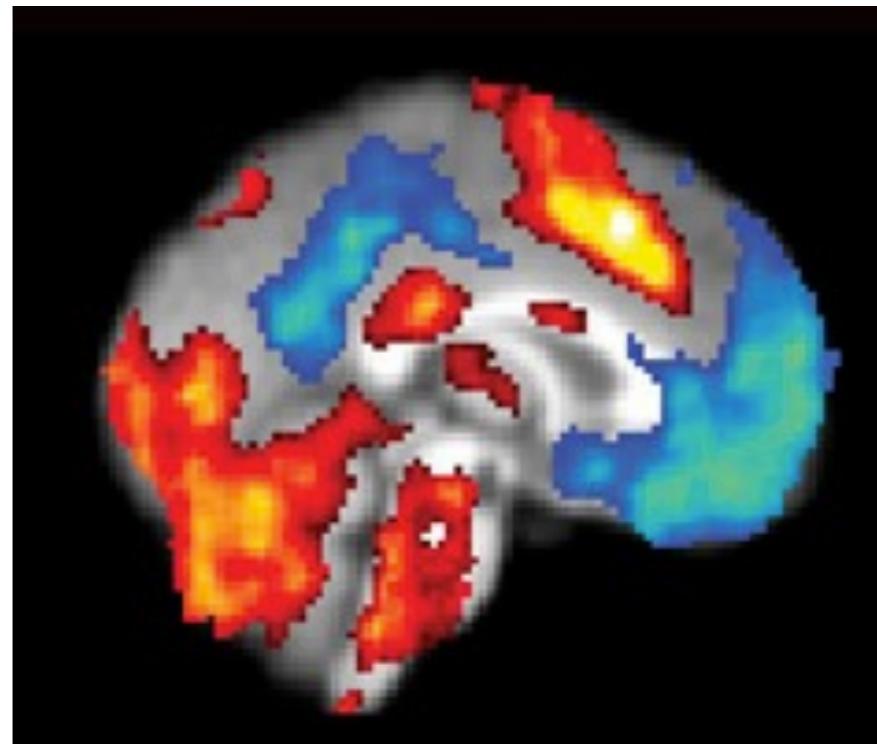




## Local analyses of regionally-specific effects

### **Mass-univariate analysis:**

Conventional analysis based on the general linear model



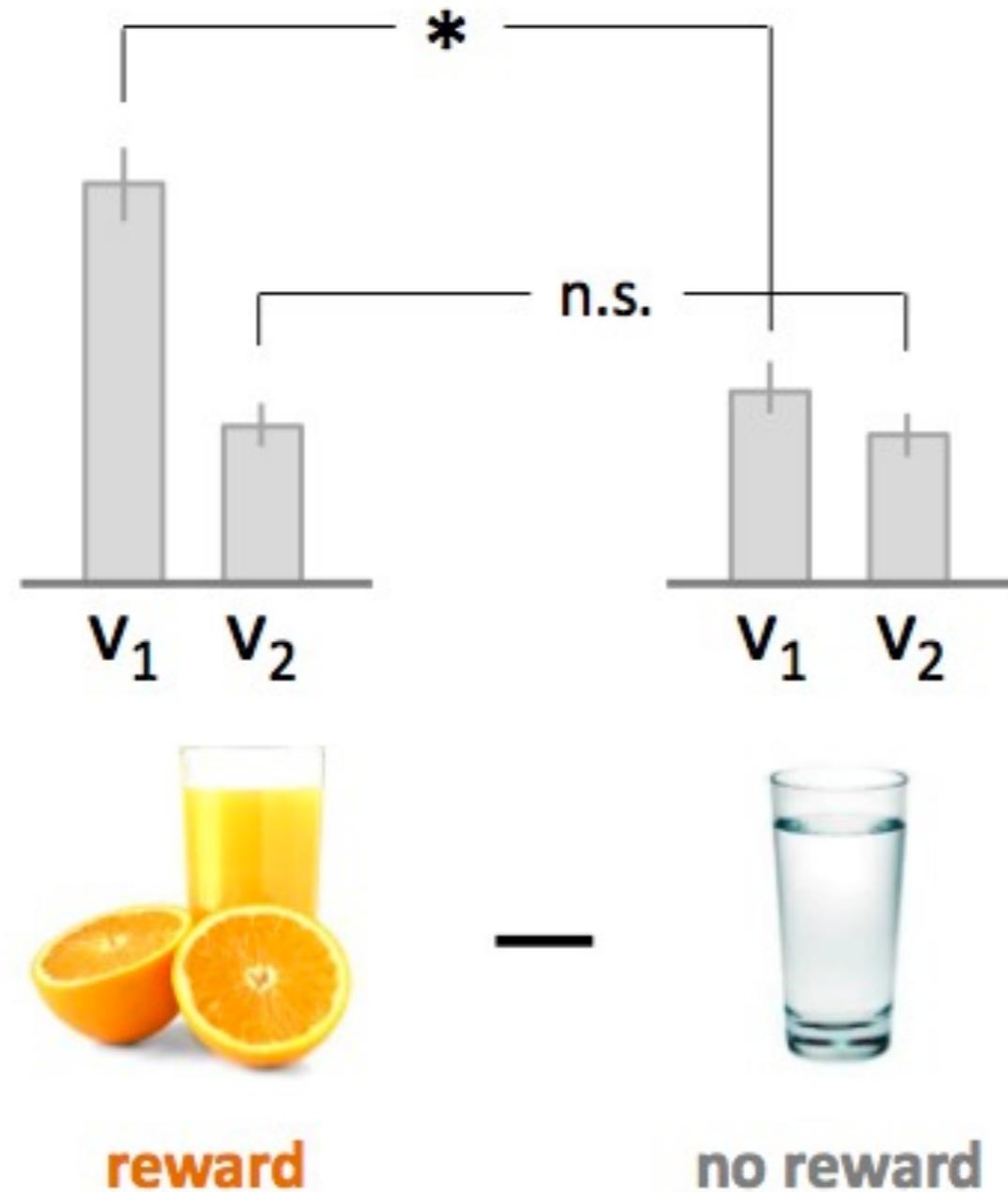
### **Multivariate analysis:**

Multivariate analysis which takes distributed patterns of activity into account

Which regions are activated during a cognitive task?



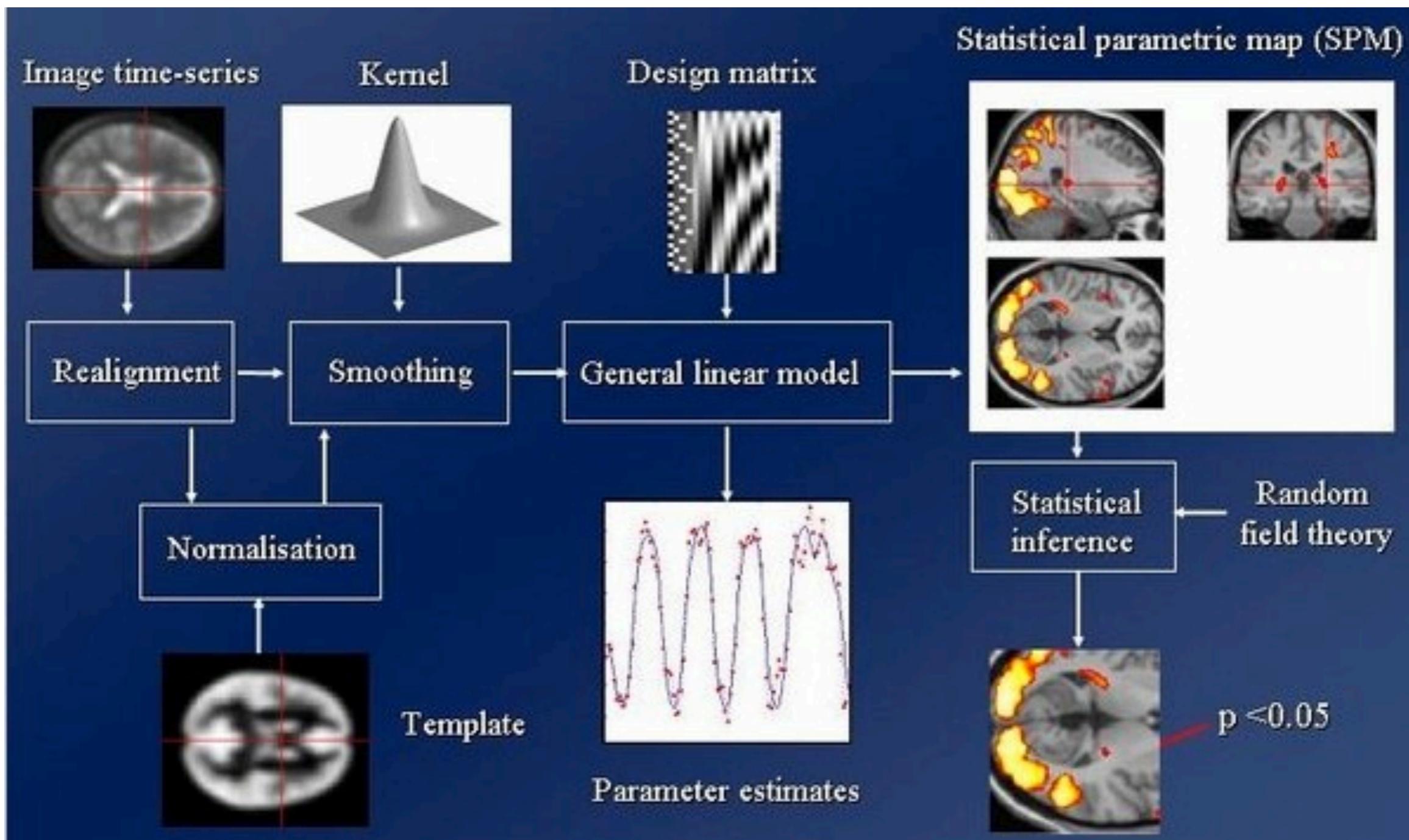
- analyzes each voxel independently
- excellent for localizing robust activations in individual voxels



Slide courtesy Kai Brodersen



# Mass-univariate analysis processing pipeline



[www.fil.ion.ucl.ac.uk/spm](http://www.fil.ion.ucl.ac.uk/spm)



## Univariate analysis of regionally-specific effects

$N \times K$

**Y**

voxel  
time-series

$N \times P$     $P \times K$

**X**  
**B**

design matrix   regression coefficients

$N \times K$

**U**

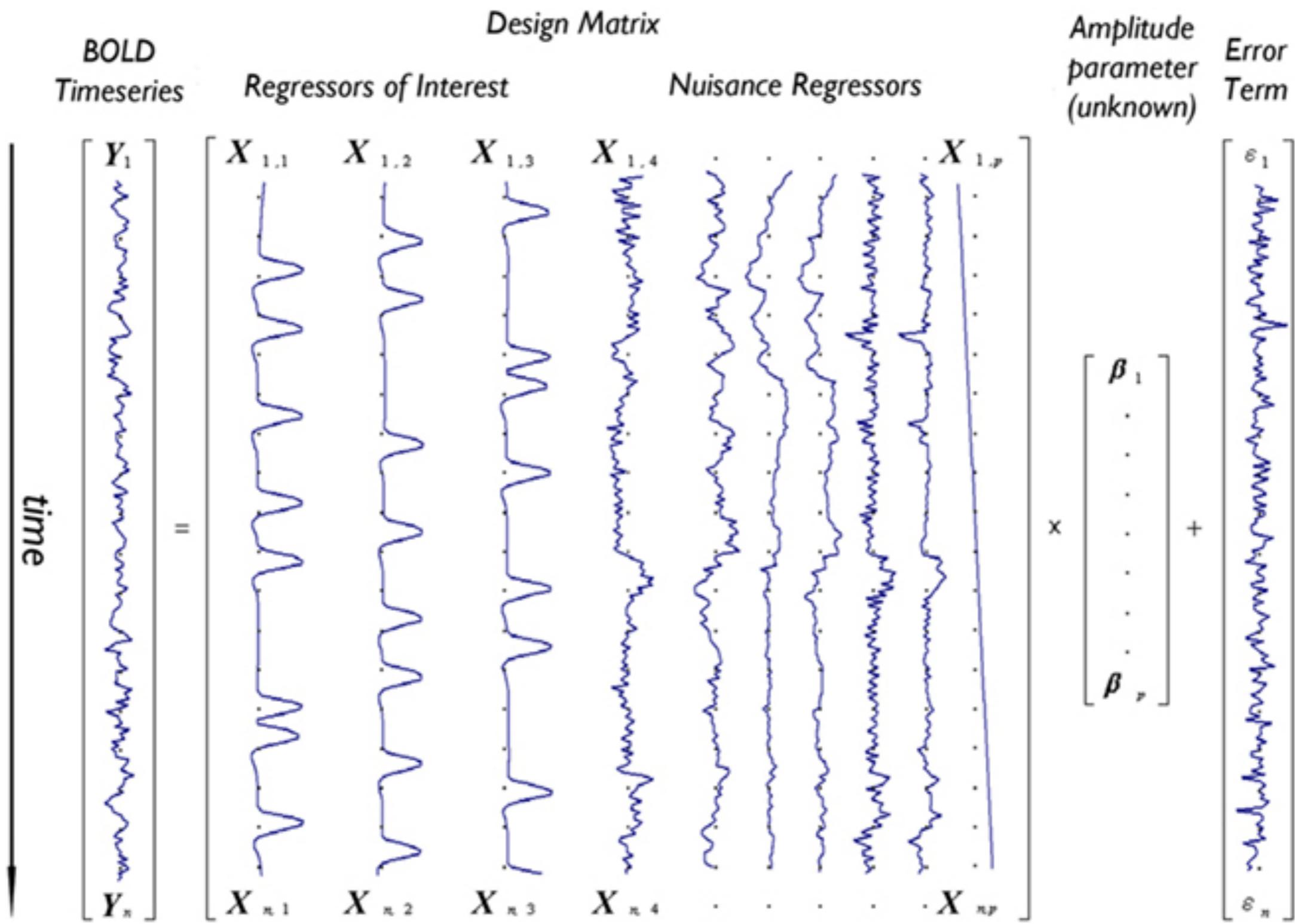
residual

$N$  = number of time points

$K$  = number of voxels

$P$  = number of regressors

## Example for one voxel





Estimation of the regression coefficients via ordinary least squares:

$$\hat{\beta} = (X^T X)^{-1} X^T y$$

Regression coefficients inform about which regressors predict a response

Interest is normally in some of the regressors; modeled using a contrast vector:

$c^T = [1 \ 0 \ 0 \dots]$  contribution of first regressor to the data

$c^T = [1 \ -1 \ 0 \dots]$  whether the first regressor is more active than the second



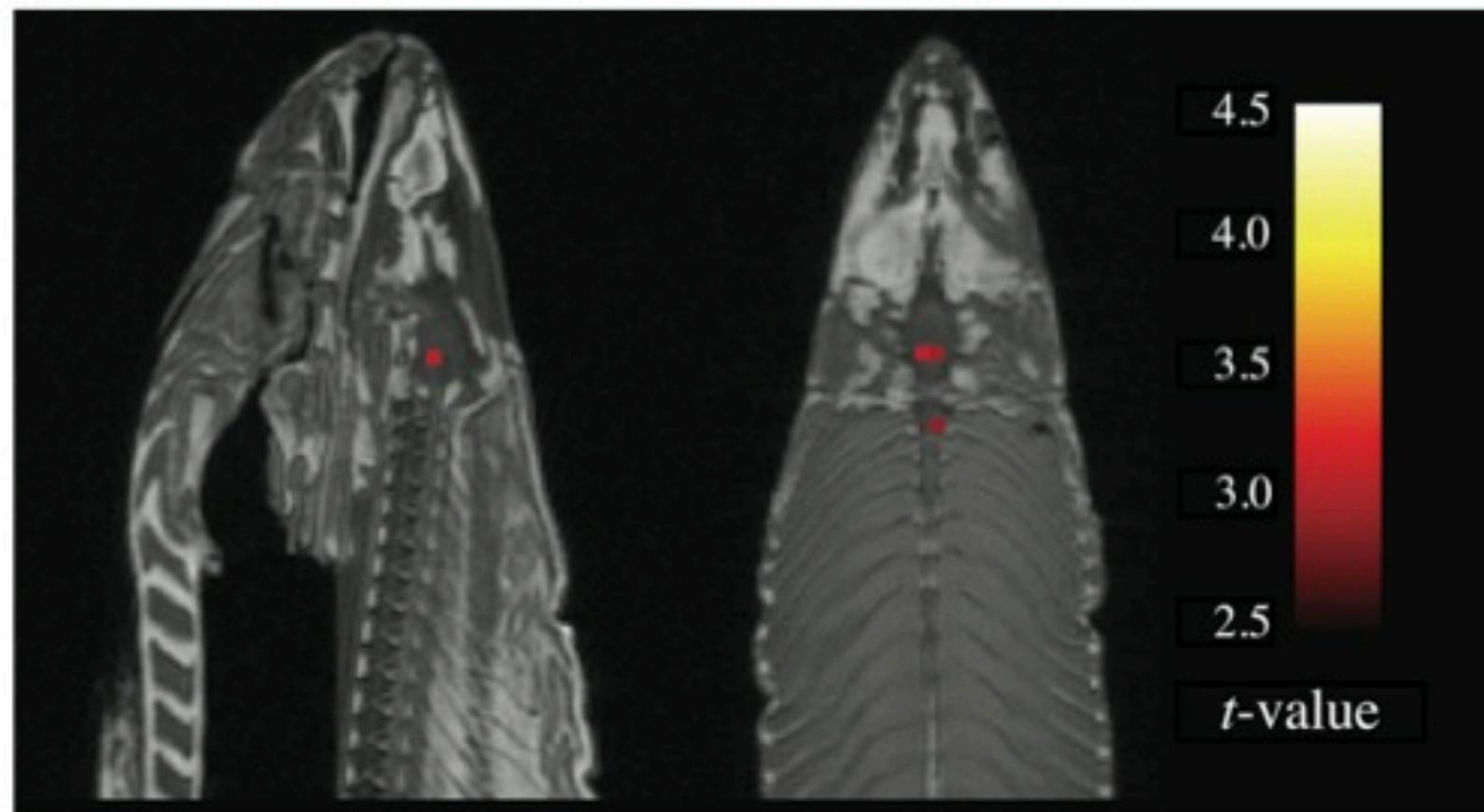
Whether a voxel is active under a contrast can (for example) be estimated using a t-statistic:

$$t_{df} = \frac{c^T \hat{\beta}}{\sqrt{\text{var}(c^T \hat{\beta})}}$$

For Gaussian errors,  $t_{df}$  approximately follows a Student's t-distribution with  $df$  degrees of freedom.

The p-value of  $t_{df}$  is then computed by testing against the null-distribution.

A voxel is declared significantly activated if the p-value is below some fixed alpha level.

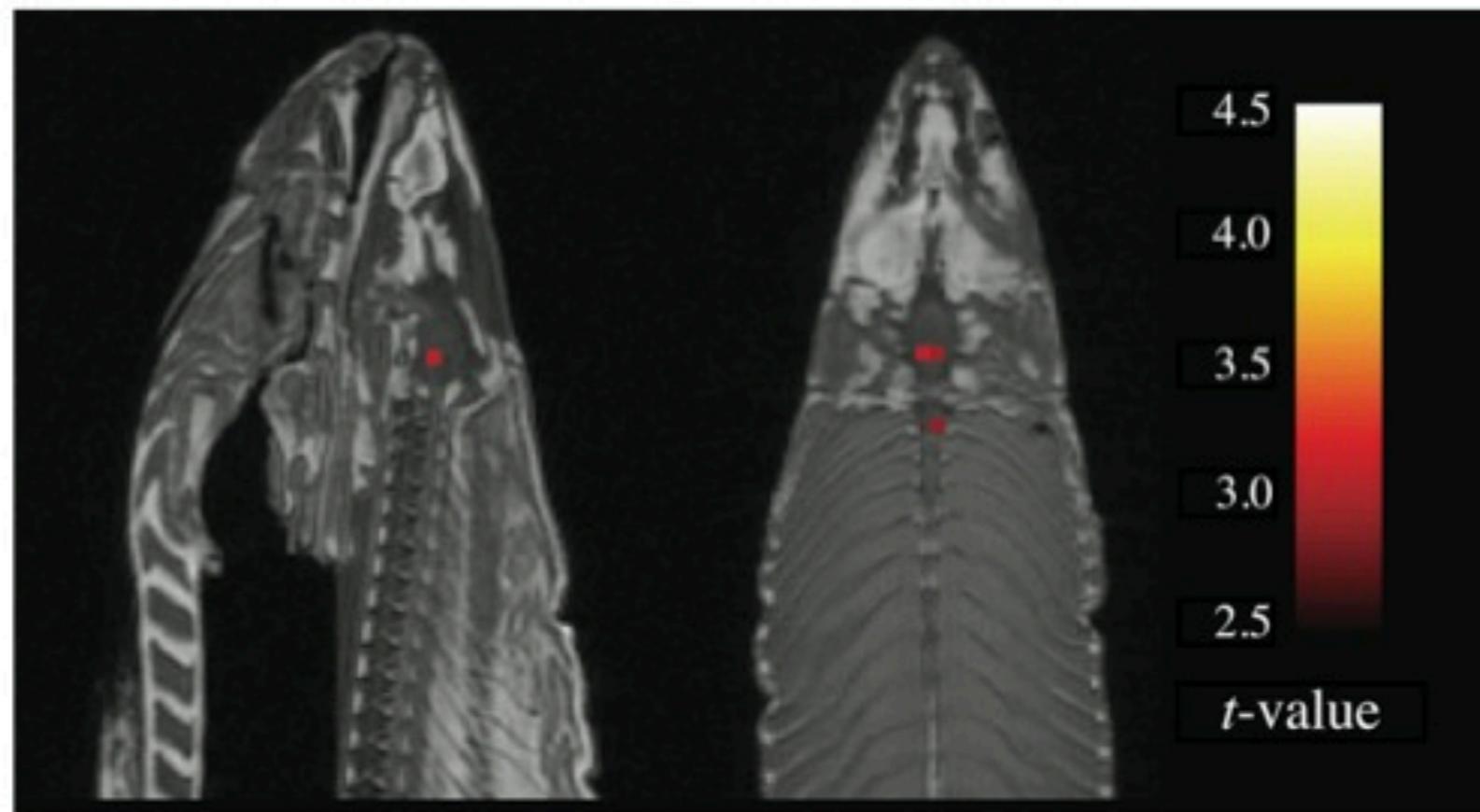


A  $t$ -contrast was used to test for regions with significant BOLD signal change during the photo condition compared to rest. The parameters for this comparison were  $t(131) > 3.15$ ,  $p(\text{uncorrected}) < 0.001$ , 3 voxel extent threshold.

## **Neural Correlates of Interspecies Perspective Taking in the Post-Mortem Atlantic Salmon: An Argument For Proper Multiple Comparisons Correction**

by: [Craig M. Bennett](#), [Abigail A. Baird](#), [Michael B. Miller](#), [George L. Wolford](#)

*Journal of Serendipitous and Unexpected Results*, Vol. 1 (2011), pp. 1-5 Key: citeulike:9170380



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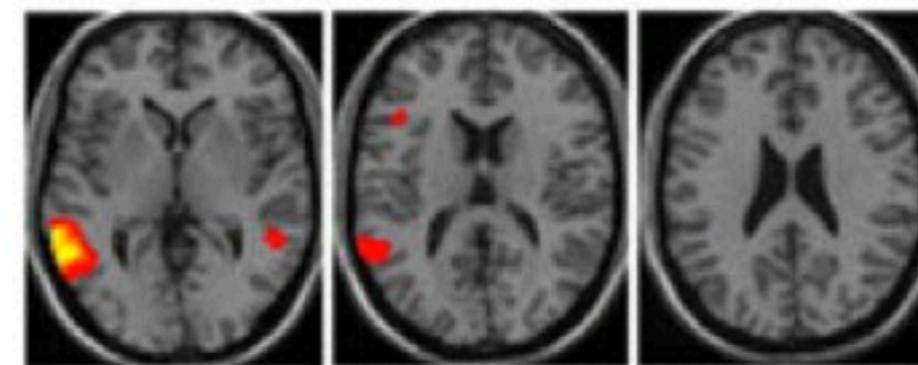
*Journal of Serendipitous and Unexpected Results*, Vol. 1 (2011), pp. 1-5 Key: citeulike:9170380

A voxel is declared significantly activated if the p-value is below some fixed alpha level, corrected for multiple comparisons.

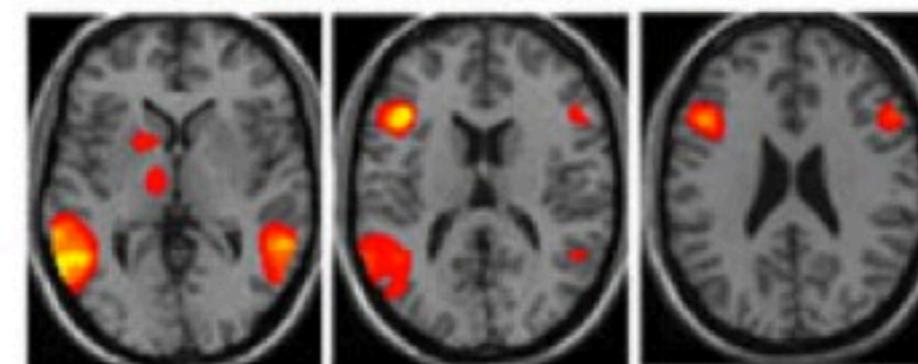
## Example



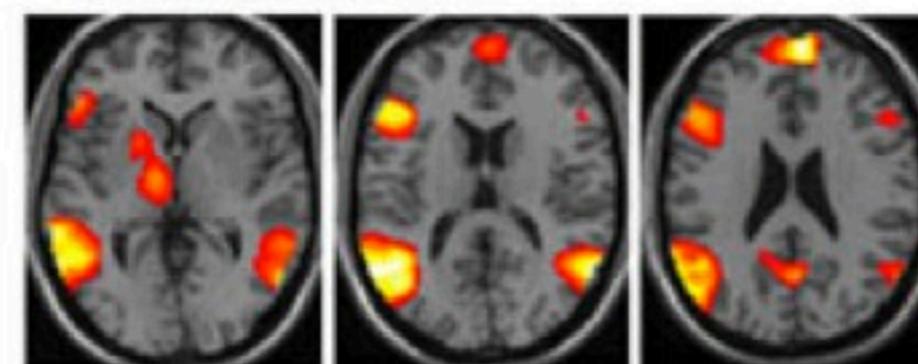
Reading Words



Reading Sentences



Reading Stories



2 mm

16 mm

24 mm

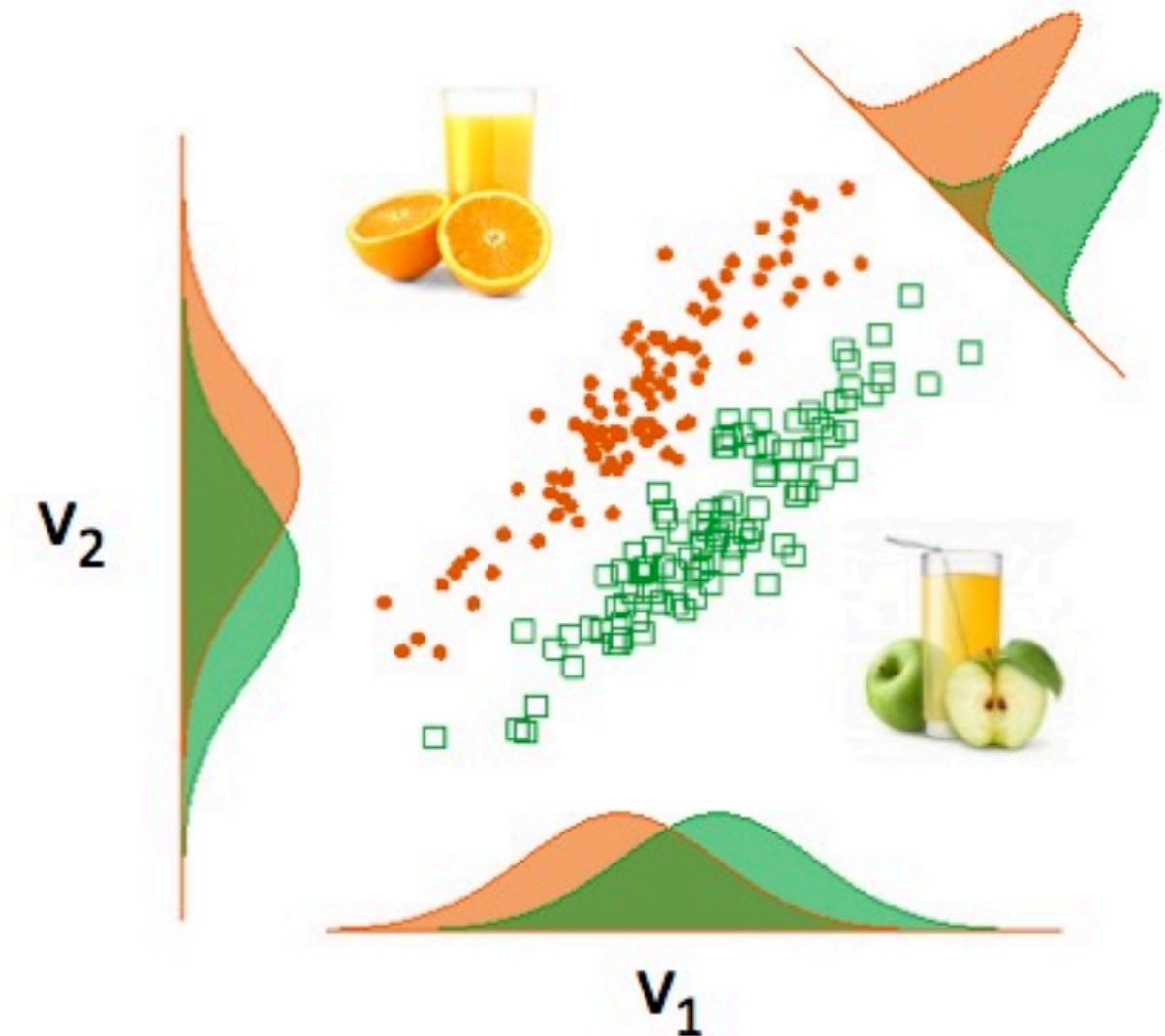
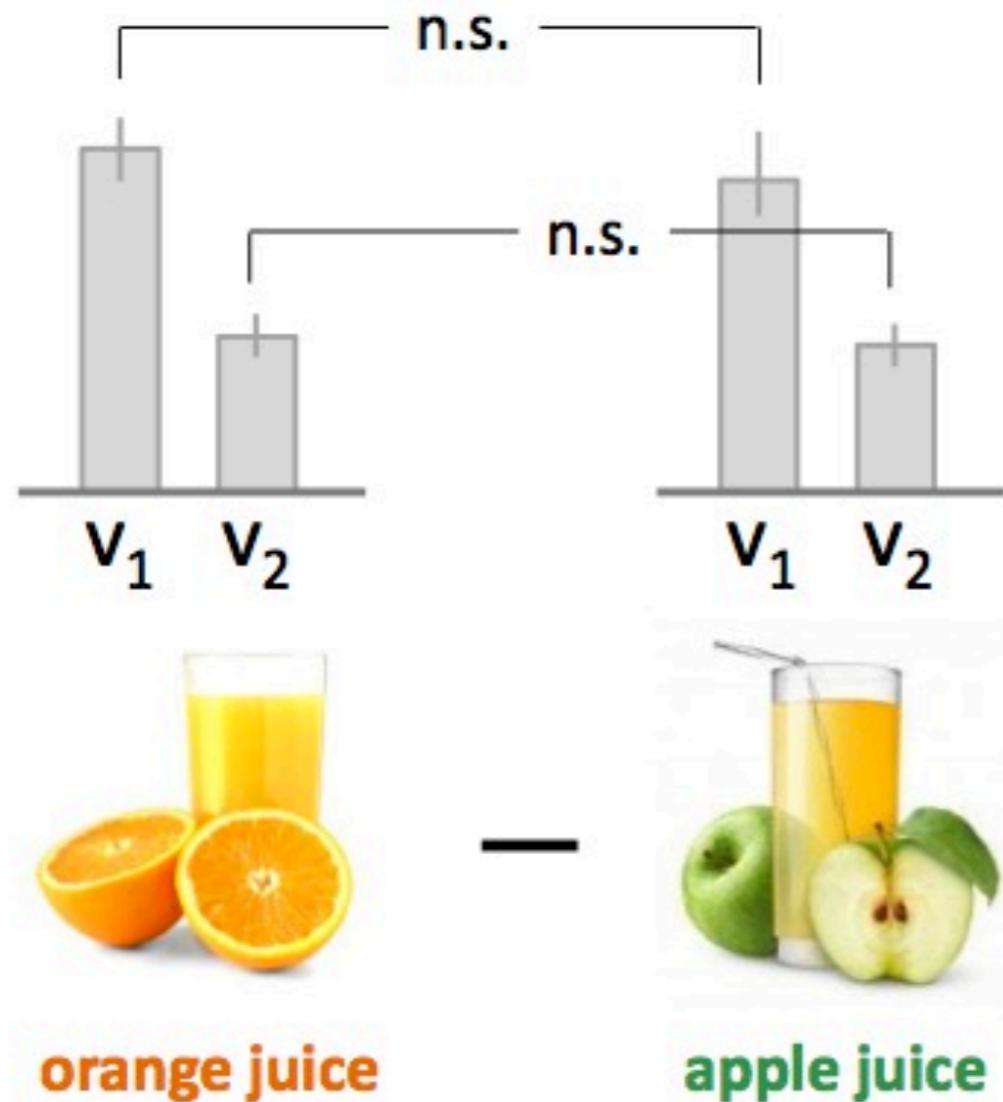
Left

Right

See Poline J-B, Brett M. The general linear model and fMRI: Does love last forever? Neuroimage. 2012 for a gentle overview.



- examine responses that are jointly encoded in multiple voxels



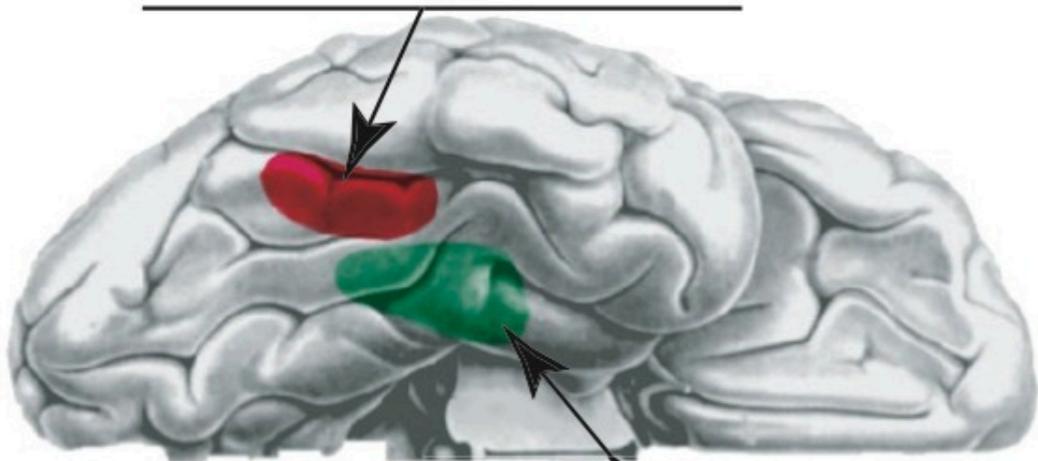
Slide courtesy Kai Brodersen



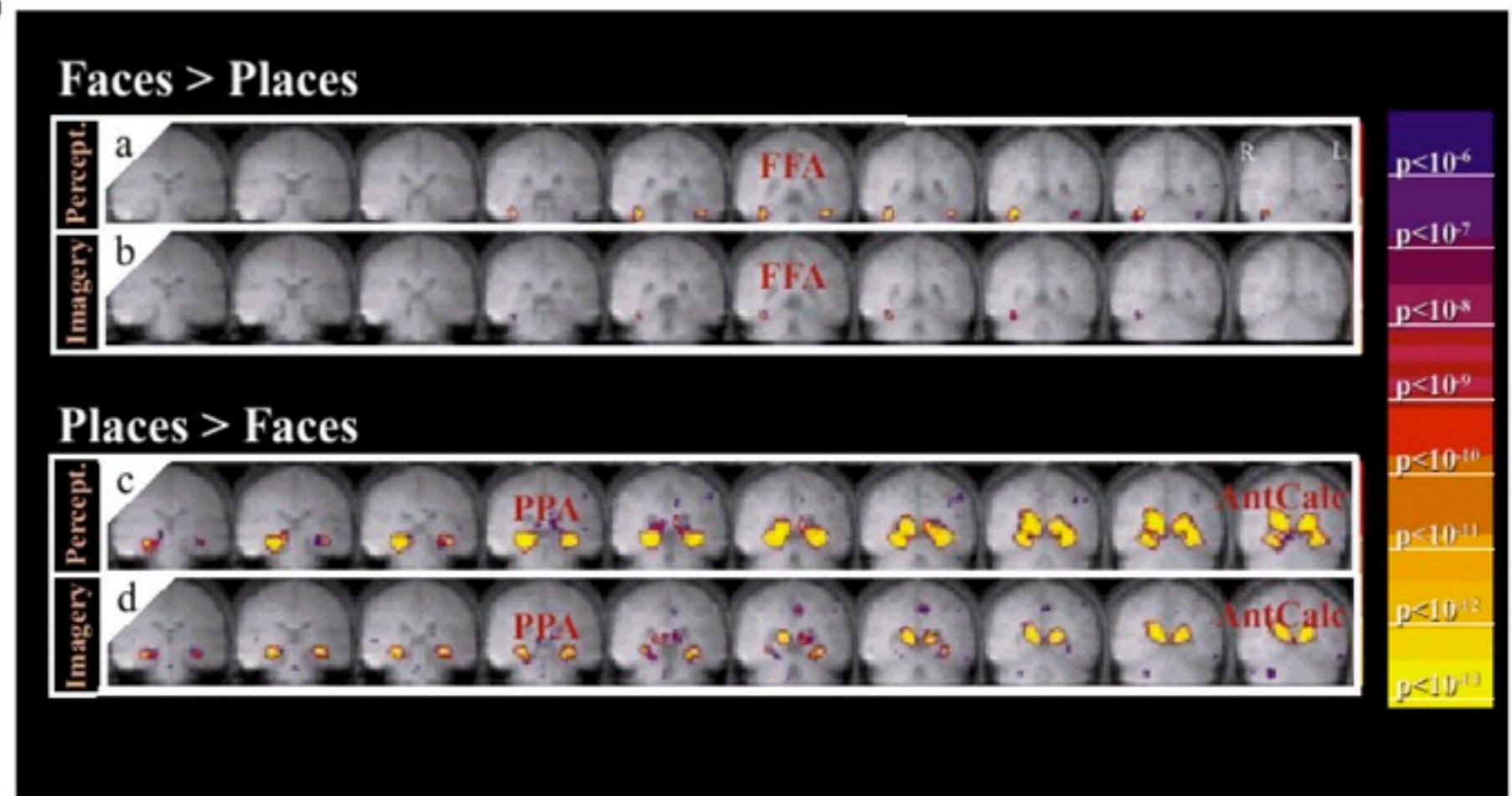
# Faces versus places



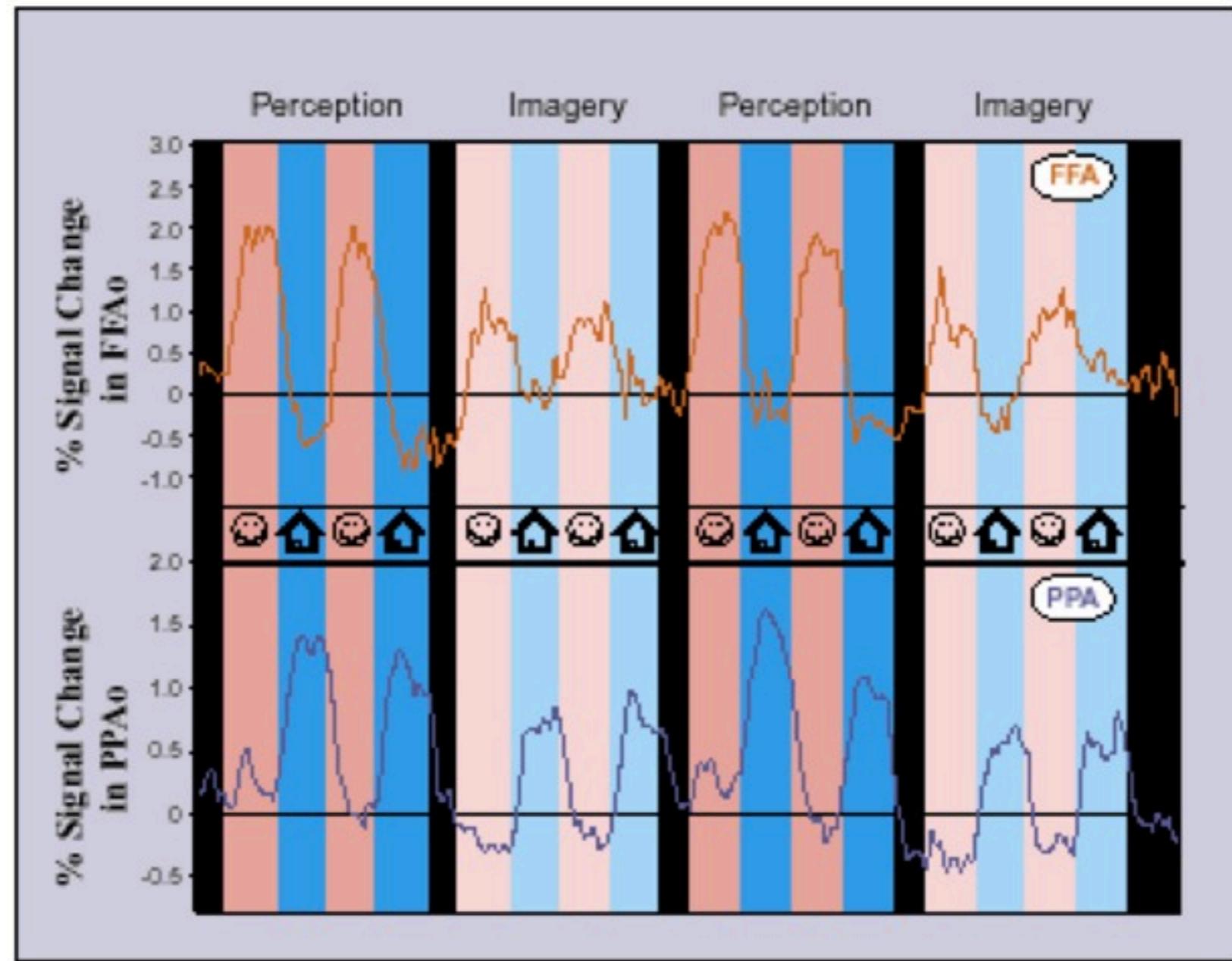
Fusiform Face Area (FFA)  
/ Visual Expertise



Parahippocampal  
Place Area (PPA)

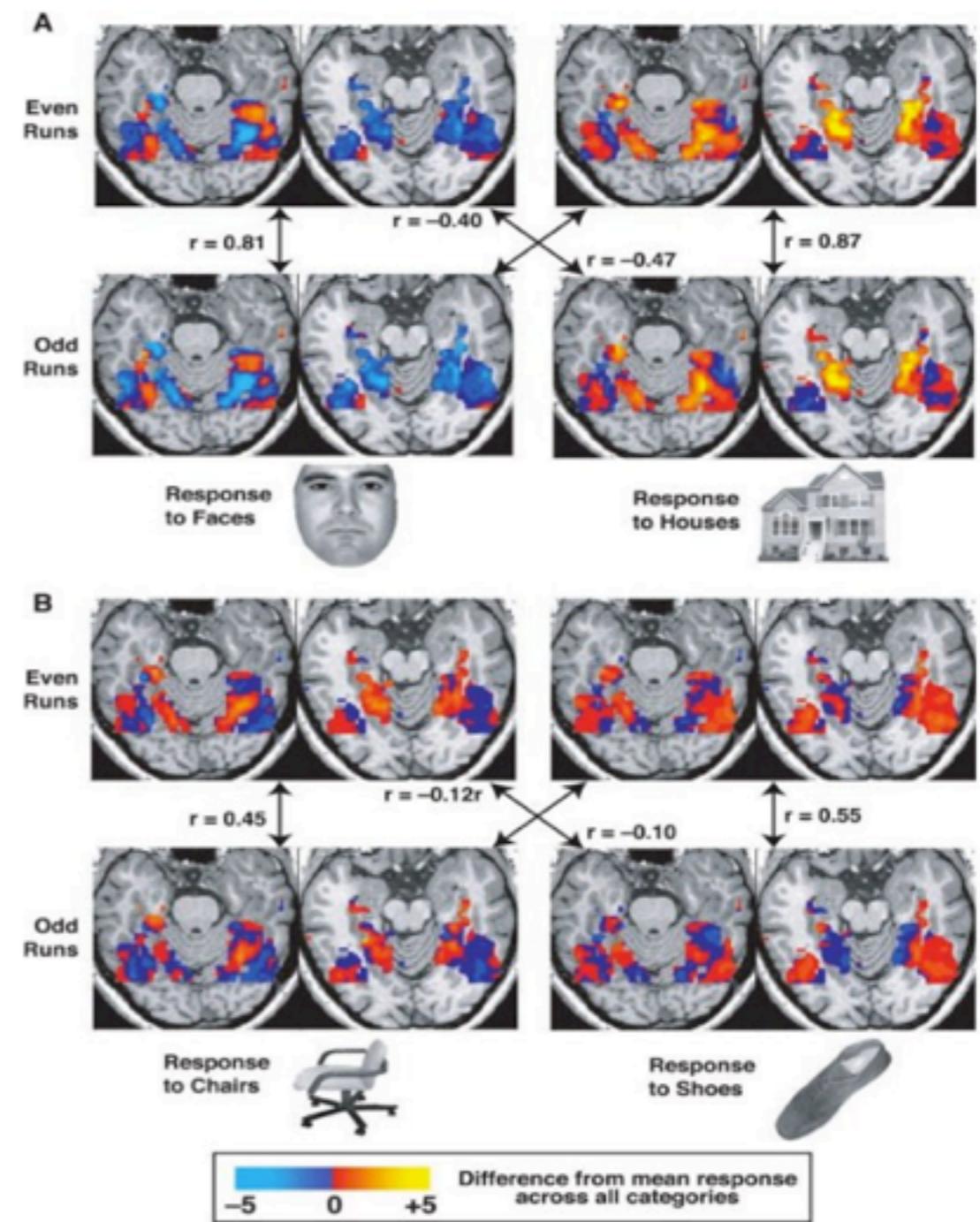
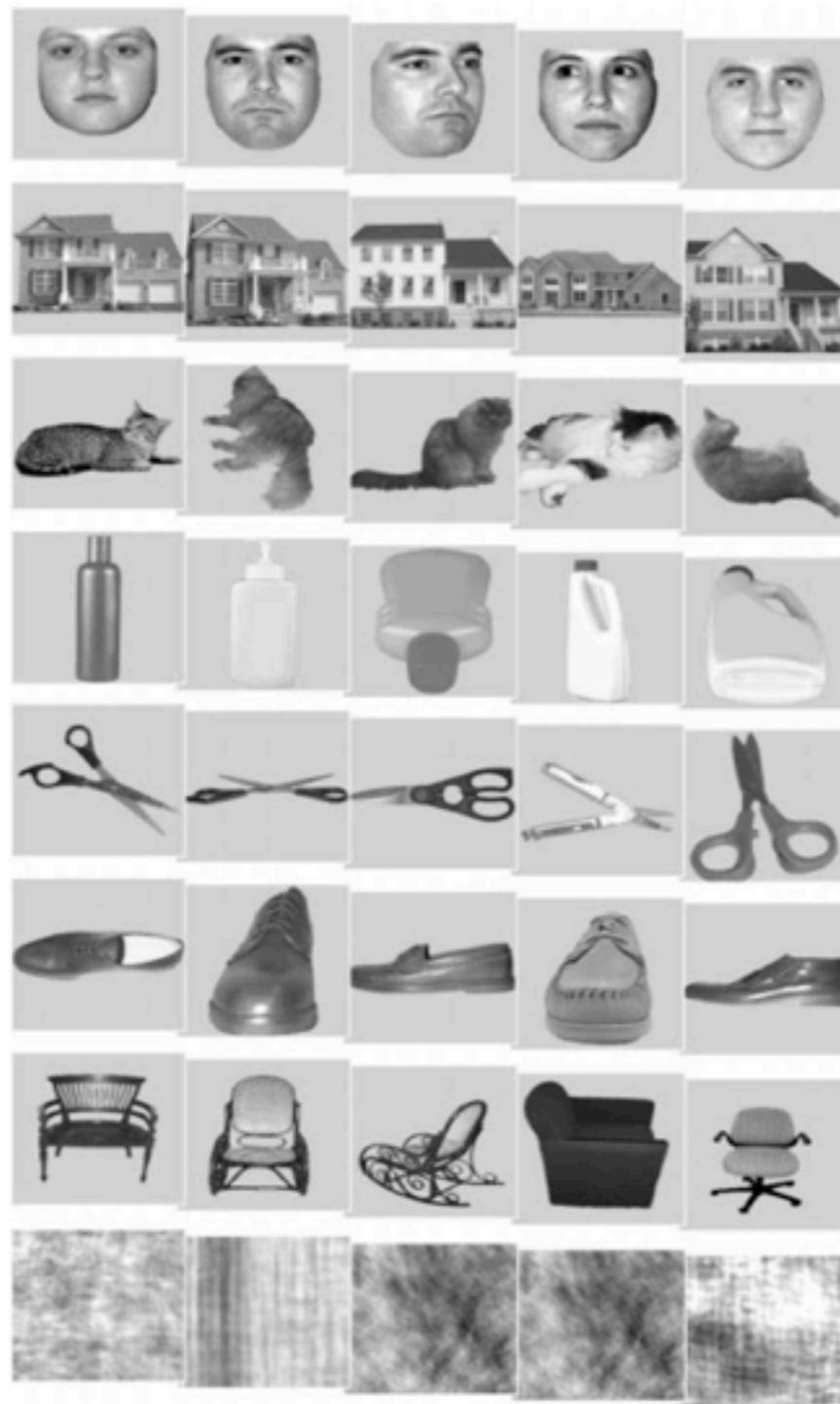


O'Craven and Kanwisher, J Cog Neu, 2000

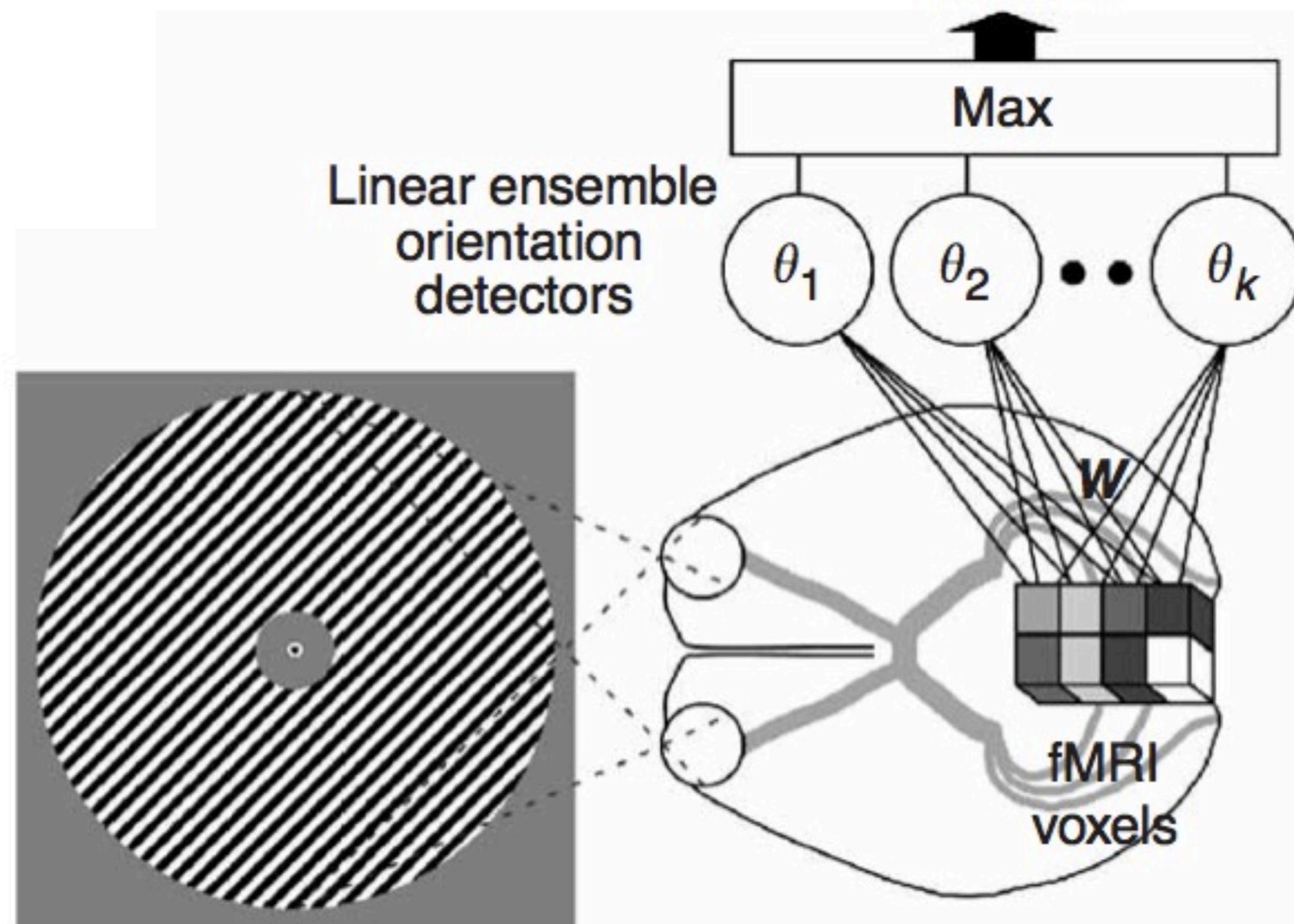


David Hume saw the relationship between percepts and images as follows: “*The difference betwixt these consists in the degrees of force and liveliness, with which they strike upon the mind . . . [Perceptions] enter with most force and violence . . . By ideas I mean the faint images of these in thinking and reasoning*” (Hume, 1739).

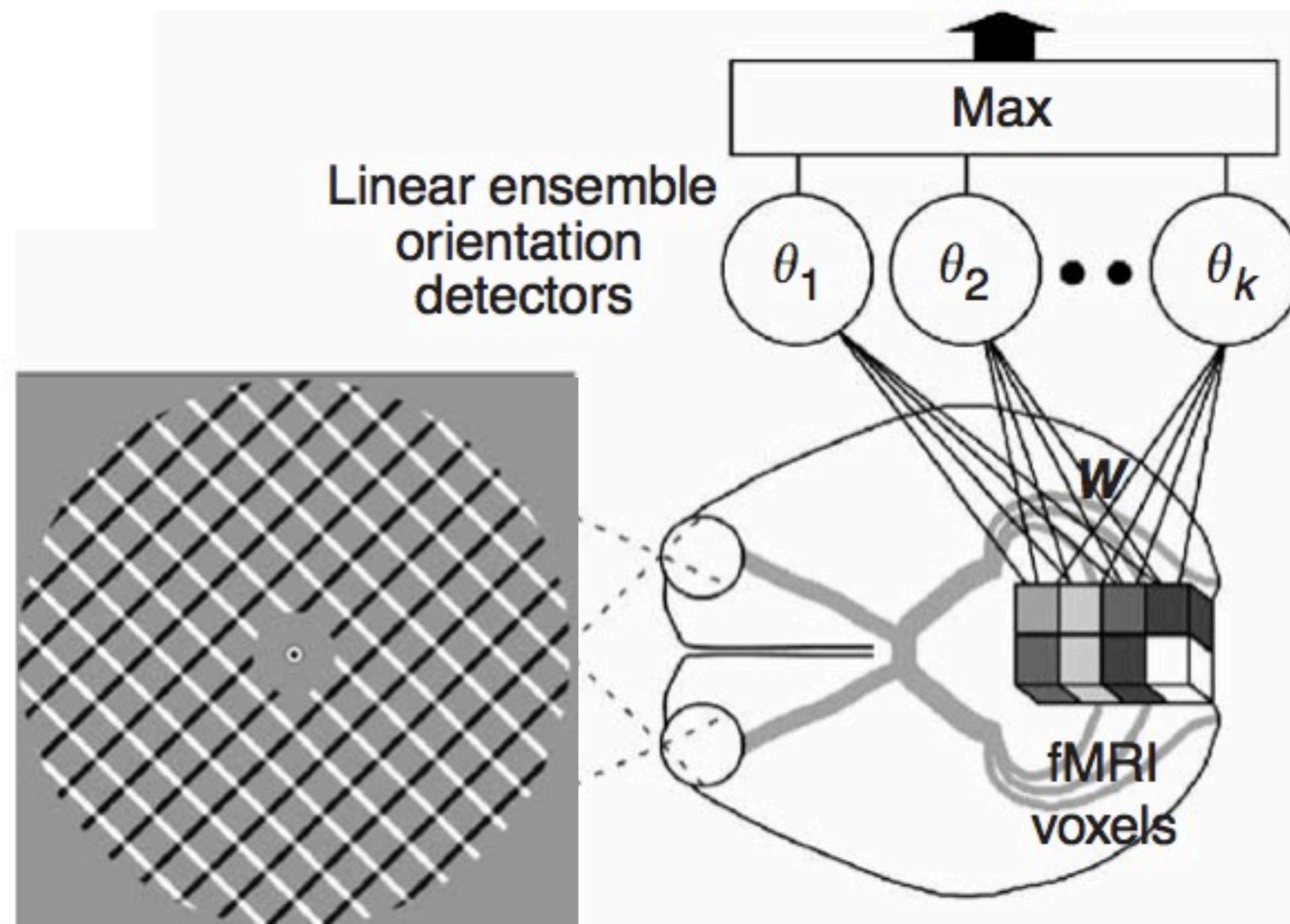
# The Haxby study



Haxby et al. Science, 2001.

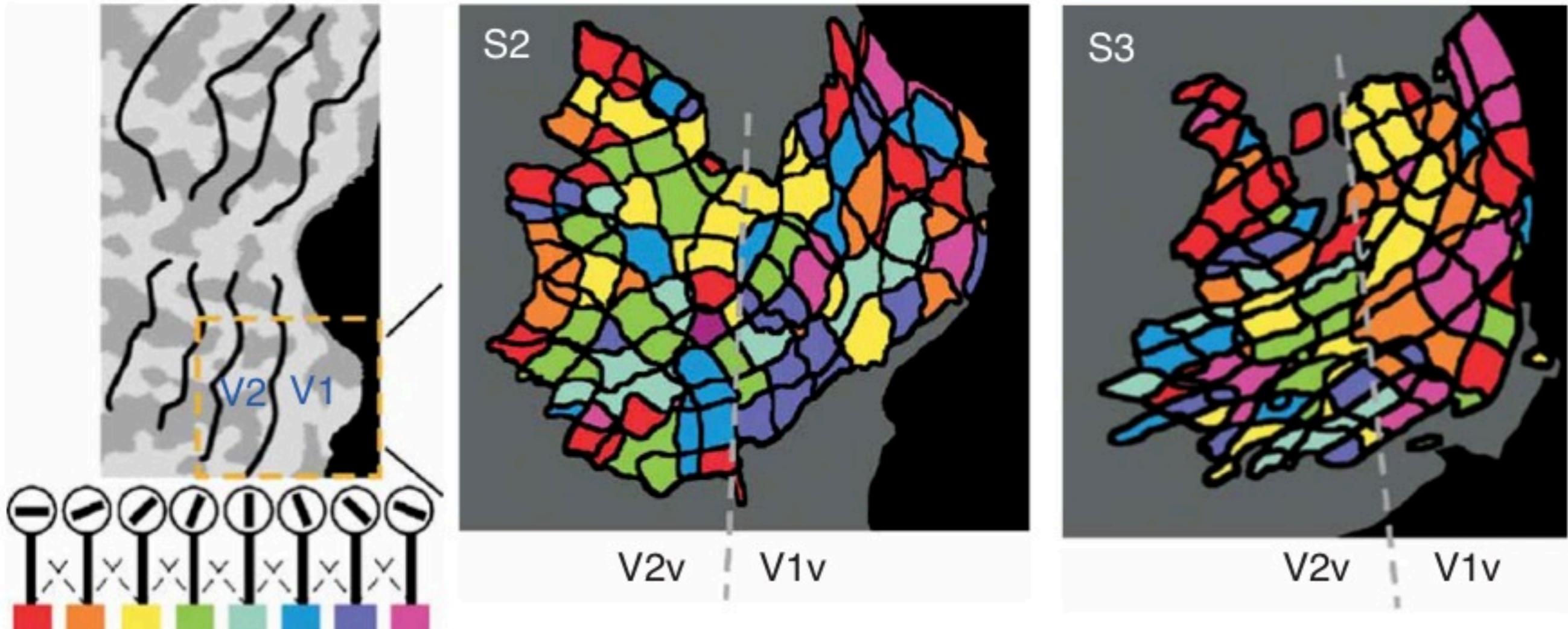


Kamitani and Tong, Nature Neuroscience, 2005



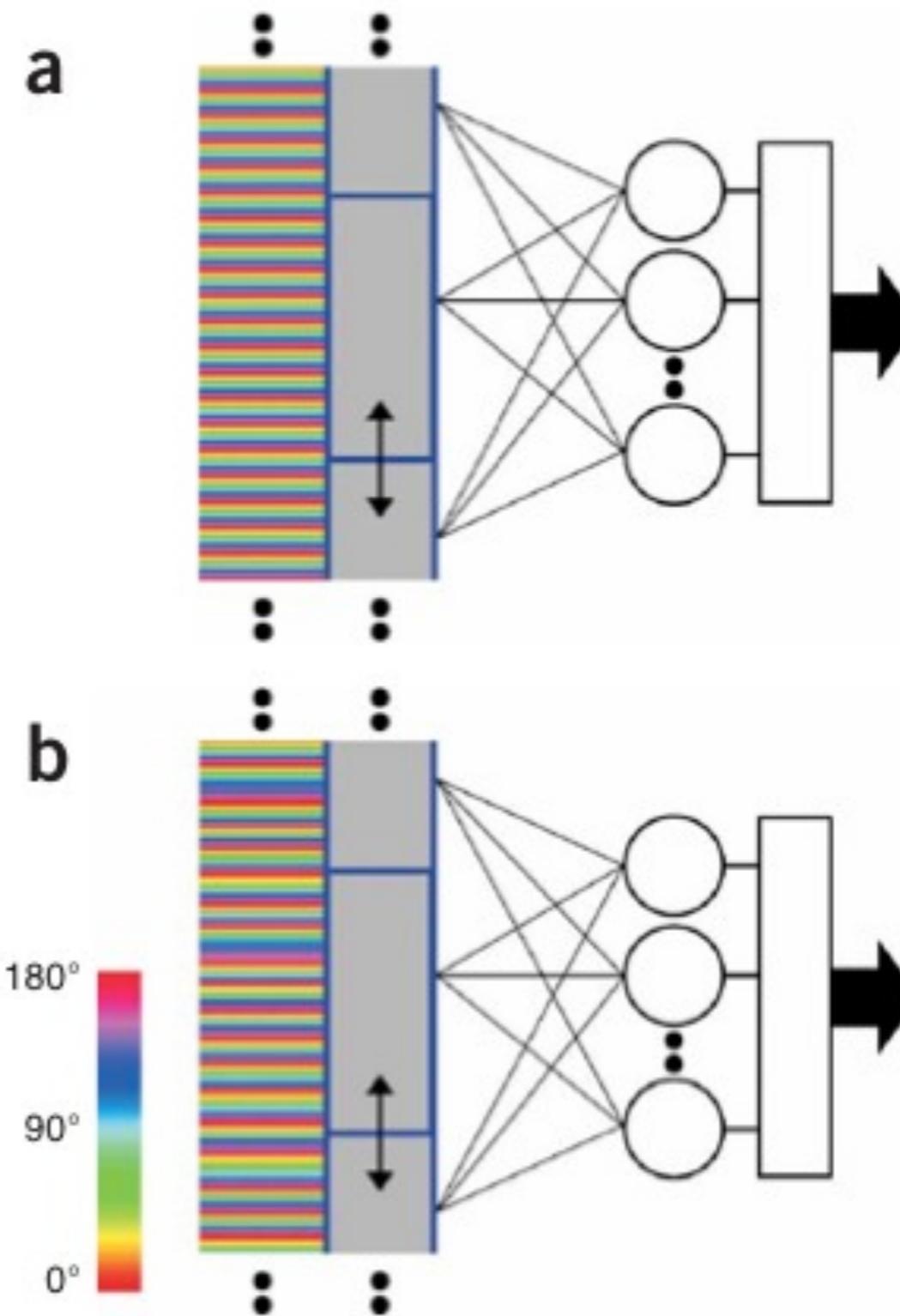
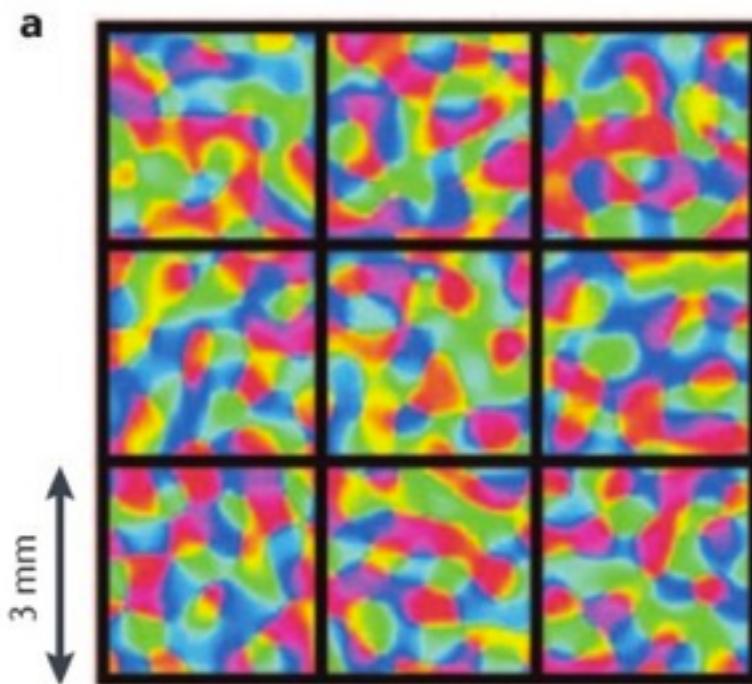
Kamitani and Tong, Nature Neuroscience, 2005

# Orientation selectivity

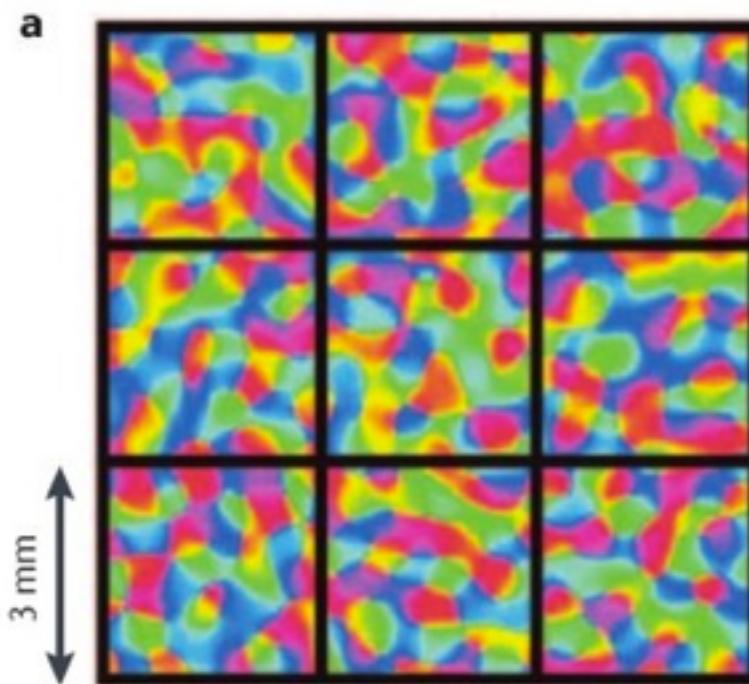


Kamitani and Tong, Nature Neuroscience, 2005

# Where does the information come from: biased sampling?

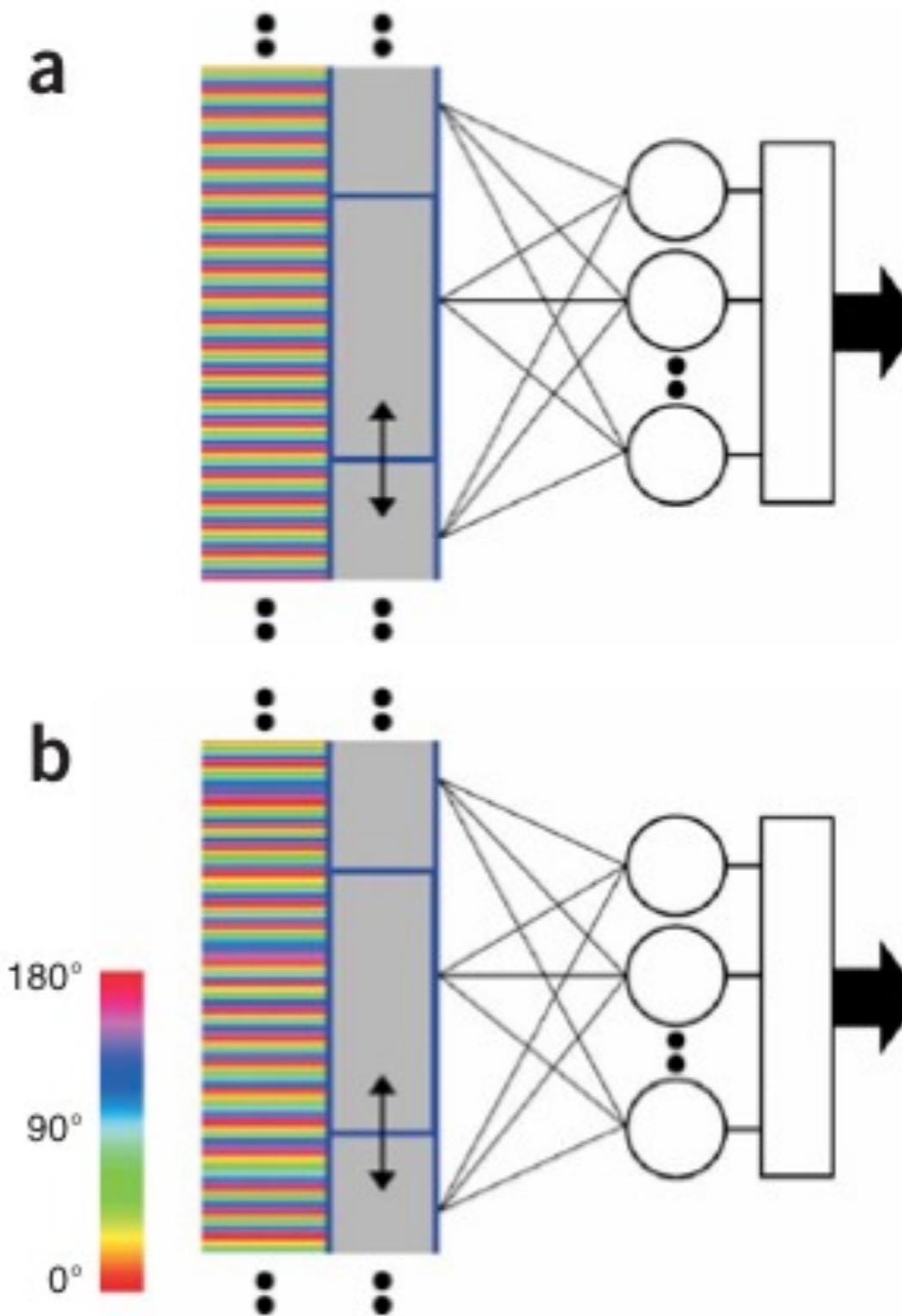


# Where does the information come from: biased sampling?

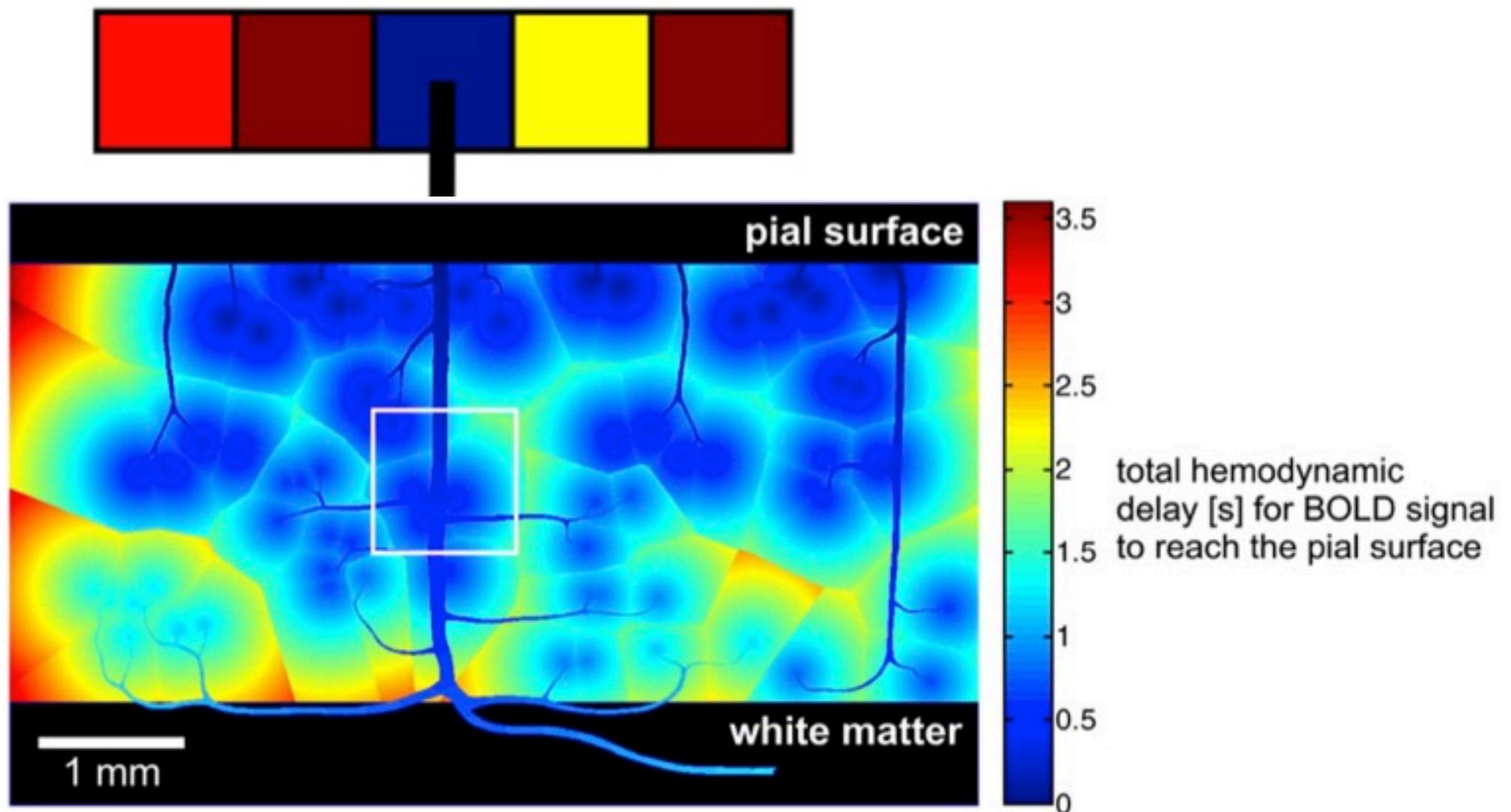


unlikely:

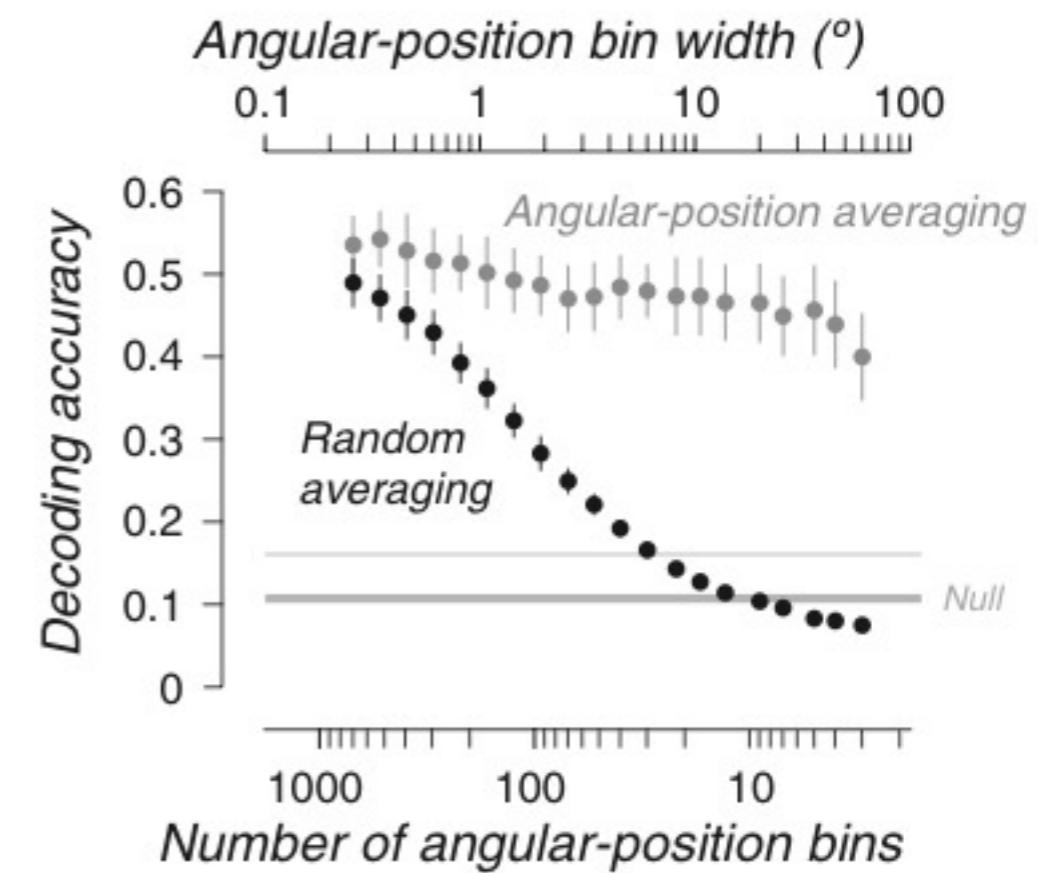
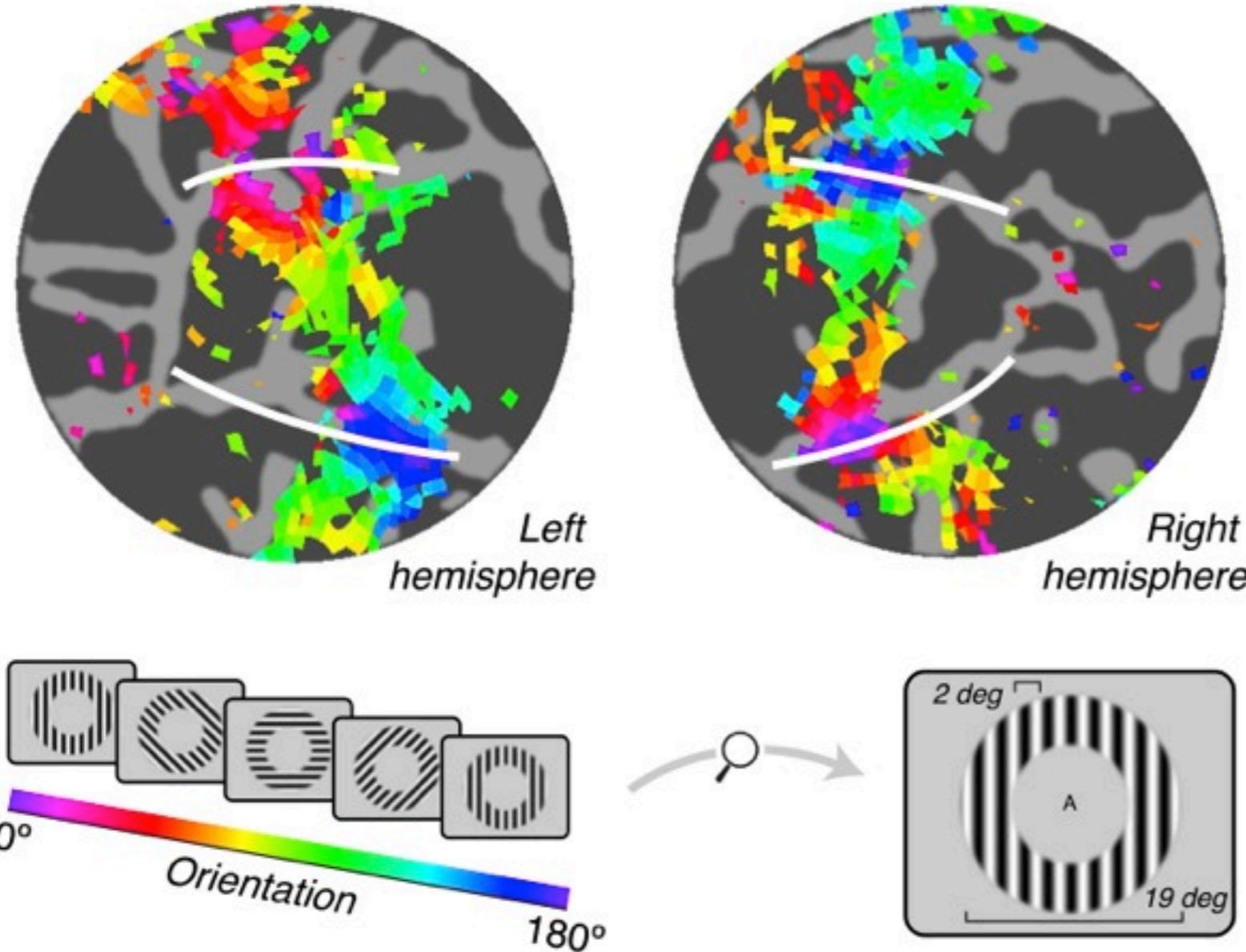
- sensitivity to draining veins
- motion artifacts

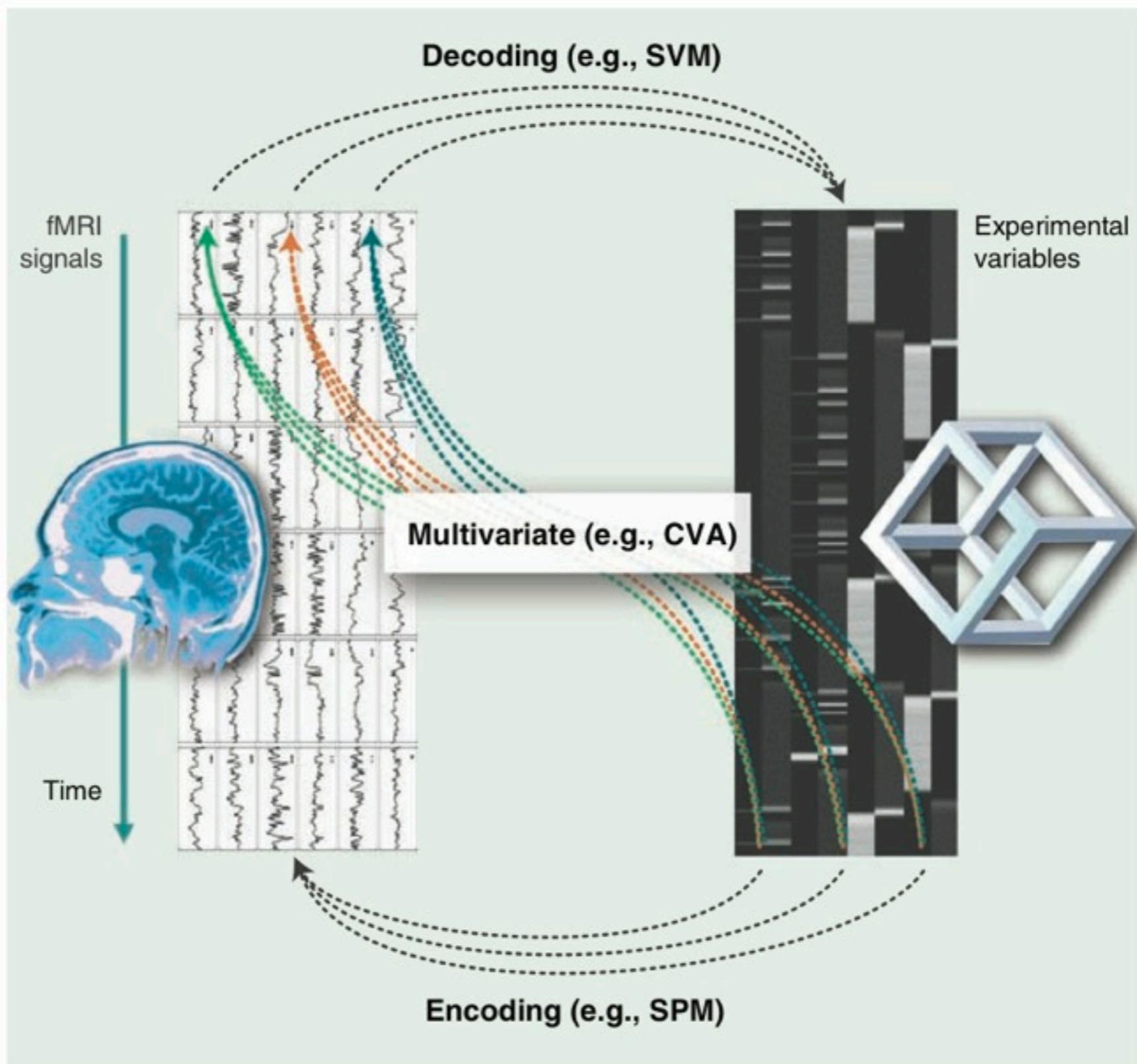


# Where does it come from: complex spatiotemporal filter?



# Where does it come from: coarse maps?

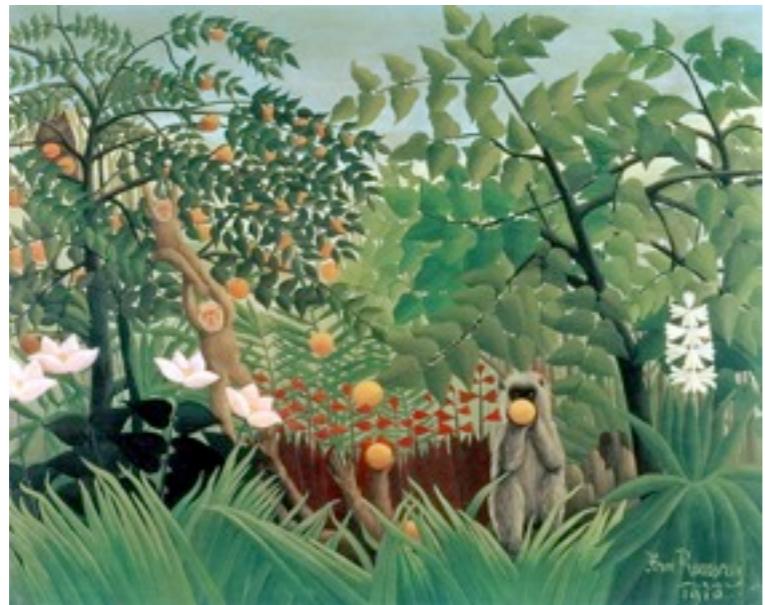




Friston, Science, 2009



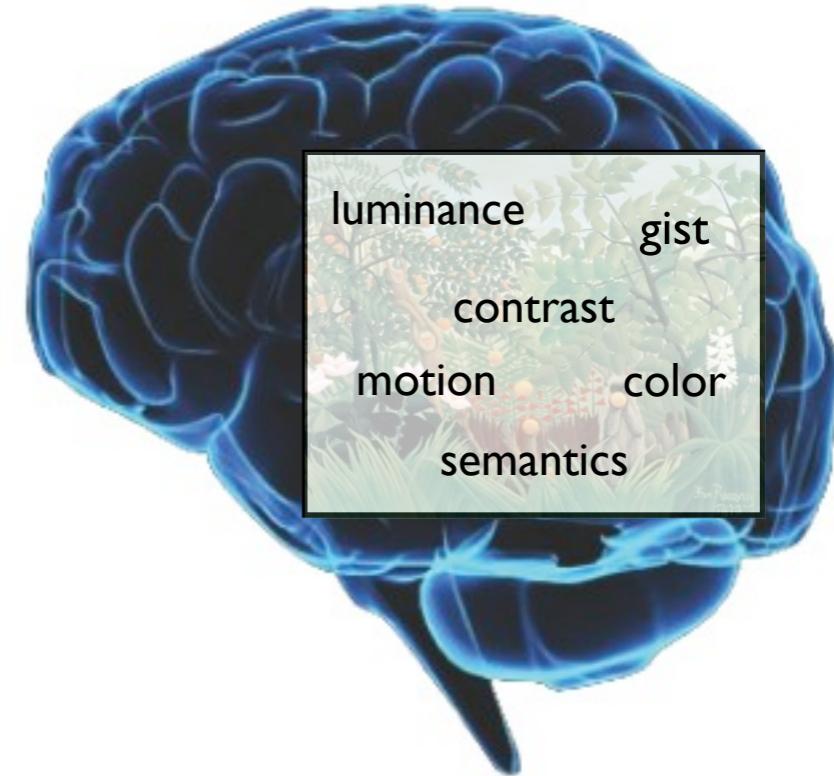
# Encoding and decoding



# Encoding and decoding



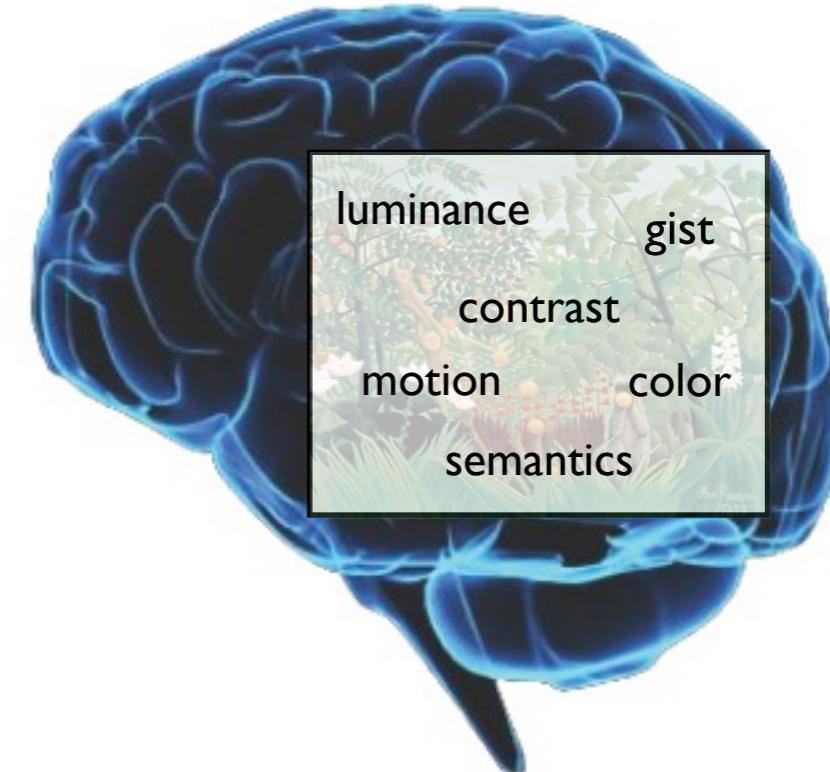
encoding



# Encoding and decoding



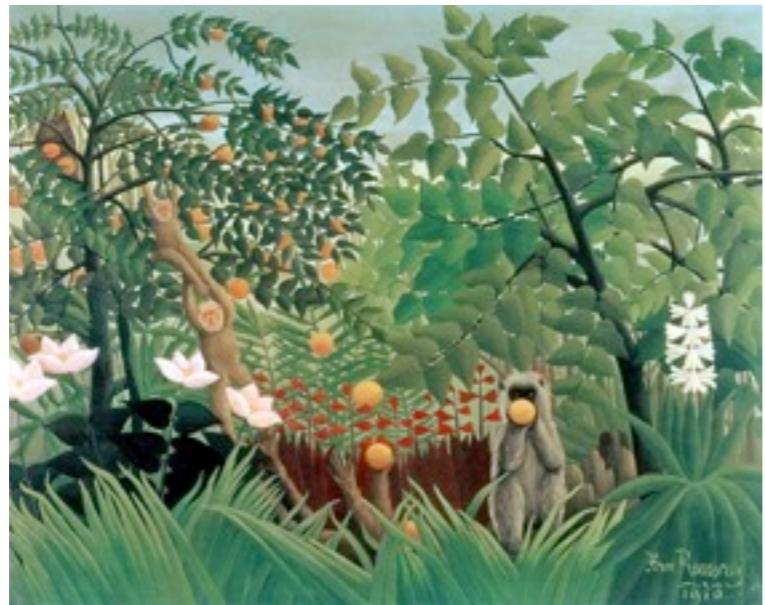
encoding



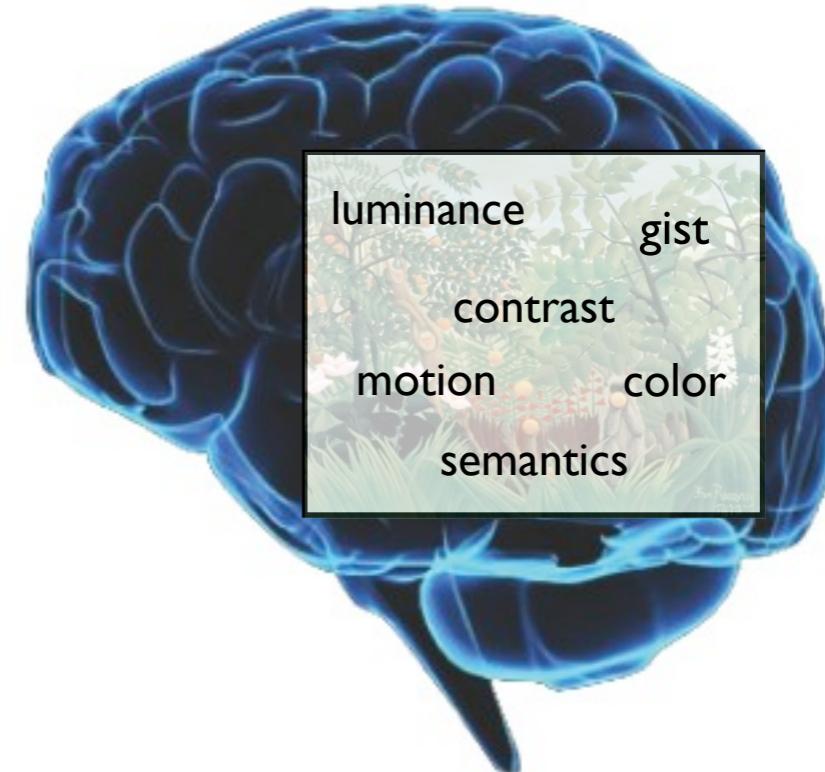
brain activity



# Encoding and decoding



encoding



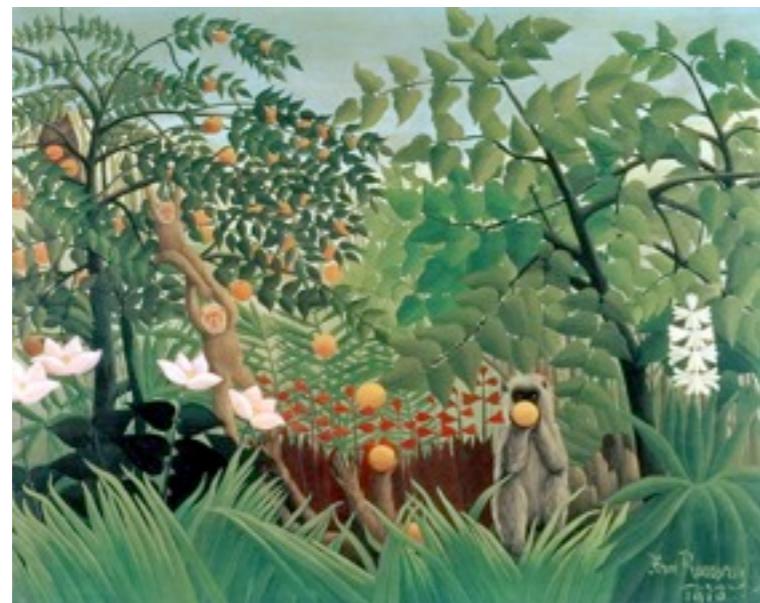
brain activity



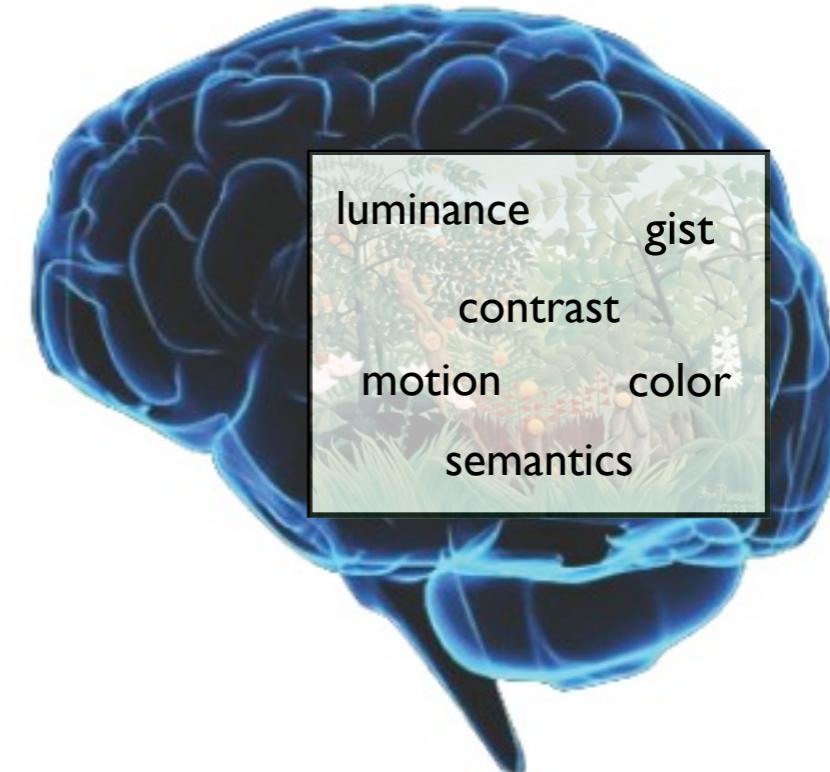
decoding



# Encoding and decoding



encoding



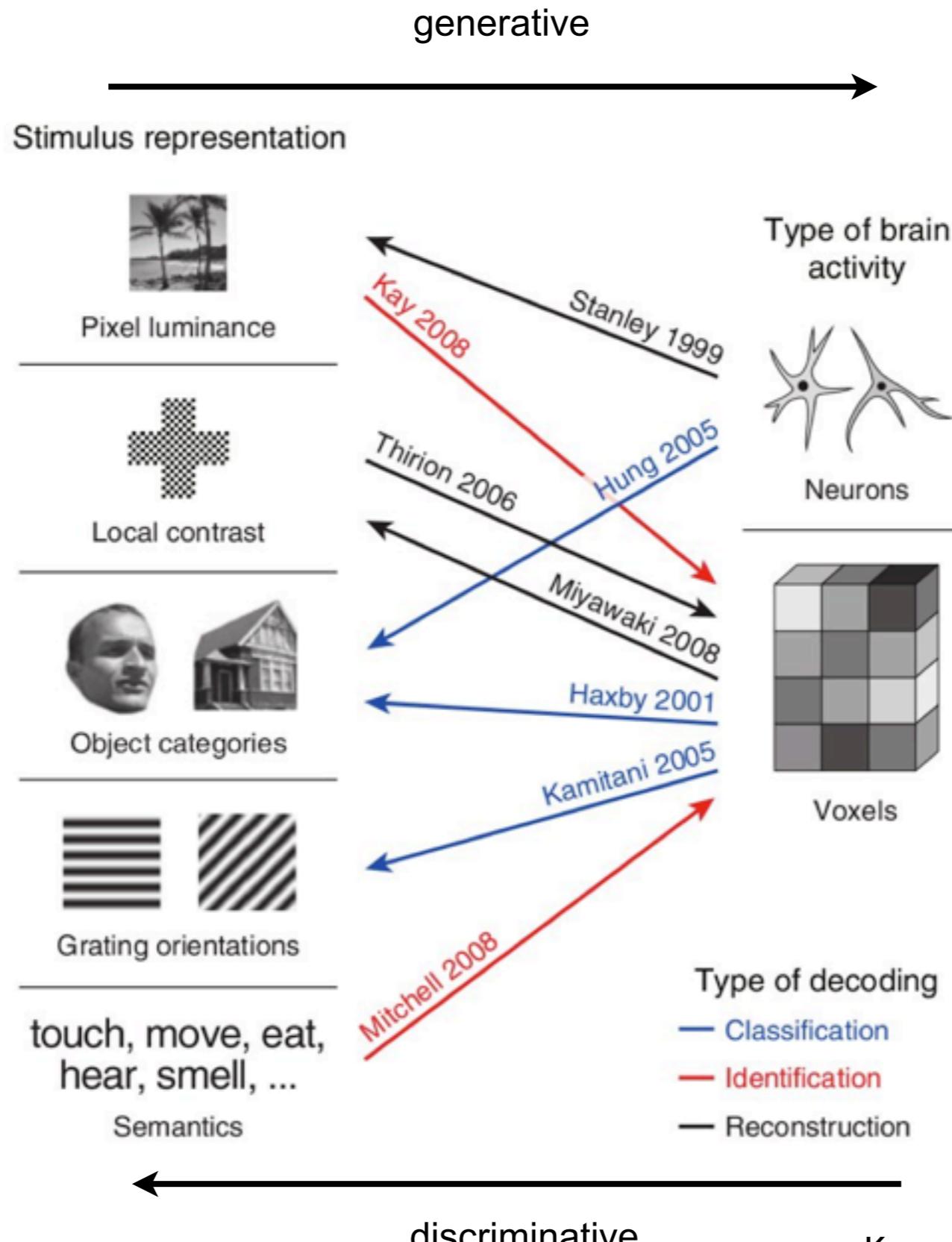
brain activity



decoding

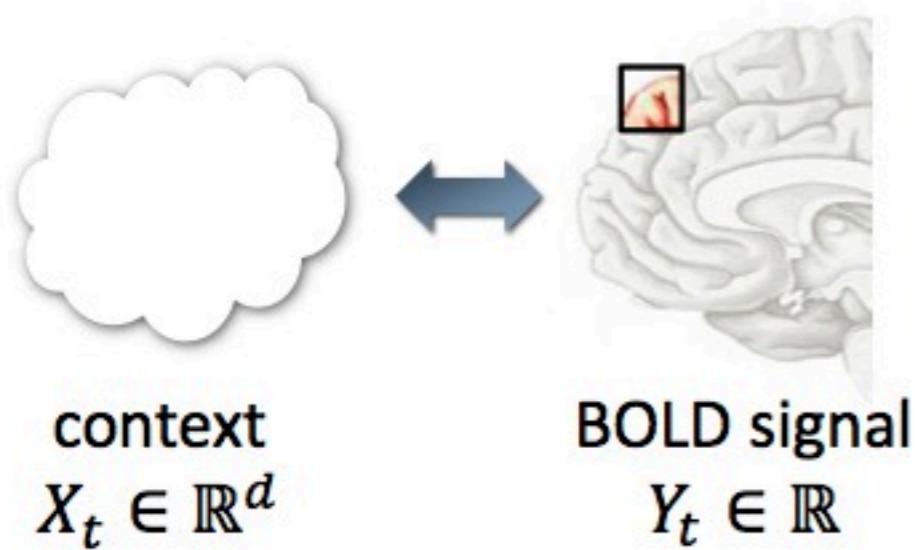


# Decoding flavors



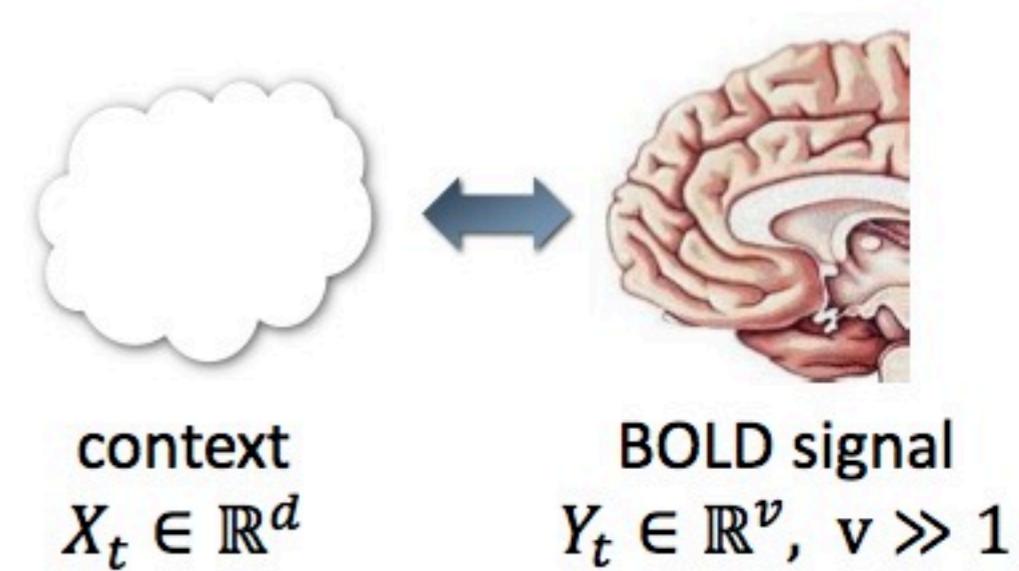


**A univariate model** considers a single voxel at a time.



Spatial dependencies between voxels are only introduced afterwards, through random field theory.

**A multivariate model** considers many voxels at once.



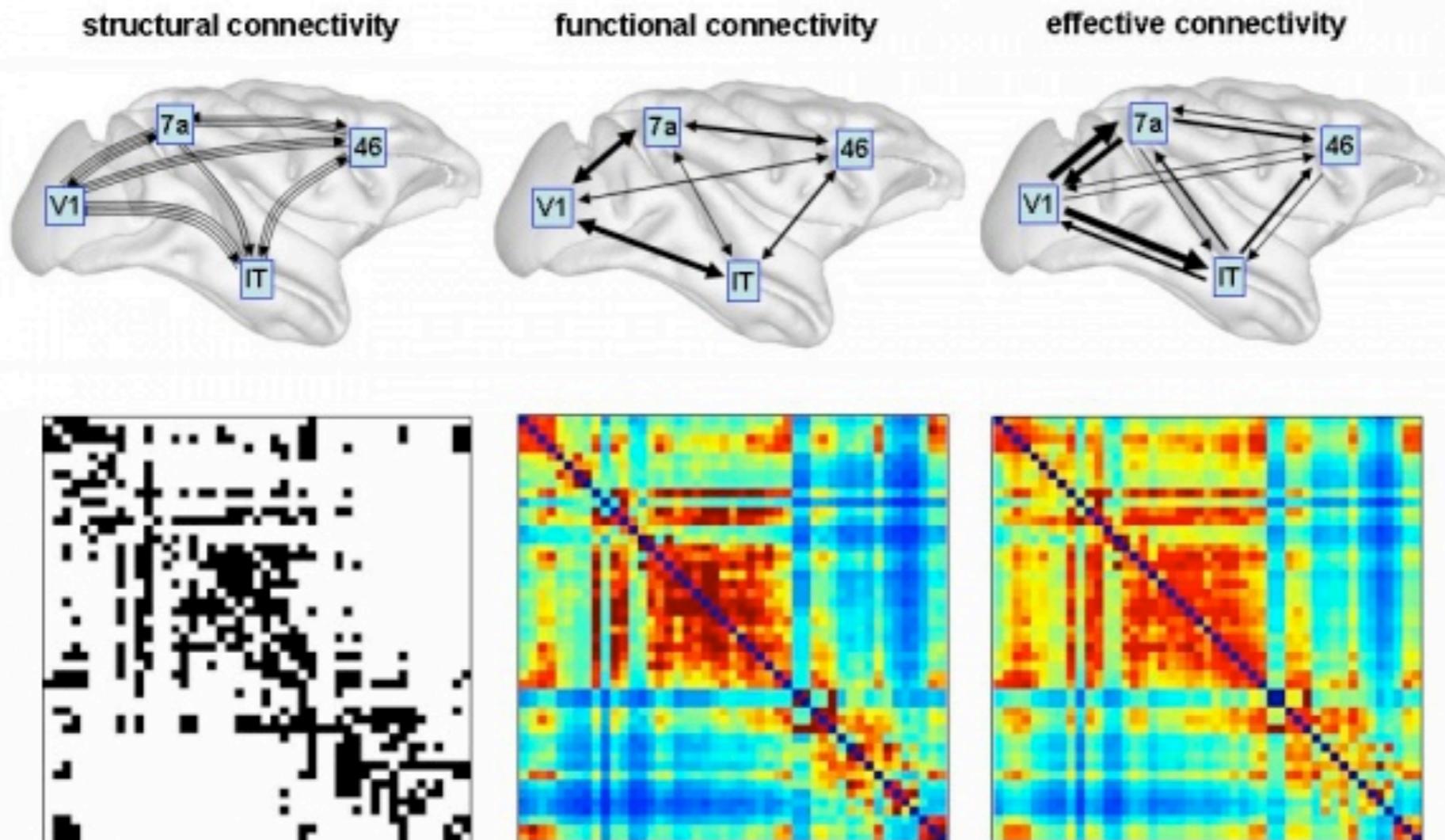
Multivariate models enable inferences on distributed responses without requiring focal activations.

Slide courtesy Kai Brodersen



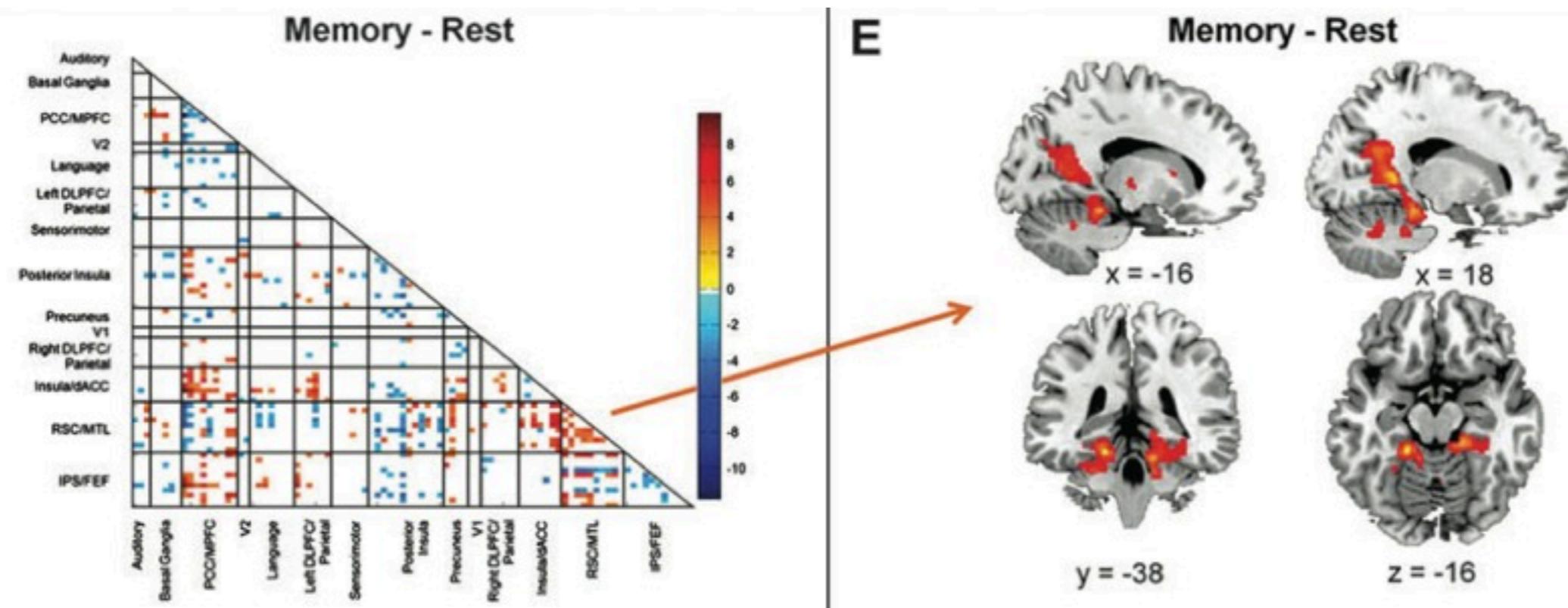


### Multivariate analyses of regional interactions



How do we estimate and describe connectivity?

# Graph-based decoding

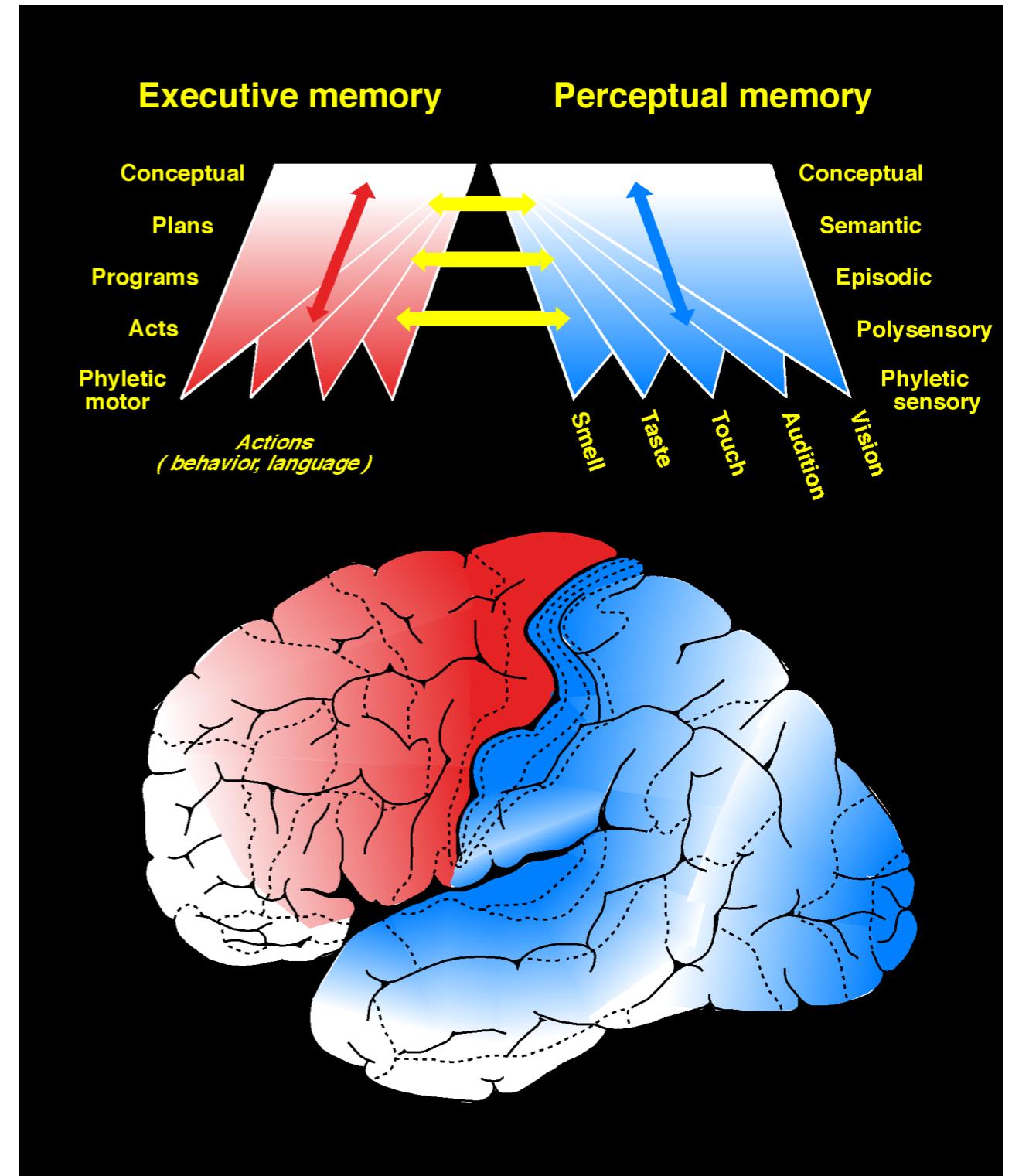


Can we predict cognitive states based on differences in connectivity?

# Distributed representations



Representations are hierarchically organized in terms of specific activity patterns within and across perceptual and executive networks.

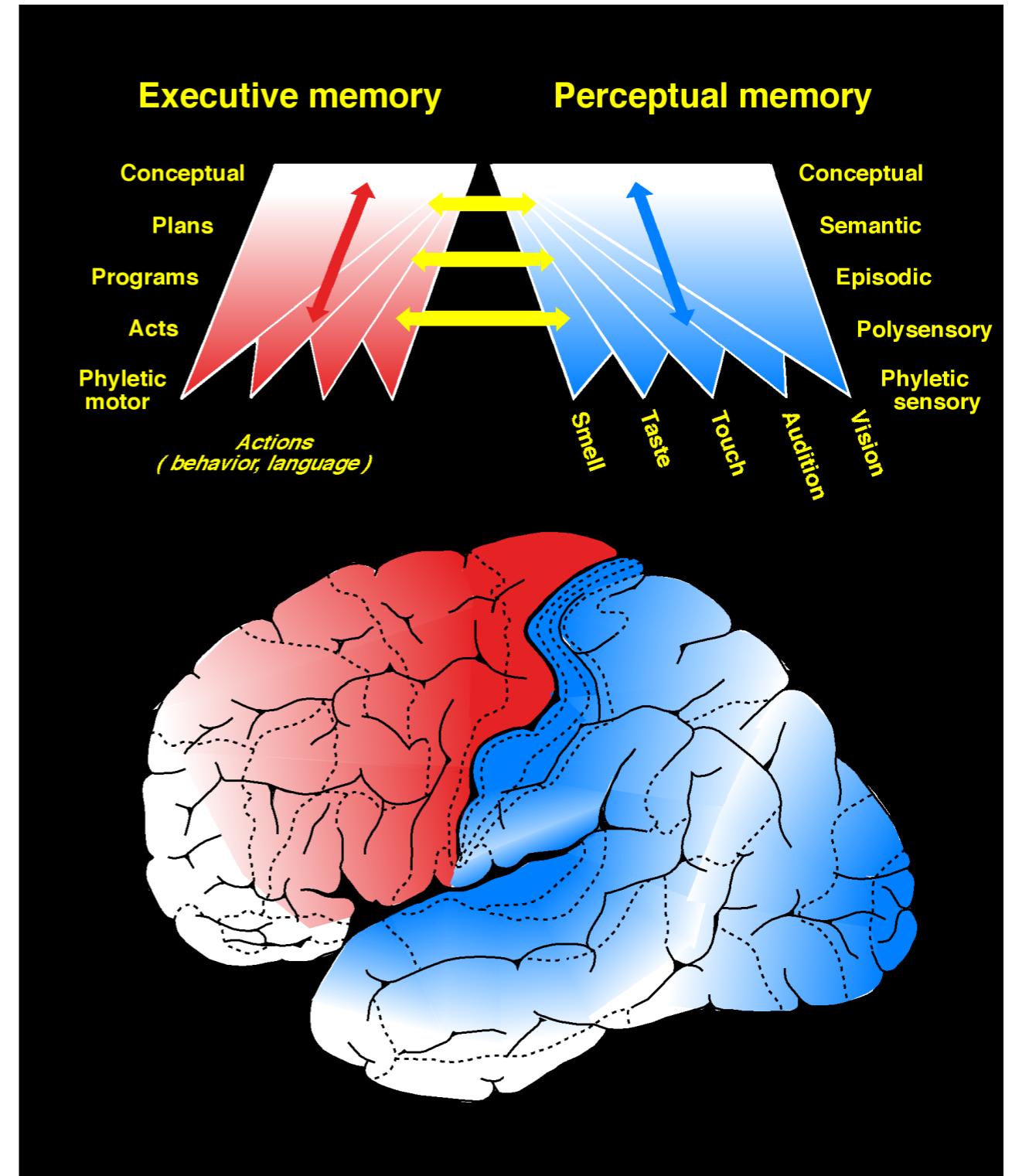


# Distributed representations



Representations are hierarchically organized in terms of specific activity patterns within and across perceptual and executive networks.

(Re-)activation of these representations involved in perception, WM maintenance, imagery, recall, decision making.



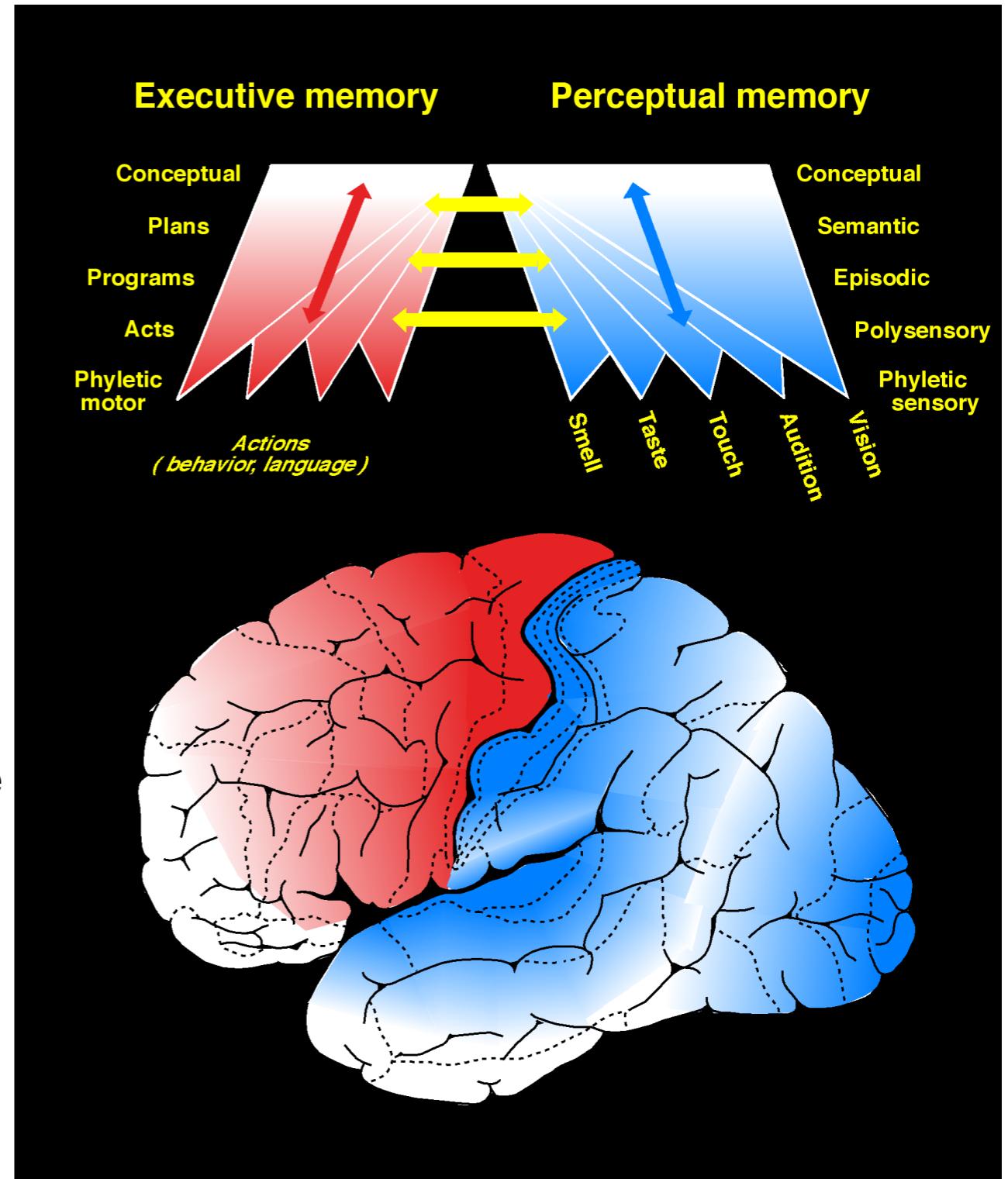


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New machine learning techniques offer new insights into the brain:

- decoding in cognitive/clinical neuroscience
- more sophisticated encoding models
- new approaches to connectivity analysis





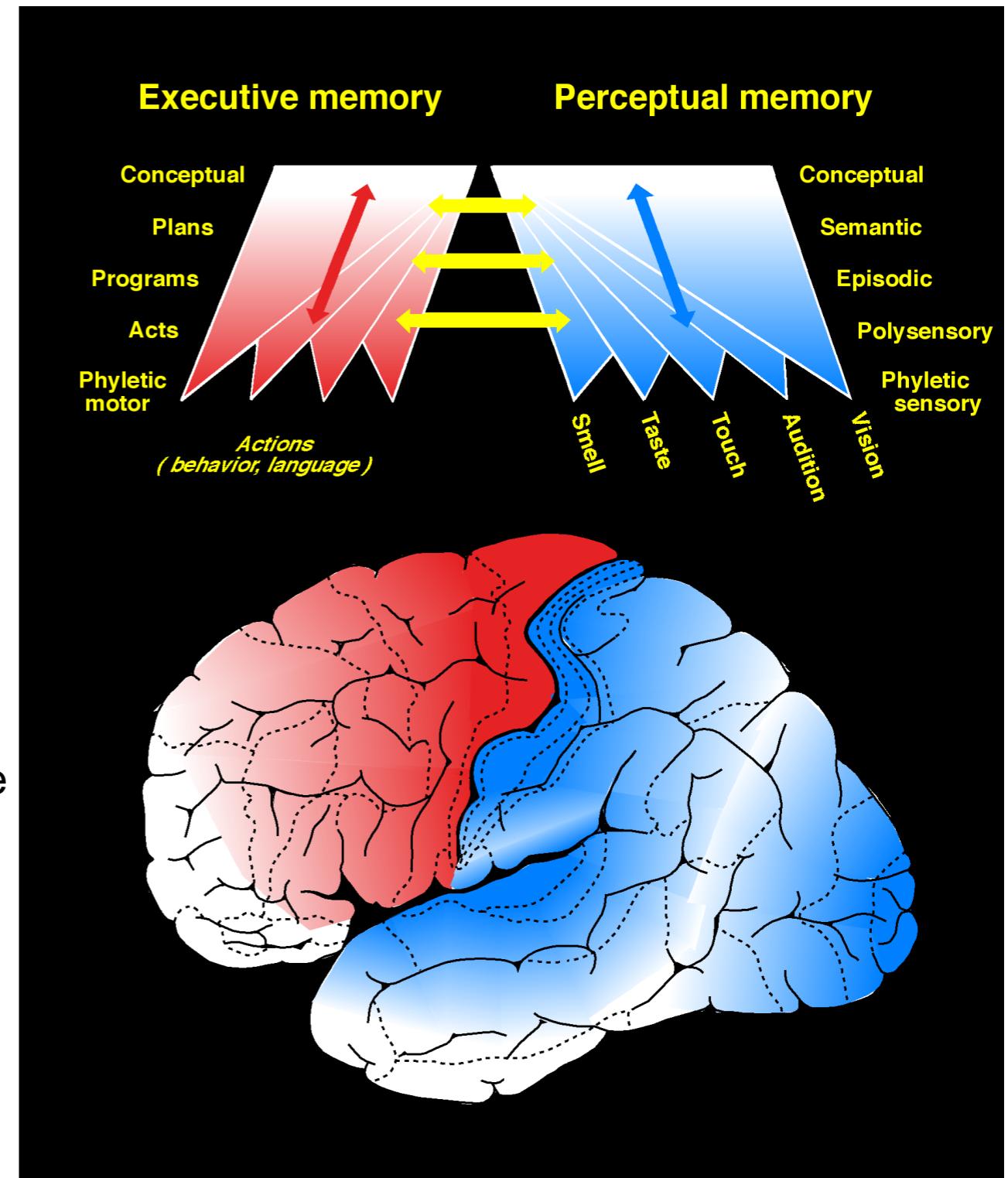
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Bayesian statistics as a lingua franca



# Course overview





## 1. Introduction





- 1. Introduction**
- 2. Discriminative decoding**



- 1. Introduction**
- 2. Discriminative decoding**
- 3. Advanced discriminative approaches**



- 1. Introduction**
- 2. Discriminative decoding**
- 3. Advanced discriminative approaches**
- 4. Generative decoding**



- 1. Introduction**
- 2. Discriminative decoding**
- 3. Advanced discriminative approaches**
- 4. Generative decoding**
- 5. Advanced generative approaches**



- 1. Introduction**
- 2. Discriminative decoding**
- 3. Advanced discriminative approaches**
- 4. Generative decoding**
- 5. Advanced generative approaches**
- 6. Connectivity analysis**



- 1. Introduction**
- 2. Discriminative decoding**
- 3. Advanced discriminative approaches**
- 4. Generative decoding**
- 5. Advanced generative approaches**
- 6. Connectivity analysis**
- 7. Bayesian connectomics**



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- 4. Generative decoding**
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- 6. Connectivity analysis**
- 7. Bayesian connectomics**
- 8. Guest lectures**



- 1. Introduction**
- 2. Discriminative decoding**
- 3. Advanced discriminative approaches**
- 4. Generative decoding**
- 5. Advanced generative approaches**
- 6. Connectivity analysis**
- 7. Bayesian connectomics**
- 8. Guest lectures**
- 9. Bayesian brain / Course conclusion**





- Paper: Friston KJ. Modalities, Modes, and Models in Functional Neuroimaging. *Science*. 2009; 326(5951):399–403.

REVIEW

# Modalities, Modes, and Models in Functional Neuroimaging

Karl J. Friston

In this, the 21st century, human-brain mapping celebrates 21 years of cognitive activation studies. This review looks at imaging neuroscience and key ideas it has pursued; some ideas portend exciting developments, and others have failed gloriously. In terms of achievements, there is much to celebrate, in the sense that it is difficult to imagine modern neuroscience without brain imaging. I will look at recent advances from the perspectives of functional segregation and integration in the brain, paying special attention to approaches that deal with the distributed and integrated nature of neuronal processing and the questions they address.



- Advantages of new statistical machine learning methods in cognitive neuroscience
- Prediction vs inference
- Mathematical form of the general linear model
- Difference between mass-univariate and multivariate analysis
- Encoding vs decoding, generative vs discriminative
- Classification, identification, reconstruction
- Functional segregation vs functional integration
- Structural, functional, effective connectivity



## Practical

Excursion to the Donders Magnetic Resonance Imaging facility!



## Lecture

Discriminative decoding  
Room TvA 4.00.19