

# M/EEG source analysis

José David López

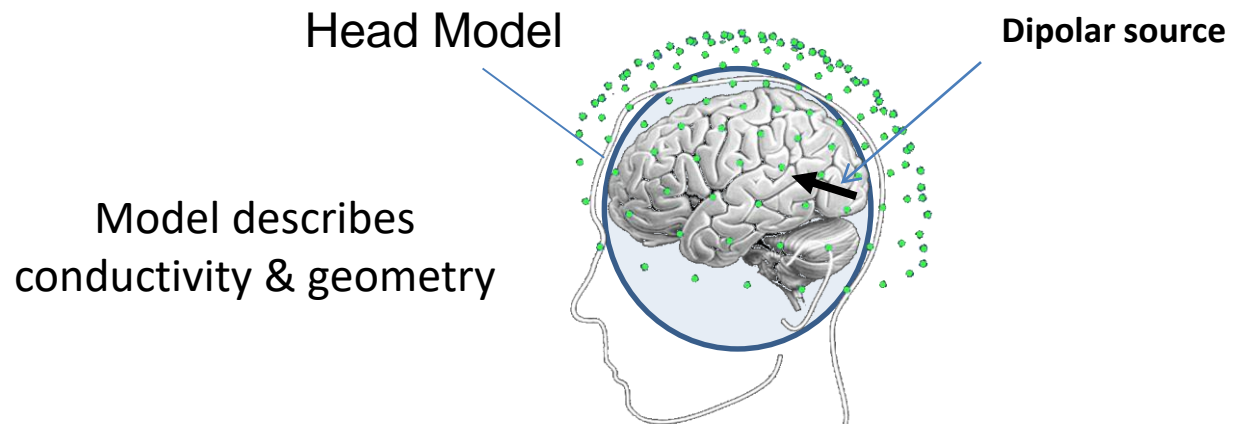
# Key points:

- What is an ill-posed inverse problem
- Prior knowledge- links to popular algorithms.
- Validation of prior knowledge/ Model evidence

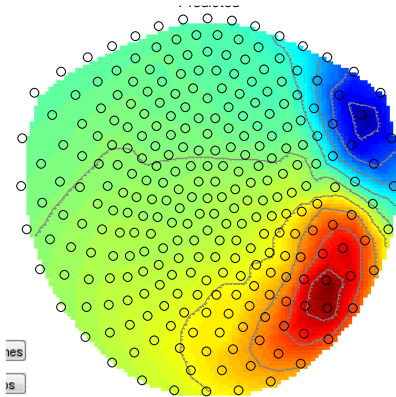
# The forward problem

***Lead field (L) is the sensitivity of the M/EEG system to a dipolar source at a particular location***

Analogy  
2+3= ?



# The Inverse problem



M/EEG  
sensors

Measurement

*Which brain sources gave rise to  
these measured data ?*

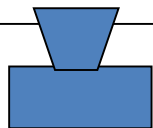
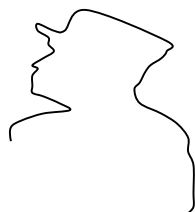
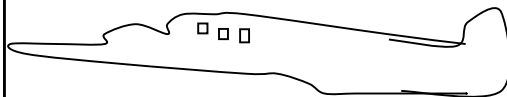
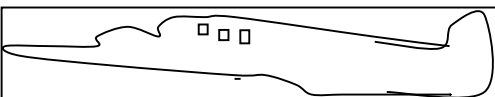
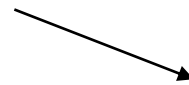
Analogy  
 $5 = ? + ?$

Inference

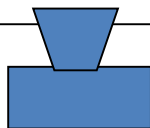


Inverse problems aren't difficult

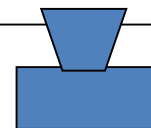




A

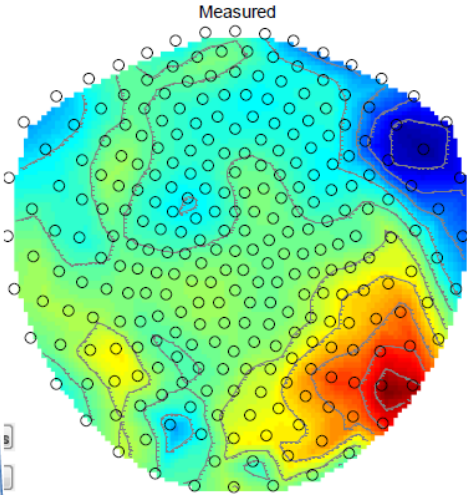


B



C

# Measurement ( $Y$ )



M/EEG sensors

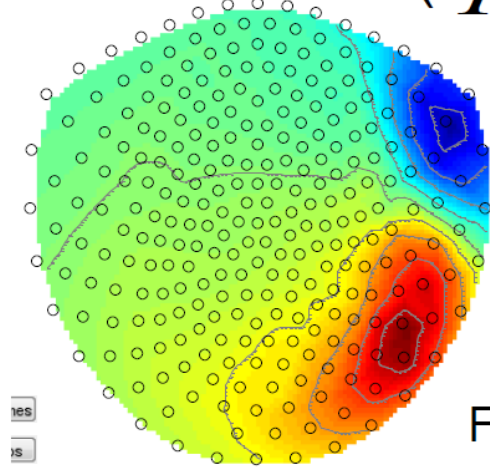
brain

?

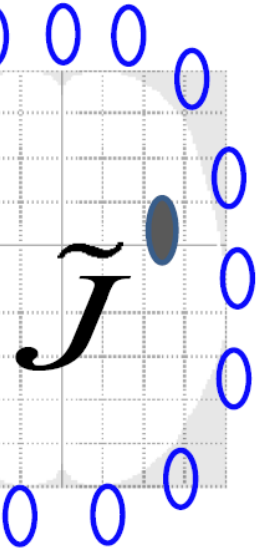
Prior info

Current density Estimate

# Prediction ( $\tilde{Y}$ )



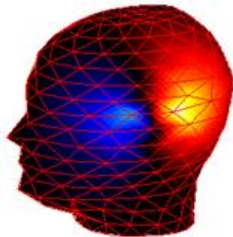
Forward problem



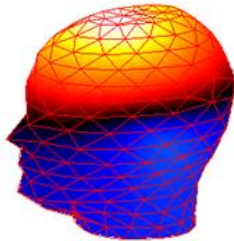
# The forward problem



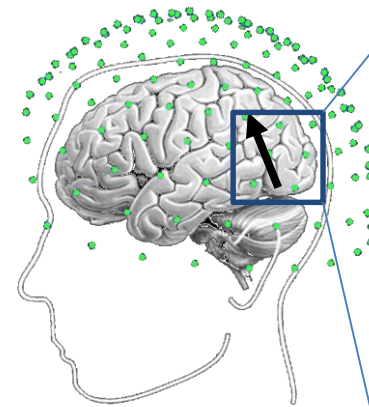
MEG



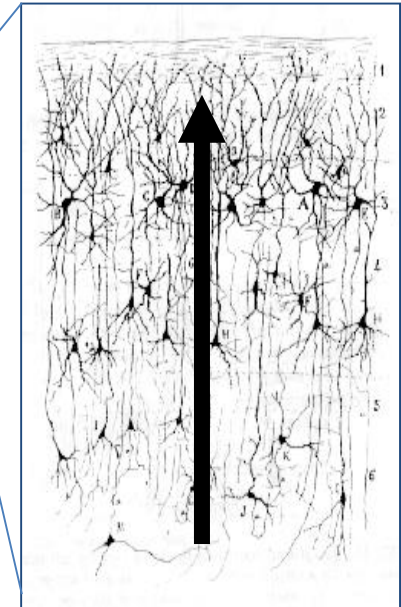
EEG



*Lead fields*



**Head tissues  
(conductivity & geometry)**



**Neurons**

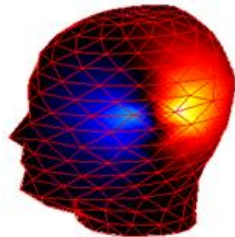




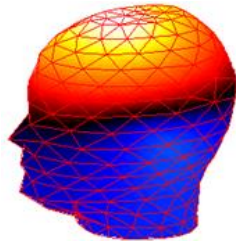
# The forward problem



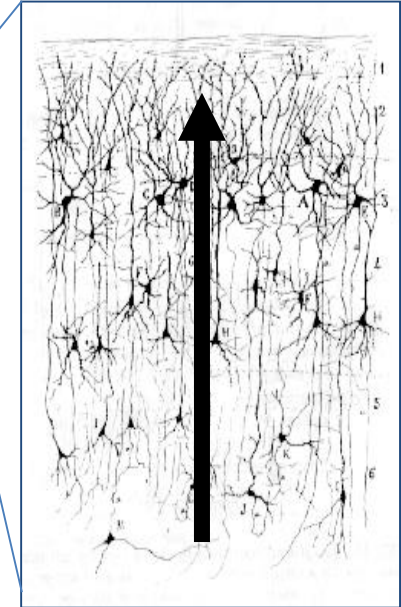
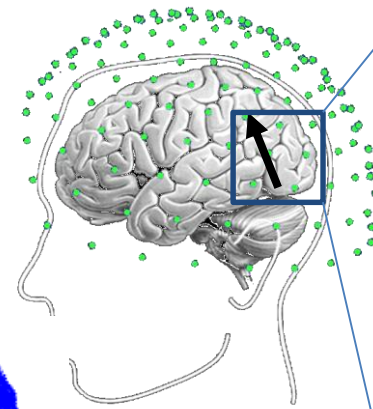
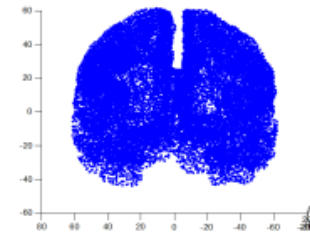
MEG



EEG



Lead fields



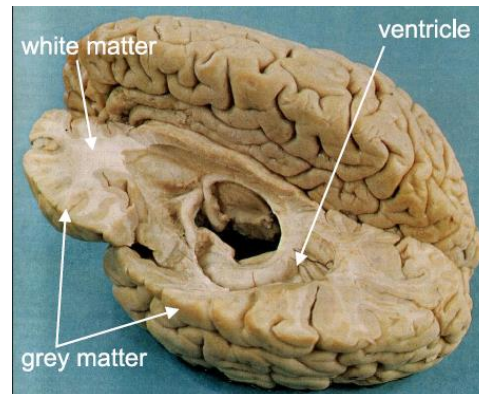
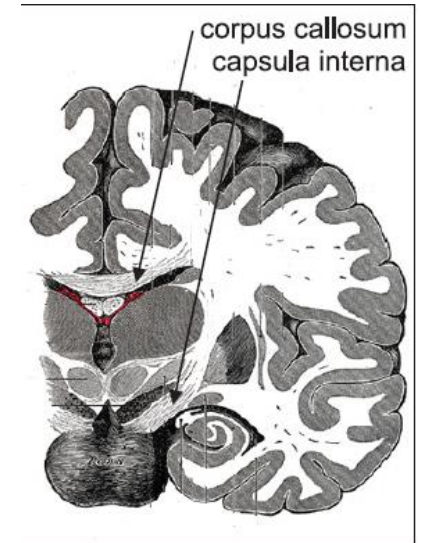
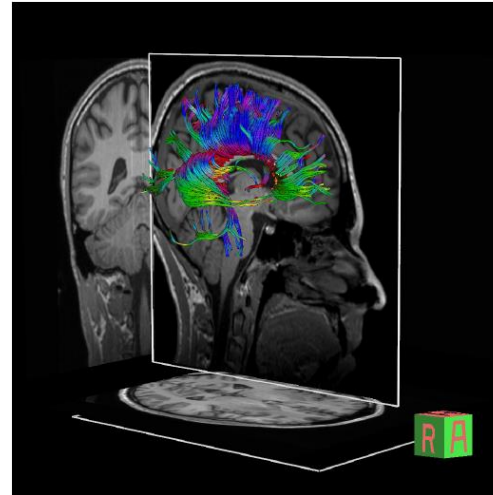
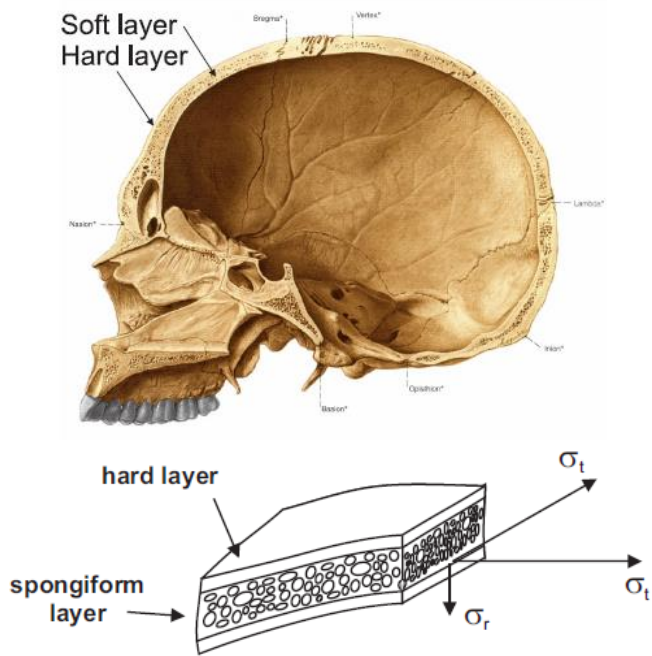
Dipolar sources



$$Y = LJ + \epsilon$$

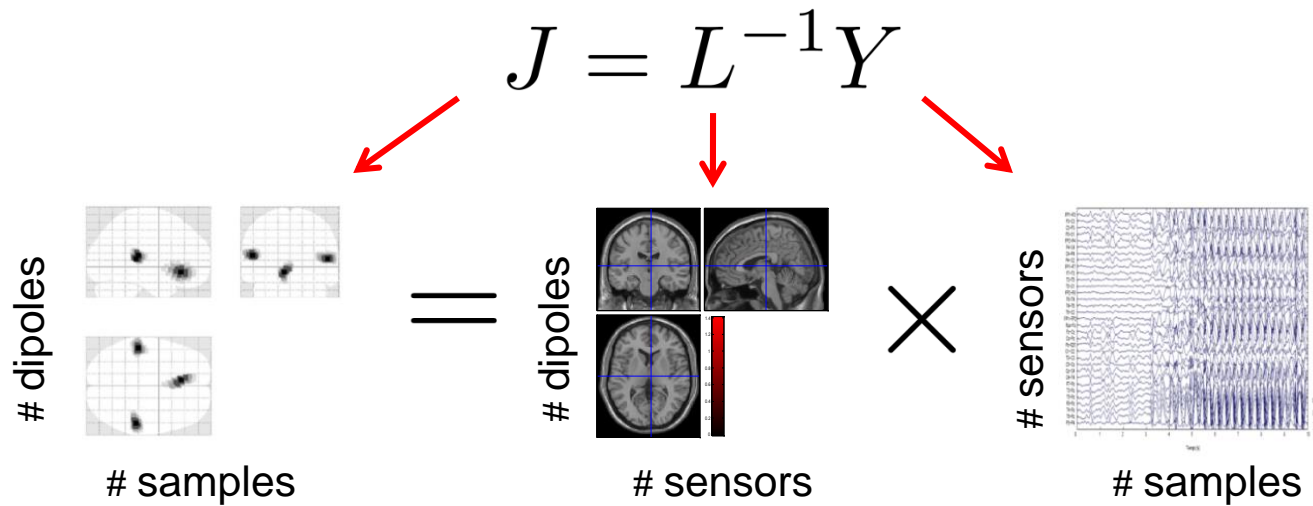
← Noise

# Head model (gain matrix)



# MEG/EEG brain imaging

With the acquired data we may recover the neural activity



But the problem is ill-posed:

NON INVERTIBLE!!!

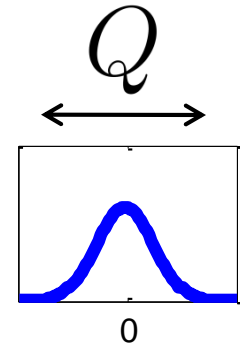
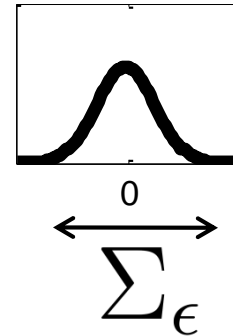


Infinite solutions!!!

# Bayesian formulation

We must include prior information:

$$Y = LJ + \epsilon$$



then we can use the Bayes' theorem:

Adjusted with the data

$$p(J|Y) = \frac{p(Y|J)p(J)}{p(Y)}$$

Assumed

Constant

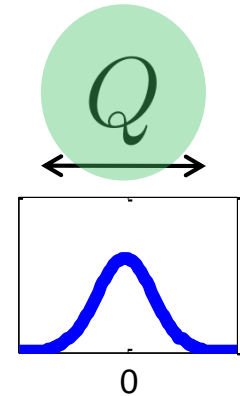
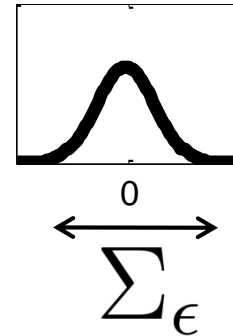
and solving for Gaussian assumptions:

$$\hat{J} = E[p(J|Y)] \rightarrow \hat{J} = QL^T(\Sigma_\epsilon + LQL^T)^{-1}Y$$

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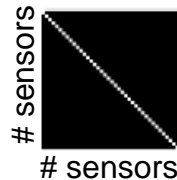
$$\hat{J} = E[p(J|Y)] \rightarrow \hat{J} = QL^T(\Sigma_\epsilon + LQL^T)^{-1}Y$$

# Prior covariance matrices

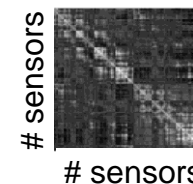
## PRIOR NOISE COVARIANCE

Independent sensor noise

$$\Sigma_{\epsilon} = h_0 I$$



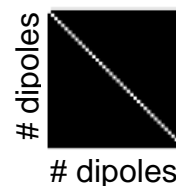
Empty room activity



## PRIOR COVARIANCE OF SOURCE SPACE ACTIVITY

Minimum norm

$$Q = I$$

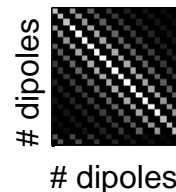


Non informative

$$\hat{J} = L(\Sigma_{\epsilon} + LL^T)^{-1}Y$$

LORETA-like:

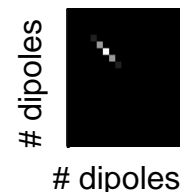
$$Q = e^{\sigma G_L}$$



Smoothed

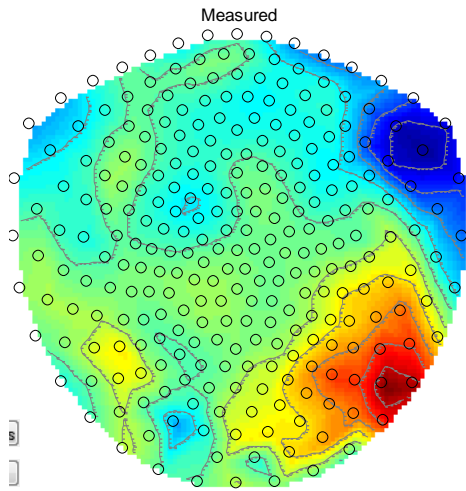
Beamformers:

$$Q = (L^T (YY^T)^{-1} L)^{-1}$$



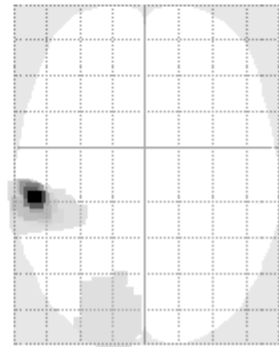
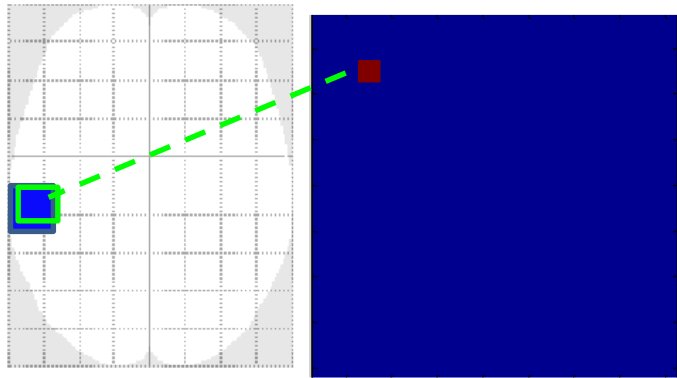
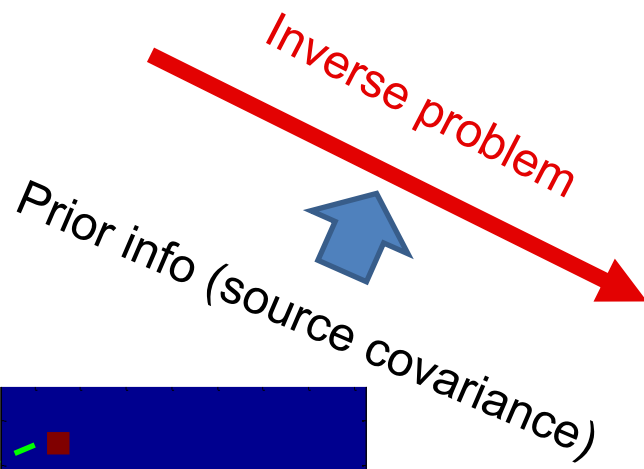
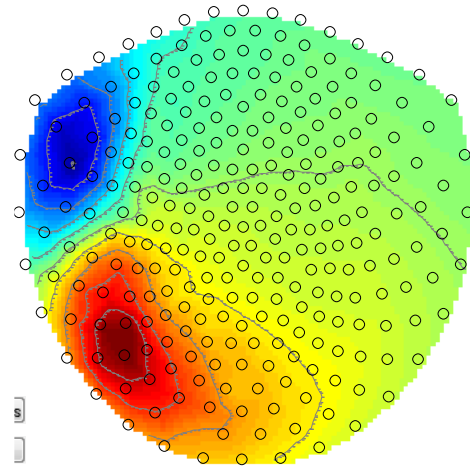
Data based

Y (measured field)



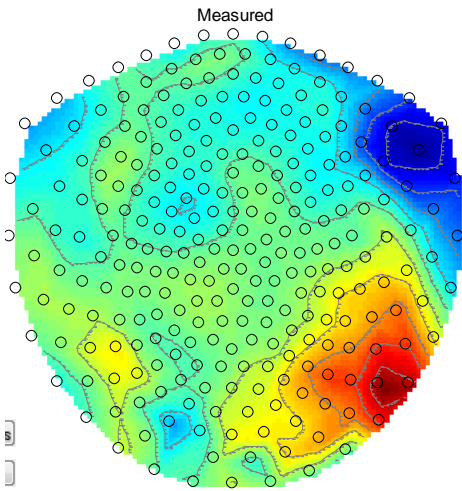
# Illustrative example

PREDICTED

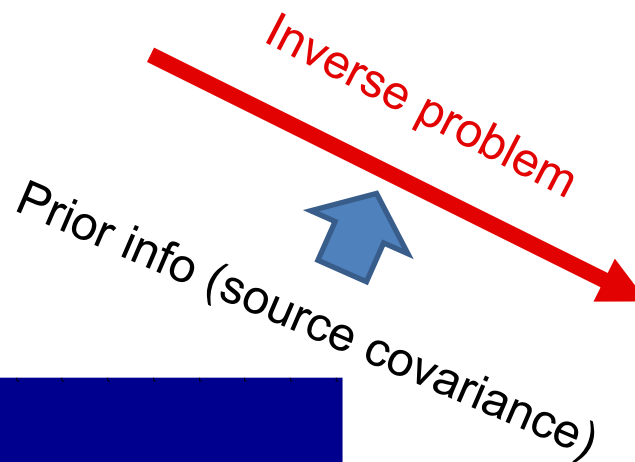


$Q$

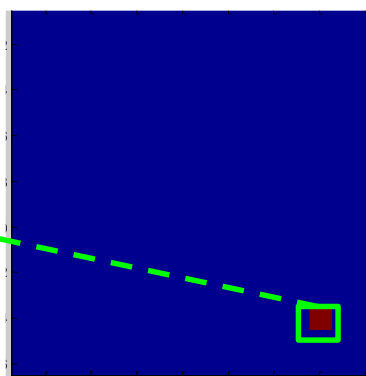
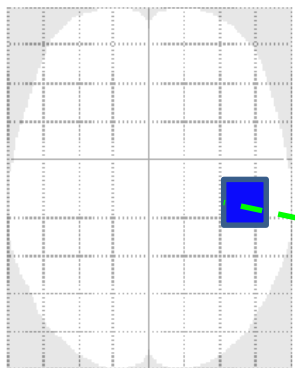
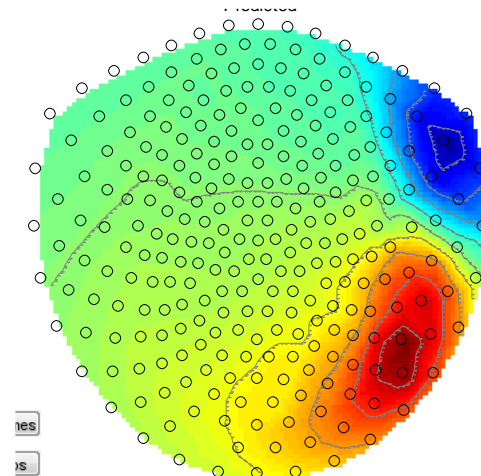
Y (measured field)



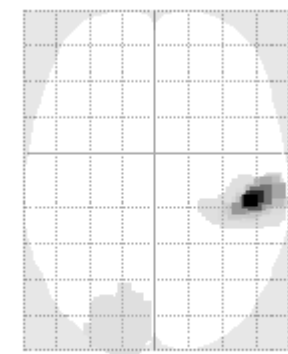
Single dipole fit



PREDICTED

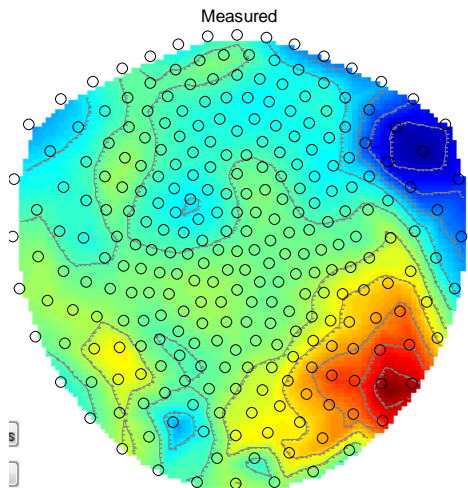


$Q$

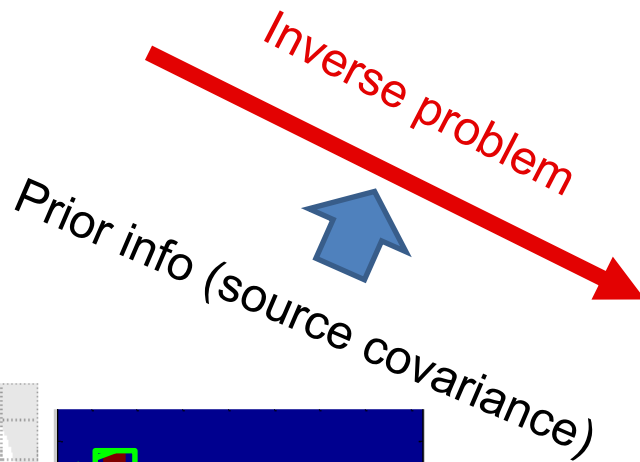




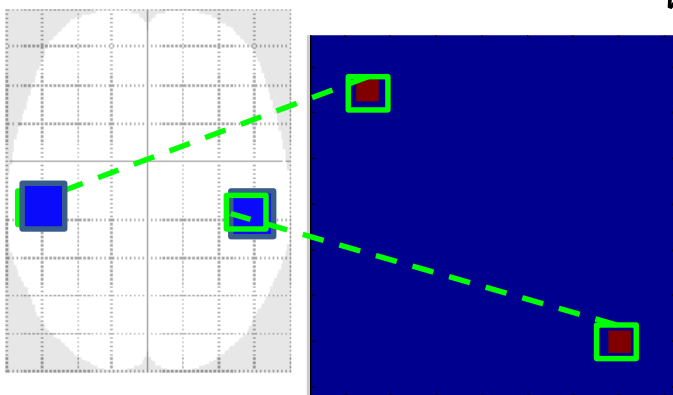
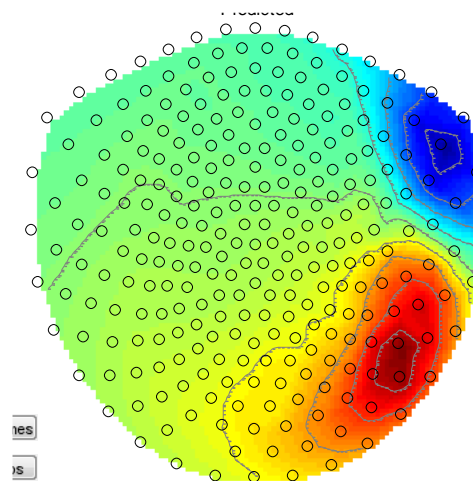
Y (measured field)



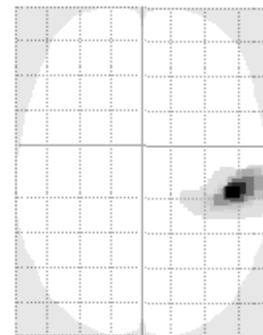
Two dipole fit



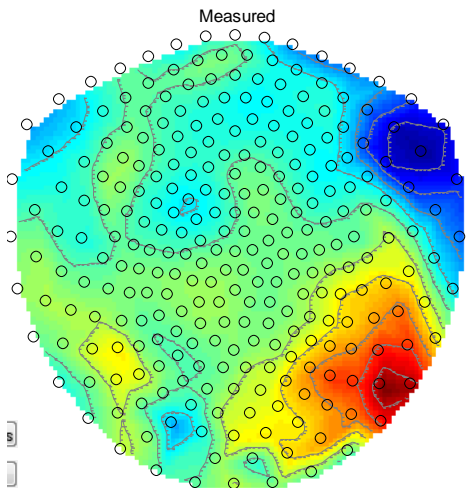
PREDICTED



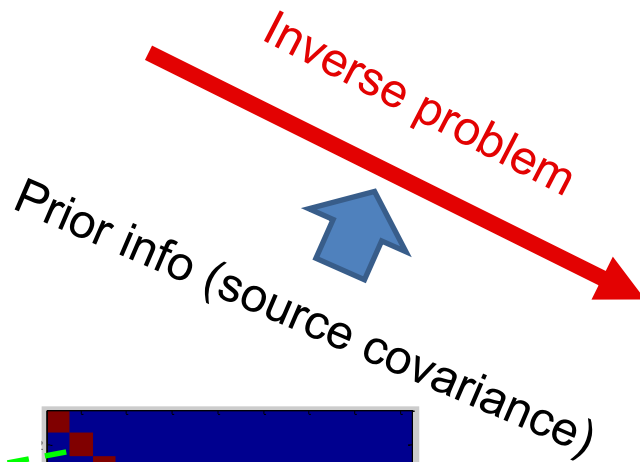
$Q$



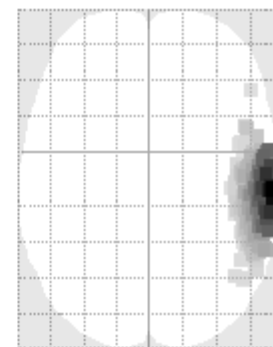
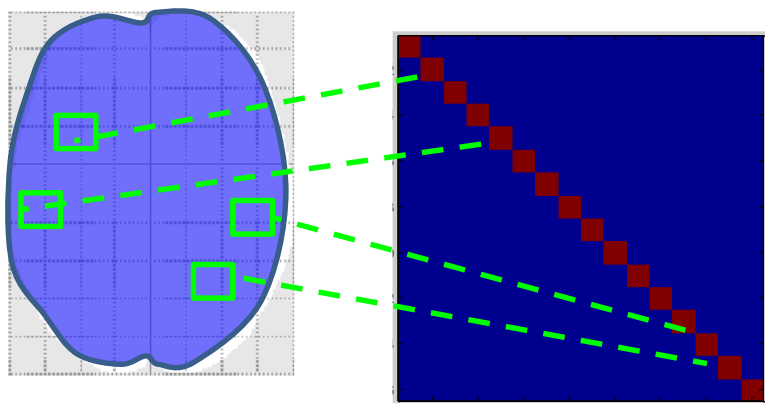
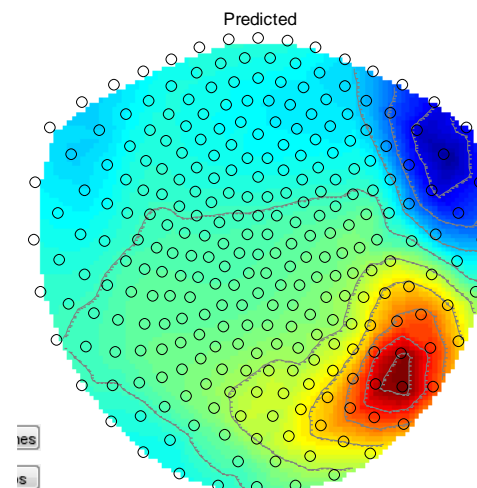
Y (measured field)



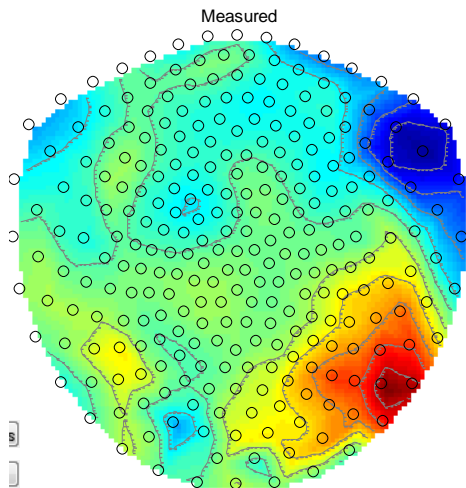
Minimum norm



PREDICTED

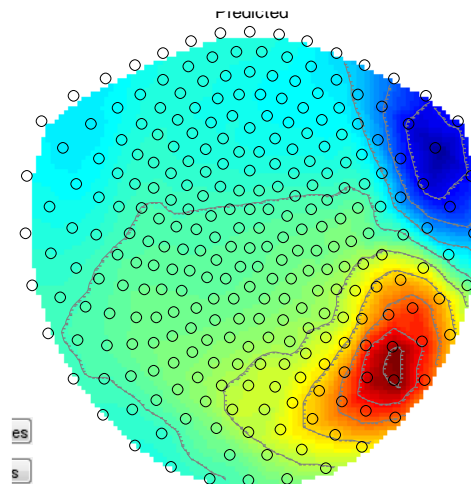


Y (measured field)



Beamformer  
(adaptive algorithm/  
Empirical )

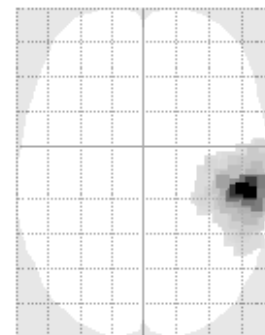
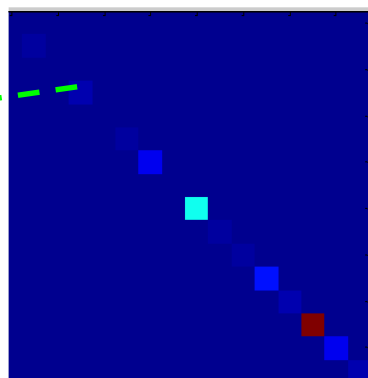
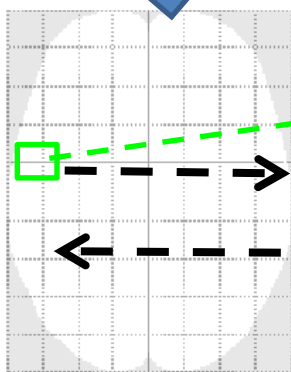
PREDICTED



Inverse problem

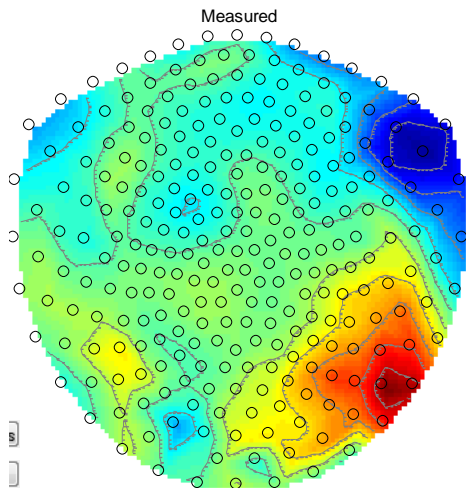
Prior info (source covariance)

Projection  
onto  
lead field\*



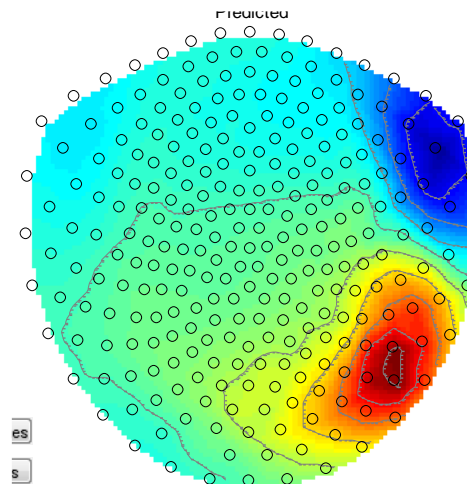
\*Assuming no correlated sources

Y (measured field)



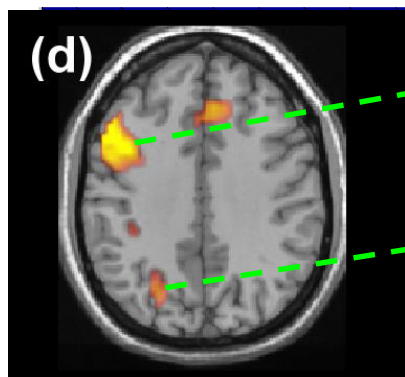
# fMRI biased dSPM (Dale et al. 2000)

PREDICTED

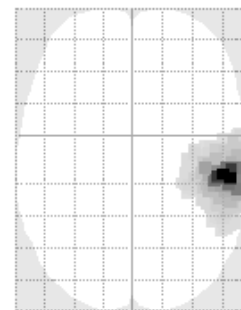
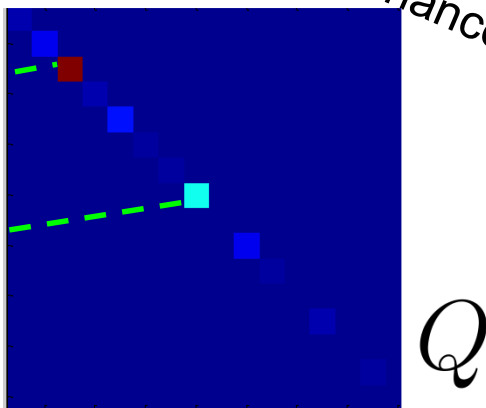


Inverse problem

Prior info (source covariance)

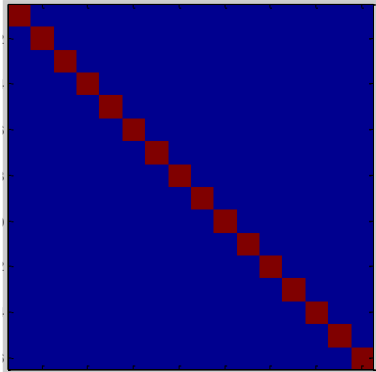


fMRI data

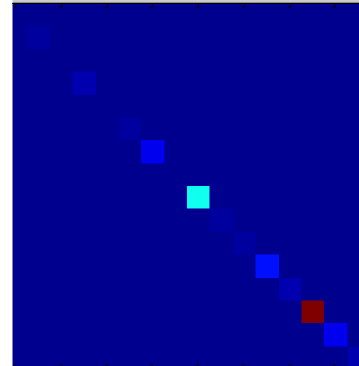


Maybe...

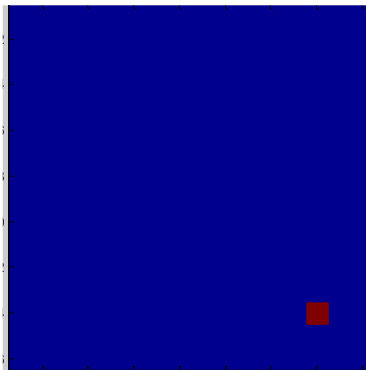
# Summary: Some popular priors



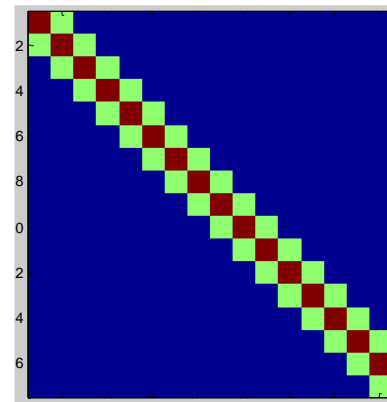
Minimum norm



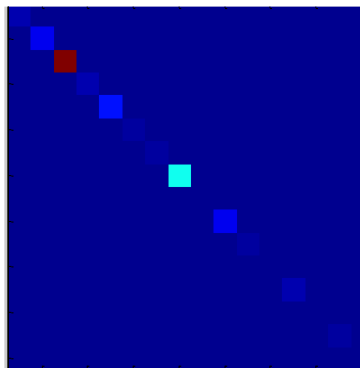
SAM, DICs  
Beamformer



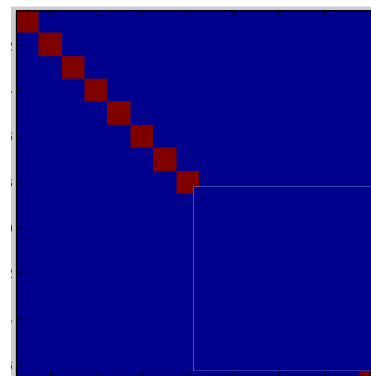
Dipole fit



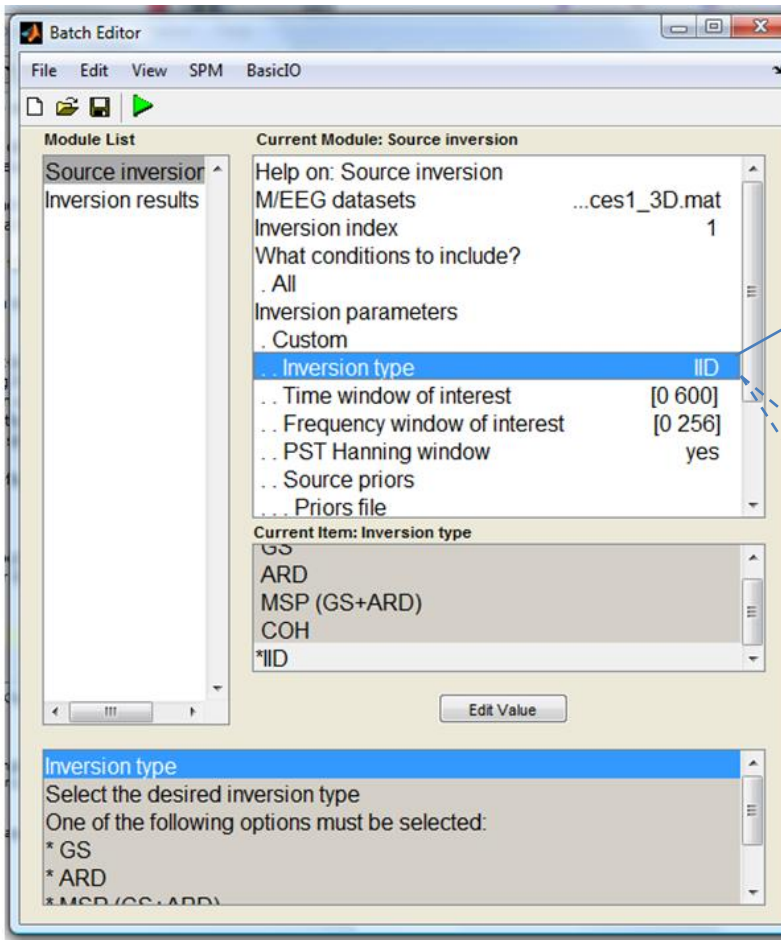
LORETA



fMRI biased  
dSPM

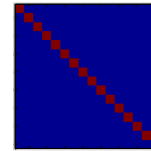


?

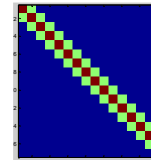


## Minimum Norm (IID

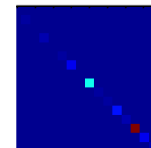
- independent and identically distributed)



## LORETA (COH- coherent)



## Empirical Bayes Beamformer (EBB)



## Multiple Sparse Priors

(MSP/ Greedy Search (GS)

Automatic relevance determination (ARD) )

# Summary

- MEG inverse problem requires prior information in the form of a source covariance matrix.
- Different inversion algorithms- SAM, DICS, LORETA, Minimum Norm, dSPM... just have different prior source covariance structure.
- Historically- different MEG groups have tended to use different algorithms/acronyms.

See

Mosher et al. 2003, Friston et al. 2008, Wipf and Nagarajan 2009, Lopez et al. 2014

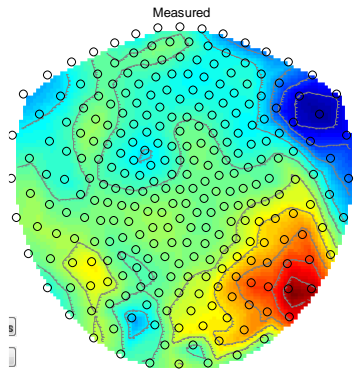
# Software

- **SPM12:** <http://www.fil.ion.ucl.ac.uk/spm/software/spm12/>
- **DAiSS-** SPM12 toolbox for Data Analysis in Source Space (beamforming, minimum norm and related methods), developed by Vladimir Litvak:  
<https://github.com/spm/DAiSS>
- **Fieldtrip :** <http://fieldtrip.fcdonders.nl/>
- **Brainstorm:** <http://neuroimage.usc.edu/brainstorm/>
- **MNE:** <http://martinos.org/mne/stable/index.html>



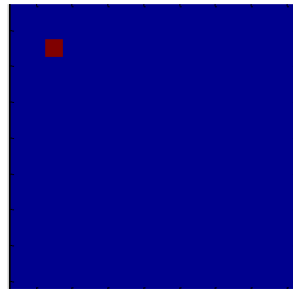
Y (measured field)

# How do we choose between priors ?

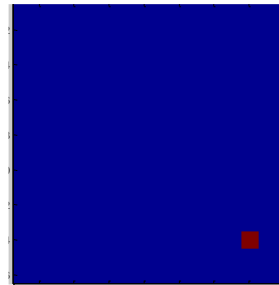


Prior

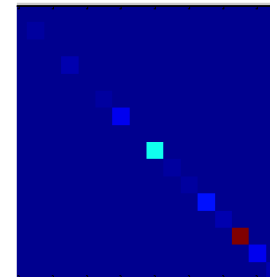
Are you drunk?



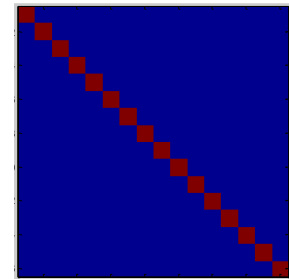
You won the lottery



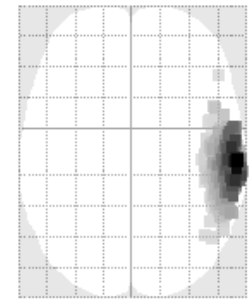
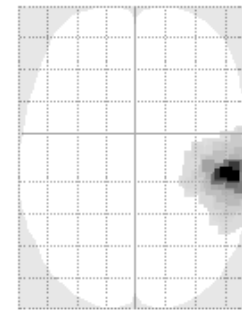
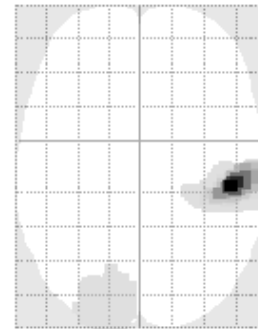
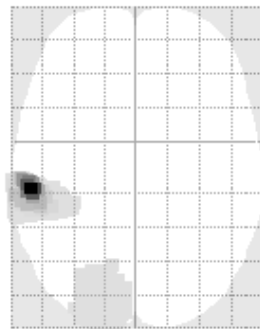
Beamformer



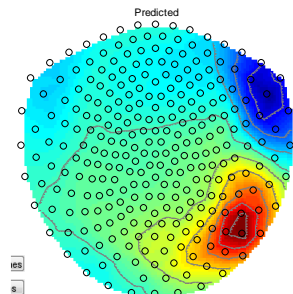
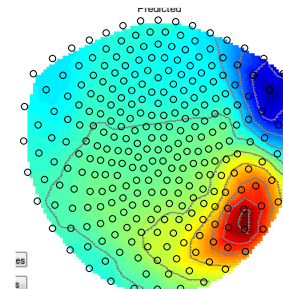
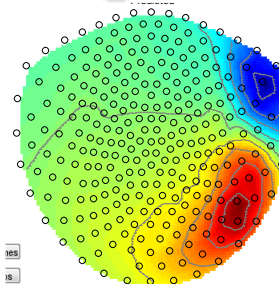
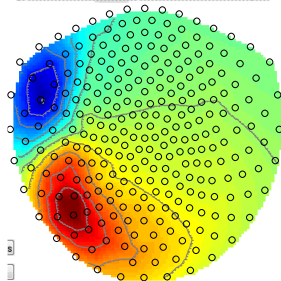
Minimum norm



Estimated Current flow



Predicted data



Variance explained

11 %

96%

97%

98%

# Negative variational free energy (1)

$$\log p(\hat{Y}) = F + KL[q(h)||p(h|Y)]$$

the divergence will be zero if the approximated distribution is equal to the posterior one:

$$q(h) = p(h|Y) \longrightarrow F = \log p(Y)$$

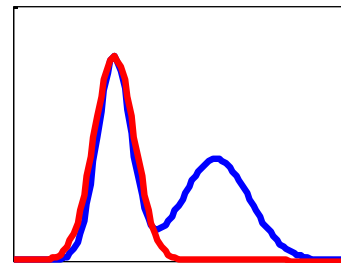
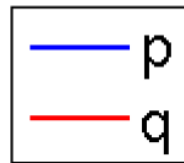
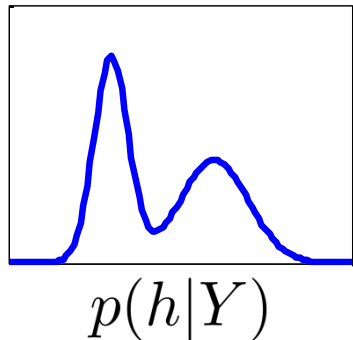
# Negative variational free energy (1)

$$\log p(\hat{Y}) = F + KL[q(h) || p(h|Y)]$$

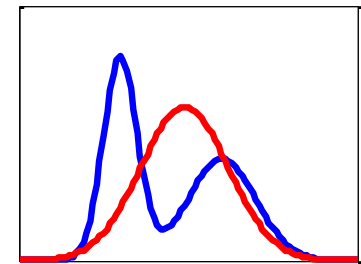
the divergence will be zero if the approximated distribution is equal to the posterior one:

$$q(h) = p(h|Y) \longrightarrow F = \log p(Y)$$

$$q_0(h) = \mathcal{N}(h; \nu, \Pi^{-1}) \longrightarrow q(h) = \mathcal{N}(h; \hat{h}, \Sigma_h)$$



Wrong



**BETTER!**

# Negative variational free energy (2)

The free energy can be expressed as:

$$F = -\frac{N_t}{2} \text{tr}(C_Y \Sigma_Y^{-1}) - \frac{N_t}{2} \log |\Sigma_Y| - \frac{N_c N_t}{2} \log(2\pi) \\ - \frac{1}{2} \text{tr} \left( (\hat{h} - \nu)^T \Pi (\hat{h} - \nu) \right) + \frac{1}{2} \log |\Pi \Sigma_h|$$

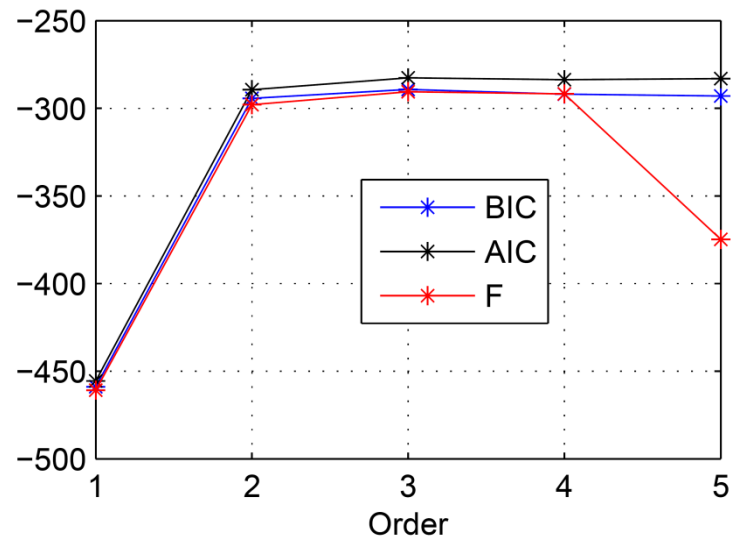
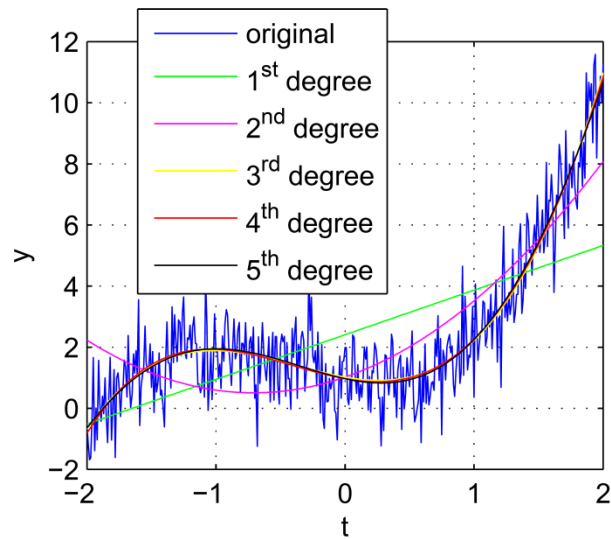
Reducing constant terms and assuming zero mean priors:

$$F = -\text{trace} \left( \frac{Y Y^T}{\Sigma_\epsilon + L Q L^T} \right) - \log |\Sigma_\epsilon + L Q L^T| \quad \rightarrow \quad \text{Accuracy}$$

$$-\text{trace}(h^T \Pi h) + \log |\Pi \Sigma_h| \quad \rightarrow \quad \text{Complexity}$$

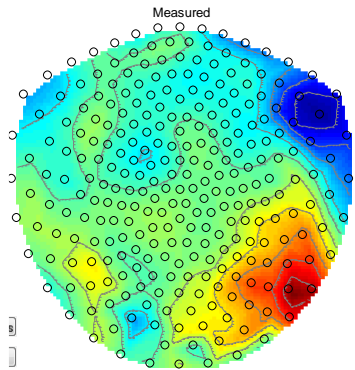
# Trade-off between accuracy and complexity

Approach	Complexity term
AIC (Akaike, 1974)	$N_q$
BIC (Schwarz, 1978)	$\frac{N_q}{2} \log N_t$
Linear function (Wipf and Nagarajan, 2009)	$h$
<i>free energy</i> (Friston et al., 2008)	$\frac{1}{2} \text{tr} \left( (h - \nu)^T \Pi (h - \nu) \right) - \frac{1}{2} \log  \Pi \Sigma_h $



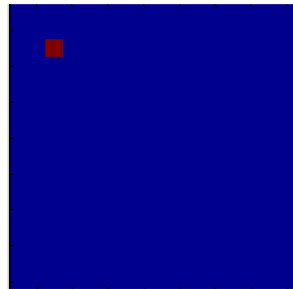
Y (measured field)

# How do we choose between priors ?

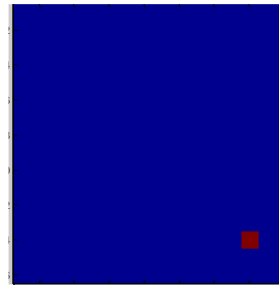


Prior

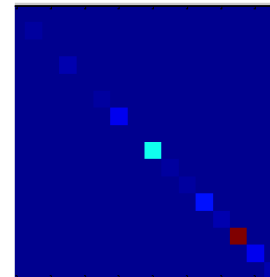
Are you drunk?



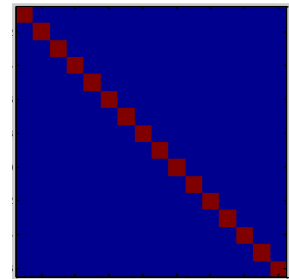
You won the lottery



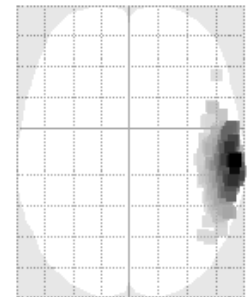
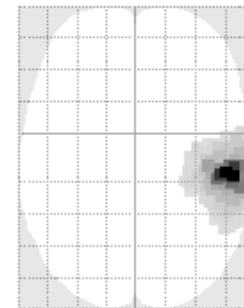
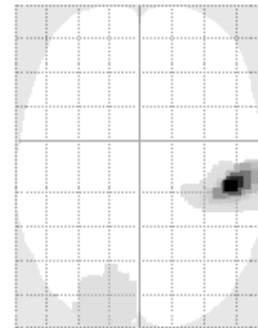
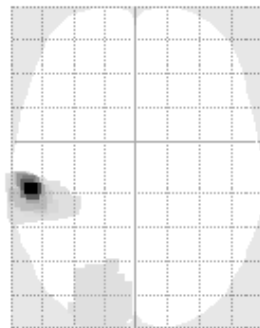
Beamformer



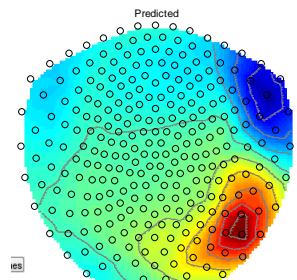
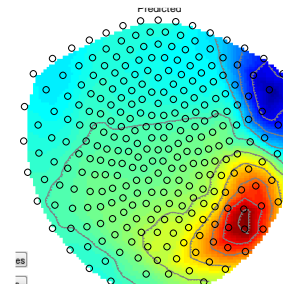
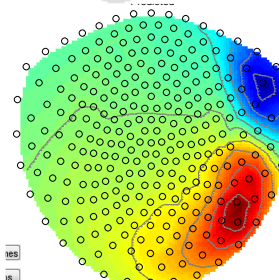
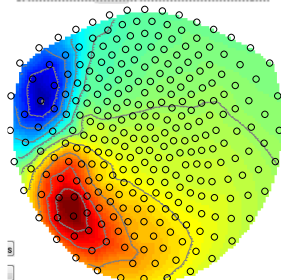
Minimum norm



Estimated Current flow



Predicted data



Variance explained

11 %

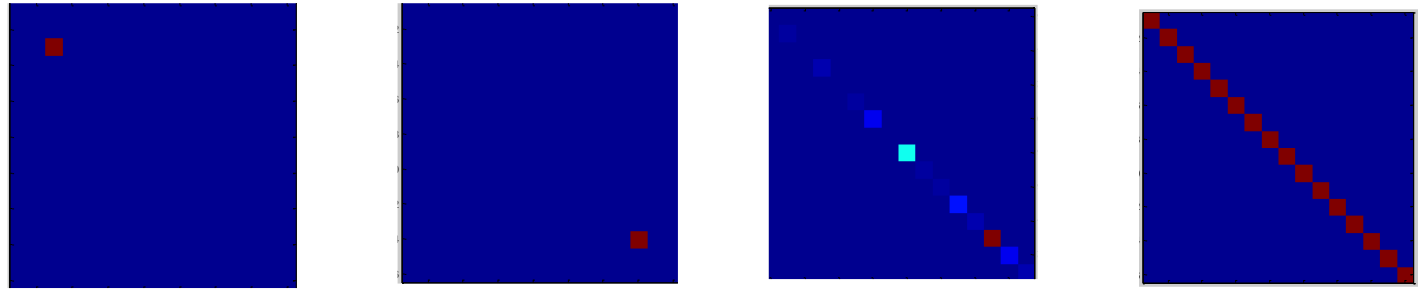
96%

97%

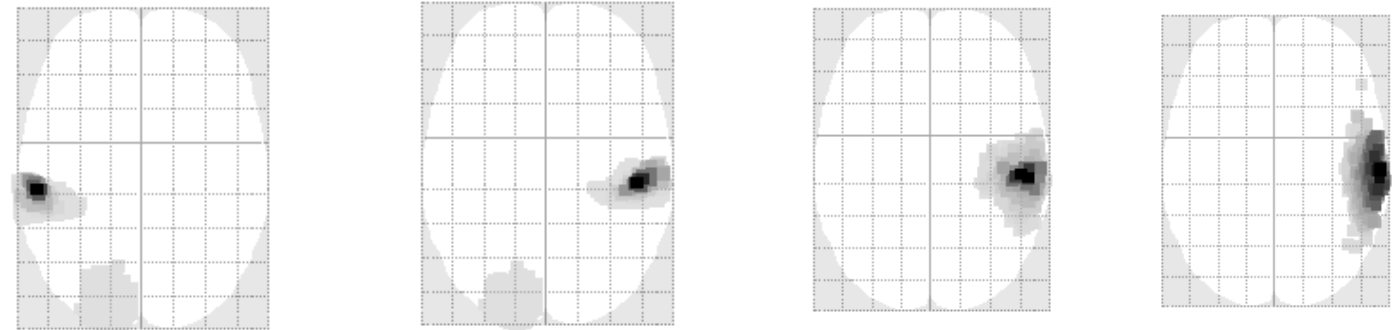
98%

# How do we choose between priors ?

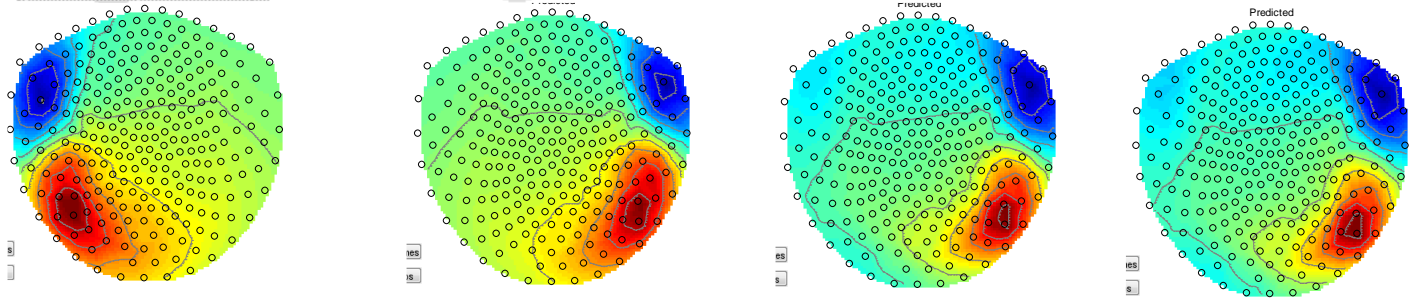
Prior



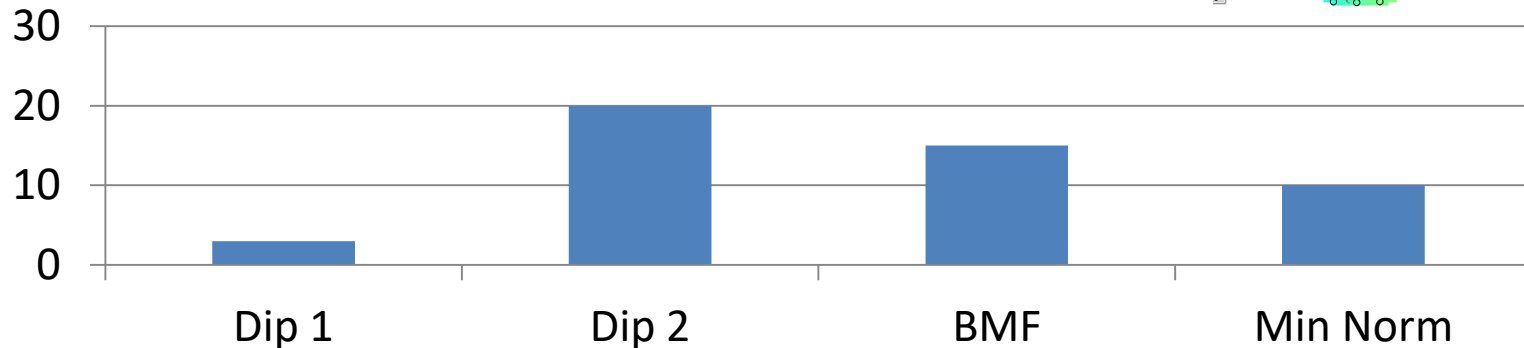
Estimated Current flow



Predicted data



Free energy  
(log model  
evidence)



# Multiple sparse priors (1)

All prior information can be included as the linear combination of a set of covariance components

$$\hat{J} = Q L^T (\Sigma_\epsilon + L Q L^T)^{-1} Y$$

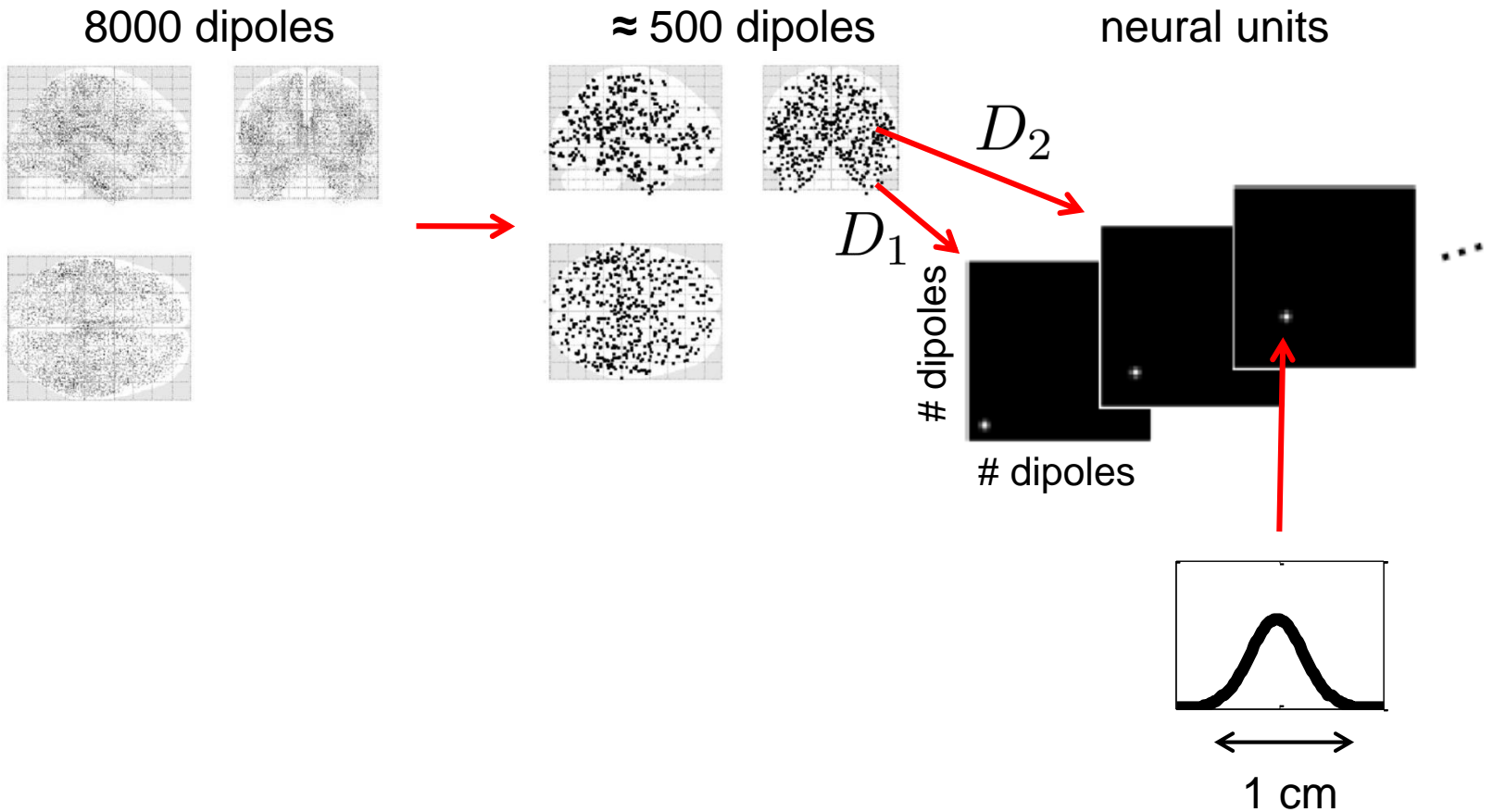
$$Q = \sum_{i=1}^{N_q} h_i D_i$$

$$D = \{D_1, \dots, D_{N_q}\}$$

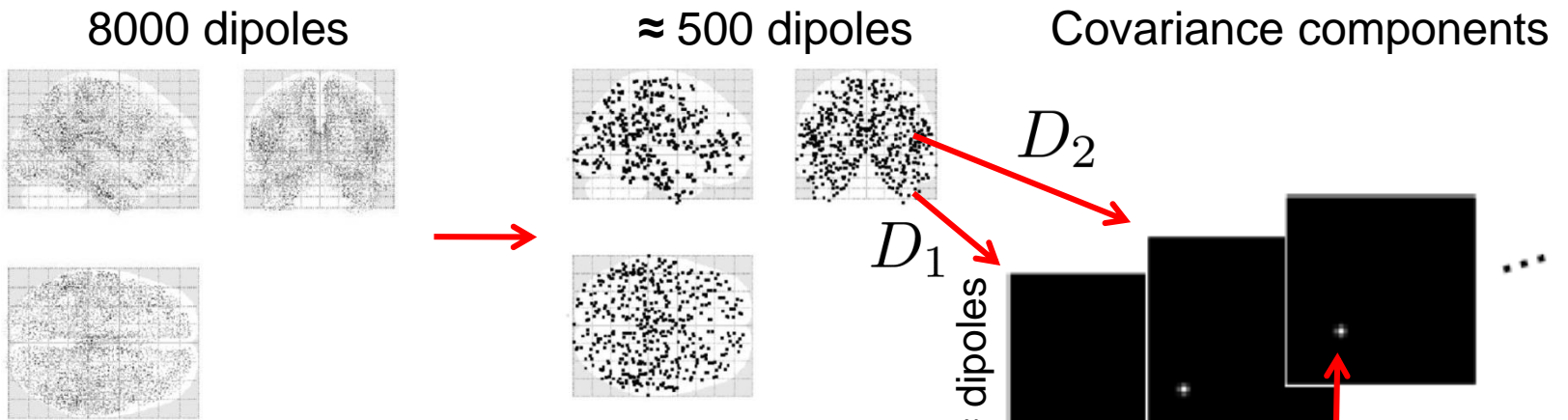
$$h = \{h_1, \dots, h_{N_q}\}$$



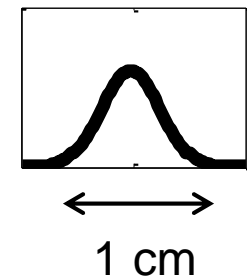
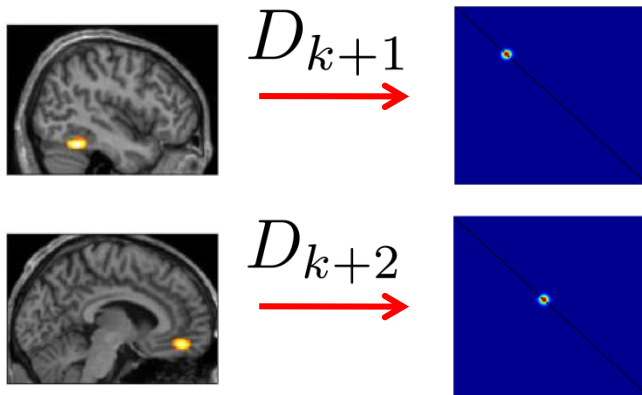
# Multiple sparse priors (2)



# Multiple sparse priors (2)

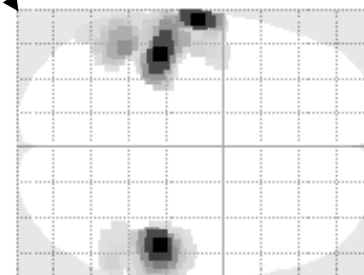
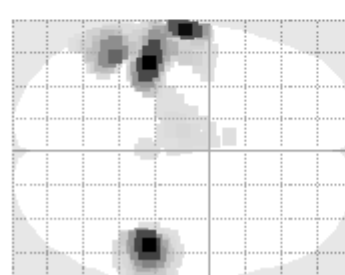
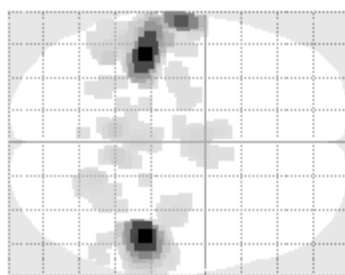
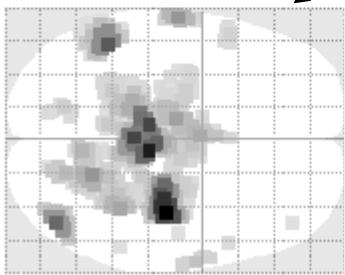
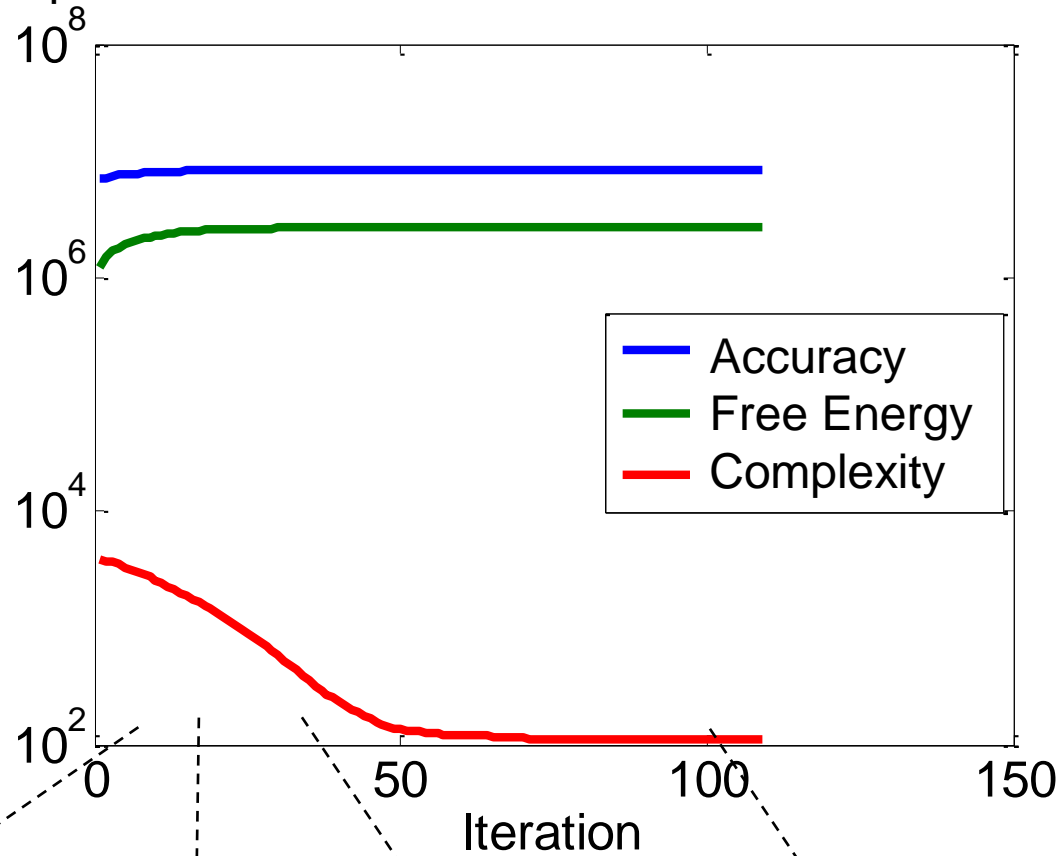


Functional imaging priors



# Multiple Sparse priors

So now construct the priors to maximise model evidence



# Key points :

- Prior knowledge- links to popular algorithms
- Validation of prior knowledge/ Model evidence

# Conclusion

- M/EEG inverse problem can be solved... If you have some prior knowledge.
- All prior knowledge encapsulated in a source covariance matrix.
- Can test between priors (or develop new priors) within a Bayesian framework.

# References

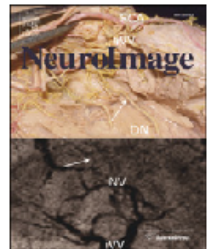
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Technical Note

## Algorithmic procedures for Bayesian MEG/EEG source reconstruction in SPM<sup>☆</sup>



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### ABSTRACT

The MEG/EEG inverse problem is ill-posed, giving different source reconstructions depending on the initial assumptions. Parametric Empirical Bayes allows one to implement most popular MEG/EEG inversion schemes

# Thank you

- Karl Friston
- Gareth Barnes
- Vladimir Litvak
- Guillaume Flandin
- Will Penny
- Jean Daunizeau
- Christophe Phillips
- Rik Henson
- Jason Taylor
- Luzia Troebinger
- Chris Mathys
- Saskia Helbling

And all SPM developers