



# Bayesian Statistics in Neuroimaging

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# Bayesian Inference

- Most statistical methods covered in introductory statistics courses are so called frequentist (or classical) methods.
- **Bayesian inference** is another approach that provides a somewhat different perspective.
- Bayesian methods have recently received a great deal of attention in fMRI research.

# Classical vs Bayesian Approach

- The frequentist (or classical) point of view:
  - Probability refers to limiting relative frequencies.
  - Parameters are fixed unknown constants. Because they do not fluctuate, no useful probability statements can be made about them.
  - Statistical procedures should be designed to have well-defined long run frequency properties.

# Classical vs Bayesian Approach

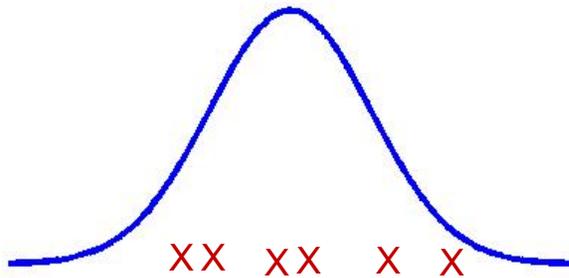
- The Bayesian point of view:
  - Probabilities describe a degree of belief.
  - Probability statements can be made about parameters, even though they are fixed constants.
  - Inferences are made about a parameter  $\theta$  by producing a probability distribution for it.

# The Bayesian Method

- Choose a probability density  $p(\theta)$ , the **prior distribution**, that expresses our beliefs about a parameter  $\theta$  before we see any data.
- Choose a statistical model  $p(y|\theta)$ , the **likelihood**, that reflects our belief about  $y$  given  $\theta$ .
- After observing  $y$ , update our beliefs and calculate the **posterior distribution**  $p(\theta|y)$ .

# Example

- Suppose we observe a sequence of observations  $y_1, \dots, y_n$  from a  $N(\mu, \sigma^2)$  distribution, where  $\mu$  is unknown and  $\sigma^2$  known.



Assume  $\mu$  is the task-induced change in brain activity and  $y_i$  is the equivalent contrast for subject  $i$ .

Likelihood:  $p(y_1, \dots, y_n | \mu) = N(\mu, \sigma^2)$

- We are interested in estimating the parameter  $\mu$ .

# Prior Distribution

- In the **Bayesian approach**  $\mathbb{W}$  can be described by a probability distribution. (**Prior Distribution**)
- The prior distribution is a subjective distribution, based on the experimenter's belief and is formulated prior to viewing the data.
- In our example assume  $p(\mathbb{W}) = N(\mathbb{W}_0, \mathbb{W}_0^2)$

# Posterior Distribution

- After a sample is taken from a population, the prior distribution can be updated using the information contained in the sample.
- The updated prior is called the **posterior distribution**. Updating is done using Bayes Rule:

$$p(\theta | y) = \frac{p(y | \theta)p(\theta)}{p(y)}$$

- Note that  $p(y)$  does not depend on .
- Hence, the posterior density is often written:

$$p(\theta | y) \propto p(y | \theta) p(\theta)$$

Posterior                  Likelihood                  Prior

# Posterior Inference

- The posterior distribution contains all current information about the parameter  $\psi$ .
- Numerical summaries (e.g., mean, median, mode) of the distribution are used to obtain point estimates of the parameter.
- We can also make probability statements about the parameter of interest and create posterior intervals.

# Example

- Let  $y_1, \dots, y_n$  be observations from a  $N(\theta, \sigma^2)$  distribution, with  $\theta$  unknown and  $\sigma^2$  known.
- Suppose we take the prior distribution of  $\theta$  to be  $N(\mu_0, \tau_0^2)$  for some choice of  $\mu_0$  and  $\tau_0^2$ .

$$p(\theta | y_1, \dots, y_n) \propto p(y_1, \dots, y_n | \theta) p(\theta)$$

$$\propto \exp\left\{-\frac{1}{2\sigma^2} \sum_{i=1}^n (y_i - \theta)^2\right\} \exp\left\{-\frac{1}{2\tau_0^2} (\theta - \mu_0)^2\right\}$$

- It can be shown that

$$\mu_n | y_1, \dots, y_n \sim N(\mu_n, \tau_n^2)$$

where

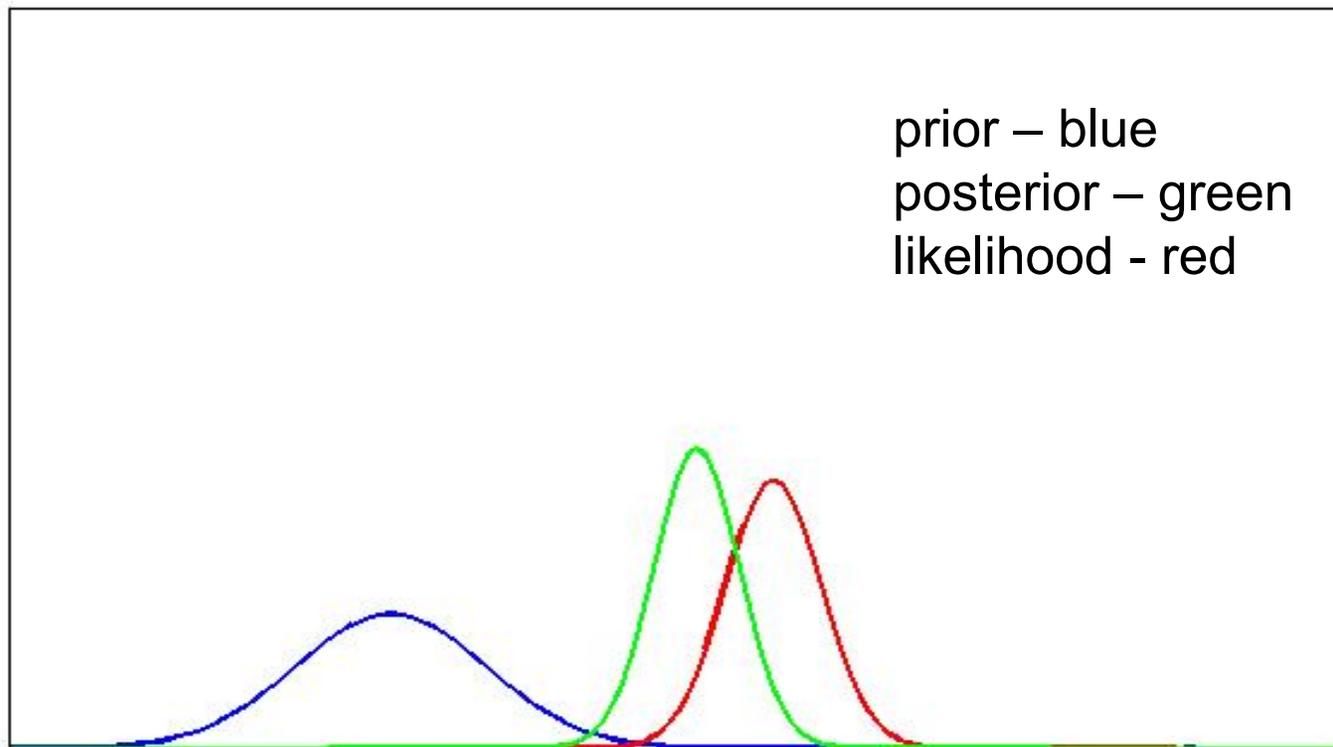
$$\mu_n = \frac{\frac{1}{\tau_0^2} \mu_0 + \frac{n}{\sigma^2} \bar{y}}{\frac{1}{\tau_0^2} + \frac{n}{\sigma^2}} \quad \text{and} \quad \frac{1}{\tau_n^2} = \frac{1}{\tau_0^2} + \frac{n}{\sigma^2}$$

- Note:  $\mu_n = w\mu_0 + (1-w)\bar{y}$

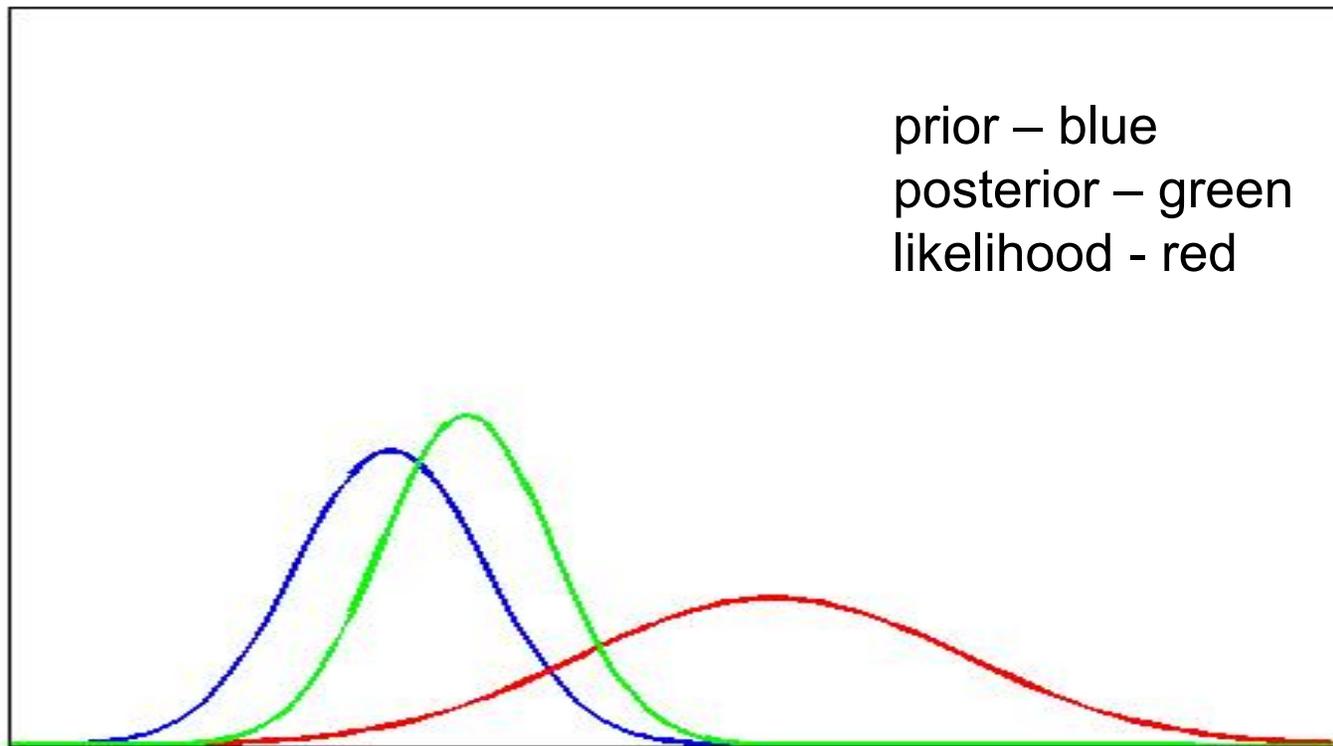
# Precision

- The inverse of the variance is called the **precision**.
- The posterior mean is expressed as a weighted average of the prior mean and the sample mean.
- The weights are proportional to the precisions.
- The posterior mean lies between the prior mean and the sample mean.

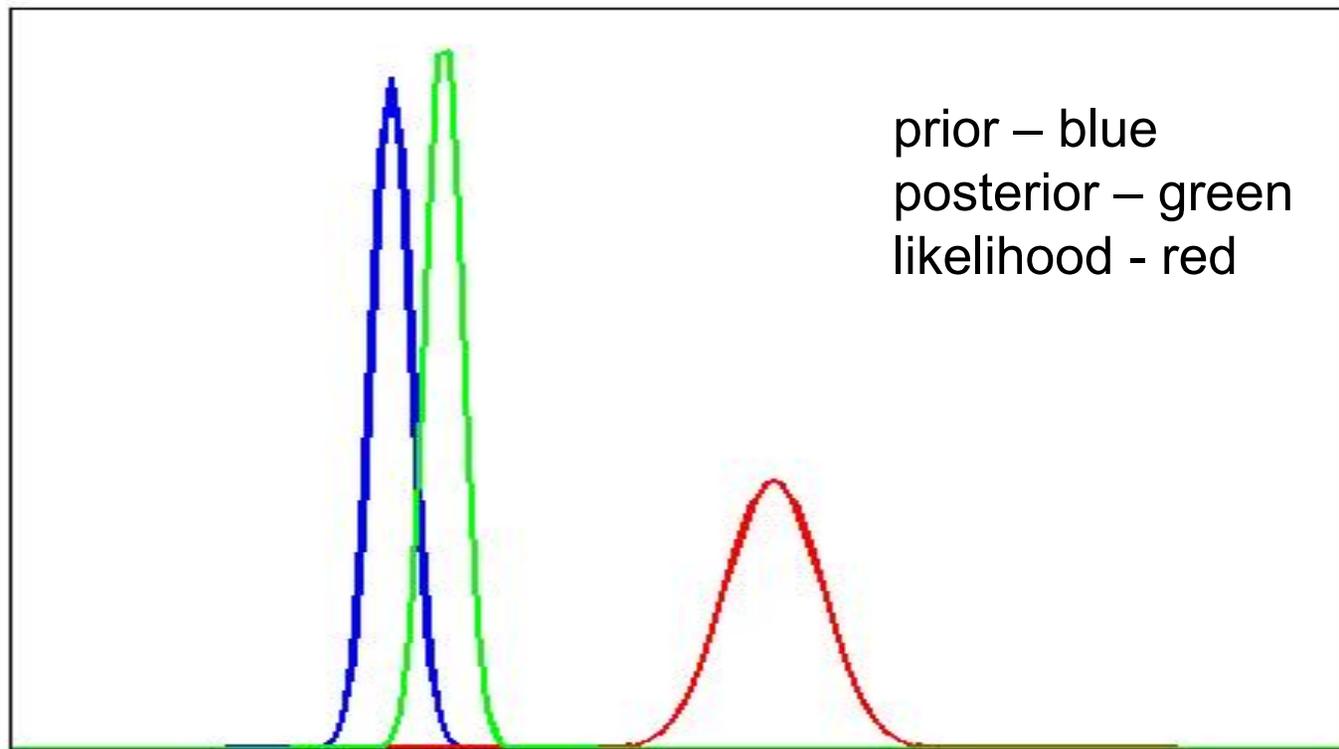
# Illustration



# Illustration



# Illustration



# Priors

- In the Bayesian framework the choice of prior is crucial.
- If we have no prior information about the parameters, we may use **non-informative priors**. These types of priors let the data speak for itself.
- One can also choose the priors in such a way that the posterior lies in the same family of distributions as the prior (**conjugate priors**).

# Posterior Computation

- The posterior distribution is the basis for all Bayesian inference.
- Even if the posterior is known, it can be difficult to obtain exact values of certain posterior quantities (e.g.,  $E(\frac{\mathbb{W}_1}{\mathbb{W}_2})$ ).
- By generating random samples from the posterior, all quantities of interest can be approximated using **Monte Carlo methods**.

# Monte Carlo Method

- Let  $g(\theta)$  be some function of  $\theta$  (e.g.,  $\log(\theta)$ ).
- Suppose we want to estimate  $E(g(\theta)|y)$ .
- Generate an i.i.d sequence  $\theta_1, \dots, \theta_N$  from the posterior distribution of  $\theta$ .
- Estimate  $E(g(\theta)|y)$  using  $\bar{g} = \frac{1}{N} \sum_{i=1}^N g(\theta_i)$

# Bayesian Computations

- Sampling from the posterior is effective when it can be implemented.
- However, it is often difficult in practice.
- For most probability distributions there is no simple way to simulate random variables of that particular distribution.

# MCMC

- **Markov-chain Monte-Carlo (MCMC)** is a method for sampling from a posterior distribution.
  - A Markov chain is generated that has the desired distribution as its **stationary distribution**.
  - The state of the chain after a large number of steps is used as a sample from the desired distribution.
  - Can be extremely computationally expensive.

# Variational Bayes

- **Variational Bayes (VB)** is an approach towards approximating the posterior density which is less computationally intensive than MCMC.
  - Received a lot of attention in fMRI research.
  - It allows one to approximate the posterior density with another density that has a more analytically tractable form.

# GLM with Priors

- Consider the standard GLM:

$$Y = X\beta + \varepsilon \quad \varepsilon \sim N(0, V)$$

- Suppose we place a prior on  $\beta$ , e.g.

$$\beta \sim N(\beta_0, \Sigma_0)$$

- It can be shown that the posterior distribution of  $\beta$  follows a normal distribution. We can use this distribution to perform inference.

- The posterior mean provides a point estimate of  $\beta$ :

$$\hat{\beta} = (X^T V^{-1} X + \Sigma_0^{-1})^{-1} (X^T V^{-1} y + \Sigma_0^{-1} \beta_0)$$

- If  $\lambda$  large then  $\hat{\beta} \approx (X^T V^{-1} X)^{-1} X^T V^{-1} y$

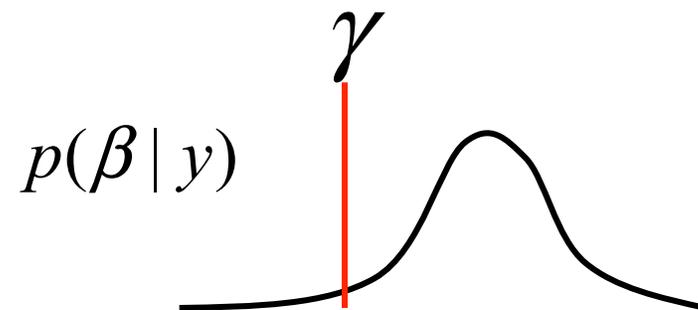
GLS estimate

- If  $\lambda = 0$  then  $\hat{\beta} = (X^T V^{-1} X + \Sigma_0^{-1})^{-1} X^T V^{-1} y$

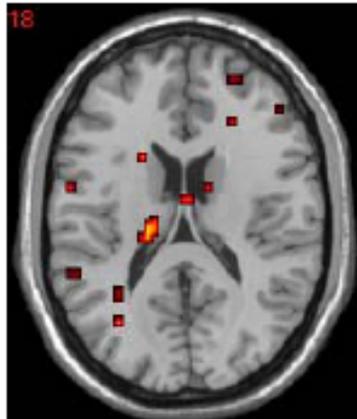
Shrinkage

# Posterior Probability Maps

- The Posterior distribution is the probability of getting an effect, given the data  $p(\beta | y)$ .
- **Posterior probability maps** are images of the probability or confidence that an activation exceeds some specified threshold, given the data  $p(\beta > \gamma | y) > \alpha$

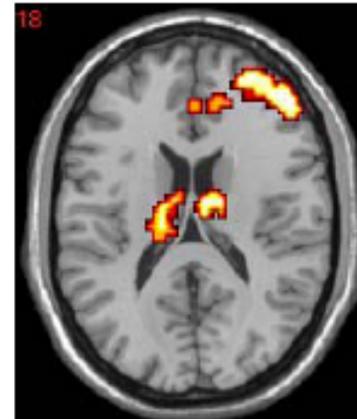


# Illustration



## Frequentist

Thresholded t-statistic map  
( $p=0.005$ , uncorrected)



## Bayesian PPM

Voxels with probabilities of  
task-related increases in  
activity exceeding  $\alpha=0.85$ .

Bayesian Spatial Hierarchical Model  
[Bowman et al., 2008, *NeuroImage*.]

# Comments

- Frequentist and Bayesian methods are answering different questions.
- To combine prior beliefs with data in a principled way use Bayesian inference.
- To construct procedures with guaranteed long run performance use frequentist methods.

# Hypothesis Testing

- In classical hypothesis testing we seek to determine whether we can reject a null hypothesis of no effect.
  - The  $p$ -value is the probability of obtaining a result as or more extreme under the assumption that the null hypothesis is true.
  - We can never accept the null hypothesis.
  - Given enough data every voxel would be significant.
- Bayesian methods allows us to derive probabilities about hypothesis of interest.
  - Not restricted to disproving the null hypothesis.
  - One can argue that it is more interesting to prove a hypothesis, than disprove a hypothesis of no effect.

# Multilevel Models

- Data sets where there is a hierarchy of nested populations are often called **multilevel**.
  - Voxels nested within subjects nested within groups.
- **Multilevel models** are extensions of regression in which data are structured in groups and coefficients can vary by group.
  - Allows information to be shared across groups.

# Multilevel GLM

$$Y = X\beta + \varepsilon \quad \varepsilon \sim N(0, V)$$

$p(y|\mathbb{W}) = N(X \mathbb{W}, V)$  represents variability within a subject.

$$\beta = \beta_g + \eta \quad \eta \sim N(0, \Sigma)$$

$p(\mathbb{W}|\mathbb{W}_g) = N(\mathbb{W}_g, \mathbb{W})$  represents variability across subjects

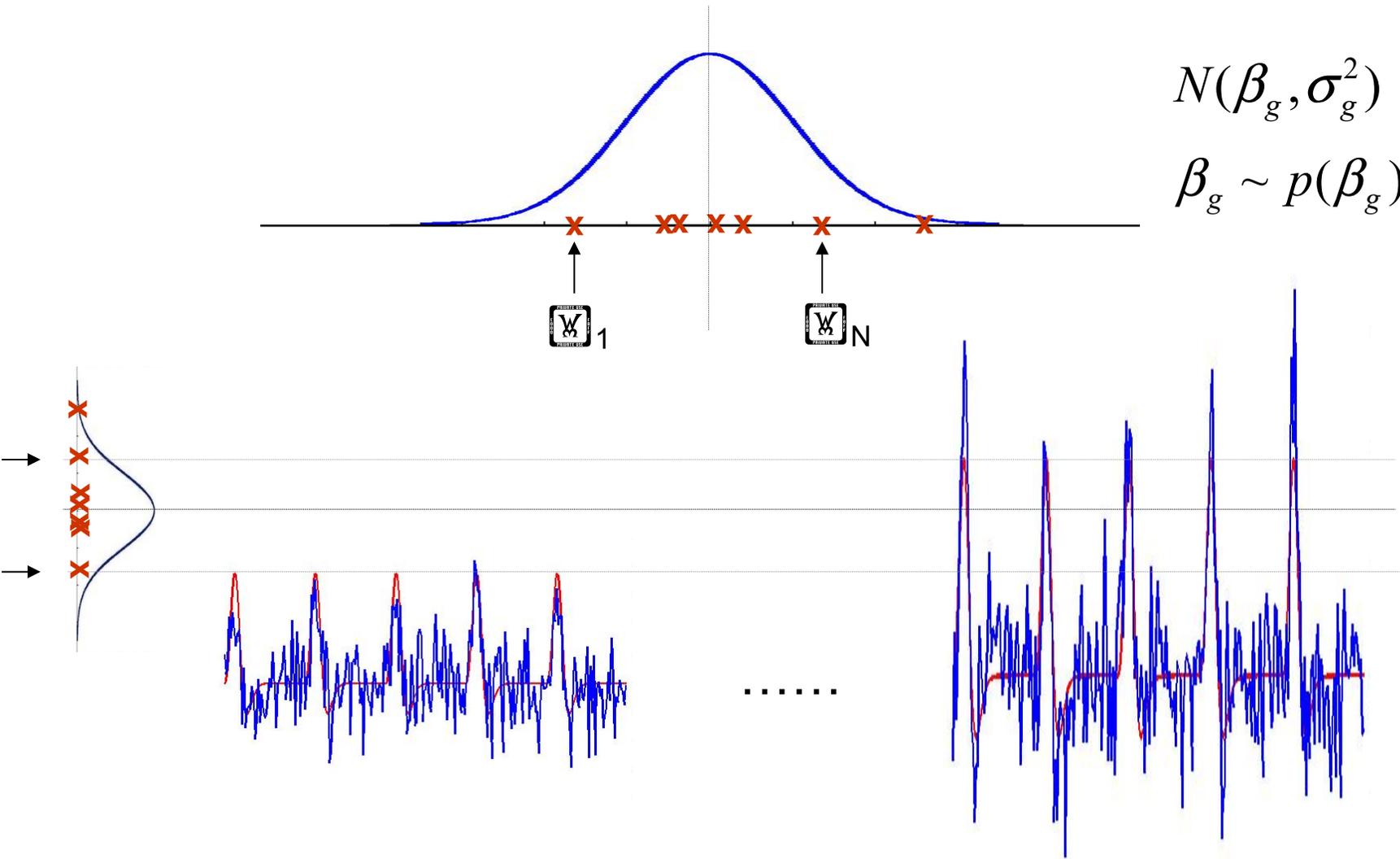
$$\beta_g \sim p(\beta_g)$$

$p(\mathbb{W}_g)$  represents information about a fixed but unknown quantity.

# Illustration

$$N(\beta_g, \sigma_g^2)$$

$$\beta_g \sim p(\beta_g)$$



# Empirical Bayes

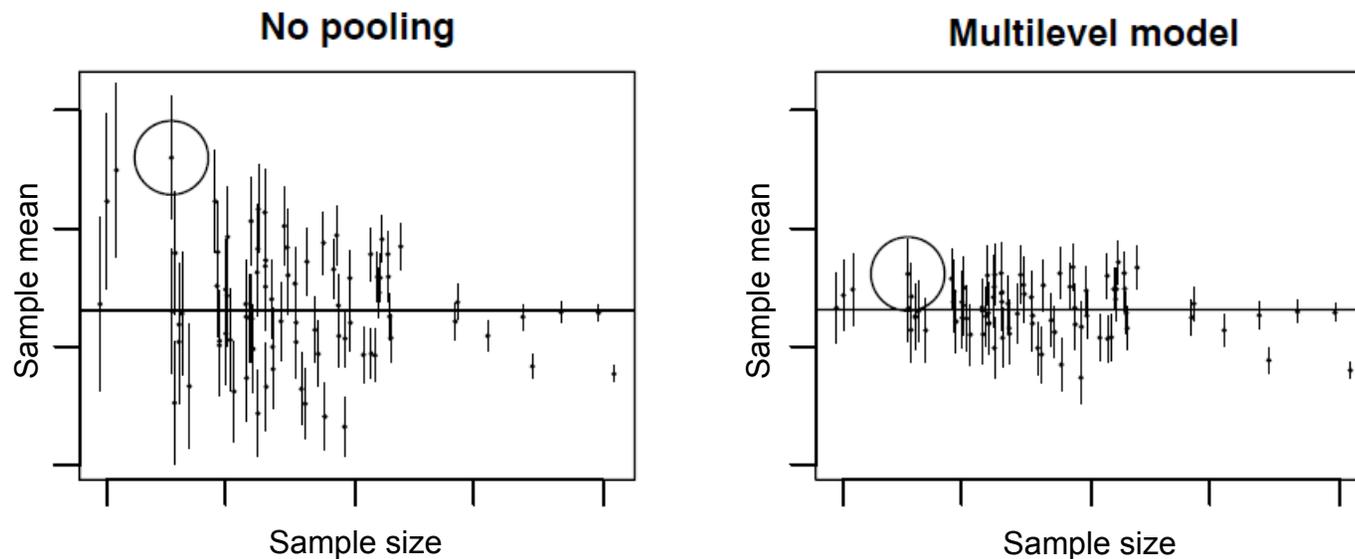
- It is common in neuroimaging to use so-called **empirical Bayes** methods.
- Here the parameters of the prior are estimated directly from the data, rather than being subject to prior specification of their own as is the case in a fully Bayesian model.

# Shrinkage

- Multilevel models allow for heterogeneity across subjects, but still consider values observed in other subjects.
- Each subject-specific estimate gets shrunk towards the overall estimate.
  - The greater the uncertainty, the more shrinkage.
  - The less the uncertainty, the more we trust that individual estimate and the less it gets shrunk.

# Example

- Radon levels of houses in 85 counties in Minnesota.



- Most differences no longer statistically significant

# Model Comparison

- Model comparison can be performed to determine whether the data favors one model over another.
- The **model evidence** is defined as

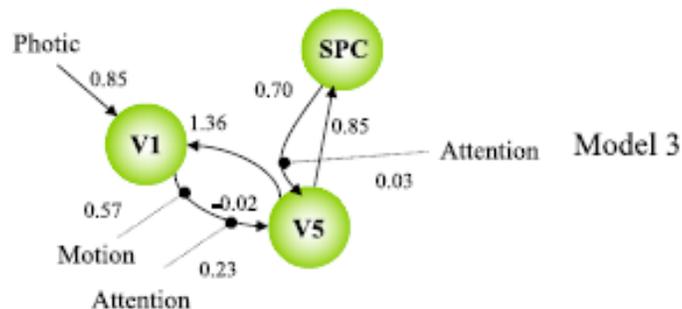
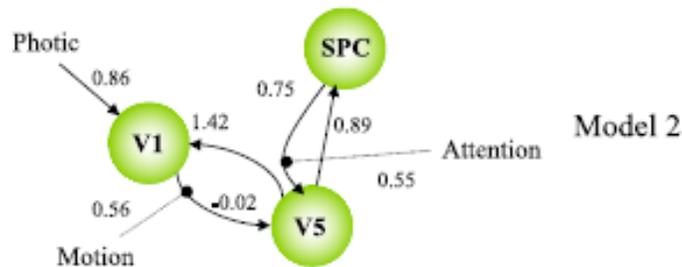
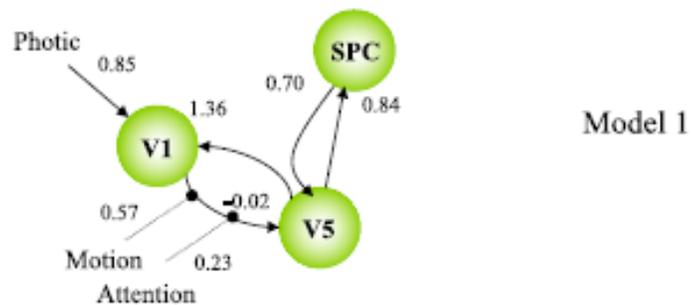
$$p(y | m) = \int p(y | \theta, m) p(\theta | m) d\theta$$

- The **Bayes factor** for comparing model i to j:

$$B_{ij} = \frac{p(y | m = i)}{p(y | m = j)}$$

If  $B_{ij}$  is large than i more likely than j.

# Example



Use Bayes factors to compare three different candidate models.

Table 6

Attention data—comparing modulatory connectivities

	$B_{12}$	$B_{13}$	$B_{32}$
AIC	3.56	2.81	1.27
BIC	3.56	19.62	0.18

Bayes factors provide consistent evidence in favor of the hypothesis embodied in model 1, that attention modulates (solely) the bottom-up connection from V1 to V5. Model 1 is preferred to models 2 and 3.

# Summary

- Bayesian methods are gaining in popularity in fMRI research.
- They allow us to calculate the probability that an activation exceeds some specific threshold, given the data.
  - Not restricted to disproving a null hypothesis.
- Requires specifying prior distributions and can be computationally expensive.