

### ♠ Conjugate Priors on Regression Parameters

- Model in matrix form:

$$\mathbf{y} = \mathbf{X}'\boldsymbol{\beta} + \boldsymbol{\epsilon}$$

- Use notation  $\mathbf{H} = \mathbf{X}'$  for comparability with traditional statistics notation:  $\mathbf{H}$  is  $n \times p$  design matrix

$$\mathbf{y} = \mathbf{H}\boldsymbol{\beta} + \boldsymbol{\epsilon}, \quad \boldsymbol{\epsilon} \sim N(0, \sigma^2 \mathbf{I}) = N(0, \phi^{-1} \mathbf{I})$$

### ♠ LSE formulæ:

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$$\hat{\boldsymbol{\beta}} = \mathbf{V}\mathbf{H}'\mathbf{y} \quad \text{and} \quad \mathbf{V}^{-1} = \mathbf{H}'\mathbf{H}$$

### ♠ (Simple) Standard Shrinkage Priors

- Conjugate normal prior centred at zero: shrinkage towards zero
- Genes centred: zero is canonical “null” hypothesis of no (partial) association
- Scale: scale of gene expression, standardised scales

$$\boldsymbol{\beta}|\phi \sim N(0, \tau\phi^{-1}\mathbf{I})$$

- Intercept: include in  $\boldsymbol{\beta}$  - different status: different scale?

$$\boldsymbol{\beta}|\phi \sim N(0, \phi^{-1}\mathbf{C}^{-1})$$

- Prior precision matrix  $\mathbf{C}$  (up to constant  $\phi$ ) is diagonal – common value  $\tau^{-1}$  for genes, possibly different (larger) for intercept
- Prior on residual variance/precision:  $\phi \sim Ga(a/2, b/2)$  with prior variance estimate  $b/a$  and degree of freedom  $a$ . Vague reference prior is special case  $a, b \rightarrow 0$

### ♠ Features of Posterior Under Shrinkage Prior

- $\boldsymbol{\beta}|\phi, \mathbf{y} \sim N(\mathbf{b}, \phi^{-1}\mathbf{B}^{-1})$
- $\mathbf{B} = \mathbf{C} + \mathbf{H}'\mathbf{H}$  and  $\mathbf{b} = \mathbf{B}^{-1}\mathbf{H}'\mathbf{y}$
- Special limiting cases: vague prior (zero precision) -  $\mathbf{B} = \mathbf{H}'\mathbf{H}$  and  $\mathbf{b} = \hat{\boldsymbol{\beta}}$
- Otherwise, shrinkage towards zero induced by prior:  $\|\mathbf{b}\| < \|\hat{\boldsymbol{\beta}}\|$
- Decision Theory: Bayes’s estimates improve expected performance in estimating  $\boldsymbol{\beta}$ 
  - quadratic (or other convex) loss functions
  - Bayesian and frequentist measures of risk in estimation
- In practical terms, shrinkage generally stabilises/regularises estimation and improves robustness in predictions
- Large  $p$  questions: LSE/likelihood methods inapplicable – need for regularisation
- Log posterior (conditional on  $\phi$ ) = log likelihood + log prior: a constant plus a term proportional to

$$(\mathbf{y} - \mathbf{H}\boldsymbol{\beta})'(\mathbf{y} - \mathbf{H}\boldsymbol{\beta}) + \boldsymbol{\beta}'\mathbf{C}\boldsymbol{\beta}$$

- Posterior mean/mode  $\mathbf{b}$  represents shrunken/regularised MLE (=LSE)

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### ♠ Marginal Posteriors for $\mathbf{b}$ and $\phi$

- $\phi|\mathbf{y} \sim Ga(a'/2, b'/2)$  with  $a' = a + n$  and  $b' = b + q$  where  $q > 0$  can be written in one of several useful forms:
  - simplest for computation:  $q = \mathbf{y}'\mathbf{e}$  where  $\mathbf{e} = \mathbf{y} - \mathbf{H}\mathbf{b}$  is the estimated residual vector
  - quadratic form representation:  $q = \mathbf{y}'\mathbf{P}'\mathbf{y}$  where  $\mathbf{P} = \mathbf{I} - \mathbf{H}\mathbf{B}^{-1}\mathbf{H}'$
  - interesting limiting case of vague reference prior: we know  $\mathbf{b} \rightarrow \hat{\beta}$  and now also  $q \rightarrow$  usual residual sum of squares in LSE regression with  $\mathbf{P} \rightarrow \mathbf{H}(\mathbf{H}'\mathbf{H})^{-1}\mathbf{H}'$
- Common point estimate of  $\sigma^2 = \phi^{-1}$  is reciprocal of posterior mean of  $\phi$ , namely  $s = b'/a'$
- $\beta|\mathbf{y} \sim T_{a'}(\mathbf{b}, s\mathbf{B}^{-1})$ 
  - credible intervals for elements of  $\beta$  are derived from Student T distribution on  $a'$  degrees of freedom
- (see matlab code: mregbayes)
- Role of  $\mathbf{C}$ , Key example:
  - $C_{1,1}$  = large (intercept term, vague prior)
  - $C_{i,i} = \tau^{-1}$  otherwise – common precision for (common scale) genes
  - Marginal likelihood function for  $\tau$  (see matlab code, examples) to assess  $\tau$ . Can be shown that marginal likelihood  $p(\mathbf{y}|\tau)$  implies

$$\log(p(\mathbf{y}|\tau)) = \text{constant} + 0.5 \log(|\mathbf{C}|/|\mathbf{B}|) - 0.5n \log(\mathbf{y}'\mathbf{e})$$

### ♠ Shrinkage Priors More Generally: Considerations

- Different degrees of shrinkage across variables (genes) via different diagonal elements in  $\mathbf{C}$
- General question of learning from data about shrinkage parameters: inference on  $\mathbf{C}$  generally
- Orthogonal designs: e.g., Factor regressions and other models: design is orthogonal (by design!)
  - $\mathbf{H}'\mathbf{H}$  is diagonal
  - so that  $\mathbf{B}$  is diagonal when  $\mathbf{C}$  is
  - elements of  $\beta$  are uncorrelated under the posterior as well as the prior (dependent only through scale  $\phi$ )