

♠ Multiple Linear Regression Model

- Recall the model in matrix form:

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

- Predictor variables in $p \times n$ matrix $\mathbf{X} = [\mathbf{x}_1, \mathbf{x}_2, \dots, \mathbf{x}_n]$ (columns are samples, rows are predictor variables)

♠ SVD of \mathbf{X}

- SVD is

$$\mathbf{X} = \mathbf{B}\mathbf{F} \quad \text{or} \quad \mathbf{X} = \mathbf{A}\mathbf{D}\mathbf{F}$$

where $\mathbf{F} = [\mathbf{f}_1, \mathbf{f}_2, \dots, \mathbf{f}_n]$ is $n \times n$ matrix of factors (columns represent samples, and rows represent factor variables)

♠ SVD Regression

- Combine the SVD with the regression model to get

$$\mathbf{y} = \mathbf{F}'\boldsymbol{\theta} + \boldsymbol{\epsilon}$$

with

- $\boldsymbol{\theta} = \mathbf{B}'\boldsymbol{\beta}$ or $\boldsymbol{\theta} = \mathbf{D}\mathbf{A}'\boldsymbol{\beta}$
- Multiple regression on the factor variables themselves as predictors
- n predictor variables, not p
- Regression parameter vector $\boldsymbol{\theta}$ to estimate
- Dimension reduction of inference/estimation problem when $p > n$, as is the case in gene expression analyses

♠ Bayesian Analysis and Stochastic Regularisation

- If $p > n$ we end up with n parameters to be estimated with n observations
- Least squares and other standard methods inapplicable: exact fit to observed data, no predictive value (“over-fitting”)
- Generally, remove some factors that do not vary or contribute much to the SVD (small values of the singular values in the \mathbf{D} matrix)
- More useful and formal solutions lie in Bayesian analysis that involves “stochastic regularisation” of the estimation problem – estimate $\boldsymbol{\theta}$ with some partial constraints on values imposed probabilistically
(*Insert two semesters of statistics in here please!*).
- Typically, reduce to a smaller number of factors and then apply Bayesian analysis to the rest
- Corresponding estimation of $\boldsymbol{\beta}$ via $\boldsymbol{\beta} = \mathbf{A}\mathbf{D}^{-1}\boldsymbol{\theta}$

♠ Software, Computation and Summary

- Point estimate analysis: iterative computation of estimates of $\boldsymbol{\theta}$ that are Bayesian *posterior modes* (EM algorithms, MAP estimation)
- Full Bayesian analysis using stochastic simulation methods (Markov chain Monte Carlo simulation, Gibbs sampling): see discussion in the *binary regression* context