

# BIOS 526 Course Summary

1. Ordinary least squares (M1):

$$\begin{aligned}\text{Cov } \mathbf{Y} &= \sigma^2 \mathbf{I} \\ \hat{\boldsymbol{\beta}}^{OLS} &= (\mathbf{X}'\mathbf{X})^{-1}\mathbf{X}'\mathbf{Y}\end{aligned}$$

2. Generalized least squares (M4, slide 16) is a formulation for any given covariance matrix:

$$\begin{aligned}\text{Cov } \mathbf{Y} &= \boldsymbol{\Sigma} \\ \hat{\boldsymbol{\beta}}^{GLS} &= (\mathbf{X}'\boldsymbol{\Sigma}^{-1}\mathbf{X})^{-1}\mathbf{X}'\boldsymbol{\Sigma}^{-1}\mathbf{Y}\end{aligned}$$

- Can apply to clustered data; in practice, use LMMs or GEEs.
- Can be used for heteroscedastic data; in practice, use GEE.
- Need to know  $\boldsymbol{\Sigma}$ ; in practice, estimated.

3. Generalized linear models (M3 part I):

$$\begin{aligned}y_i &\overset{ind}{\sim} P(\mathbf{x}'_i\boldsymbol{\beta}) \\ g(E(y_i)) &= \mathbf{x}'_i\boldsymbol{\beta}\end{aligned}$$

- logistic regression for 0, 1 response.  $\text{Var}(y_i) = E(y_i) \{1 - E(y_i)\}$ .
- Poisson regression for count data.  $\text{Var}(y_i) = E(y_i)$ . Watch out for overdispersion.

4. Generalized linear mixed models (M2, M3 part II):

Random intercept model:

$$\begin{aligned}y_{ij} &\sim P(\mathbf{x}'_{ij}\boldsymbol{\beta} + \theta_i) \\ g\{E(y_{ij}|\theta_i)\} &= \mathbf{x}'_{ij}\boldsymbol{\beta} + \theta_i \\ \theta_i &\overset{iid}{\sim} N(0, \tau^2)\end{aligned}$$

- Handle repeated measurements / longitudinal / clustered data.
- For Gaussian, interpretation of  $\beta$  not impacted by conditional versus marginal (the estimates of  $\beta$  from LMM and GEE are different but usually similar, in some cases GEE with exchangeable correlation structure and random intercept LMM have equivalent  $\hat{\beta}$ ).
- For logistic, interpretation of  $\beta$  in GLMM (conditional model) is different from the interpretation in a GEE (the estimates of  $\beta$  from GLMM and GEE are different).
- Use if interested in subject-specific predictions (shrinkage towards population effects).
- Can use if no overdispersion in logistic or Poisson, no heteroscedasticity in Gaussian.

5. Generalized estimating equations (M4):

$$y_{ij} \sim P(\mathbf{x}'_{ij}\beta)$$

$$\text{Cov}(\mathbf{y}_i) = \mathbf{D}_i^{1/2} \mathbf{R}(\alpha) \mathbf{D}_i^{1/2}$$

where  $\mathbf{R}(\alpha)$  is the working correlation and  $\mathbf{D}_i$  is a diagonal matrix with diagonal elements equal to the variance determined by the likelihood.

- Handle repeated measurements / longitudinal / clustered data.
- Use robust standard errors.
- Use if heteroscedasticity and/or overdispersion (valid inference, unlike GLMM).
- Marginal inference (no random effects).

6. Generalized additive models (M5):

$$g(E(y_i)) = \beta_0 + s_1(x_{i1}) + \cdots + s_j(x_{ip})$$

- Handle non-linear effects.
- Can incorporate random effects for longitudinal / repeated measures / clustered data.
- Can generalize interactions from linear models to bivariate splines, e.g.,  $s(x_{i1}, x_{i2})$ , i.e., 2D surfaces.
- Estimate  $s(x_{ip})$  using either cross-validation or mixed model formulation of spline coefficients.

7. Bias-Variance Tradeoff (M5, slides 33-43, M6 II slides 5-6)

- $MSE(\hat{f}(x)) = Var(\hat{f}(x)) + Bias(\hat{f}(x))^2$
- Fewer parameter: more bias, less variance

- More parameters: less bias, more variance
- Use cross-validation or generalized cross-validation to approximately minimize the MSE

8. Principal component analysis (M6 I): uses the singular value decomposition on standardized data:

$$\mathbf{X}_{scaled} = \mathbf{U}\mathbf{D}\mathbf{V}'$$

- Lower dimensional representation using first  $l$  left eigenvectors.
- Can use in principal component regression when have issues with multicollinearity.

9. Ridge Regression (L2-norm regularization) (M5 part II, M6 part II):

$$\hat{\boldsymbol{\beta}}^{Ridge} = (\mathbf{X}'\mathbf{X} + \lambda\mathbf{I})^{-1}\mathbf{X}'\mathbf{Y}$$

- Regularization method with a nice closed form.
- Can use when lots of covariates,  $p > n$ .
- Use for shrinking spline coefficients in GAMs (used in MGCV).

10. Lasso (L1-norm regularization) (M6 part II):

$$\hat{\boldsymbol{\beta}}^{Lasso} = \underset{\boldsymbol{\beta}}{\operatorname{argmin}} - \sum_{i=1}^n \ell(y_i; \mathbf{x}'_i \boldsymbol{\beta}) + \lambda \|\boldsymbol{\beta}\|_1$$

- Regularization that results in variable selection by setting many coefficients equal to 0.

11. Elastic net (L1-norm and L2-norm regularization) (M6 part II):

$$\hat{\boldsymbol{\beta}}^{Elnet} = \underset{\boldsymbol{\beta}}{\operatorname{argmin}} - \sum_{i=1}^n \ell(y_i; \mathbf{x}'_i \boldsymbol{\beta}) + \lambda \sum_{j=1}^p \left( \alpha |\beta_j| + \frac{(1-\alpha)}{2} \beta_j^2 \right).$$

- A good choice when predictors are correlated.
- Use for variable selection.

12. Bayesian statistics (M7):

- (a) Choose a prior  $P(\boldsymbol{\theta})$ .
  - (b) Choose a likelihood  $P(\mathbf{Y}|\boldsymbol{\theta})$ .
  - (c) Collect data and calculate posterior  $P(\boldsymbol{\theta}|\mathbf{Y})$ .
- Connection between regularized regression and priors on the parameters.
  - In Bayesian regression with a Gaussian likelihood and a Gaussian prior, equivalent to ridge regression
  - In Bayesian regression with a Laplace (i.e., double exponential) prior, the maximum a posteriori estimate corresponds to the Lasso solution.
  - Parameters are random variables.
  - In practice, obtain samples from the posterior distribution.
  - Calculate credible intervals from quantiles of a posterior sample.