### Correlated Data Overview

### FW8051 Statistics for Ecologists

Department of Fisheries, Wildlife and Conservation Biology



# Learning Objectives

 Be able to model correlated binary and correlated count data using generalized linear mixed effect models (GLMMs) and generalized estimating equations (GEEs)

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- Be able to interpret parameters in linear and generalized linear mixed effects models
- Be able to describe models and their assumptions using equations and text and match parameters in these equations to estimates in computer output.

# Correlated Data Methods in Ecology

#### For data that are normally distributed:

- Linear mixed effects model
- Generalized Least Squares

### For count or binary data:

- Generalized linear mixed effects models (GLMMS)
- Generalized Estimating Equations (GEE)

# Correlated Data Methods in Ecology

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Fieberg, J., Rieger, R.H., Zicus, M. C., Schildcrout, J. S. 2009. Regression modelling of correlated data in ecology: subject specific and population averaged response patterns. Journal of Applied Ecology 46:1018-1025.

# Mallard Nesting structures



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# Mallard Nesting structures



**Research Questions**: which types of cylinders are best (single or double)? Where should they be placed?

### Mallard Data

- 110 nest structures placed in 104 wetlands
- Structure type (single versus double) was chosen randomly at each location
  - 53 single-cylinders, 57 double-cylinders
- Occupancy (0,1) and clutch sizes were recorded in 1997, 1998, 1999

Zicus, M.C., J. Fieberg and D. P. Rave. 2003. Does mallard clutch size vary with landscape composition: a different view. Wilson Bulletin 114:409-413.

Zicus, M. C., D. P. Rave, and J. Fieberg. 2006. Cost effectiveness of single-vs. double-cylinder over-water nest structures. Wildlife Society Bulletin 34:647-655.

Zicus, M. C., Rave, D. P., Das, A., Riggs, M. R., and Buitenwerf, M. L. (2006). Influence of land use on mallard nest-structure occupancy. The Journal of wildlife management, 70(5), 1325-1333.

# Example Clutch Size Data

 $Y_{ij}$  = clutch size for the  $i^{th}$  structure during year j

 ${\tt Init.Date}_{ij} = {\tt nest}$  initiation date (Julian day) for the  $i^{th}$  structure during year j

I (deply=2)  $_{i}$  = 0 if  $i^{th}$  structure is a single cylinder, 1 if double cylinder

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### **Model**

$$Y_{ij} = (\beta_0 + b_{0i}) + \beta_1 \text{Init.Date}_{ij} + \beta_2 \text{I(deply=2)}_i + \epsilon_{ij}$$

$$\epsilon_{ij} \sim N(0, \sigma^2)$$

$$b_{0i} \sim N(0, \tau^2)$$

Assume  $\epsilon_{ij}$  and  $b_{0i}$  are independent.

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$$\begin{aligned} Y_{ij} &= (\beta_0 + b_{0i}) + \beta_1 \texttt{Init.Date}_{ij} + \beta_2 \texttt{I} \, (\texttt{deply=2})_{\,i} + \epsilon_{ij} \\ &\epsilon_{ij} \sim N(0, \sigma^2) \\ &b_{0i} \sim N(0, \tau^2) \end{aligned}$$

Assume  $\epsilon_{ij}$  and  $b_{0i}$  are independent.

Similar to including "structure" as a series of dummy variables with the added assumption that the parameters are drawn from a normal distribution

# Generalized Least Squares

For normally distributed response data, we can fit correlated data models without having to resort to random effects:

$$\begin{aligned} \text{Clutch size}_{ij} &= \beta_0 + \beta_1 \text{Init.Date}_{ij} + \beta_2 \text{I (deply=2)}_{i} + \epsilon_{ij} \\ &\epsilon_{ij} \sim N(0,\Omega) \end{aligned}$$

 $\Omega$  = Var/Cov matrix for  $\epsilon$ . We no longer assume the errors,  $\epsilon_{ij}$ , are independent!

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Example: compound symmetric covariance matrix for data within each cluster:

$$\Omega = \begin{bmatrix} \Sigma_i & 0 & \cdots & 0 \\ 0 & \ddots & \ddots & \vdots \\ \vdots & \ddots & \ddots & 0 \\ 0 & \cdots & 0 & \Sigma_i \end{bmatrix} \text{ with } \Sigma_i = \begin{bmatrix} \sigma^2 & \rho\sigma^2 & \cdots & \rho\sigma^2 \\ \rho\sigma^2 & \ddots & \ddots & \vdots \\ \vdots & \ddots & \ddots & \rho\sigma^2 \\ \rho\sigma^2 & \cdots & \rho\sigma^2 & \sigma^2 \end{bmatrix}$$

Correlation between:

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This model is equivalent to a linear mixed effects model with random intercepts for each structure!

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### 2 Options:

- Start with generalized linear models (logistic or Poisson regression) and add random coefficients (intercepts, slopes)
- Use generalized estimating equations or cluster-level bootstrap (recognizing clusters serve as independent units)

### Option 1: Generalized linear mixed effects models

Option 2: Generalized Estimating equations

Systematic component: 
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- $f(y_i|x_i)$  is in the exponential family (includes normal, Poisson, binomial, gamma, inverse Gaussian)
- $f(y_i|x_i)$  that describes unmodeled variation about  $\mu_i = E[Y_i|X_i]$

## Generalized Linear Mixed Effects Models

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•  $Y_{ij}|b_{0i},b_{1i} \sim f(y_{ij}|b_{0i},b_{1i})$ , with  $f(y_i|b_{0i},b_{1i})$  in the exponential family (e.g., Poisson, binomial).

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- $b_{0i}$  and  $b_{1i}$  allow intercepts and parameters to vary among groups.
- Usually assume:  $(b_{0i}, b_{1i}) \sim N(0, D)$

## Conditional models

#### Poisson-normal model:

- $Y_{ij}|b_i \sim \text{Poisson}(\lambda_{ij})$
- $log(\lambda_{ij}) = (\beta_0 + b_{0i}) + (\beta_1 + b_{1i})x_{ij}$
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### Logistic-normal model:

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Frequentist analysis: the b's are unobserved random variables... we do not estimate them! Rather, we estimate  $\beta_0, \beta_1, D$ .

# Non-linear models: Modeling Occupancy Probability

Logistic-normal model:

$$\begin{aligned} Y_{ij}|b_i \sim \text{Bernoulli}(p_{ij})\\ \log[p_{ij}/(1-p_{ij})|b_i] &= \beta_0 + b_{0i} + \beta_1 VOM_{ij} + \beta_2 \text{I (deply=2)}_i\\ b_{0i} \sim N(0,\tau^2) \end{aligned}$$

Structures have different "propensities" of being occupied:

• Depending on visual obstruction (VOM), structure type, and other unmeasured characteristics ( $b_{0i}$ ) associated with the structure and the landscape in which it is placed.

### Parameter Estimation

To estimate parameters using Maximum Likelihood, we need to determine:

• the distribution of Y (not  $Y|b_i$ )

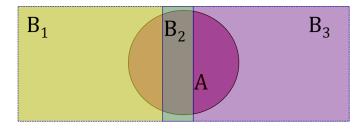
### Parameter Estimation

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Total law of Probability If events  $B_1, B_2, \dots, B_k$  are mutually exclusive and together make up all possibilities, then:

$$P(A) = \sum_{i} P(A|B_i)P(B)$$



## Unconditional Model and Likelihood

To determine the distribution of  $Y_{ij}$ , we integrate over the random effects:

$$L(Y_{ij}|\beta,D) = \int f(Y_{ij}|b_i)f(b_i)db_i$$

- $\beta$  = fixed effects parameters in  $f(Y_{ij}|b_i)$
- D are variance parameters of the random effects distribution,  $f(b_i)$

# Normally Distributed Data: Linear Mixed Effects Models

$$Y_{ij}|b \sim N(\mu_{ij}, \sigma^2)$$
  

$$\mu_{ij} = X_{ij}\beta + Z_ib$$
  

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If we average over (or integrate out) the random effects (*b*), we get the marginal Distribution of *Y*:

$$Y \sim MVN(X\beta, \Omega)$$
  
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$$0 \quad \cdots \quad 0$$

$$0 \quad \cdots \quad \vdots$$

$$\vdots \quad \cdots \quad 0$$

$$0 \quad \cdots \quad 0 \quad 1$$

$$(n \times n)$$

#### $Y \sim MVN(X\beta, \Omega)$

#### Random Intercept Model:

$$\Omega = \begin{bmatrix} \Sigma_i & 0 & \cdots & 0 \\ 0 & \ddots & \ddots & \vdots \\ \vdots & \ddots & \ddots & 0 \\ 0 & \cdots & 0 & \Sigma_i \end{bmatrix} \text{ with } \Sigma_i = \begin{bmatrix} \tau^2 + \sigma^2 & \tau^2 & \cdots & \tau^2 \\ \tau^2 & \ddots & \ddots & \vdots \\ \vdots & \ddots & \ddots & \tau^2 \\ \tau^2 & \cdots & \tau^2 & \tau^2 + \sigma^2 \end{bmatrix}$$

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GLS Model with compound-symmetric correlation structure:

$$\Omega = \begin{bmatrix} \Sigma_i & 0 & \cdots & 0 \\ 0 & \ddots & \ddots & \vdots \\ \vdots & \ddots & \ddots & 0 \\ 0 & \cdots & 0 & \Sigma_i \end{bmatrix} \text{ with } \Sigma_i = \begin{bmatrix} \sigma^2 & \rho\sigma^2 & \cdots & \rho\sigma^2 \\ \rho\sigma^2 & \ddots & \ddots & \vdots \\ \vdots & \ddots & \ddots & \rho\sigma^2 \\ \rho\sigma^2 & \cdots & \rho\sigma^2 & \sigma^2 \end{bmatrix}$$

$$Y_{ij}|b_i \sim Bernouli(p_{ij})$$

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• 
$$f(Y_{ij}|b_i) = p_{ij}^{Y_{ij}} (1 - p_{ij})^{1 - Y_{ij}}$$
  
•  $f(b_i) = \frac{1}{\sqrt{2\pi}\sigma_b} e^{\frac{(b_{0i} - 0)^2}{2\sigma_b^2}}$ 

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$$L(Y_{ij}|\beta, D) = \int f(Y_{ij}|b_i)f(b_i)db_i$$

$$\int \left[ \frac{exp(\beta_0 + b_{0i} + \beta_1 x_{ij})}{1 + exp(\beta_0 + b_{0i} + \beta_1 x_{ij})} \right]^{Y_{ij}} \left[ \frac{1}{1 + exp(\beta_0 + b_{0i} + \beta_1 x_{ij})} \right]^{1 - Y_{ij}} \frac{1}{\sqrt{2\pi}\sigma_b} e^{\frac{(b_{0i} - 0)^2}{2\sigma_b^2}} db_{0i}$$

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No closed-form solution!

How do we use maximum likelihood to estimate parameters?

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- Use numerical integration (can be slow, difficult, particularly with multiple random effects)
- Add priors, and use Bayesian techniques

### Numerical Integration: glmer

#### nAGQ

- specifies number of points per axis for evaluating the adaptive Gauss-Hermite approximation to the log-likelihood.
- default = 1 (Laplace approximation)
- values > 1 produce greater accuracy in the evaluation of the log-likelihood at the expense of speed.
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See also mixed\_model in the GLMMadaptive package.

## Linear versus Generalized Linear Mixed Effects Models

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Parameters in generalized linear mixed effects models have a "subject-specific", but not "population-average" interpretation.

Clutch size = 
$$(\beta_0 + b_{0i}) + \beta_1$$
Init.Date<sub>ij</sub> +  $\beta_2$ I (deply=2)<sub>i</sub> +  $\epsilon_{ij} \sim N(0, \sigma^2)$ 

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How does clutch size vary with nest initiation date and structure type for a "typical" structure (i.e., one with  $b_{0i} = 0$ )?

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  $E[Y|X,b_{0i}=0]=eta_0+eta_1$ Init.Date $+eta_2$ I(deply=2)

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Fixed effects parameters have both population-averaged and subject-specific interpretations!

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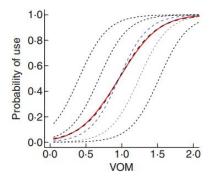
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is  $\beta_2$  meaningful? What if the explanatory variable was "sex" in a model where animals were repeatedly sampled?

### Interpretation of Parameters: GLMMS



 $E[Y_{ij}|X]$  (red curve) is no longer the same as  $E[Y_{ij}|X,b_{0i}=0]$  (blue curve)!

$$Y_{ij}|b_{0i}, b_{1i} \sim f(y_{ij}|b_{0i}, b_{1i})$$
  
 $(b_{0i}, b_{1i}) \sim N(0, D)$ , with

 $f(y_i|b_{0i},b_{1i})$  given by Poisson, binomial, negative binomial.

How can we quantify how E[Y|X] changes with X (as opposed to  $E[Y|X,b_i]$  or  $E[Y|X,b_i=0]$ )?

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- mixed\_model + marginal\_coefs in GLMMadaptive package to estimate equivalent "marginal coefficients" (based on Hedeker et al. 2018). See simlogit file on Canvas.

Hedeker, D., du Toit, S. H., Demirtas, H. and Gibbons, R. D. (2018), A note on marginalization of regression parameters from mixed models of binary outcomes. Biometrics 74, 354-361.

#### **GLMM** Resources on Canvas

#### Readings:

- Bolker et al. 2008. Generalized linear mixed models: a practical guide for ecology and evolution
- Bolker et al. 2013: Strategies for fitting nonlinear ecological models in R, AD Model Builder, and BUGS

#### Useful Links

- GLMM wiki
- GLMMS worked examples

Kery (Ch. 16 and 19), Zuur et al. Ch 13.

Species Richness versus NAP and Exposure (low)

Open up the template RichnessGLMM\_t.R:

• Fit generalized linear mixed effects models using glmer

Option 1: Generalized linear mixed effects models

Option 2: Generalized Estimating equations

### Generalized Estimating Equations

- motivation (least squares, maximum likelihood...)
- assumptions and implementation of GEE approach

For Normally distributed data:

$$L(\mu, \sigma^2 | y_1, y_2, \dots, y_n) = \prod_{i=1}^n \frac{1}{\sqrt{2\pi}\sigma} \exp\left(-\frac{(y_i - \mu)^2}{2\sigma^2}\right)$$

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With linear regression, we assume  $Y_i \sim N(\beta_0 + x_i\beta_1, \sigma^2)$ , so...

$$L(\beta_0, \beta_1, \sigma | x) = \prod_{i=1}^n \frac{1}{\sqrt{2\pi}\sigma} \exp\left(-\frac{(y_i - \beta_0 + x_i \beta_1)^2}{2\sigma^2}\right)$$
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$$\Rightarrow log L = -nlog(\sigma) - \frac{n}{2}log(2\pi) - \sum_{i=1}^n \frac{(y_i - \beta_0 + x_i \beta_1)^2}{2\sigma^2}$$

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$$\begin{split} L(\beta_0,\beta_1,\sigma|x) &= \prod_{i=1}^n \frac{1}{\sqrt{2\pi}\sigma} \exp\left(-\frac{(y_i-\beta_0+x_i\beta_1)^2}{2\sigma^2}\right) \\ &= \frac{1}{(\sqrt{2\pi}\sigma)^n} \exp\left(-\sum_{i=1}^n \frac{(y_i-\beta_0+x_i\beta_1)^2}{2\sigma^2}\right) \\ \Rightarrow logL &= -nlog(\sigma) - \frac{n}{2}log(2\pi) - \sum_{i=1}^n \frac{(y_i-\beta_0+x_i\beta_1)^2}{2\sigma^2} \\ \Rightarrow \text{maximizing logL} \Rightarrow \text{minimizing } \sum_{i=1}^n \frac{(y_i-\beta_0+x_i\beta_1)^2}{2\sigma^2} \\ \text{or, equivalently } \sum_{i=1}^n (y_i-\beta_0-x_i\beta_1)^2 \end{split}$$

#### Least squares

$$\sum_{i=1}^{n} (y_i - E[Y_i|X_i])^2 = \sum_{i=1}^{n} (y_i - (\beta_0 + x_{1i}\beta_1 + \dots))^2$$

Least squares leads to the following set of equations for estimating parameters (take the derivative and set = 0):

$$2\sum_{i=1}^{n} \frac{\partial E[Y_i|X_i]}{\partial \beta} (Y_i - E[Y_i|X_i]) = 0, \text{ or}$$
$$\sum_{i=1}^{n} X(Y_i - E[Y_i|X_i]) = 0$$

#### Generalized Linear Models

Maximum Likelihood estimators are found by solving:

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#### Poisson Regression:

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GEE: 
$$\hat{\beta}$$
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 $R_i$  = working correlation model that describes within subject correlation.

• Examples include exchangeable (equal correlation among all observations), Ar(1) (time series), unstructured

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- Fixed effects parameters have a population-averaged interpretation (how does y change across the population of structures that have different values of x)
- Fit using geeglm in geepack library

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Uses "robust" (or "sandwich") standard errors, treating clusters as independent observational units

•  $\hat{\beta} \pm 1.96SE$  gives valid CIs (for large numbers of clusters)

Species Richness versus NAP and Exposure (low)

Open up the template RichnessGEEt.R:

• Fit generalized estimating equations using independence and exchangeable working correlation structures