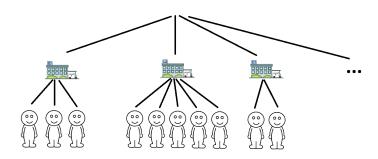
# Lecture 13: Clustered Data, An Introduction to Applied Mixed Models

Author: Nick Reich / Transcribed by Gregory Guranich, edited by Bing Miu

Course: Categorical Data Analysis (BIOSTATS 743)

#### Clustered Data

- Clustered Data:
- Hierarchy, nested populations
- Longitudinal, Correlated observations or sets .eg repeated measure
- Example: (Hierarchy of nested populations)
- ▶ Patient  $\in$  Hospital  $\in$  Region



#### Clustered Data

- Example: (Longitudinal)
- ▶ Repeated measurements on the same unit of observation
  - Patients have multiple temperature measurements over time
  - ► Temperature ∈ Patient ∈ Clinic

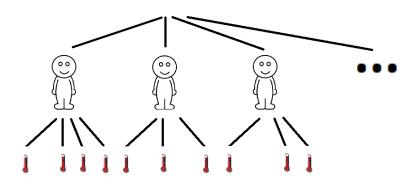
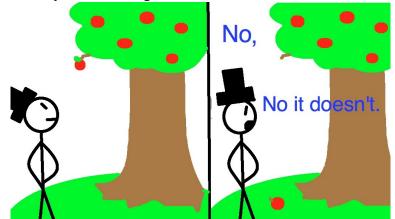


Figure 1: Alkema 2017

## Implications of Clustered Data

- Observations in clusters violates our assumptions of independence
- Clustered data are less informative and less generalizable than independent data

► **Example:** Subjects within a household are more similar and will vary less. Similar genetics and behavior



## **Example Notation**

Mixed models, also known as multilevel or hierarchical models, are used to model cluster data. We will use the following notation to model such data.

 $Y_{ij} =$ Response variable

 $Trt_{ij} = Treatment variable$ 

## Example Notation 2

$$j=1,\ 2,\ ...,\ n$$
  $i=1,\ 2,\ ...,\ n_i$   $N=\sum_i n_i={
m total\ number\ of\ groups}$   $Treatment\ Trt_{ij}=egin{cases} 1,\ if\ Trt_{ij}\ {
m was\ treated} \ 0,\ {
m otherwise} \end{cases}$ 

Treatment jj	Region j	Mortality Count	
1	1	72	<b>)</b> =
0	1	256	<b>&gt;</b> <i>y</i> <sub>1</sub>
1	1	91	<b>)</b>
0	2	11	
1	2	12	
1	2	7	$\bar{y}_2$
1	2	13	32

## Example to begin with

► Let *Y<sub>ij</sub>* denotes the counts of mortality

$$Y_{ij} \sim Poisson(\lambda_{ij}, P_i)$$
  
 $log(\lambda_{ii}) = \beta_0 + \beta_1 Trt_{ii} + \beta_2 X_i$ 

- ▶ We want to make inference about  $\beta_1$ . What is the treatment effect?
- ▶ Observations are not iid, need "adjustment"
- How can we account for variance within groups?
  - 1. Marginal models with generalized estimating equations (GEE) for variance adjustment
  - 2. generalized linear mixed models (GLMM's)

#### GLMM's Models

- Account for modeling for individual and group level variation in estimating group-level coefficients
- ► Allow the proper measure of variation in individual level regression coefficients
- For point estimates, Shrinkage (relative to sample size) of parameters toward group means
- Rule of thumb
  - number of groups greater than 5
  - substainial variation among groups

#### **GLMMS Model Notation**

lacktriangle where eta is a fixed effect and  $\mu_i$  are varying (random) effects

$$g(E[y_{ij}|\mu_i]) = X_{ij}^T \beta + Z_{ij}^T \mu_i$$
$$\mu_i \sim N(0, G_\theta)$$

- Agresti uses i to represent the group,  $Y_{ij}$  is the jth observation in ith group
- lacktriangle So here we have *i* different  $\mu$  which are draws from a normal
- ▶ The  $\mu_i$  have a common distribution
- ▶ Notice  $\beta$  has no subscript,  $\beta$  is fixed
- ightharpoonup eta is a global estimate and  $\mu_i$  is a group specific estimate
- ▶ g is a glm link
- m heta is a parameter that governs the distribution of the random effect

#### Estimation

$$L(\beta, \theta) = f(\vec{y}|\beta, \theta)$$
$$= \int f(\vec{y}|\vec{\mu})f(\vec{\mu})du$$

- where we integrate across our marginal  $f(\hat{\mu})$
- these problems usually do not have closed form solutions
- ▶ How to approximate? Use Bayesian MCMC or HMC algorithmms to sample from posterior distribution of  $\beta$  and  $\theta$
- approximate with Laplace methods (some coefficients are subject to penalty terms)
- Degrees of Freedom: Approximated by estension of the Hat matrix
  - some closed for solutions (ex: beta-binomial conjugate in next lecture)

$$tr(H) = p$$
$$p \le tr(H) \le p + q$$

### Example

▶ Lets say we are looking at treatment(spinal implants) to relieve back pain. Rows represent visit 1 and columns represent visit 2

visit1/visit2	success	failure
success	63	16
failure	12	35

Figure 3:

ightharpoonup pain indicator  $Y_{ij}$  and group indicator  $X_{ij}$ 

$$y_{ij} = \begin{cases} 1 & \text{patient j at visit i has no pain} \\ 0 & \text{patient j at visit i has pain} \end{cases}$$

$$x_{ij} = \begin{cases} 1 & i=2; \text{ patient's second visit} \\ 0 & i=1; \text{ pateint's first visit} \end{cases}$$

# Example (possible models)

logistic-normal model

$$logit(P(Y_{ij} = 1 | \mu_i)) = \alpha + \beta X_{ij} + \mu_i$$
$$\mu_i \sim N(0, \sigma_{\mu}^2)$$

► Similar notation for bayesians

$$Y_i \sim N(\pi, \sigma_y^2)$$
  
 $logit(\pi) = \alpha_i + \beta X_{ij}$   
 $\alpha_i \sim N(\beta_0, \sigma_\alpha)$   
 $\beta_0 \sim N(0, 100)$   
 $\sigma_\alpha \sim U(0, 5)$ 

# Example (interpretation)

$$logit(P(Y_{ij} = 1 | \mu_i)) = \alpha + \beta X_{ij} + \mu_i$$
$$\mu_i \sim N(0, \sigma_{\mu}^2)$$

- lacktriangledown  $lpha\sim\log$  odds of pain free at visit 1
- $\blacktriangleright$   $\beta \sim$  change in log odds of being pain free comparing visit 2 to visit 1

# Example (interpretation)

▶ With mixed effect  $\mu_i$  our intercept can now vary

