Comparing GLMM and GEE

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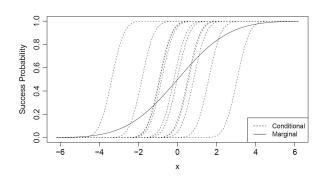
Review

Recall that the two methods covered that can accomodate correlated responses are:

- GLMM is a subject specific model, modeling how the predictors affect an individual subject's response
- GEE is a marginal model, modeling how the predictors affect the average of the subject's response

Comparison

- When $\hat{\sigma}_u = 0$ in GLMM or $\hat{\rho} = 0$ in GEE, estimates and standard errors same as treating repeated observations as independent
- When $\hat{\sigma}_u$ is large, estimated β 's from random effects logit model usually larger than from marginal GEE model. They are estimating different things



Example (Teratology Overdispersions)

Female rats on iron-deficient diets assigned to four groups:

- placebo
- 2 iron injections on days 7 and 10
- iron injections on days 0 and 7
- iron injections weekly

Then they are made pregnant and sacrificed after 3 weeks. The response is whether fetus is dead or alive and the *cluster* is the litter.

Notation:

- GRP = group,
- LS = litter size,
- ND = number dead in litter

 $logit[P(fetus t in litter i dead)] = \alpha + \beta_2 z_{i2} + \beta_3 z_{i3} + \beta_4 z_{i4}$

$$z_{ij} = egin{cases} 1 & ext{if litter } i ext{ in group } j \ 0 & ext{otherwise} \end{cases}$$

We will fit regular binomial GLM, i.e. assuming independence.

> terat.binom=glm(cbind(ND,N-ND)~GRP, family=binomial)

> summary(terat.binom)

Coefficients:

Estimate Std. Error z value Pr(>|z|)

(Intercept) 1.1440 0.1292 8.855 < 2e-16 ***
GRP2 -3.3225 0.3308 -10.043 < 2e-16 ***

GRP3 -4.4762 0.7311 -6.122 9.22e-10 ***
GRP4 -4.1297 0.4762 -8.672 < 2e-16 ***

Null deviance: 509.43 on 57 degrees of freedom

Residual deviance: 173.45 on 54 degrees of freedom

- > X2=sum(resid(terat.binom,type="pearson")^2);X2
- [1] 154.707
- > 1-pchisq(X2,df.residual(terat.binom)) # GoF Pearson
 [1] 1.187217e-11
- [1] 1,10,21,0 11
- > X2/df.residual(terat.binom) # Evidence of overdispersion
 [1] 2.864945
 - Poor fit (GoF)
 - Overdispersion, and standard errors invalid (too small) due to overdispersion
 - Inter-litter variability that cannot be accounted for. Fetuses are more alike within litters than across litters, even within the same treatment group

Possible solutions:

- GEE: models marginal (population averaged) effect of treatment
- GLMM: models litter-specific effect
- At least two other approaches not discussed (thoroughly) in this class:
 - Quasi-binomial: simplified version of GEE
 - Beta-binomial: parametric mixture model, analogous to negative-binomial for count data. Motivation similar to GLMM

- > # Big cor matrix (17 x 17), but all off-diag entries
- > terat.gee\$working.correlation[1,2]
- [1] 0.8051211

```
> terat.glmm <- glmer((Resp == "Dead") ~ GRP + (1|Litter),</pre>
        data = teratbnry, family = binomial)
> summary(terat.glmm)
Random effects:
Groups Name Variance Std.Dev.
Litter (Intercept) 2.284 1.511
Number of obs: 607, groups: Litter, 58
Fixed effects:
         Estimate Std. Error z value Pr(>|z|)
4.5396 0.7345 6.181 6.39e-10 ***
GRP2
    5.8833 1.1754 5.005 5.58e-07 ***
GRP3
           5.6062 0.9076 6.177 6.54e-10 ***
GRP4
```

	Binomial ML	GEE	GLMM
(Intercept)	1.14 (0.13)	1.21 (0.27)	1.81 (0.33)
GRP2	-3.32 (0.33)	-3.37 (0.43)	-4.54 (0.68)
GRP3	-4.48 (0.73)	-4.58 (0.62)	-5.88 (1.18)
GRP4	-4.13 (0.48)	-4.25 (0.6)	-5.61 (0.86)

- SEs for binomial ML fit invalid (because of lack of fit)
- GEE estimates are similar to binomial but with larger SEs. Estimate marginal (population averaged) effects
- GLMM estimates are larger in magnitude. Estimate conditional (within litter) effects

It seems that there are differences between groups 2,3,4 with the base group 1. As an exercise compare groups 2 and 3 for the GLMM model.

Hint: Create pairwise CI for $\beta_i - \beta_j$ for which you will need the estimated covariances $Cov(\hat{\beta}_i, \hat{\beta}_i)$ and remember to use Bonferonni adjustment.

We learned

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