Mixed models in R using the Ime4 package Part 5: Generalized linear mixed models

Douglas Bates

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Generalized Linear Mixed Models

- When using linear mixed models (LMMs) we assume that the response being modeled is on a continuous scale.
- Sometimes we can bend this assumption a bit if the response is an
 ordinal response with a moderate to large number of levels. For
 example, the Scottish secondary school test results in the mlmRev
 package are integer values on the scale of 1 to 10 but we analyze
 them on a continuous scale.
- However, an LMM is not suitable for modeling a binary response, an ordinal response with few levels or a response that represents a count.
 For these we use generalized linear mixed models (GLMMs).
- To describe GLMMs we return to the representation of the response as an n-dimensional, vector-valued, random variable, \mathcal{Y} , and the random effects as a q-dimensional, vector-valued, random variable, \mathcal{B} .

Parts of LMMs carried over to GLMMs

- Random variables
 - ${oldsymbol{\mathcal{Y}}}$ the response variable
 - ${\cal B}$ the (possibly correlated) random effects
 - ${\mathcal U}$ the orthogonal random effects, such that ${\mathcal B}=\Lambda_{ heta}{\mathcal U}$
- Parameters
 - $oldsymbol{eta}$ fixed-effects coefficients
 - σ the common scale parameter (not always used)
 - $m{ heta}$ parameters that determine $\mathrm{Var}(m{\mathcal{B}}) = \sigma^2 m{\Lambda}_{ heta} m{\Lambda}_{ heta}^\mathsf{T}$
- Some matrices
 - ${m X}$ the $n \times p$ model matrix for ${m eta}$
 - $m{Z}$ the $n \times q$ model matrix for $m{b}$
 - P fill-reducing $q \times q$ permutation (from Z)
 - $m{\Lambda}_{ heta}$ relative covariance factor, s.t. $\mathsf{Var}(m{\mathcal{B}}) = \sigma^2 m{\Lambda}_{ heta} m{\Lambda}_{ heta}^\mathsf{T}$

The conditional distribution, ${oldsymbol {\mathcal Y}}|{oldsymbol {\mathcal U}}$

- For GLMMs, the marginal distribution, $\mathcal{B} \sim \mathcal{N}\left(\mathbf{0}, \mathbf{\Sigma}_{\theta}\right)$ is the same as in LMMs except that σ^2 is omitted. We define $\mathcal{U} \sim \mathcal{N}(\mathbf{0}, \mathbf{I}_q)$ such that $\mathcal{B} = \mathbf{\Lambda}_{\theta} \mathcal{U}$.
- For GLMMs we retain some of the properties of the conditional distribution for a LMM

$$(\mathcal{Y}|\mathcal{U}=u)\sim\mathcal{N}\left(\mu_{\mathcal{Y}|\mathcal{U}},\sigma^2I
ight)$$
 where $\mu_{\mathcal{Y}|\mathcal{U}}(u)=Xeta+Zoldsymbol{\Lambda}_{ heta}u$ Specifically

- ▶ The conditional distribution, $\mathcal{Y}|\mathcal{U}=u$, depends on u only through the conditional mean, $\mu_{\mathcal{Y}|\mathcal{U}}(u)$.
- ▶ Elements of \mathcal{Y} are *conditionally independent*. That is, the distribution, $\mathcal{Y}|\mathcal{U}=u$, is completely specified by the univariate, conditional distributions, $\mathcal{Y}_i|\mathcal{U}, i=1,\ldots,n$.
- ► These univariate, conditional distributions all have the same form. They differ only in their means.
- GLMMs differ from LMMs in the form of the univariate, conditional distributions and in how $\mu_{\mathcal{Y}|\mathcal{U}}(u)$ depends on u.

Some choices of univariate conditional distributions

- Typical choices of univariate conditional distributions are:
 - The Bernoulli distribution for binary (0/1) data, which has probability mass function

$$p(y|\mu) = \mu^y (1-\mu)^{1-y}, \quad 0 < \mu < 1, \quad y = 0, 1$$

- Several independent binary responses can be represented as a binomial response, but only if all the Bernoulli distributions have the same mean.
- ▶ The *Poisson* distribution for count (0, 1, ...) data, which has probability mass function

$$p(y|\mu) = e^{-\mu} \frac{\mu^y}{y!}, \quad 0 < \mu, \quad y = 0, 1, 2, \dots$$

• All of these distributions are completely specified by the conditional mean. This is different from the conditional normal (or Gaussian) distribution, which also requires the common scale parameter, σ .

The link function, g

- When the univariate conditional distributions have constraints on μ , such as $0 < \mu < 1$ (Bernoulli) or $0 < \mu$ (Poisson), we cannot define the conditional mean, $\mu_{\boldsymbol{\mathcal{Y}}|\boldsymbol{\mathcal{U}}}$, to be equal to the linear predictor, $\boldsymbol{X}\boldsymbol{\beta} + \boldsymbol{X}\boldsymbol{\Lambda}_{\boldsymbol{\theta}}\boldsymbol{u}$, which is unbounded.
- We choose an invertible, univariate *link function*, g, such that $\eta=g(\mu)$ is unconstrained. The vector-valued link function, g, is defined by applying g component-wise.

$$\boldsymbol{\eta} = \boldsymbol{g}(\boldsymbol{\mu})$$
 where $\eta_i = g(\mu_i), \quad i = 1, \dots, n$

• We require that g be invertible so that $\mu=g^{-1}(\eta)$ is defined for $-\infty<\eta<\infty$ and is in the appropriate range $(0<\mu<1$ for the Bernoulli or $0<\mu$ for the Poisson). The vector-valued inverse link, g^{-1} , is defined component-wise.

"Canonical" link functions

- There are many choices of invertible scalar link functions, g, that we could use for a given set of constraints.
- For the Bernoulli and Poisson distributions, however, one link function arises naturally from the definition of the probability mass function. (The same is true for a few other, related but less frequently used, distributions, such as the gamma distribution.)
- To derive the canonical link, we consider the logarithm of the probability mass function (or, for continuous distributions, the probability density function).
- For distributions in this "exponential" family, the logarithm of the probability mass or density can be written as a sum of terms, some of which depend on the response, y, only and some of which depend on the mean, μ , only. However, only one term depends on **both** y and μ , and this term has the form $y \cdot g(\mu)$, where g is the canonical link.

The canonical link for the Bernoulli distribution

The logarithm of the probability mass function is

$$\log(p(y|\mu)) = \log(1-\mu) + y\log\left(\frac{\mu}{1-\mu}\right), \ 0 < \mu < 1, \ y = 0, 1.$$

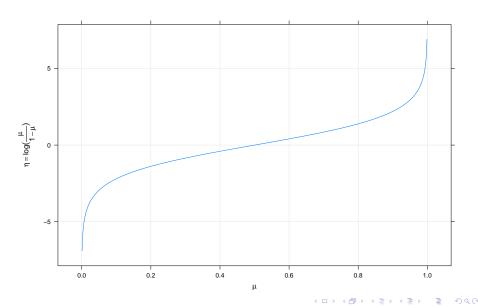
Thus, the canonical link function is the logit link

$$\eta = g(\mu) = \log\left(\frac{\mu}{1-\mu}\right).$$

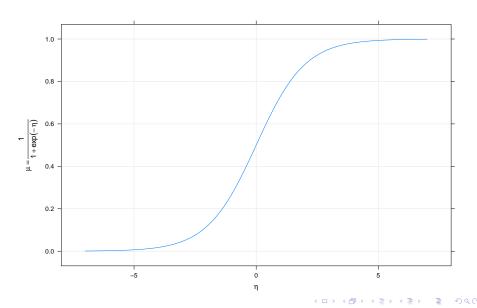
- Because $\mu=P[\mathcal{Y}=1]$, the quantity $\mu/(1-\mu)$ is the odds ratio (in the range $(0,\infty)$) and g is the logarithm of the odds ratio, sometimes called "log odds".
- The inverse link is

$$\mu = g^{-1}(\eta) = \frac{e^{\eta}}{1 + e^{\eta}} = \frac{1}{1 + e^{-\eta}}$$

Plot of canonical link for the Bernoulli distribution



Plot of inverse canonical link for the Bernoulli distribution



The canonical link for the Poisson distribution

The logarithm of the probability mass is

$$\log(p(y|\mu)) = \log(y!) - \mu + y\log(\mu)$$

• Thus, the canonical link function for the Poisson is the log link

$$\eta = g(\mu) = \log(\mu)$$

The inverse link is

$$\mu = g^{-1}(\eta) = e^{\eta}$$

The canonical link related to the variance

- For the canonical link function, the derivative of its inverse is the variance of the response.
- For the Bernoulli, the canonical link is the logit and the inverse link is $\mu=g^{-1}(\eta)=1/(1+e^{-\eta}).$ Then

$$\frac{d\mu}{d\eta} = \frac{e^{-\eta}}{(1+e^{-\eta})^2} = \frac{1}{1+e^{-\eta}} \frac{e^{-\eta}}{1+e^{-\eta}} = \mu(1-\mu) = \text{Var}(\mathcal{Y})$$

• For the Poisson, the canonical link is the log and the inverse link is $\mu=g^{-1}(\eta)=e^{\eta}.$ Then

$$\frac{d\mu}{d\eta} = e^{\eta} = \mu = \text{Var}(\mathcal{Y})$$



The unscaled conditional density of $\mathcal{U}|\mathcal{Y}=y$

 As in LMMs we evaluate the likelihood of the parameters, given the data, as

$$L(\boldsymbol{ heta}, oldsymbol{eta} | oldsymbol{y}) = \int_{\mathbb{R}^q} [oldsymbol{\mathcal{Y}} | oldsymbol{\mathcal{U}}](oldsymbol{y} | oldsymbol{u}) [oldsymbol{\mathcal{U}}](oldsymbol{u}) \ d oldsymbol{u},$$

- The product $[\mathcal{Y}|\mathcal{U}](y|u)[\mathcal{U}](u)$ is the unscaled (or *unnormalized*) density of the conditional distribution $\mathcal{U}|\mathcal{Y}$.
- The density $[\mathcal{U}](u)$ is a spherical Gaussian density $\frac{1}{(2\pi)^{q/2}}e^{-\|u\|^2/2}$.
- The expression $[\mathcal{Y}|\mathcal{U}](y|u)$ is the value of a probability mass function or a probability density function, depending on whether $\mathcal{Y}_i|\mathcal{U}$ is discrete or continuous.
- The linear predictor is $g(\mu_{\mathcal{Y}|\mathcal{U}}) = \eta = X\beta + Z\Lambda_{\theta}u$. Alternatively, we can write the conditional mean of \mathcal{Y} , given \mathcal{U} , as

$$oldsymbol{\mu_{\mathcal{Y}|\mathcal{U}}(u) = g^{-1} \left(oldsymbol{X}oldsymbol{eta} + oldsymbol{Z}oldsymbol{\Lambda}_{ heta} oldsymbol{u}
ight)}$$

The conditional mode of $\mathcal{U}|\mathcal{Y}=y$

• In general the likelihood, $L(\theta, \beta|y)$ does not have a closed form. To approximate this value, we first determine the *conditional mode*

$$ilde{u}(oldsymbol{y}|oldsymbol{ heta},oldsymbol{eta}) = rg\max_{oldsymbol{u}}[oldsymbol{\mathcal{Y}}|oldsymbol{\mathcal{U}}](oldsymbol{y}|oldsymbol{u})[oldsymbol{\mathcal{U}}](oldsymbol{u})$$

using a quadratic approximation to the logarithm of the unscaled conditional density.

 This optimization problem is (relatively) easy because the quadratic approximation to the logarithm of the unscaled conditional density can be written as a penalized, weighted residual sum of squares,

$$ilde{m{u}}(m{y}|m{ heta},m{eta}) = rg \min_{m{u}} \left\| \left[m{W}^{1/2}(m{\mu}) \left(m{y} - m{\mu}_{m{\mathcal{Y}}|m{\mathcal{U}}}(m{u})
ight)
ight]
ight\|^2$$

where $W(\mu)$ is the diagonal weights matrix. The weights are the inverses of the variances of the \mathcal{Y}_i .

The PIRLS algorithm

- ullet Parameter estimates for generalized linear models (without random effects) are usually determined by iteratively reweighted least squares (IRLS), an incredibly efficient algorithm. PIRLS is the penalized version. It is iteratively reweighted in the sense that parameter estimates are determined for a fixed weights matrix $oldsymbol{W}$ then the weights are updated to the current estimates and the process repeated.
- For fixed weights we solve

$$\min_{oldsymbol{u}} \left\| \left[egin{aligned} oldsymbol{W}^{1/2} \left(oldsymbol{y} - oldsymbol{\mu}_{oldsymbol{\mathcal{Y}} | oldsymbol{\mathcal{U}}}(oldsymbol{u})
ight) \\ -oldsymbol{u} \end{aligned}
ight]^2$$

as a nonlinear least squares problem with update, δ_u , given by

$$\boldsymbol{P}\left(\boldsymbol{\Lambda}_{\theta}^{\mathsf{T}}\boldsymbol{Z}^{\mathsf{T}}\boldsymbol{M}\,\boldsymbol{W}\boldsymbol{M}\,\boldsymbol{Z}\boldsymbol{\Lambda}_{\theta}+\boldsymbol{I}_{q}\right)\boldsymbol{P}^{\mathsf{T}}\boldsymbol{\delta}_{\boldsymbol{u}}=\boldsymbol{\Lambda}_{\theta}^{\mathsf{T}}\boldsymbol{Z}^{\mathsf{T}}\boldsymbol{M}\,\boldsymbol{W}(\boldsymbol{y}-\boldsymbol{\mu})-\boldsymbol{u}$$

where $\pmb{M}=d\pmb{\mu}/d\pmb{\eta}$ is the (diagonal) Jacobian matrix. Recall that for the canonical link, $\pmb{M}=\mathsf{Var}(\pmb{\mathcal{Y}}|\pmb{\mathcal{U}})=\pmb{W}^{-1}$.

The Laplace approximation to the deviance

ullet At convergence, the sparse Cholesky factor, $oldsymbol{L}$, used to evaluate the update is

$$oldsymbol{L} oldsymbol{L}^\mathsf{T} = oldsymbol{P} \left(oldsymbol{\Lambda}_{ heta}^\mathsf{T} oldsymbol{Z}^\mathsf{T} oldsymbol{M} oldsymbol{W} oldsymbol{M} oldsymbol{Z} oldsymbol{\Lambda}_{ heta} + oldsymbol{I}_q
ight) oldsymbol{P}^\mathsf{T}$$

or

$$oldsymbol{L} oldsymbol{L}^\mathsf{T} = oldsymbol{P} \left(oldsymbol{\Lambda}_{ heta}^\mathsf{T} oldsymbol{Z}^\mathsf{T} oldsymbol{M} oldsymbol{Z} oldsymbol{\Lambda}_{ heta} + oldsymbol{I}_q
ight) oldsymbol{P}^\mathsf{T}$$

if we are using the canonical link.

- The integrand of the likelihood is approximately a constant times the density of the $\mathcal{N}(\tilde{\pmb{u}}, \pmb{L} \pmb{L}^\mathsf{T})$ distribution.
- On the deviance scale (negative twice the log-likelihood) this corresponds to

$$d(\boldsymbol{\beta}, \boldsymbol{\theta}|\boldsymbol{y}) = d_g(\boldsymbol{y}, \boldsymbol{\mu}(\tilde{\boldsymbol{u}})) + ||\tilde{\boldsymbol{u}}||^2 + \log(|\boldsymbol{L}|^2)$$

where $d_q(\boldsymbol{y}, \boldsymbol{\mu}(\tilde{\boldsymbol{u}}))$ is the GLM deviance for \boldsymbol{y} and $\boldsymbol{\mu}$.

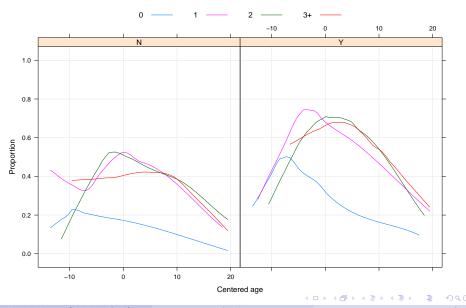
Modifications to the algorithm

- Notice that this deviance depends on the fixed-effects parameters, $\boldsymbol{\beta}$, as well as the variance-component parameters, $\boldsymbol{\theta}$. This is because $\log(|\boldsymbol{L}|^2)$ depends on $\boldsymbol{\mu}_{\boldsymbol{\mathcal{Y}}|\boldsymbol{\mathcal{U}}}$ and, hence, on $\boldsymbol{\beta}$. For LMMs $\log(|\boldsymbol{L}|^2)$ depends only on $\boldsymbol{\theta}$.
- In practice we begin by optimizing w.r.t. u and β simultaneously, evaluating the Laplace approximation and optimizing this w.r.t. θ . Then we use a "pure" Laplace approximation which is optimized w.r.t. both β and θ .
- The second stage can be suppressed with the optional argument nAGQ
 another argument verbose = 2 shows the two stages explicitly
- Another approach is adaptive Gauss-Hermite quadrature (AGQ). This
 has a similar structure to the Laplace approximation but is based on
 more evaluations of the unscaled conditional density near the
 conditional modes. It is only appropriate for models in which the
 random effects are associated with only one grouping factor

Contraception data

- One of the data sets in the "mlmRev" package, derived from data files available on the multilevel modelling web site, is from a fertility survey of women in Bangladesh.
- One of the (binary) responses recorded is whether or not the woman currently uses artificial contraception.
- Covariates included the woman's age (on a centered scale), the number of live children she had, whether she lived in an urban or rural setting, and the district in which she lived.
- ullet Instead of plotting such data as points, we use the 0/1 response to generate scatterplot smoother curves versus age for the different groups.

Contraception use versus age by urban and livch



Comments on the data plot

- These observational data are unbalanced (some districts have only 2 observations, some have nearly 120). They are not longitudinal (no "time" variable).
- Binary responses have low per-observation information content (exactly one bit per observation). Districts with few observations will not contribute strongly to estimates of random effects.
- Within-district plots will be too imprecise so we only examine the global effects in plots.
- The comparisons on the multilevel modelling site are for fits of a model that is linear in age, which is clearly inappropriate.
- The form of the curves suggests at least a quadratic in age.
- The urban versus rural differences may be additive.
- It appears that the livch factor could be dichotomized into "0" versus "1 or more".

Preliminary model using Laplacian approximation

```
Generalized linear mixed model fit by maximum likelihood ['merMod']
Family: binomial
Formula: use ~ age + I(age^2) + urban + livch + (1 | district)
  Data: Contraception
     AIC BIC logLik deviance
2388.785 2433.323 -1186.392 2372.785
Random effects:
Groups Name Variance Std.Dev.
district (Intercept) 0.2253 0.4747
Number of obs: 1934, groups: district, 60
Fixed effects:
            Estimate Std. Error z value
(Intercept) -1.0152756 0.1739721 -5.836
age 0.0035135 0.0092106 0.381
I(age^2) -0.0044867 0.0007228 -6.207
urbanY 0.6844018 0.1196840 5.718
livch1 0.8018807 0.1618666 4.954
livch2 0.9010171 0.1847709 4.876
livch3+ 0.8994111 0.1854007 4.851
```

Comments on the model fit

- This model was fit using the Laplacian approximation to the deviance.
- There is a highly significant quadratic term in age.
- The linear term in age is not significant but we retain it because the age scale has been centered at an arbitrary value (which, unfortunately, is not provided with the data).
- The urban factor is highly significant (as indicated by the plot).
- Levels of livch greater than 0 are significantly different from 0 but may not be different from each other.

Reduced model with dichotomized livch

```
Generalized linear mixed model fit by maximum likelihood ['merMod']
Family: binomial
Formula: use ~ age + I(age^2) + urban + ch + (1 | district)
  Data: Contraception
     AIC
              BIC logLik deviance
2385.241 2418.645 -1186.621 2373.241
Random effects:
Groups Name Variance Std.Dev.
district (Intercept) 0.2242 0.4735
Number of obs: 1934, groups: district, 60
Fixed effects:
           Estimate Std. Error z value
(Intercept) -0.987377 0.167529 -5.894
   0.006170 0.007826 0.788
age
I(age^2) -0.004558 0.000714 -6.384
urbanY 0.680226 0.119476 5.693
```

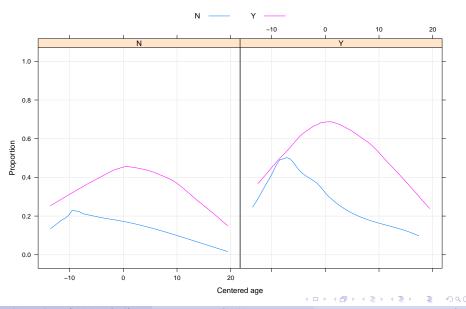
chY 0.846208 0.147027 5.755

Comparing the model fits

A likelihood ratio test can be used to compare these nested models.

- The large p-value indicates that we would not reject cm2 in favor of cm1 hence we prefer the more parsimonious cm2.
- The plot of the scatterplot smoothers according to live children or none indicates that there may be a difference in the age pattern between these two groups.

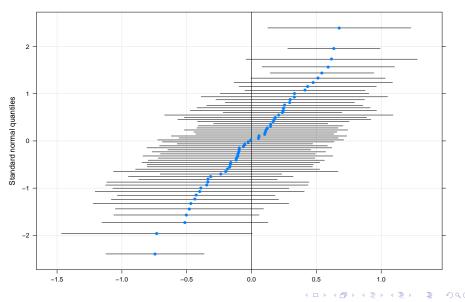
Contraception use versus age by urban and ch



Allowing age pattern to vary with ch

```
Generalized linear mixed model fit by maximum likelihood ['merMod']
Family: binomial
Formula: use ~ age * ch + I(age^2) + urban + (1 | district)
  Data: Contraception
     AIC BIC logLik deviance
2379.181 2418.153 -1182.591 2365.181
Random effects:
Groups Name Variance Std.Dev.
district (Intercept) 0.2231 0.4723
Number of obs: 1934, groups: district, 60
Fixed effects:
            Estimate Std. Error z value
(Intercept) -1.3230260 0.2144340 -6.170
age -0.0472742 0.0218378 -2.165
chY 1.2105520 0.2069791 5.849
I(age<sup>2</sup>) -0.0057574 0.0008358 -6.889
urbanY 0.7139915 0.1202574 5.937
age:chY 0.0683363 0.0254331 2.687
```

Prediction intervals on the random effects



Extending the random effects

 We may want to consider allowing a random effect for urban/rural by district. This is complicated by the fact the many districts only have rural women in the study

```
district
urban
                                           8
                                                    10
                                                         11
                                                              12
                                                                  13
                                                                                 16
        54
             20
                                     18
                                          35
                                               20
                                                    13
                                                         21
                                                              23
                                                                  16
                                                                       17
                                                                                 18
        63
                       11
                                 7
                                           2
                                                3
                                      0
                                                     0
                                                          0
                                                               6
                                                                      101
                                                                                  2
      district
                                               25
urban
        17
             18
                  19
                       20
                           21
                                22
                                     23
                                          24
                                                    26
                                                         27
                                                              28
                                                                  29
                                                                       30
                                                                            31
                                                                                 32
                                     15
                                                    13
        24
             33
                  22
                       15
                            10
                                20
                                          14
                                               49
                                                         39
                                                             45
                                                                  25
                                                                       45
                                                                            27
                                                                                 24
```

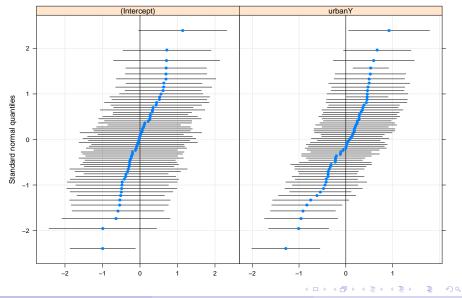
Including a random effect for urban by district

```
Generalized linear mixed model fit by maximum likelihood ['merMod']
Family: binomial
Formula: use ~ age * ch + I(age^2) + urban + (urban | district)
  Data: Contraception
     AIC
              BIC logLik deviance
2371.530 2421.636 -1176.765 2353.530
Random effects:
Groups Name Variance Std.Dev. Corr
district (Intercept) 0.3783 0.6150
        urbanY 0.5261 0.7253 -0.793
Number of obs: 1934, groups: district, 60
Fixed effects:
            Estimate Std. Error z value
(Intercept) -1.3441450 0.2227632 -6.034
age -0.0461824 0.0219444 -2.105
chY 1.2116385 0.2082354 5.819
I(age^2) -0.0056514 0.0008431 -6.703
urbanY 0.7901052 0.1600452 4.937
age:chY 0.0664711 0.0255671 2.600
```

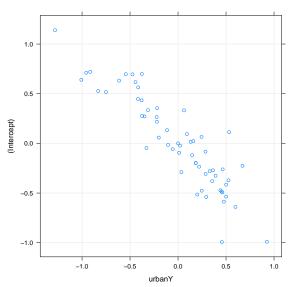
Significance of the additional random effect

- The additional random effect is highly significant in this test.
- Most of the prediction intervals still overlap zero.
- A scatterplot of the random effects shows several random effects vectors falling along a straight line. These are the districts with all rural women or all urban women.

Prediction intervals for the bivariate random effects



Scatter plot of the BLUPs



Nested simple, scalar random effects versus vector-valued

```
Generalized linear mixed model fit by maximum likelihood ['merMod']
 Family: binomial
Formula: use ~ age * ch + I(age^2) + urban + (1 | urban:district) + (1 |
  Data: Contraception
              BIC logLik deviance
2370.464 2415.003 -1177.232 2354.464
Random effects:
Groups Name Variance Std.Dev.
urban:district (Intercept) 0.30987 0.5567
district (Intercept) 0.01155 0.1075
Number of obs: 1934, groups: urban:district, 102; district, 60
Fixed effects:
            Estimate Std. Error z value
(Intercept) -1.3406181 0.2210382 -6.065
age -0.0461495 0.0220190 -2.096
chY 1.2129371 0.2089635 5.805
I(age^2) -0.0056309 0.0008437 -6.674
urbanY 0.7833789 0.1691559 4.631
age:chY 0.0664924 0.0256365 2.594
```

Using the interaction term only

```
Generalized linear mixed model fit by maximum likelihood ['merMod']
Family: binomial
Formula: use ~ age * ch + I(age^2) + urban + (1 | urban:district)
  Data: Contraception
     AIC BIC logLik deviance
2368.475 2407.446 -1177.237 2354.475
Random effects:
Groups Name Variance Std.Dev.
urban:district (Intercept) 0.3229 0.5683
Number of obs: 1934, groups: urban:district, 102
Fixed effects:
            Estimate Std. Error z value
(Intercept) -1.3406928 0.2211336 -6.063
age -0.0461540 0.0220213 -2.096
chY 1.2128990 0.2089889 5.804
I(age<sup>2</sup>) -0.0056264 0.0008437 -6.669
urbanY 0.7866579 0.1705857 4.612
age:chY 0.0664658 0.0256387 2.592
```

Comparing models with random effects for interactions

```
> anova(cm6,cm5,cm4)

Data: Contraception
Models:
cm6: use ~ age * ch + I(age^2) + urban + (1 | urban:district)
cm5: use ~ age * ch + I(age^2) + urban + (1 | urban:district) + (1 |
cm5: district)
cm4: use ~ age * ch + I(age^2) + urban + (urban | district)
    Df AIC BIC logLik deviance Chisq Chi Df Pr(>Chisq)
cm6  7  2368.5  2407.4 -1177.2  2354.5
cm5  8  2370.5  2415.0 -1177.2  2354.5  0.0107  1  0.9177
cm4  9  2371.5  2421.6 -1176.8  2353.5  0.9338  1  0.3339
```

- The random effects seem to best be represented by a separate random effect for urban and for rural women in each district.
- The districts with only urban women in the survey or with only rural women in the survey are naturally represented in this model.

Showing the optimization stages

```
> cm6 <- glmer(use ~ age*ch + I(age^2) + urban + (1|urban:dist;
+
                Contraception, binomial, verbose=2L)
npt = 3, n = 1
rhobeg = 0.2, rhoend = 2e-07
  0.020: 5: 2354.74;0.600000
 0.0020: 8: 2354.59; 0.572147
0.00020: 11: 2354.59;0.567988
2.0e-05: 13: 2354.59;0.567275
2.0e-06: 14: 2354.59;0.567275
2.0e-07: 15: 2354.59;0.567273
At return
18: 2354.5894: 0.567273
npt = 9, n = 7
rhobeg = 2e-04, rhoend = 2e-07
2.0e-05: 11: 2354.54;0.567273 -1.30421 -0.0446451 1.18092 -0.005675
2.0e-06: 24: 2354.52;0.567257 -1.30425 -0.0444421 1.18090 -0.005619
2.0e-07: 300: 2354.51:0.567979 -1.30454 -0.0436418 1.18099 -0.005617
At return
```

Conclusions from the example

- Again, carefully plotting the data is enormously helpful in formulating the model.
- Observational data tend to be unbalanced and have many more covariates than data from a designed experiment. Formulating a model is typically more difficult than in a designed experiment.
- A generalized linear model is fit with the function glmer() which requires a family argument. Typical values are binomial or poisson
- Profiling is not provided for GLMMs at present but will be added.
- We use likelihood-ratio tests and z-tests in the model building.

A word about overdispersion

- In many application areas using "pseudo" distribution families, such as quasibinomial and quasipoisson, is a popular and well-accepted technique for accommodating variability that is apparently larger than would be expected from a binomial or a Poisson distribution.
- This amounts to adding an extra parameter, like σ , the common scale parameter in a LMM, to the distribution of the response.
- It is possible to form an estimate of such a quantity during the IRLS algorithm but it is an artificial construct. There is no probability distribution with such a parameter.
- I find it difficult to define maximum likelihood estimates without a probability model. It is not clear how this "distribution which is not a distribution" could be incorporated into a GLMM. This, of course, does not stop people from doing it but I don't know what the estimates from such a model would mean.

Summary

- GLMMs allow for the conditional distribution, $\mathcal{Y}|B=b$, to be other than a Gaussian. A Bernoulli (or, more generally, a binomial) distribution is used to model binary or binomial responses. A Poisson distribution is used to model responses that are counts.
- The conditional mean depends upon the linear predictor, $X\beta + Zb$, through the inverse link function, g^{-1} .
- The conditional mode of the random effects, given the observed data,
 y, is determined through penalized iteratively reweighted least squares (PIRLS).
- We optimize the Laplace approximation at the conditional mode to determine the mle's of the parameters. In some simple cases, a more accurate approximation, adaptive Gauss-Hermite quadrature (AGQ), can be used instead, at the expense of greater computational complexity.