

Teratology - Alternative Binary Models

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First the teratology data are loaded. The data set exists in two different versions, "teratology" shows the aggregated count data whereas "teratology2" includes the original data.

```
> library(catdata)
> data(teratology)
> data(teratology2)
```

For the first two models, the simple independence model and the quasi-likelihood model, the aggregated data are needed.

```
> attach(teratology)
```

The simple and naive independence model is fitted by the following command.

```
> mLogit <- glm(cbind(D,L) ~ as.factor(Grp), family=binomial())
> summary(mLogit)
```

Call:

```
glm(formula = cbind(D, L) ~ as.factor(Grp), family = binomial())
```

Deviance Residuals:

Min	1Q	Median	3Q	Max
-4.4295	-0.9750	-0.0285	1.4024	2.7826

Coefficients:

	Estimate	Std. Error	z value	Pr(> z)
(Intercept)	1.1440	0.1292	8.855	< 2e-16 ***
as.factor(Grp)2	-3.3225	0.3308	-10.043	< 2e-16 ***
as.factor(Grp)3	-4.4762	0.7311	-6.122	9.22e-10 ***
as.factor(Grp)4	-4.1297	0.4762	-8.672	< 2e-16 ***

Signif. codes:

0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1

(Dispersion parameter for binomial family taken to be 1)

```
Null deviance: 509.43 on 57 degrees of freedom
Residual deviance: 173.45 on 54 degrees of freedom
AIC: 252.92
```

Number of Fisher Scoring iterations: 5

Now the quasi-likelihood model is fitted. The coefficients are the same as in the independence model before, only the standard errors have to be multiplied by $\sqrt{\hat{\phi}}$.

```
> mQuasi <- glm(cbind(D,L) ~ as.factor(Grp), family=quasibinomial(link="logit"))
> summary(mQuasi)

Call:
glm(formula = cbind(D, L) ~ as.factor(Grp), family = quasibinomial(link = "logit"))

Deviance Residuals:
    Min      1Q  Median      3Q      Max 
-4.4295 -0.9750 -0.0285  1.4024  2.7826 

Coefficients:
            Estimate Std. Error t value Pr(>|t|)    
(Intercept)  1.1440    0.2187   5.231 2.81e-06 ***
as.factor(Grp)2 -3.3225    0.5600  -5.933 2.18e-07 ***
as.factor(Grp)3 -4.4762    1.2375  -3.617 0.000656 ***
as.factor(Grp)4 -4.1297    0.8061  -5.123 4.14e-06 ***
---
Signif. codes:  0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1

(Dispersion parameter for quasibinomial family taken to be 2.864945)

Null deviance: 509.43 on 57 degrees of freedom
Residual deviance: 173.45 on 54 degrees of freedom
AIC: NA
```

Number of Fisher Scoring iterations: 5

The next model to be fitted is a GEE with independence correlation structure. For that purpose the library "gee" is loaded.

```
> library(gee)
```

Now we use the original data set "teratology2".

```
> detach(teratology)
> attach(teratology2)
```

The GEE is fitted by the following command. The coefficients are again equal to those from the independence model, the standard errors for the independence models can be found in the column "Naive S.E.". The new standard errors from the GEE are those in the column "Robust S.E.".

```
> mGee <- gee(y ~ as.factor(Grp), id=Rat, family=binomial)
```

```

(Intercept) as.factor(Grp)G2 as.factor(Grp)G3
      1.143981       -3.322513      -4.476184
as.factor(Grp)G4
      -4.129663

> summary(mGee)

GEE: GENERALIZED LINEAR MODELS FOR DEPENDENT DATA
gee S-function, version 4.13 modified 98/01/27 (1998)

Model:
Link: Logit
Variance to Mean Relation: Binomial
Correlation Structure: Independent

Call:
gee(formula = y ~ as.factor(Grp), id = Rat, family = binomial)

Summary of Residuals:
      Min        1Q     Median        3Q        Max
-0.75840979 -0.10169492 -0.03448276  0.24159021  0.96551724

Coefficients:
            Estimate Naive S.E.    Naive z
(Intercept)   1.143981  0.1296195  8.825688
as.factor(Grp)G2 -3.322513  0.3319396 -10.009393
as.factor(Grp)G3 -4.476185  0.7335486 -6.102098
as.factor(Grp)G4 -4.129663  0.4778032 -8.643020
            Robust S.E.  Robust z
(Intercept)    0.2758667  4.146861
as.factor(Grp)G2  0.4400582 -7.550168
as.factor(Grp)G3  0.6104577 -7.332507
as.factor(Grp)G4  0.5763810 -7.164814

Estimated Scale Parameter: 1.006633
Number of Iterations: 1

Working Correlation
 [,1] [,2] [,3] [,4] [,5] [,6] [,7] [,8] [,9] [,10]
[1,]    1    0    0    0    0    0    0    0    0    0
[2,]    0    1    0    0    0    0    0    0    0    0
[3,]    0    0    1    0    0    0    0    0    0    0
[4,]    0    0    0    1    0    0    0    0    0    0
[5,]    0    0    0    0    1    0    0    0    0    0
[6,]    0    0    0    0    0    1    0    0    0    0
[7,]    0    0    0    0    0    0    1    0    0    0
[8,]    0    0    0    0    0    0    0    1    0    0
[9,]    0    0    0    0    0    0    0    0    1    0
[10,]   0    0    0    0    0    0    0    0    0    1

```

```

[11,] 0 0 0 0 0 0 0 0 0 0
[12,] 0 0 0 0 0 0 0 0 0 0
[13,] 0 0 0 0 0 0 0 0 0 0
[14,] 0 0 0 0 0 0 0 0 0 0
[15,] 0 0 0 0 0 0 0 0 0 0
[16,] 0 0 0 0 0 0 0 0 0 0
[17,] 0 0 0 0 0 0 0 0 0 0
[,11] [,12] [,13] [,14] [,15] [,16] [,17]
[1,] 0 0 0 0 0 0 0
[2,] 0 0 0 0 0 0 0
[3,] 0 0 0 0 0 0 0
[4,] 0 0 0 0 0 0 0
[5,] 0 0 0 0 0 0 0
[6,] 0 0 0 0 0 0 0
[7,] 0 0 0 0 0 0 0
[8,] 0 0 0 0 0 0 0
[9,] 0 0 0 0 0 0 0
[10,] 0 0 0 0 0 0 0
[11,] 1 0 0 0 0 0 0
[12,] 0 1 0 0 0 0 0
[13,] 0 0 1 0 0 0 0
[14,] 0 0 0 1 0 0 0
[15,] 0 0 0 0 1 0 0
[16,] 0 0 0 0 0 1 0
[17,] 0 0 0 0 0 0 1

```

For the following beta-binomial model the library "VGAM" with its function "vglm" and the data set "teratology" is needed.

```

> library(VGAM)
> detach(teratology2)
> attach(teratology)

```

Furthermore we construct the Variable N as sum of all fetuses in one litter. We will use N to make a subset with $N > 1$ for the beta-binomial model.

```
> N <- D + L
```

Now the beta-binomial model is fitted.

```

> mBetaBin <- vglm(cbind(D,L) ~ as.factor(Grp), family=betabinomial, subset=N>1)
> summary(mBetaBin)

Call:
vglm(formula = cbind(D, L) ~ as.factor(Grp), family = betabinomial,
      subset = N > 1)

Pearson residuals:
          Min        1Q    Median        3Q       Max
logitlink(mu) -2.659 -0.6689 -0.1732  0.9008  1.403

```

```

logitlink(rho) -1.495 -0.9686  0.2121  0.6938  3.907

Coefficients:
            Estimate Std. Error z value Pr(>|z|)
(Intercept):1     1.3451    0.2444   5.505 3.70e-08 ***
(Intercept):2    -1.1402    0.3234  -3.525 0.000423 ***
as.factor(Grp)2  -3.0868    0.5209  -5.926 3.10e-09 ***
as.factor(Grp)3  -3.8647    0.8634  -4.476 7.61e-06 ***
as.factor(Grp)4  -3.9197    0.6838  -5.732 9.92e-09 ***
---
Signif. codes:
0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1

Names of linear predictors: logitlink(mu), logitlink(rho)

Log-likelihood: -93.2973 on 109 degrees of freedom

Number of Fisher scoring iterations: 8

Warning: Hauck-Donner effect detected in the following estimate(s):
'as.factor(Grp)3', 'as.factor(Grp)4'

```

For the following two mixed models again the original data are required.

```

> detach(teratology)
> attach(teratology2)

```

With the function "glmmPQL" from the "MASS"-library a mixed model is fitted by penalized quasi-likelihood, the mixed model contains random intercepts but no random slopes.

```

> mMixPql<- glmmPQL(y ~ as.factor(Grp), random=~1 | Rat, family=binomial)
> summary(mMixPql)

Linear mixed-effects model fit by maximum likelihood
Data: NULL
AIC BIC logLik
NA NA     NA

Random effects:
Formula: ~1 | Rat
          (Intercept) Residual
StdDev:    1.455986 0.8006729

Variance function:
Structure: fixed weights
Formula: ~invwt

Fixed effects: y ~ as.factor(Grp)
                Value Std.Error DF t-value p-value
(Intercept)      1.687058 0.3057620 549  5.517553      0
as.factor(Grp)G2 -4.130280 0.6141687  54 -6.724993      0

```

```

as.factor(Grp)G3 -5.274063 0.9808235 54 -5.377178      0
as.factor(Grp)G4 -5.109542 0.7474739 54 -6.835747      0
Correlation:
          (Intr) a.(G)G2 a.(G)G3
as.factor(Grp)G2 -0.498
as.factor(Grp)G3 -0.312  0.155
as.factor(Grp)G4 -0.409  0.204   0.128

Standardized Within-Group Residuals:
      Min       Q1       Med       Q3       Max
-3.5217300 -0.3475892 -0.1548062  0.2821502  5.1647080

Number of Observations: 607
Number of Groups: 58

```

In order to fit a mixed model by maximum likelihood we load the library "glmmML".

```
> library(glmmML)
```

For a mixed model to be fitted by Gauss–Hermite quadrature we need the function "glmmML" with the option "method='ghq'", the favoured number of quadrature points is determined by the option "n.points".

```

> mGaussH <- glmmML(y ~ as.factor(Grp), cluster=Rat, method = "ghq", n.points = 14,
+                      boot = 0)
> summary(mGaussH)

Call: glmmML(formula = y ~ as.factor(Grp), cluster = Rat, method = "ghq",      n.points = 14)

              coef se(coef)     z Pr(>|z|)
(Intercept) 1.802  0.3623  4.976 6.50e-07
as.factor(Grp)G2 -4.515  0.7360 -6.134 8.58e-10
as.factor(Grp)G3 -5.855  1.1899 -4.921 8.63e-07
as.factor(Grp)G4 -5.594  0.9191 -6.087 1.15e-09

Scale parameter in mixing distribution: 1.533 gaussian
Std. Error: 0.2878

LR p-value for H_0: sigma = 0: 2.7e-15

Residual deviance: 435.6 on 602 degrees of freedom           AIC: 445.6

```

Again we change the data set, for the discrete mixture model "teratology" is required.

```
> detach(teratology2)
> attach(teratology)
```

For discrete mixture models the library "flexmix" with its functions "flexmix" and "stepFlexmix" can be used.

```
> library(flexmix)
```

In "stepFlexmix" the procedure is run several times, the maximum likelihood solution is returned. The favoured number of iterations can be specified by the option "nrep". The number of components is determined by the option "k".

Due to random processes the results of different runs of "stepFlexmix" will differ slightly.

```
> library(stats4)

> mDiscmix <- stepFlexmix(cbind(D,L) ~ 1, k = 2, nrep=5,
+                           model = FLXMRglmfix(family = "binomial", fixed = ~as.factor(Grp)))

2 : * * * * *

> summary(mDiscmix)

Call:
stepFlexmix(cbind(D, L) ~ 1, model = FLXMRglmfix(family = "binomial",
fixed = ~as.factor(Grp)), k = 2, nrep = 5)

      prior size post>0 ratio
Comp.1   0.3    15     53 0.283
Comp.2   0.7    43     53 0.811

'log Lik.' -92.89085 (df=6)
AIC: 197.7817   BIC: 210.1444

> parameters(mDiscmix)

          Comp.1    Comp.2
coef.as.factor(Grp)2 -4.3109413 -4.310941
coef.as.factor(Grp)3 -5.5103569 -5.510357
coef.as.factor(Grp)4 -5.0833859 -5.083386
coef.(Intercept)     -0.2096181  2.461400
```