



JOHNS HOPKINS
BLOOMBERG SCHOOL
of PUBLIC HEALTH

Lecture 10

Models for longitudinal / clustered binary responses Introduction to count outcomes and log-linear regression

Lecture 9 Review:

- ▶ Marginal Models for clustered binary outcomes

- ▶ Define the distribution of Y

$$\begin{aligned}Y_{ij} &\sim \text{Bernoulli}(\mu_{ij}) \\E(Y_{ij}) &= \Pr(Y_{ij} = 1) = \mu_{ij} \\Var(Y_{ij}) &= \mu_{ij}(1 - \mu_{ij}) \\Corr(Y_{ij}, Y_{ik}) &= f(\alpha, j, k)\end{aligned}$$

- ▶ Linear model: Identity link

$$\mu_{ij} = X'_{ij}\beta$$

- ▶ Logistic model: logit link

$$\text{logit}(\mu_{ij}) = X'_{ij}\beta$$

- ▶ Estimation uses generalized estimating equations

- ▶ Add robust variance estimate to protect against misspecification of variance of the outcome and correlation model



Lecture 9 Review:

- Marginal Model: OR = 1.72

	Time = 0	Time = 1
Y = 0	61	46
Y = 1	50	65

- Conditional Model: OR = 10.89

	Y1 = 0	Y1 = 1
Y0 = 0	39	22
Y0 = 1	7	43



Two example studies

- ▶ Placebo-controlled trial to improve respiratory function
 - ▶ 111 patients
 - ▶ Baseline + 4 follow-ups
 - ▶ Compare the change in odds from baseline to follow-up across the active treatment vs. placebo groups.
- ▶ Matched case-control study looking at effect of exogenous estrogens on the risk of endometrial cancer
 - ▶ 63 matched sets: one case + 4 controls
 - ▶ Alive in same community at the time of diagnosis for the case, age within 1 year, same marital status and entered community at roughly the same time
 - ▶ Do women who use estrogens, have a history of gall-bladder disease or hypertension at increased risk of endometrial cancer?



Conditional Models

- ▶ Random effects logistic regression model:



Conditional Models

$$\begin{aligned}\text{logit}[Pr(Y_{ij} = 1 | post_{ij}, trtmnt01_i, b_i)] &= \beta_{0i}^c + \beta_1^c I(post_{ij} > 0) + \beta_2^c I(post_{ij} > 0) trtmnt01_i \\ &= \beta_0^c + b_i + \beta_1^c I(post_{ij} > 0) + \beta_2^c I(post_{ij} > 0) trtmnt01_i\end{aligned}$$

where $b_i \sim N(0, \sigma^2)$ and the covariates are independent of b_i .

Interpretation:

- β_{0i}^c : defines a patient specific log-odds of a good respiratory response at baseline
- $\beta_{0i}^c = \beta_0^c + b_i$, where $b_i \sim N(0, \sigma^2)$: β_0^c is the log-odds of a good respiratory response for the average patient (i.e. $b_i = 0$)
- $\beta_{0i}^c = \beta_0^c + b_i$, where $b_i \sim N(0, \sigma^2)$: b_i represents the deviation from this average log-odds of a good respiratory response for patient i



Example: Logistic regression with random intercept

$$\begin{aligned}\text{logit}[Pr(Y_{ij} = 1 | post_{ij}, trtmnt01_i, b_i)] &= \beta_{0i}^c + \beta_1^c I(post_{ij} > 0) + \beta_2^c I(post_{ij} > 0) trtmnt01_i \\ &= \beta_0^c + b_i + \beta_1^c I(post_{ij} > 0) + \beta_2^c I(post_{ij} > 0) trtmnt01_i\end{aligned}$$

where $b_i \sim N(0, \sigma^2)$ and the covariates are independent of b_i .

$$\mu_{ij}^c = \frac{\exp(\beta_0^c + b_i + \beta_1^c I(post_{ij} > 0) + \beta_2^c I(post_{ij} > 0) trtmnt01_i)}{1 + \exp(\beta_0^c + b_i + \beta_1^c I(post_{ij} > 0) + \beta_2^c I(post_{ij} > 0) trtmnt01_i)}$$

- ▶ Slopes are log [ratio of individual odds]!



Example: Random intercept logistic model in R using glmer

```
ri.fit = glmer(r~post + postXtrt+(1|id),data=data,family="binomial",nAGQ=7)
summary(ri.fit)
```

Random effects:

##	Groups	Name	Variance	Std.Dev.
----	--------	------	----------	----------

##	id	(Intercept)	6.49	2.55
----	----	-------------	------	------

Number of obs: 555, groups: id, 111

##

Fixed effects:

##		Estimate	Std. Error	z value	Pr(> z)
----	--	----------	------------	---------	----------

##	(Intercept)	-0.4212	0.3667	-1.15	0.25
----	-------------	---------	--------	-------	------

##	post	-0.0834	0.3683	-0.23	0.82
----	------	---------	--------	-------	------

##	postXtrt	1.9452	0.4850	4.01	6.1e-05 ***
----	----------	--------	--------	------	-------------

- ▶ Intercept: For the average or typical patient (i.e. $b_i = 0$), the probability of a good response is

$$\frac{\exp(-0.42)}{1+\exp(-0.42)} = 0.40$$

- ▶ You can compute baseline probability of a good response for any patient by: $\frac{\exp(-0.42+b_i)}{1+\exp(-0.42+b_i)}$



Example: Interpretation

```
ri.fit = glmer(r~post + postXtrt+(1|id),data=data,family="binomial",nAGQ=7)
summary(ri.fit)
```

```
## Random effects:
```

```
## Groups Name          Variance Std.Dev.
```

```
## id      (Intercept) 6.49      2.55
```

```
## Number of obs: 555, groups: id, 111
```

```
##
```

```
## Fixed effects:
```

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

```
## (Intercept) -0.4212      0.3667  -1.15    0.25
```

```
## post        -0.0834      0.3683  -0.23    0.82
```

```
## postXtrt     1.9452      0.4850   4.01 6.1e-05 ***
```



Example: Interpretation

```
ri.fit = glmer(r~post + postXtrt+(1|id),data=data,family="binomial",nAGQ=7)
summary(ri.fit)
```

```
## Random effects:
```

```
## Groups Name          Variance Std.Dev.
```

```
## id      (Intercept) 6.49      2.55
```

```
## Number of obs: 555, groups: id, 111
```

```
##
```

```
## Fixed effects:
```

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

```
## (Intercept) -0.4212      0.3667  -1.15    0.25
```

```
## post        -0.0834      0.3683  -0.23    0.82
```

```
## postXtrt     1.9452      0.4850   4.01 6.1e-05 ***
```



Comparison of marginal and conditional slope terms

Compare the marginal (β) and conditional (β^c) parameter estimates.

```
cbind(summary(fit.exch)$coeff[,1],summary(ri.fit)$coeff[,1])
```

```
##           [,1]      [,2]
## (Intercept) -0.1989 -0.42120
## post        -0.0410 -0.08343
## postXtrt     1.0083  1.94525
```

- Recall our discussion of confounding: Assume b_i is independent of covariates (as we do in random effects models)

Marginal model: $\text{logit}[Pr(Y_{ij}|X_{ij})] = \beta_0 + \beta_1 X_{ij}$

Conditional model: $\text{logit}[(Pr(Y_{ij}|X_{ij}, b_i))] = \beta_0^c + \beta_1^c X_{ij} + b_i$

In general:

- β = change in log population odds per unit change in X
- β^c = change in cluster-specific log odds per unit change in X

Estimation: Random effects logistic regression model

► Basic Idea:

$$f(Y|\beta^c, D) = \int f(Y, b_i|\beta^c, D)db_i = \int f(Y|b_i, \beta^c, D)f(b_i|D)db_i$$

$$\begin{aligned} L(y|\beta^c, D) &= \prod_{i=1}^m \int \prod_{j=1}^{n_i} (\mu_{ij}^c(\beta^c, b_i))^{y_{ij}} (1 - \mu_{ij}^c(\beta^c, b_i))^{1-y_{ij}} f(b_i|D)db_i \\ &= \prod_{i=1}^m \int Pr(y_{i1}, \dots, y_{in_i}|\beta^c, b_i)Pr(b_i|D)db_i \end{aligned}$$

It can be shown that:

$$\frac{\partial \log(L(y|\beta^c, D))}{\partial \beta^c} = \sum_{i=1}^m \sum_{j=1}^{n_i} X'_{ij} (y_{ij} - E_{b_i|y}(\mu_{ij}^c(b_i, \beta^c)))$$

Estimation: Random effects logistic regression model

$$\begin{aligned} L(y|\beta^c, D) &= \prod_{i=1}^m \int \prod_{j=1}^{n_i} (\mu_{ij}^c(\beta^c, b_i))^{y_{ij}} (1 - \mu_{ij}^c(\beta^c, b_i))^{1-y_{ij}} f(d_i|D) db_i \\ &= \prod_{i=1}^m \int Pr(y_{i1}, \dots, y_{in_i} | \beta^c, b_i) Pr(b_i | D) db_i \end{aligned}$$

- ▶ Solving the likelihood function requires estimation of the integral
- ▶ This is typically estimated via numerical methods
- ▶ Gaussian quadrature
- ▶ Adaptive gaussian quadrature
- ▶ Requires a number of quadrature points: $nAGQ = 7$



Special case: Matched Case-Control Study

- Consider the likelihood function for the logistic regression model with random intercept

$$\text{logit}[\mu_{ij}^c] = X_{ij}'\beta^c + b_i \quad b_i \sim N(0, \sigma^2).$$

$$L(y|\beta^c, \sigma^2) = \prod_{i=1}^m \int \frac{\exp \left[\left(\sum_{j=1}^{n_i} y_{ij} X_{ij} \right)' \beta^c + y_i^+ b_i \right]}{\prod_{j=1}^{n_i} (1 + \exp(X_{ij}'\beta^c + b_i))} f(b_i|\sigma^2) db_i$$

$$y_i^+ = \sum_{j=1}^{n_i} y_{ij} \text{ is sufficient for } b_i, \text{ i.e. } \Pr(y_{ij}|y_i^+, b_i) \text{ does not depend on } b_i$$



Special case: Matched Case-Control Study

► Data:

- Case: $Y_{i1} = 1, X_{i1}$
- Control: $Y_{i0} = 0, X_{i0}$

► Model:

$$Pr(Y_{ij} = 1 | X_{ij}, b_i) = \frac{\exp(X_{ij}'\beta^c + b_i)}{1 + \exp(X_{ij}'\beta^c + b_i)}$$

- Goal is to estimate parameters for X without making assumptions about distribution of b

$$CL(Y_i | \beta^c) = \prod_{i=1}^m [Pr(Y_{i0} = 0 | X_{i0}, y_i^+ = 1) Pr(Y_{i1} = 1 | X_{i1}, y_i^+ = 1)]$$



Special case: Matched Case-Control Study

$$\begin{aligned}Pr(Y_{i1} = 1|X_{i1}, Y_i^+ = 1, b_i) &= \frac{Pr(Y_{i1}=1 \text{ and } Y_i^+=1|b_i)}{Pr(Y_i^+=1|b_i)} \\&= \frac{Pr(Y_{i1}=1 \text{ and } Y_{i0}=0|b_i)}{Pr(Y_{i1}=1 \text{ and } Y_{i0}=0|b_i) + Pr(Y_{i1}=0 \text{ and } Y_{i0}=1|b_i)} \\&= \frac{Pr(Y_{i1}=1|b_i) \times Pr(Y_{i0}=0|b_i)}{Pr(Y_{i1}=1|b_i) \times Pr(Y_{i0}=0|b_i) + Pr(Y_{i1}=0|b_i) \times Pr(Y_{i0}=1|b_i)} \\&= \frac{\left(\frac{\exp(X_{i1}\beta^c + b_i)}{1 + \exp(X_{i1}\beta^c + b_i)} \times \frac{1}{1 + \exp(X_{i0}\beta^c + b_i)} \right)}{\frac{\exp(X_{i1}\beta^c + b_i)}{1 + \exp(X_{i1}\beta^c + b_i)} \times \frac{1}{1 + \exp(X_{i0}\beta^c + b_i)} + \frac{1}{1 + \exp(X_{i1}\beta^c + b_i)} \times \frac{\exp(X_{i0}\beta^c + b_i)}{1 + \exp(X_{i0}\beta^c + b_i)}} \\&= \frac{\exp(X_{i1}\beta^c + b_i)}{\exp(X_{i1}\beta^c + b_i) + \exp(X_{i0}\beta^c + b_i)} \\&= \frac{\exp(X_{i1}\beta^c)}{\exp(X_{i1}\beta^c) + \exp(X_{i0}\beta^c)} \\&= \frac{\exp((X_{i1} - X_{i0})\beta^c)}{1 + \exp((X_{i1} - X_{i0})\beta^c)}\end{aligned}$$

Special case: Matched Case-Control Study

$$CL(Y|\beta^c) = \prod_{i=1}^m \left[\frac{\exp((X_{i1} - X_{i0})\beta^c)}{1 + \exp((X_{i1} - X_{i0})\beta^c)} \right]^1$$

- ▶ Marginal logistic regression:
 - ▶ No intercept
 - ▶ Responses $Y = 1$
 - ▶ Covariates:

$$(X_{11} - X_{10}, X_{21} - X_{20}, \dots, X_{m1} - X_{m0})$$



Example: Endometrial cancer example

Matched case-control study looking at effect of exogenous estrogens on the risk of endometrial cancer

- ▶ 63 matched sets: one case + 4 controls
 - ▶ Alive in same community at the time of diagnosis for the case, age within 1 year, same marital status and entered community at roughly the same time
 - ▶ Do women who use estrogens, have a history of gall-bladder disease or hypertension at increased risk of endometrial cancer?
-
- ▶ I will do the analysis using the case + 1 matched control
 - ▶ You will revisit this data in Problem Set 3 using all the participants.



Example: Endometrial cancer example

```
dat = read.table("./endometrial.txt")
names(dat) = c("set", "case", "age", "ageg", "est", "gall", "hyp", "obesity", "nonestdrug")
dat$est = dat$est - 1
dat$gall = dat$gall - 1
dat$hyp = dat$hyp - 1
dat$obesity[dat$obesity==3] = NA
dat$obesity = dat$obesity - 1
dat$nonestdrug = dat$nonestdrug - 1
dat$firstctrl = unlist(tapply(dat$set, dat$set, FUN=function(x) c(0,1,rep(0,length(x)-2))))

tapply(dat$est, dat$case, mean)

##      0      1
## 0.5040 0.8889

tapply(dat$gall, dat$case, mean)

##      0      1
## 0.09524 0.26984

tapply(dat$hyp, dat$case, mean)

##      0      1
## 0.3254 0.4127
```



Example: Endometrial cancer example

```
library(survival)

## Warning: package 'survival' was built under R version 3.6.3

## Fit the conditional logistic model with
## all three exposures using only 1st control
fit1=clogit(case~est+gall+hyp+ strata(set), data=subset(dat,case==1|firstctrl==1))
summary(fit1)$coeff

##              coef exp(coef)  se(coef)      z      Pr(>|z|)
## est    2.2479841  9.4686292  0.6255817  3.5934304  0.0003263528
## gall   0.6907726  1.9952565  0.6157373  1.1218625  0.2619209223
## hyp   -0.1333443  0.8751637  0.4455392 -0.2992874  0.7647207469

## Drop hypertension from the model
fit1=clogit(case~est+gall+strata(set),
  data=subset(dat,case==1|firstctrl==1))
summary(fit1)$coeff

##              coef exp(coef)  se(coef)      z      Pr(>|z|)
## est    2.209052   9.107077  0.6097099  3.623120  0.0002910712
## gall   0.694732   2.003172  0.6156339  1.128482  0.2591162174
```



Example: Endometrial cancer example

```
## Add the interactions
fit1.int=clogit(case~est*gall+strata(set),
               data=subset(dat,case==1|firstctrl==1))
summary(fit1.int)$coeff

##              coef exp(coef) se(coef)      z    Pr(>|z|)
## est          2.671060 14.4552809 0.7533387  3.545629 0.0003916766
## gall          2.292397  9.8986370 1.2224136  1.875304 0.0607509226
## est:gall     -2.141460  0.1174832 1.3700403 -1.563064 0.1180376313

# Compute the synergistic effect
coeff.sum = sum(fit1.int$coefficients)
var.sum = t(c(1,1,1)) %*% vcov(fit1.int) %*% c(1,1,1)
exp(coeff.sum)

## [1] 16.81038

exp(coeff.sum-1.96*sqrt(var.sum))

##           [,1]
## [1,] 2.855665

exp(coeff.sum+1.96*sqrt(var.sum))

##           [,1]
## [1,] 98.95735
```



Example: Endometrial cancer example

- ▶ In summary, both estrogen use and history of gall bladder disease were found to increase the risk of endometrial cancer. Furthermore, these risk factors were found to be non-additive. That is, on the log odds scale, the risk associated with having both risk factors is only marginally greater than the risk associated with having a single risk factor. However, on the odds scale this translates to a substantive increase in risk. One way to interpret the findings is below.
- ▶ The estimated odds of being a case for subjects with only estrogen use are 14.5 (95% CI: 3.1 to 71.4) times the odds of being a case for subjects with neither estrogen use or history of gallbladder disease.
- ▶ The estimated odds of being a case for subjects with only a history of gall bladder disease are 9.9 (95%CI: 0.95 to 104.8) times the odds of being a case for subjects with neither estrogen use or history of gallbladder disease.
- ▶ Finally, the estimated odds of being a case for subjects with both estrogen use and gall bladder disease are 16.8 (95% CI: 2.9 to 99.0) times the odds of being a case for subjects with neither estrogen use or history of gallbladder disease. This is approximately double the odds ratio from either risk factor alone.



Log-linear models for count variables

- ▶ Count variable
 - ▶ Takes on values of non-negative integers
 - ▶ 0, 1, 2, ..., 3321, 10001,
- ▶ Examples
 - ▶ Number of non-accidental deaths per day in Chicago
 - ▶ Number of days of work missed due to illness within a year
 - ▶ Number of myocardial infarctions (MIs) among patients at risk for MI
- ▶ Notice anything? Counts of things occurring within a given time range or group of eligible persons



Log-linear models for count variables

- ▶ Characteristics of count variables
 - ▶ Non-negative integers
 - ▶ Variability tends to increase as mean increases
 - ▶ Effects of predictors tend to be multiplicative (reflecting relative changes not absolute change)

EXAMPLE: Numbers of Non-accidental Death per Day in Chicago, 1987-1994

Season	Mean	Variance	Variance/Mean
Winter (Dec-Feb)	122	177.6	1.45
Summer (June-Aug)	107	128.4	1.20



Poisson process

- ▶ Poisson process defines how observations of events of interest occur over time or space
- ▶ Imagine a range of time $[0, T]$ and breaking that range of time into small bins $[t, t+dt]$
- ▶ $\Pr(\text{Event occurs in } [t, t+dt]) = \lambda dt$
- ▶ $\Pr(2 \text{ or more events occur in } [t, t+dt]) \sim 0$
- ▶ Memoryless property: chance of an event in one interval is independent of the chance of an event in a future interval
- ▶ In a Poisson process, the event times in an interval $[0, T]$ are uniformly distributed, that is, have equal chance of occurring anywhere in the part of the interval.



Poisson process

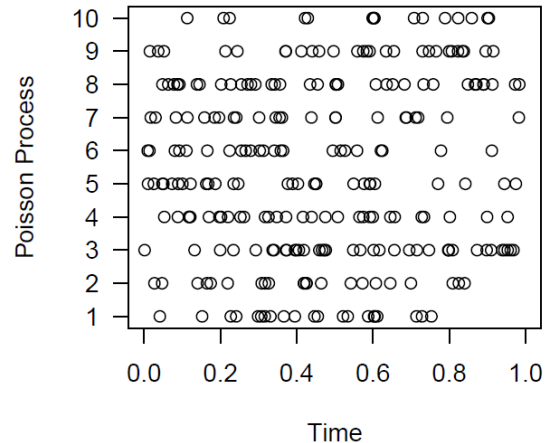
- ▶ The number of events X occurring in the interval $[0, T]$ follows a Poisson distribution

- ▶ Probability mass function: $P(X = x) = \frac{e^{-\lambda} \lambda^x}{x!}$

See page 3 of Lecture 10 handout for derivation.

- ▶ The mean and variance of X is λT

10 Realizations of Poisson Process



Log-linear model

- ▶ First formulation -> we will assume exposure time is the same for all observations!

- ▶ General form:

$$Y_i \sim P(\mu_i), i = 1, \dots, n \text{ independent}$$

$$\log(E(Y_i)) = \log(\mu_i) = \beta_0 + \beta_1 X_{i1} + \dots + \beta_p X_{ip}$$

- ▶ Interpretation:



Log-linear model

- ▶ First formulation -> we will assume exposure time is the same for all observations!
- ▶ Hypothetical example: a study of insulin-dependent diabetic patients followed for 4 weeks after acquiring an insulin pump. The patients record and report the total number of hypoglycemic episodes during the 4 week follow-up.
- ▶ The goal of the analysis is to compare the total number of hypoglycemic episodes for male and female diabetic patients



Example: Same exposure time

$$\text{Log}(E(Y_i)) = \text{Log}(\mu_i) = \beta_0 + \beta_1 \text{male}_i$$

```
set.seed(1346)
N = 100
male = rbinom(N,1,0.5)
Y= rpois(N,exp(log(12)+0.2*male))
summary(glm(Y~male,family="poisson"))$coefficients
```

##	Estimate	Std. Error	z value	Pr(> z)
## (Intercept)	2.5176965	0.04016096	62.690141	0.0000000000
## male	0.1956729	0.05421405	3.609266	0.0003070652

- $\hat{\beta}_0$ is the logarithm of the mean number of hypoglycemic episodes during the 4-week follow-up among females. The mean number of hypoglycemic episodes among females during the follow-up is $\exp(\hat{\beta}_0) = \exp(2.52) = 12.4$.
- $\hat{\beta}_0 + \hat{\beta}_1$ is the logarithm of the mean number of hypoglycemic episodes during the 4-week follow-up among males. The mean number of hypoglycemic episodes among males during the follow-up is $\exp(\hat{\beta}_0 + \hat{\beta}_1) = \exp(2.52 + 0.20) = 15.2$.



Example: Same exposure time

$$\text{Log}(E(Y_i)) = \text{Log}(\mu_i) = \beta_0 + \beta_1 \text{male}_i$$

```
set.seed(1346)
N = 100
male = rbinom(N,1,0.5)
Y= rpois(N,exp(log(12)+0.2*male))
summary(glm(Y~male,family="poisson"))$coefficients
```

##	Estimate	Std. Error	z value	Pr(> z)
## (Intercept)	2.5176965	0.04016096	62.690141	0.0000000000
## male	0.1956729	0.05421405	3.609266	0.0003070652

- $\hat{\beta}_1$ is the difference in the log mean number of hypoglycemic episodes during the 4 week follow-up comparing males to females OR the log relative mean number of hypoglycemic episodes during the 4 week follow-up comparing males to females.
- $\exp(\hat{\beta}_1) = \exp(0.20) = 1.22$ represents the relative mean number of hypoglycemic episodes comparing males to females. The mean number of hypoglycemic episodes during the 4-week follow-up is 22% greater for males compared to females.



Log-linear model

- ▶ Second formulation -> we will NOT assume exposure time is the same for all observations!
- ▶ Hypothetical example: a study of insulin-dependent diabetic patients followed up to 4 weeks after acquiring an insulin pump.
- ▶ Now suppose that not all patients were able to be followed for the entire 4-week period; patients were followed from **10 to 28 days**. Patients report the number of hypoglycemic episodes within the duration of the patient's specific follow-up.
- ▶ The goal of the analysis is to compare the total number of hypoglycemic episodes for male and female diabetic patients



Example: Variable exposure time

$Y_i \sim P(\mu_i) = P(N_i \lambda_i), i = 1, \dots, n$ independent

$$\begin{aligned}\text{Log}(E(Y_i)) &= \text{Log}(\mu_i) \\ &= \text{Log}(N_i \lambda_i) \\ &= \text{Log}(N_i) + \text{Log}(\lambda_i) \\ &= \text{Log}(N_i) + \beta_0 + \beta_1 \text{male}_i\end{aligned}$$

- for patient i , the expected number of hypoglycemic episodes is $N_i \lambda_i$ where N_i is the total follow-up time in days for patient i and λ_i is the risk of a hypoglycemic episode per unit time / per day.
- β_0 is the logarithm of the risk of a hypoglycemic episode in a day for females.
- $\beta_0 + \beta_1$ is the logarithm of the risk of a hypoglycemic episode in a day for males.
- $\exp(\beta_1)$ is the relative risk of a hypoglycemic episode in a day comparing males to females OR the relative expected number of hypoglycemic episodes comparing males and females who have the same duration of follow-up.



Example: Variable exposure time

$$\log(E(Y_i)) = \log(\mu_i) = \log(N_i \lambda_i) = \log(N_i) + \beta_0 + \beta_1 \text{male}_i$$

```
##              Estimate Std. Error   z value    Pr(>|z|)
## (Intercept) -0.2752677 0.03603750 -7.638368 2.199923e-14
## male         0.1142061 0.05012278  2.278527 2.269520e-02

expected.Y = fit$fitted
predicted.lambda = exp(fit$coefficients[1] + male*fit$coefficients[2])
head(cbind(N,Y,male,expected.Y,predicted.lambda))

##      N  Y male expected.Y predicted.lambda
## 1 17 19    1   14.47107      0.8512397
## 2 22 18    0   16.70611      0.7593688
## 3 19 16    1   16.17355      0.8512397
## 4 19 15    1   16.17355      0.8512397
## 5 22 13    0   16.70611      0.7593688
## 6 25 18    1   21.28099      0.8512397
```



Example: Variable exposure time

$$\log(E(Y_i)) = \log(\mu_i) = \log(N_i \lambda_i) = \log(N_i) + \beta_0 + \beta_1 \text{male}_i$$

##	Estimate	Std. Error	z value	Pr(> z)
## (Intercept)	-0.2752677	0.03603750	-7.638368	2.199923e-14
## male	0.1142061	0.05012278	2.278527	2.269520e-02

► Interpret β_0

► Interpret β_1



Estimation: Maximum likelihood estimation

The likelihood function is:

$$L(\beta|Y) = \prod_{i=1}^n \frac{e^{-\mu_i} \mu_i^{y_i}}{y_i!}$$

The log-likelihood is:

$$\log L(\beta|Y) = \sum_{i=1}^n (-\mu_i) + y_i \log(\mu_i) - \log(y_i!)$$

The score equation is:

$$\frac{\partial \log L(\beta|Y)}{\partial \beta} = \sum_{i=1}^n \left(-\frac{\partial \mu_i}{\partial \beta} \right) + y_i \frac{\partial \log(\mu_i)}{\partial \beta}$$

$$= \sum_{i=1}^n (-\mu_i X_i') + y_i X_i'$$

$$= \sum_{i=1}^n X_i' (y_i - \mu_i)$$

$$\hat{\beta} \sim N(\beta, (X' \text{diag}(\hat{\mu}) X)^{-1})$$



Robust variance estimation

Count data is almost always over-dispersed, i.e. $Var(Y_i) > E(Y_i)$.

Solution: Assume $E(Y_i|X_i) = \mu_i = N_i e^{X_i^T \beta}$ and $Var(Y_i|X_i) = \mu_i \phi$.

We can estimate ϕ by:

$$\hat{\phi} = \sum_{i=1}^n \frac{(y_i - \hat{\mu}_i)^2}{\hat{\mu}_i} \bigg/ (n - p)$$

which is the Pearson residual estimate of ϕ .

Alternatively, you can use the deviance estimator as:

$$\hat{\phi} = 2 \sum_{i=1}^n [Y_i \log(Y_i / \mu_i) - (Y_i - \mu_i)] \bigg/ (n - p)$$

Either is fine for computing the robust variance estimate.



Example: Robust variance estimation

- ▶ Daily non-accidental deaths in Chicago, 1987 – 1994
- ▶ Log-linear model for daily deaths as a function of:
 - ▶ PM10
 - ▶ Current temperature + average of prior three days (natural spline 3 df)
 - ▶ Time: year, season, month
- ▶ Data are overdispersed; greater variance than expected by Poisson model



Example: Robust variance estimation

```
fit.poisson.year = glm(total~ pm10+ns(temp,3)+ns(avgtemp,3)+as.factor(year),  
  data=data,family="poisson")
```

```
fit.robust.year = glm(total~ pm10+ns(temp,3)+ns(avgtemp,3)+as.factor(year),  
  data=data,family="quasipoisson")
```

##	Poisson beta	Poisson SE	Robust beta	Robust SE
## 1	0.00349	0.00104	0.00349	0.00116
## 2	0.00229	0.00107	0.00229	0.00117
## 3	0.00178	0.00111	0.00178	0.00118



Next time....

- ▶ Estimation of excess deaths after Hurricane Maria

