# More On Logistic Regression Week 9

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# 1 Other Logistic Regression Models

#### 1.1 Session Overview

- New Packages!
- Other Logistic Regression Models
  - Types
  - Rationale
  - Requirements
  - Commands
- Examples
  - Model Development
  - Diagnostics
  - Interpretations
- Visualization

### 1.2 Packages

library(survival)

• For conditional logistic regression

library(nnet)

• For multinomial logistic regression

library(MASS)

• For ordinal logistic regression

library(gtsummary)

library(car)

# 1.3 Logistic Regression for Other Outcome Types

- Conditional Logistic
- For situations with non-independence (i.e. matched studies)
- Multinomial/Polytomous Logistic
- For outcomes with more than 2 categories with no assumed order
- Ordinal Logistic
- For outcomes with more than 2 ordered categories

# 2 Conditional Logistic Regression

#### 2.1 Overview

- An extension of binary logistic regression
- For matched data with a binary outcome
- The unit of analysis is the *group/match*, not the individual
- Permits adjustment for variables not involved in direct matching

## Unmatched Sample

	Event	Non-even
Exposed	a	b
Unexposed	$\mathbf{c}$	d

$$OR = \frac{(a)(d)}{(b)(c)}$$

## 1:1 Matched Sample

	Non-event Exposed	Non-event Unexposed
Event	W	X
Exposed Event Unexposed	Y	Z

$$OR = \frac{(X)}{(Y)}$$

# 2.2 Regression Characteristics

- Conditional logit utilizes the exact conditional likelihood
- There is no valid baseline term (b0) in conditional logistic regression
- You cannot adjust for variables used in the matching process
- Conditional logit results are inherently biased toward the null

#### 2.3 Data Requirements

- A binary outcome coded 0 vs. 1
- Covariates of interest
- Basic assumptions of normality apply for continuous variables
- Frequencies of categories in factor variables must be sufficient (i.e. > 10)
- A matching/grouping identifier (group)

patientid	age	sex	x2	x3	group	outcome	
patientid	age	sex	x2	x3	group	outcome	
A_123456	25	Μ	243	A	1	0	
B_298298	25	Μ	125	G	1	1	
$C_{222444}$	49	$\mathbf{F}$	284	В	2	0	
D_554457	49	$\mathbf{F}$	96	В	2	1	
E_000456	18	$\mathbf{F}$	101	$\mathbf{C}$	3	0	
F_234292	18	$\mathbf{F}$	192	$\mathbf{C}$	3	1	
$G_{245221}$	62	Μ	204	Y	4	0	
H_501100	62	$\mathbf{M}$	222	Z	4	1	

#### 2.4 Commands

from library(survival)

```
model.clogit <- clogit(outcome ~ x1 + x2 + ... + xn + strata(group_variable), data = df)
summary(model.clogit)</pre>
```

• Why is clogit in the survival package?

At its most basic level and with a binary outcome, a conditional logit has the same likelihood formula as a stratified Cox model where time is a constant. The clogit command is essentially a *wrapper* for the coxph command with pre-specified arguments.

By default, only the beta estimates are stored in model.clogit. The ORs will have to be exponentiated manually.

- Coefficients are stored in the object as: clogit.res\$coef
- Confidence intervals are stored as: clogit.res\$confint

```
expConvert <- cbind("Odds Ratio" = exp(coef(model.clogit)), exp(confint(model.clogit)))</pre>
```

The cbind() command is from base R and will create a mini data frame to hold the exponentiated ORs and 95% CIs.

cbind stands for "Column Bind" and will append columns to data frames.

# 3 Multinomial Logistic Regression

#### 3.1 Overview

- An extension of binary logistic regression
- Outcome variable is nominal with >2 categories
- Categories do not have a logical order to them
- Observations can only experience **one category** of the outcome
- Many names for this: Polytomous, Polychotomous, Multiclass
- Still utilizes Maximum Likelihood Estimation

#### 3.2 Basic Modeling Assumptions

- 1. One reference category for the outcome  $(event_0)$
- 2. An outcome category (where R=1) to assess to the reference (event<sub>R</sub>) and:
- At least one other outcome category (where R > 1) to assess to the reference (event<sub>R</sub>)
- 3. A common list of independent variables  $(x_1, x_2, ..., x_k)$
- 4. A unique baseline probability  $(b_{0,R})$  specific to the outcome category that is being analyzed
- 5. Unique estimates of association  $(b_{1.R}, b_{2.R}, ..., b_{k.R})$  specific to the outcome category and independent variables being analyzed

$$ln(\frac{p_{event_R}}{p_{event_0}}) = b_{0.R} + b_{1.R}(x_1) + b_{2.R}(x_2) + \ldots + b_{k.R}(x_k)$$

# 3.3 Example Model

Outcome: Recategorized df.ed\$disposition

- Reference category: HOME (event<sub>0</sub>)
- Outcome category 1: ADMITTED (event<sub>1</sub>)
- Outcome category 2: OTHER ( $event_2$ ) basically, everything else (ELOPED, LEFT AGAINST MEDICAL ADVICE, LEFT WITHOUT BEING SEEN, OTHER, TRANSFER)
- Independent variables: racewhite, highpain, los, gender

Comparison 1: Discharge to HOME vs. ADMITTED

$$ln(\frac{p_{event_1}}{p_{event_0}}) = b_{0.1} + b_{1.1}(racewhite) + b_{2.1}(highpain) + b_{3.1}(los) + b_{4.1}(gender)$$

Comparison 2: Discharge to Care Facility vs. Routine Home Discharge

$$ln(\frac{p_{event_2}}{p_{event_0}}) = b_{0.2} + b_{1.2}(racewhite) + b_{2.2}(highpain) + b_{3.2}(los) + b_{4.2}(gender)$$

#### 3.4 Analysis Details

Resembles two independent binary logistic regression models with the same reference category. HOWEVER:

In binary logistic regression:

$$1.00 = p + q$$

Where p = probably of the event and q = probability of not having the event and both must equal to 100%.

For polychotomous logistic regression:

$$1.00 = p_0 + p_1 + p_2 + \dots + p_R$$

Therefore, the likelihood function (and log-likelihood) is based on the probability of all outcome categories and their inputs and respective coefficients.

#### 3.5 Regression Caveats

- There are no specific diagnostic tools for multicollinearity or outlier assessment for multinomial logits
- Can use the vif() commands out of library(car) for each pairwise outcome set (manually)
- Typically requires a very large sample due to multiple equations
- Can take a long time to compute
- Convergence toward a solution is adversely affected by low cell counts or poorly-powered samples
- The Independence of Irrelevant Alternatives Assumption
- Assumes that removal of outcome categories does not affect the odds of retained outcome categories

## 3.6 Preparation

First, check the distribution of your outcome to confirm its structure:

```
table(df$outcome)
```

By default, R will use the first value it shows as the reference category. Revise the reference group as needed:

```
df$outcome <- relevel(df$outcome, ref = "REFERENCE")</pre>
```

Covariates should be appropriately treated before proceeding with the regression

#### 3.7 Commands

Commands for the primary mlogit analysis come from nnet.

```
library(nnet)
model <- multinom(outcome ~ var1 + var2 + ... + varN, data=df)
summary(model)</pre>
```

The outcome variable should be already set as a factor. The lowest category is the default reference unless it has been releveled

summary (model) displays the *coefficients* and *standard errors* for each variable for each outcome category (vs. the reference)

#### 3.8 Model Development

```
library(nnet)

df <- df %>%
    mutate(disp3cat = case_when(
        disposition == "HOME" ~ "HOME",
        disposition == "ADMITTED" ~ "ADMITTED",
        .default = "OTHER"
    ))

df <- df %>%
    mutate(across(c("disp3cat", "racewhite", "arrival_transport", "gender"), as.factor))

table(df$disp3cat)
```

```
ADMITTED HOME OTHER 150 60 12
```

```
df$disp3cat <- relevel(df$disp3cat, ref = "HOME")</pre>
mlogit.m1 <- multinom(disp3cat ~ racewhite + lnlos + highpain + gender, data = df)</pre>
# weights: 18 (10 variable)
initial value 243.891928
iter 10 value 166.399395
final value 165.942351
converged
summary(mlogit.m1)
Call:
multinom(formula = disp3cat ~ racewhite + lnlos + highpain +
   gender, data = df)
Coefficients:
        (Intercept) racewhite1
                                   lnlos
                                            highpain
                                                      genderM
ADMITTED
           4.726817 -0.2381286 -0.6649349 0.08767236 0.6667535
OTHER
           1.611070 -1.0402787 -0.5116694 -0.01617277 1.0502466
Std. Errors:
                                                   genderM
        (Intercept) racewhite1
                                  lnlos highpain
ADMITTED
           OTHER
           2.667780 0.7269330 0.4441803 0.7047875 0.7229142
Residual Deviance: 331.8847
```

The multinom() function doesn't automatically show p-values, and the output only contains raw coefficients and standard errors.

#### 3.9 Using gtsummary for Display

AIC: 351.8847

```
library(gtsummary)
mlogit.m1 %>% tbl_regression(exponentiate = TRUE)
```

- i Multinomial models have a different underlying structure than the models gtsummary was designed for.
- \* Functions designed to work with `tbl\_regression()` objects may yield unexpected results.

Characteristic	OR	95% CI	p-value	
ADMITTED				
racewhite				
0		<u>—</u>		
1	0.79	0.38,  1.62	0.5	
lnlos	0.51	0.32,0.82	0.005	
highpain	1.09	0.55,  2.18	0.8	
gender				
$\mathbf{F}$		<u>—</u>		
M	1.95	0.97,  3.93	0.062	
OTHER				
racewhite				
0		<u>—</u>		
1	0.35	0.09, 1.47	0.2	
lnlos	0.60	0.25, 1.43	0.2	
highpain	0.98	0.25,3.92	>0.9	
gender				
F	_			
${ m M}$	2.86	0.69, 11.8	0.15	

 $\overline{\text{Abbreviations: CI} = \text{Confidence Interval, OR} = \overline{\text{Odds Ratio}}}$ 

# 4 Ordinal Logistic Regression

#### 4.1 Overview

- For categorical outcomes that had an order to to them
- Must have >2 categories
- Also known as proportional odds regression (not to be confused with proportional hazards regression)
- Only one model equation is generated and only one set of coefficients are estimated

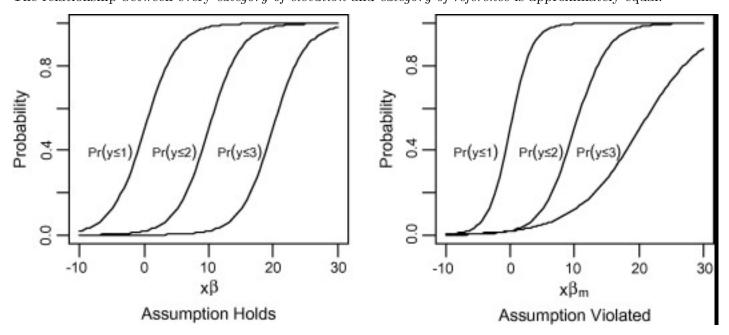
#### 4.2 Modeling Structure

- 1. Any number of ordinal outcome categories (event<sub>R</sub>), where R > 1
- 2. Reference outcome category is all categories that precede the one of interest  $(event_{1-R})$  for each comparison
- 3. A common list of independent variables  $(x_1, x_2, ..., x_n)$
- 4. Unique baseline probabilities  $(b_{0.R})$  specific to the outcome categories being analyzed
- 5. One set of estimates  $(b_1, b_2, ..., b_n)$  for the entire model

$$ln(\frac{p_{event_R}}{p_{event_{1-R}}}) = b_{0.R} + b_1(x1) + b_2(x2) + \ldots + b_n(xn)$$

#### 4.3 Proportional Odds Assumption

The relationship between every category of elevation and category of reference is approximately equal.



We can test the assumption for each risk factor using the Cochran-Mantel-Haenszel test.

Involves calculation of the probability of the outcome when increasing from a *lower category* to the *next category* in the order based on the independent variables

#### 4.4 Commands

From library (MASS)

```
library(MASS)
model <- polr(outcome ~ x1 + x2 + x3, data=df, Hess=TRUE)
summary(model)</pre>
```

- polr() is the command for ordinal logistic regression
- outcome ~ x1 + x2 + x3 is the standard formula notation in R
- Hess = TRUE includes the Hessian matrix; required for generating standard errors

#### 4.5 Modeling Demonstration

```
library(MASS)
Attaching package: 'MASS'
The following object is masked from 'package:gtsummary':
    select
The following object is masked from 'package:dplyr':
    select
table(df$opioidlvl)
 Extreme
             Mild Moderate
                                None
      20
               23
                        23
                                 156
df$opioidlvl <- factor(df$opioidlvl, levels = c("None", "Mild", "Moderate", "Extreme"), ordered = TR
ologit.m1 <- polr(opioidlvl ~ lnlos + gender, data = df, Hess=TRUE)</pre>
summary(ologit.m1)
```

Characteristic	OR	95% CI
lnlos	1.39	0.91, 2.16
gender		
$\mathbf{F}$	_	_
M	4.45	2.44, 8.35

Abbreviations: CI = Confidence Interval, OR = Odds Ratio

#### Call:

polr(formula = opioidlvl ~ lnlos + gender, data = df, Hess = TRUE)

#### Coefficients:

Value Std. Error t value lnlos 0.3292 0.2211 1.489 genderM 1.4920 0.3131 4.766

#### Intercepts:

 Value
 Std. Error t value

 None|Mild
 3.5860 1.3397 2.6767

 Mild|Moderate
 4.2099 1.3468 3.1259

 Moderate|Extreme
 5.1651 1.3665 3.7797

Residual Deviance: 387.0998

AIC: 397.0998

tbl\_regression(ologit.m1, exponentiate = TRUE)

#### 4.6 Testing the Proportional Odds Assumption

Used to be an involved process that required you to generate the dummy variables and evaluate each association in normal binary logits.

Now, you can use the car package!

#### library(car)

Loading required package: carData

Attaching package: 'car'

The following object is masked from 'package:dplyr':

recode

```
The following object is masked from 'package:purrr': some
```

```
poTest(ologit.m1)
```

```
Tests for Proportional Odds
polr(formula = opioidlvl ~ lnlos + gender, data = df, Hess = TRUE)
        b[polr] b[>None] b[>Mild] b[>Moderate] Chisquare df Pr(>Chisq)
Overall
                                                     6.70 4
                                                                   0.15
                                                     5.97 2
lnlos
          0.329
                   0.241
                            0.189
                                         0.811
                                                                   0.05 .
                                         2.045
          1.492
                   1.441
                            1.534
                                                     1.00 2
                                                                   0.61
genderM
```

Signif. codes: 0 '\*\*\*' 0.001 '\*\*' 0.01 '\*' 0.05 '.' 0.1 ' ' 1

## 4.7 Testing Multicollinearity

Collinearity can be evaluated with the variance inflation factor vif() from the car package

```
library(car)
vif(ologit.m1)
```

lnlos gender
1.000113 1.000113