Introduction to Logistic Regression

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Today's Lecture

Logistic regression

[Note: more on logistic regression can be found in Kaplan, Chapter 16 and the OpenIntro Statistics textbook, Chapter 8. These slides are based, in part, on the slides from OpenIntro.]

Regression so far ...

At this point we have covered:

- Simple linear regression
 - Relationship between numerical response and a numerical or categorical predictor
- Multiple regression
 - Relationship between numerical response and multiple numerical and/or categorical predictors

What we haven't covered is what to do when the response is not continuous (i.e. categorical, count data, etc.)

Example - Birdkeeping and Lung Cancer

A 1972 - 1981 health survey in The Hague, Netherlands, discovered an association between keeping pet birds and increased risk of lung cancer. To investigate birdkeeping as a risk factor, researchers conducted a case-control study of patients in 1985 at four hospitals in The Hague (population 450,000). They identified 49 cases of lung cancer among the patients who were registered with a general practice, who were age 65 or younger and who had resided in the city since 1965. They also selected 98 controls from a population of residents having the same general age structure.

Ramsey, F.L. and Schafer, D.W. (2002). The Statistical Sleuth: A Course in Methods of Data Analysis (2nd ed)

Example - Birdkeeping and Lung Cancer - Data

```
library(Sleuth3)
birds = case2002
head(birds)

## LC FM SS BK AG YR CD

## 1 LungCancer Male Low Bird 37 19 12

## 2 LungCancer Male Low Bird 41 22 15

## 3 LungCancer Male High NoBird 43 19 15

## 4 LungCancer Male Low Bird 46 24 15

## 5 LungCancer Male Low Bird 49 31 20

## 6 LungCancer Male High NoBird 51 24 15
```

```
LC Whether subject has lung cancer
FM Sex of subject
S Socioeconomic status
BK Indicator for birdkeeping
AG Age of subject (years)
YR Years of smoking prior to diagnosis or examination
CD Average rate of smoking (cigarettes per day)
```

NoCancer is the reference response (0 or failure), LungCancer is the non-reference response (1 or success) - this matters for interpretation.

Example - Birdkeeping and Lung Cancer - Data

What types of associations do you expect to see between the predictors below and lung cancer? Might you expect any interactions to be present?

LC Whether subject has lung cancer

FM Sex of subject

SS Socioeconomic status

BK Indicator for birdkeeping

AG Age of subject (years)

YR Years of smoking prior to diagnosis or examination

CD Average rate of smoking (cigarettes per day)

Interpreting linear regressions of binary data

We can use linear regression for binary data, and for *very simple models* it gives reasonable and interpretable output.

 $IC \sim 1$

```
birds$LCnum <- as.numeric(birds$LC=="LungCancer")</pre>
sum(birds$LCnum)
## [1] 49
nrow(birds)
## [1] 147
summary(lm(LCnum ~ 1, data=birds))$coef
##
                Estimate Std. Error t value Pr(>|t|)
## (Intercept) 0.3333333 0.03901372 8.544004 1.574829e-14
```

Interpreting linear regressions of binary data

We can use linear regression for binary data, and for *very simple models* it gives reasonable and interpretable output.

$$LC \sim FM$$

```
mod1 <- lm(LCnum ~ FM, data=birds)
round(summary(mod1)$coef, 3)

## Estimate Std. Error t value Pr(>|t|)
## (Intercept) 0.333 0.079 4.214 0
## FMMale 0.000 0.091 0.000 1
```

What is this model's estimated probability of lunc cancer for men? for women?

Interpreting linear regressions of binary data

But if the model gets too complicated, then it can produce some tricky results.

$$LC \sim FM + YR$$

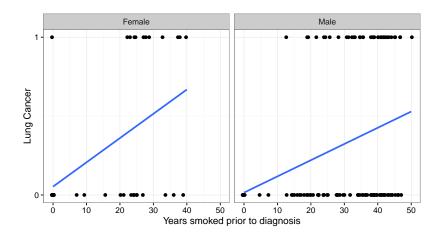
```
mod2 <- lm(LCnum ~ FM + YR, data=birds)
round(summary(mod2)$coef, 3)

## Estimate Std. Error t value Pr(>|t|)
## (Intercept) 0.119 0.092 1.294 0.198
## FMMale -0.150 0.094 -1.592 0.114
## YR 0.012 0.003 4.044 0.000
```

What is this model's estimated probability of lunc cancer for men who have never smoked? for women who never smoked?

Lung cancer as a function of smoking years

```
(p <- ggplot(birds, aes(x=YR, y=as.numeric(LC=="LungCancer")*1)) +
   geom_jitter(height=0) + geom_smooth(method="lm", se=FALSE) +
   ylab("Lung Cancer") + xlab("Years smoked prior to diagnosis") +
   scale_y_continuous(breaks=c(0,1)) +
   facet_grid(.~FM))</pre>
```



Odds

Odds are another way of quantifying the probability of an event, commonly used in gambling (and logistic regression).

For some event E,

$$odds(E) = \frac{P(E)}{P(E^c)} = \frac{P(E)}{1 - P(E)}$$

Similarly, if we are told the odds of E are x to y then

$$odds(E) = \frac{x}{y} = \frac{x/(x+y)}{y/(x+y)}$$

which implies

$$P(E) = x/(x + y), \quad P(E^c) = y/(x + y)$$

Odds Ratios

Odds Ratios compare the odds of an event in two different groups. For some outcome of interest (say, disease) in two groups, (e.g. exposed and unexposed),

$$OR = \frac{P(\mathsf{disease}|\mathsf{exposed})/[1-P(\mathsf{disease}|\mathsf{exposed})]}{P(\mathsf{disease}|\mathsf{unexposed})/[1-P(\mathsf{disease}|\mathsf{unexposed})]}$$

Facts about Odds Ratios

- ORs have range of $(0, \infty)$.
- ullet OR = 1 means no difference between the groups.
- They have a multiplicative scale: e.g. OR = 0.5 and OR = 2 both indicate that one group has twice the odds of another.
- This means that the log OR is on an additive scale of odds (This is important for logistic regression!).
- OR is not a ratio of probabilities.

Unadjusted association btw lung cancer and sex

```
library(epitools)
birds$LC <- relevel(birds$LC, ref="NoCancer")</pre>
(tmp <- with(birds, table(FM, LC)))</pre>
         T.C
##
## FM NoCancer LungCancer
## Female
                24
            74
                          37
## Male
oddsratio(tmp)$measure
##
          odds ratio with 95% C.I.
## FM estimate lower upper
  Female 1.0000000
                          NA
##
                                   MΑ
    Male 0.9954866 0.4516538 2.280673
##
```

Do men have different odds of lung cancer compared to women, without adjustment for possible confounders?

Unadjusted association btw lung cancer and birdkeeping

```
birds$BK <- relevel(birds$BK, ref="NoBird")</pre>
(tmp <- with(birds, table(BK, LC)))</pre>
##
           LC
            NoCancer LungCancer
##
    NoBird
                  64
                              16
                 34
                             33
##
    Bird
oddsratio(tmp)$measure
           odds ratio with 95% C.I.
##
## BK
          estimate lower
                             upper
##
     NoBird 1.000000
                          MΑ
                                    MΑ
     Bird 3.827991 1.86773 8.124253
##
```

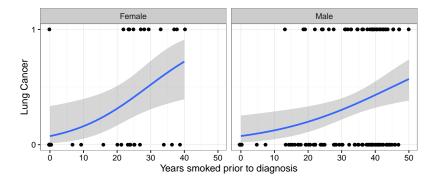
Do birdkeepers have different odds of lung cancer compared to non-birdkeepers, without adjustment for possible confounders?

Lung cancer as a function of years smoked

Modeling the log-odds is one solution to the problem of linearity.

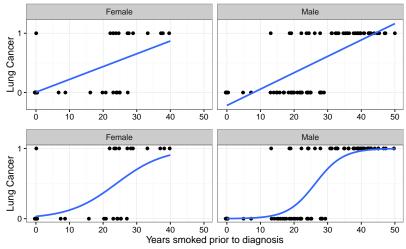
$$\log odds(LC) \sim FM + YR$$

```
(p <- ggplot(birds, aes(x=YR, y=LCnum)) +
   geom_jitter(height=0) + facet_grid(.~FM) +
   stat_smooth(method='glm', method.args=list(family='binomial')) +
   ylab("Lung Cancer") + xlab("Years smoked prior to diagnosis") +
   scale_y_continuous(breaks=c(0,1)))</pre>
```



A more drastic example: why we use the log-odds

I dropped individuals who smoked >30 years prior to diagnosis who did not have LC.



Generalized linear models: defined

All generalized linear models have the following three characteristics:

- 1. A probability distribution describing the outcome variable
 - e.g. $Y \sim \mathsf{Bernoulli}(p) \longrightarrow \mathbb{E}[Y|p] = p$.
- 2. A linear model
- 3. **A link function** that relates the linear model to the parameter of the outcome distribution
 - $g(\mathbb{E}[Y]) = g(p) = \eta$ or $\mathbb{E}[Y] = p = g^{-1}(\eta)$

MLR is a special case of a GLM

For continuous outcome, we often do this

- 1. A probability distribution describing the outcome variable
 - $Y|X \sim \text{Normal}(\mu, \sigma^2) \longrightarrow \mathbb{E}[Y|X] = \mu.$
- 2. A linear model
- A link function that relates the linear model to the parameter of the outcome distribution
 - $g(\mathbb{E}[Y|X]) = g(\mu) = \mu = \eta$

$$g(\mathbb{E}[Y|X]) = E[Y|X] = \mu = \beta_0 + \beta_1 X_1 + \dots + \beta_p X_p$$

Logistic regression: a common GLM for 0/1 outcome data

- 1. A probability distribution describing the outcome variable
 - $Y|X \sim \text{Bernoulli}(p) \longrightarrow \mathbb{E}[Y|X] = Pr(Y = 1|X) = p$.
- 2. A linear model
- A link function that relates the linear model to the parameter of the outcome distribution
 - $g(\mathbb{E}[Y|X]) = g(p) = logit(p) = log \frac{p}{1-p} = \eta$

$$g(\mathbb{E}[Y|X]) = logit[Pr(Y=1|X)] = \beta_0 + \beta_1 X_1 + \dots + \beta_p X_p$$

Logistic regression has log(odds) as the link

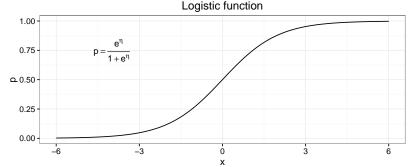
A logistic regression model can be defined as follows:

$$Y_i | \mathbf{x}_i \sim \mathsf{Bernoulli}(p_i)$$

$$\mathbb{E}[Y_i | \mathbf{x}_i] = Pr(Y_i = 1 | \mathbf{x}_i) = p_i$$

$$g(p_i) = logit(p_i) = \log \frac{p_i}{1 - p_i}$$

$$logit(\mathbb{E}[Y_i | \mathbf{x}_i]) = logit(p_i) = \eta = \beta_0 + \beta_1 X_{i1} + \dots + \beta_p X_{ip}$$



Example - Birdkeeping and Lung Cancer - Model

$$logitPr(LC = 1 | \mathbf{x}) = \beta_0 + \beta_1 BK + \beta_2 FM + \beta_3 SS + \beta_4 AG + \beta_5 YR + \beta_6 CD$$

Example - Birdkeeping and Lung Cancer - Interpretation

```
summary(lm1)$coef
##
                 Estimate Std. Error z value Pr(>|z|)
   (Intercept) -1.27063830 1.82530568 -0.6961236 0.4863514508
  BKBird
           1.36259456 0.41127585 3.3130916 0.0009227076
## FMMale
             -0.56127270 0.53116056 -1.0566912 0.2906525319
## SSLow
             -0.10544761 0.46884614 -0.2249088 0.8220502474
## AG
              -0.03975542 0.03548022 -1.1204952 0.2625027758
           0.07286848 0.02648741 2.7510612 0.0059402544
## YR.
               0.02601689 0.02552400 1.0193110 0.3080553359
## CD
```

Keeping all other predictors constant then,

- ► The odds ratio of getting lung cancer for bird keepers vs non-bird keepers is exp(1.3626) = 3.91.
- ► The odds ratio of getting lung cancer for an additional year of smoking is exp(0.0729) = 1.08.

What the numbers do not mean ...

The most common mistake made when interpreting logistic regression is to treat an odds ratio as a ratio of probabilities.

Bird keepers are *not* 4x more likely to develop lung cancer than non-bird keepers.

This is the difference between relative risk and an odds ratio.

$$RR = \frac{P(\mathsf{disease}|\mathsf{exposed})}{P(\mathsf{disease}|\mathsf{unexposed})}$$

$$OR = \frac{P(\mathsf{disease}|\mathsf{exposed})/[1-P(\mathsf{disease}|\mathsf{exposed})]}{P(\mathsf{disease}|\mathsf{unexposed})/[1-P(\mathsf{disease}|\mathsf{unexposed})]}$$

To match or not to match

Case-control studies are common for (rare) binary outcomes

- lacktriangle Randomly selected controls \longrightarrow vanilla logistic regression
- lacktriangle Matched controls \longrightarrow conditional logistic regression

Conditional logistic regression

- Accounts for the fact that you have "adjusted" for some variables in the design.
- Calculates an OR for each matched-set/pair, then "averages" across sets
- Forfeits ability to estimate effects of matched variables, but design can substantially improve power.
- Implemented in R with clogit().

Important notes about GLMs

On logistic regression in particular...

- There are other link functions for binary data (e.g. probit, cloglog).
- Other, less parametric methods may be appropriate here too –
 e.g. CART, random forests, classification algorithms.

Beyond the scope of this course, but interesting topics...

- How are logistic models (and other GLMs) fitted?
- Can we perform the same kind of model diagnostics to determine whether a model provides a good fit to data?
- ROC curves and classification rules