## Logistic regression: Probabilities and odds ratios

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#### Introduction

- An important distinction between linear and logistic regression is that the regression coefficients in logistic regression are not terribly meaningful on their own
- In linear regression, a coefficient  $\beta_j=1$  means that if you change  $x_j$  by 1, the expected value of Y will go up by 1 (very interpretable)
- In logistic regression, a coefficient  $\beta_j=1$  means that if you change  $x_j$  by 1, the log of the odds that Y occurs will go up by 1 (much less interpretable)

# Introduction (cont'd)

- When outcomes are categorical, it is much easier to think about the outcome in terms of probabilities and odds ratios
- These are the quantities that are usually reported and described in an analysis, rather than the regression coefficients themselves
- Today's lecture is about estimating, constructing confidence intervals, and carrying out hypothesis tests for these quantities

## Estimating probabilities

- We have already talked about estimation of probabilities based on the fit of a logistic regression model:
  - (1) Given a vector of explanatory variables  $\mathbf{x}$ , calculate the linear predictor  $\hat{\eta} = \mathbf{x}^T \hat{\boldsymbol{\beta}}$
  - (2) Estimate the probability based on

$$\hat{\pi} = \frac{e^{\hat{\eta}}}{1 + e^{\hat{\eta}}}$$

• It should be noted that, since maximum likelihood estimates are invariant to transformation,  $\hat{\pi}$  may also be considered the maximum likelihood estimate of  $\pi$ 

## Confidence intervals for probabilities

- Construction of confidence intervals proceeds similarly
- Using the fact that

$$\frac{\mathbf{x}^T \widehat{\boldsymbol{\beta}} - \mathbf{x}^T \boldsymbol{\beta}}{\widehat{\mathrm{SE}}} \stackrel{.}{\sim} z,$$

where 
$$\widehat{\mathrm{SE}} = \sqrt{\mathbf{x}^T (\mathbf{X}^T \mathbf{W} \mathbf{X})^{-1} \mathbf{x}}$$

• We can then construct a confidence interval for  $\eta$ :

$$(L, U) = (\hat{\eta} - z_{\alpha/2}\widehat{SE}, \hat{\eta} + z_{\alpha/2}\widehat{SE})$$

• A  $(1-\alpha)$  confidence interval for  $\pi$  is therefore

$$\left(\frac{e^L}{1+e^L}, \frac{e^U}{1+e^U}\right)$$

## Hypothesis testing

• In principle, one could carry out hypothesis tests of  $H_0: \pi = \pi_0$  based on this approach as well: calculate

$$\eta_0 = \log\left(\frac{\pi_0}{1 - \pi_0}\right)$$

and then test  $H_0: \eta = \eta_0$  based on the fact that, under  $H_0$ ,

$$\frac{\hat{\eta} - \eta_0}{\widehat{SE}} \stackrel{.}{\sim} z$$

• In practice, however, it is quite rare to have a hypothesis about a specific type of subject that you are interested in testing

## Limitations of estimating probabilities

- Probabilities have the advantage that they are particularly easy to interpret (everyone knows what a probability is)
- However, working with probabilities is inconvenient for logistic regression in many ways
- First of all, the model is not linear in probability
- Suppose  $\beta_j=1$  and  $x_j$  changes by 1; how does that affect the probability?
- Well, if we started out at 50%, it goes up to 73%; but if we start at 90%, it only goes up to 96%

## Limitations of estimating probabilities (cont'd)

- Another way of putting this is that in order to calculate the change in probability as one explanatory variable changes, you have to specify all the explanatory variables
- This complicates (when lots of variables are present, greatly complicates) the most attractive feature of an additive model: the ability to describe what happens when you change one thing and leave the rest the same

## Logistic regression and case-control studies

- This has particularly important consequences for case-control studies
- Theorem: In a case-control study, the maximum likelihood estimates  $\widehat{eta}_1,\ldots,\widehat{eta}_{p-1}$  as well as their approximate sampling distributions are equivalent to that obtained from the logistic regression model, under the assumptions that
  - (a) The model is correct
  - (b) The selection of cases and controls is independent of the explanatory variables

### Case-control assumptions

- The assumption that selection of cases and controls is independent of the explanatory variables is actually a pretty big assumption, often violated in actual case-control studies
- This is a major source of bias
- For example, case-control studies have found links between childhood leukemia and exposure to electromagnetic fields (EMF)

### Electromagnetic field example

- However, subsequent investigations have indicated that this is due entirely to case-control bias
- Families with low socioeconomic status are more likely to live near electromagnetic fields
- Families with low socioeconomic status are also less likely to participate in studies as controls
- Socioeconomic status does not affect the participation of cases, however (cases are usually eager to participate)
- This results in a spurious association between EMF and leukemia

#### The intercept

- Another important thing to note from the preceding theorem is that it makes no claim about  $\hat{\beta}_0$
- This is because the case-control sampling affects the likelihood with respect to  $\beta_0$ , as the sampling probabilities are absorbed into the intercept:

$$\beta_0^{\text{Case control}} = \beta_0^{\text{General population}} + \log \left( \frac{\tau_1}{\tau_0} \right),$$

where  $\tau_1$  and  $\tau_0$  are the selection probabilities for cases and controls, respectively

- In the usual situation where we intentionally oversample cases, this leads to overestimation of  $\beta_0$
- Thus, it is not possible to estimate the  $\beta_0$  that describes the general population (unless we know  $\tau_1$  and  $\tau_0$ ), and therefore not possible to estimate probabilities either

#### Odds ratios: introduction

- Given that the regression coefficients are difficult to interpret, but that estimation of probabilities has a number of drawbacks, how should we summarize and report our model?
- It turns out that odds ratios provide a readily interpretable and easily estimated compromise, without any of the drawbacks of estimating probabilities

#### Estimation of odds ratios

ullet Note that the odds of the event occurring for a subject with vector of explanatory variables  ${f x}$  is

$$\frac{\pi}{1-\pi} = \exp(\mathbf{x}^T \boldsymbol{\beta})$$

• Thus, the odds ratio for comparing two subjects, one with explanatory variables  $x_1$  and the other with  $x_2$ , is

$$\frac{\pi_2/(1-\pi_2)}{\pi_1/(1-\pi_1)} = \frac{\exp(\mathbf{x}_2^T \boldsymbol{\beta})}{\exp(\mathbf{x}_1^T \boldsymbol{\beta})}$$
$$= \exp\{(\mathbf{x}_2 - \mathbf{x}_1)^T \boldsymbol{\beta}\})$$

# Estimation of odds ratios (cont'd)

• In particular, consider the odds ratio for what happens when  $x_j$  changes by an amount  $\delta_j$ , while the rest of the explanatory variables remain the same:

$$OR = \exp(\delta_j \beta_j)$$

- This is exactly what we need: all the other variables vanish and our estimate depends only on the  $\beta_j$  and the change in  $x_j$
- We can therefore estimate this odds ratio, as well as carry out inference, based entirely on  $\hat{\beta}_j$
- In particular, we don't even need to be able to estimate  $\beta_0$ , so we can apply these results to case-control studies

### Estimation of odds ratios: example

 For example, what is the odds ratio for a member of the Donner party failing to survive the winter with a 10-year increase in age?

$$OR = \exp(10 \cdot 0.0325) = 1.4$$
 (Male)  
 $OR = \exp(10 \cdot 0.1941) = 7.0$  (Female)

 Note that these odds ratios apply to any 10-year difference (50 vs. 40, 30 vs. 20, etc.) due to the linearity assumption

#### Confidence intervals for odds ratios

- As with probabilities, confidence intervals for odds ratios can be obtained by first obtaining confidence intervals in terms of  $\hat{\beta}$  and then transforming
- The confidence intervals for the slope of the effect of age on the linear predictors are

$$(-0.0366, 0.1016)$$
 (Male)  $(0.0227, 0.3654)$  (Female)

• The confidence intervals for the odds ratios of dying (again, with a change of 10 years in age) are:

$$e^{10\cdot(L,U)}=(0.7,2.8)$$
 (Male) 
$$e^{10\cdot(L,U)}=(1.3,38.6)$$
 (Female)

### Hypothesis tests

- Hypothesis tests of odds ratios (and for that matter, probabilities) are directly equivalent to tests of regression coefficients, since all the following are equivalent:
  - $\beta_i = 0$
  - Odds ratio = 1
  - Difference in probabilities = 0
  - Ratio of probabilities (relative risk) = 1

## Hypothesis tests: Example

ullet For example, the test of  $H_0:eta_{\mathtt{Age}|\mathtt{Male}}=0$  yields

$$z = \frac{-0.03248}{0.03527} = -0.921,$$

and 
$$p = 2\Phi^{-1}(-0.921) = 0.36$$

- Meanwhile, the test of  $H_0: \beta_{\texttt{Age}|\texttt{Female}} = 0$  yields p = .03
- The tests are relatively low-powered because there are only 45 subjects; hence the relatively high p-values despite the large estimated effects
- This phenomenon was apparent from the wide confidence intervals as well

#### Confidence intervals: Wald vs. likelihood ratio

- This is not the only way (indeed, it isn't even the best way) to approach inference for GLMs
- The approach we have discussed today is known as the Wald approach, after the statistician Abraham Wald
- This approach, however, is based on an approximation to the likelihood at  $\widehat{\beta}$ ; it does not necessarily provide accurate information about values of  $\beta$  far from  $\widehat{\beta}$
- Next time, we will discuss an alternative approach to testing and constructing confidence intervals based on likelihood ratios and see that it suffers less from the approximation problem