## Last Class!

Revisit interactions + Intro to Poisson Regression

#### Announcements

Do you all get requests for the formal course evaluations?

- Remaining grading
  - Homework 4, 5, and 6 redos
  - Project Analysis, Report, and Presentation
- Anything I'm missing?

#### Class 17 Learning Objectives

Not as formal today:

1. Calculate odds ratios for variables (categorical x continuous) involved in interactions

 If we want, we can look at categorical x categorical and continuous x continuous

2. Run and interpret a simple Poisson Regression

#### Odds Ratio in the Presence of Interaction (I)

 When interaction exists between a risk factor (F) and another variable (X), the estimate of the odds ratio for F depends on the value of X

- When an interaction term (F\*X) exists in the model
  - $OR_F \neq exp(\beta_F)$  in general

• Assume we want to compute the odds ratio for  $(F = f_1)$  and  $F = f_0$ , the correct model-based estimate is

$$\widehat{OR}_F = \exp(\widehat{g}(F = f_1, x) - \widehat{g}(F = f_0, x))$$

#### Odds Ratio in the Presence of Interaction (II)

We may write the two logits for given x as below:

$$\hat{g}(f_1, x) = \hat{\beta}_0 + \hat{\beta}_1 f_1 + \hat{\beta}_2 x + \hat{\beta}_3 f_1 \times x$$

$$\hat{g}(f_0, x) = \hat{\beta}_0 + \hat{\beta}_1 f_0 + \hat{\beta}_2 x + \hat{\beta}_3 f_0 \times x$$

The difference in two logits is:

$$\hat{g}(f_1, x) - \hat{g}(f_0, x) = \hat{\beta}_1(f_1 - f_0) + \hat{\beta}_3 x(f_1 - f_0)$$

Therefore,

$$\widehat{OR}(F = f_1, F = f_0, X = x) = exp[\widehat{\beta}_1(f_1 - f_0) + \widehat{\beta}_3 x(f_1 - f_0)]$$

#### Odds Ratio in the Presence of Interaction (III)

To find the confidence interval for the  $\widehat{OR}$ , we need to find confidence interval for  $\hat{g}(f_1, x) - \hat{g}(f_0, x)$  first

$$Var(\hat{g}(f_1, x) - \hat{g}(f_0, x))$$

$$= (f_1 - f_0)^2 \times Var(\hat{\beta}_1) + [x(f_1 - f_0)]^2 \times var(\hat{\beta}_3) + 2x(f_1 - f_0)^2 \times cov(\hat{\beta}_1, \hat{\beta}_3)$$

 We can follow the same procedure as we did previously to construct 95% CI for  $\hat{g}(f_1, x) - \hat{g}(f_0, x)$ , denoted by  $[L_{a01}, U_{a01}]$ .

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$$\widehat{OR} = \exp(\widehat{g}(f_1, x) - \widehat{g}(f_0, x))$$

 $\widehat{OR} = \exp(\widehat{g}(f_1, x) - \widehat{g}(f_0, x))$ The 95% CI for  $\widehat{OR}$  is:  $\left[\exp(L_{g01}), \exp(U_{g01})\right]$ 

#### Example: GLOW – Computing OR under interaction

Step 1: The two sets of values of the covariates are (priorfrac = 1, Age = a) compared to (priorfrac = 0, Age = a)

**Step 2:** Substituting these values into the general expression

**Step 3:** Taking the difference in the two functions

**Step 4:** Exponentiating the result

**Step 1:** The two sets of values of the covariates are (priorfrac = 1, Age = a) compared to (priorfrac = 0, Age = a)

Substituting these values into the general expression Step 2:

$$logit(\pi(\mathbf{x})) = \hat{\beta}_0 + \hat{\beta}_1 PF + \hat{\beta}_2 Age + \hat{\beta}_3 PF \times Age$$

logit(
$$\pi(PF = 1, Age = a)$$
) =  $\hat{\beta}_0 + \hat{\beta}_1 \times 1 + \hat{\beta}_2 a + \hat{\beta}_3 \times 1 \times a$   
=  $\hat{\beta}_0 + \hat{\beta}_1 + \hat{\beta}_2 a + \hat{\beta}_3 a$ 

logit(
$$\pi(PF = 0, Age = a)$$
) =  $\hat{\beta}_0 + \hat{\beta}_1 \times 0 + \hat{\beta}_2 a + \hat{\beta}_3 \times 0 \times a$   
=  $\hat{\beta}_0 + \hat{\beta}_2 a$ 

Taking the difference in the two functions Step 3:

$$[logit(\pi(PF = 1, Age = a))] - [logit(\pi(PF = 0, Age = a))]$$
$$= [\hat{\beta}_0 + \hat{\beta}_1 + \hat{\beta}_2 a + \hat{\beta}_3 a] - [\hat{\beta}_0 + \hat{\beta}_2 a]$$
$$= \hat{\beta}_1 + \hat{\beta}_3 a$$

Exponentiating the Step 4: result

Model	Variable	Coeff.	Std. Err.	z	p	95% C	CI
3	$\widehat{eta}_1$ PRIORFRAC	4.961	1.8102	2.74	0.006	1.413,	8.509
	$\hat{\beta}_2$ AGE	0.063	0.0155	4.04	< 0.001	0.032,	0.093
	$\hat{\beta}_3$ PRIORFRAC × AGE	-0.057	0.0250	-2.29	0.022	-0.106, -0.106	-0.008
	$\hat{\beta}_0$ Constant	-5.689	1.0841	-5.25	< 0.001	-7.814, -	-3.565

$$\widehat{OR}[(PF = 1, Age = a), (PF = 0, Age = a)] = \exp(\widehat{\beta}_1 + \widehat{\beta}_3 a)$$
  
=  $\exp(4.961 - 0.057a)$ 

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If we let a = 60, i.e., compute OR for age = 60, then

• 
$$\widehat{OR}_{a=60} = \exp(4.961 - 0.057 * 60) = 4.669$$

If we let a = 70, i.e., compute OR for age = 70, then

• 
$$\widehat{OR}_{a=70} = \exp(4.961 - 0.057 * 70) = 2.64$$

#### Example: GLOW – Table of ORs

Sometimes a Table listing the estimated odds ratio and 95%
 CI at different value of X will also be helpful and informative

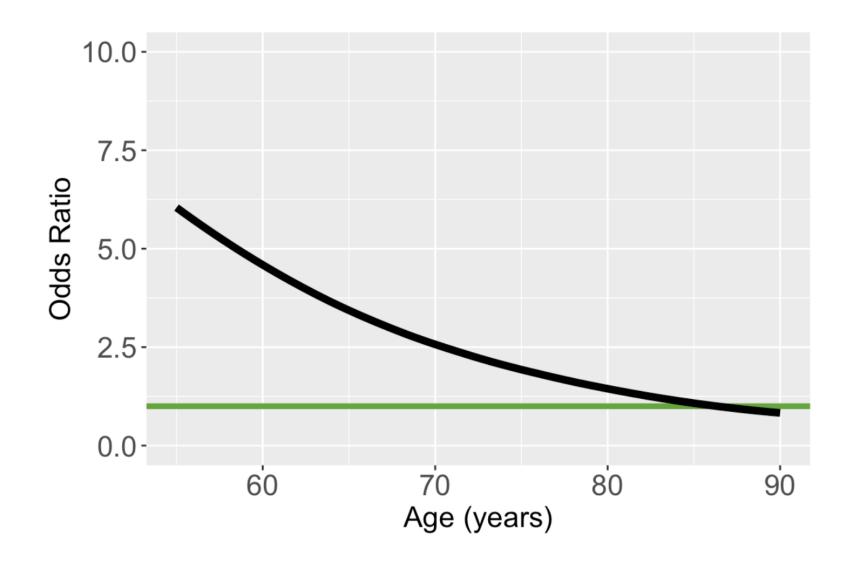
Table 3.13 Estimated Odds Ratios for Prior Fracture as a Function of Age from Model 3 in Table 3.12

Age	Odds Ratio	95% CI
55	6.1	2.38, 15.53
60	4.6	2.20, 9.49
65	3.4	1.96, 5.99
70	2.6	1.63, 4.06
75	1.9	1.20, 3.11
80	1.4	0.79, 2.65

#### Plot of Odds Ratio Involving Interaction Term

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 Odds ratio comparing prior fracture to no prior fracture



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#### Graphically Showing Interaction Revisited

• One easy way to see the nature of the interaction between F and X is to plot

the two logit functions

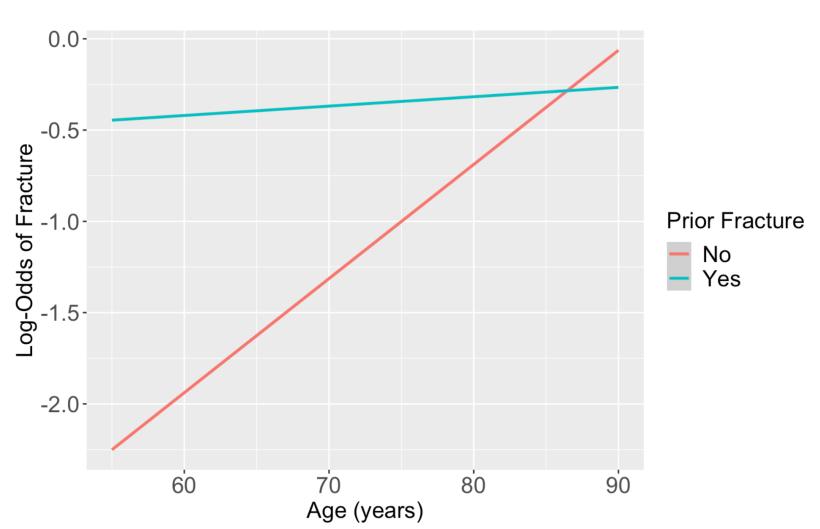
Using the GLOW fracture example:

$$\log it(\pi(PF = 1, Age = a))$$

$$= \hat{\beta}_0 + \hat{\beta}_1 + \hat{\beta}_2 a + \hat{\beta}_3 a$$

$$= (\hat{\beta}_0 + \hat{\beta}_1) + (\hat{\beta}_2 + \hat{\beta}_3)a$$

logit
$$(\pi(PF = 0, Age = a))$$
  
=  $\hat{\beta}_0 + \hat{\beta}_2 a$ 



# Simple Poisson Regression

#### Poisson Distribution (I)

This distribution is often used to model count data

- Examples:
  - Distribution of number of deaths due to lung cancer
  - Distribution of number of individuals diagnosed with leukemia
  - Distribution of number of hospitalizations

#### Poisson Distribution (II)

• The probability function of Poisson distribution:

$$P(Y = y | \mu) = \frac{e^{-\mu} \mu^y}{y!}$$

- Where y are non-negative integers y = 0, 1, 2, ...
- Where  $\mu$  is the mean of Y, that is  $E(Y) = \mu$
- And also,  $Var(Y) = \mu$
- For a Poisson distribution,  $Y \sim Poisson(\mu)$ 
  - Range:  $[0, \infty)$

#### Poisson Distribution (III)

• If we look at the probability of *y* events <u>in a time period *t*</u> for a Poisson random variable, we could write:

$$P(Y = y | \mu) = \frac{e^{-\mu} \mu^y}{y!}$$

- Where y are non-negative integers y = 0, 1, 2, ...
- And  $\mu = \lambda t$ , where  $\lambda$  is the expected number of events per unit time
- Then  $\mu$  is the expected number of events over time t

#### Poisson Distribution (IV)

- What does  $\lambda$  represent here?
  - A rate, the expected number of events in a given population over a given period time
    - Example: Number of patient arrivals into the Emergency Room *per hour*
  - The Poisson distribution is the prototype for assigning probabilities of observing any number of events

# Poll Everywhere Question 1

#### Why Person-Years? (I)

- In the example of number of patient arrivals, an event does not conclude the study
  - If someone arrives within the first minute of the study, then we keep counting
  - We may be able to study the association of arrivals with qualities of the hospital, but we can't measure qualities of the individuals arriving
- What happens if we want to measure qualities of the individual?
  - We can measure a hospitalization rate

#### Why Person-Years? (II)

- If we are measuring at the individual level and counting something that is "terminal" then our count will always be 0 or 1
  - Example: Number of individuals diagnosed with leukemia
    - This only happens once, so how do we measure the rate here?
- Since rate involves the counts and time we can use the time to diagnosis to estimate the rate
  - Often expressed in units such as events per thousand person-years

#### Calculating Person-Year

- One person-year is a unit of time defined as one person being followed for one year
- Person-years for a sample of n subjects is calculated as the total years followed for the n subjects, where each subject could have different follow-up time
- Example: suppose we have 5 subjects, two of the subjects were followed for 2 years, and two of them are followed for 3 years and the fifth subject was followed for 3.8 years

```
person - years
= 2 people × 2 years + 2 people × 3 years
+ 1 person × 3.8 years = 13.8 person - years
```

### Calculating Rate (II)

 Suppose that we observe one event during the follow-up period, then

Rate of event = 
$$\frac{\text{\# events}}{\text{person - years}} = \frac{1 \text{ event}}{13.8 \text{ person - years}}$$
  
= 0.072 events per person - year  
= 72 events per 1000 person - years

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### Stop here?

#### If we do:

• Good tutorial in R: <a href="https://www.dataquest.io/blog/tutorial-poisson-regression-in-r/">https://www.dataquest.io/blog/tutorial-poisson-regression-in-r/</a>

#### Review: Simple Logistic Regression

- Let Y is the dependent variable of interest and x is a predictor variable,
  - In simple logistic regression, we have

$$\log\left(\frac{\pi(x_i)}{1-\pi(x_i)}\right) = \beta_0 + \beta_1 x_i$$

where 
$$\pi(x_i) = P(Y_i = 1|x_i)$$

#### Simple Poisson Regression Model

What do we model in a Poisson regression?

- Log of conditional mean of Y given x
  - Let Y be a Poisson count for a given unit of time, then  $\mu(x) = \lambda(x)$
  - In a simple Poisson regression, we have

$$\ln(\mu(x_i)) = \ln(\lambda(x_i)) = \beta_0 + \beta_1 x_i$$

So this is also called a log-linear model

#### Parameter Interpretation (I)

• In simple Poisson regression:

$$\ln(\mu(x_i)) = \ln(\lambda(x_i)) = \beta_0 + \beta_1 x_i$$

- When x is a binary variable: How do we interpret  $\beta_0$  and  $\beta_1$ ?
  - When  $x_i = 0$ :

$$\ln(\mu(x_i=0)) = \ln(\lambda(x_i=0)) = \beta_0$$

- $\mu(x_i = 0) = \exp(\beta_0)$ : the mean (rate) of  $Y_i$  when  $x_i = 0$
- When  $x_i = 1$ :

$$\ln(\mu(x_i=1)) = \ln(\lambda(x_i=1)) = \beta_0 + \beta_1$$

•  $\mu(x_i = 1) = \exp(\beta_0 + \beta_1)$ : the mean (rate) of  $Y_i$  when  $x_i = 1$ 

#### Parameter Interpretation (II)

- When x is a binary variable: How do we interpret  $\beta_0$  and  $\beta_1$ ?
  - By subtraction, we have

$$\beta_1 = \ln\left(\frac{E(Y_i|x_i=1)}{E(Y_i|x_i=0)}\right) = \ln\left(\frac{\lambda(x_i=1)}{\lambda(x_i=0)}\right)$$

- $\beta_1$ : Log-rate ratio
- And  $\exp(\beta_1)$  is the rate ratio

#### Further reading / tutorials on Poisson regression

- Good tutorial in R: <a href="https://www.dataquest.io/blog/tutorial-poisson-regression-in-r/">https://www.dataquest.io/blog/tutorial-poisson-regression-in-r/</a>
- When people are followed for different amounts of time, we should include an offset
  - Poisson Regression Modeling Using Rate Data: section from above site that discusses offsets
- We can use Wald test and LRT in the same way as logistic regression to test our coefficients and variables

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#### Wrap-up

• 4-minute exit ticket

Thanks for a great quarter!

#### **Class 17 Exit Ticket**



https://forms.office.c om/r/5q7YxDi58s

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