UC Irvine ISI-BUDS Day 12

Zhaoxia Yu

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Study Goals

Review of LM

Regression
Poisson

Regression

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Review of LM

Logistic Regression

Poisson Regression

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Review of LM

- ► GI M
 - Logistic Regression
 - Poisson Regression
 - Multinomial Regression

► The assumption of independent observations

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Regression
The Assumpti

The Assumption of Independence

Suppose

$$Y = \beta_0 + x_1 \times \beta_1 + \ldots + x_p \times \beta_p + \epsilon,$$

where

- the regressand Y is the response / outcome / dependent / endogenous variable
- ▶ the regressors (x_1, \dots, x_p) are the p covariates / independent / explanatory variables
- the random term ϵ has a zero mean and variance $\sigma^2 > 0$
- the intercept is β_0 , the other p coefficients are β_1, \dots, β_p

Logistic Regression

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Consider the *i*th observation:

$$Y_i = \beta_0 + x_{i1} \times \beta_1 + \ldots + x_{ip} \times \beta_p + \epsilon_i, i = 1, \ldots, n$$

- Basic assumptions
 - ► $E(\epsilon_i) = 0$, which is equivalent to $E(Y_i|X_i) = \beta_0 + x_{i1} \times \beta_1 + \dots + x_{in} \times \beta_n$
 - Var $(\epsilon_i) = \sigma^2$. Note, this is equivalent to say $Var(Y_i|X_i) = \sigma^2$.
 - $(\epsilon_1, \dots, \epsilon_n)$ are mutually independent
- ▶ If $(\epsilon_1, \dots, \epsilon_n)$ are i.i.d. $N(0, \sigma^2)$, we can derive t-tests and F-tests
- ▶ Question: what if the assumptions are violated?

Logistic Regression

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► A motivating example: Consider a binary response variable, i.e., *Y_i* takes values of 0 or 1.

▶ Is LM a good choice for this problem?

Consider the Alzheimer dataWe create a binary variable

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Logistic Regression

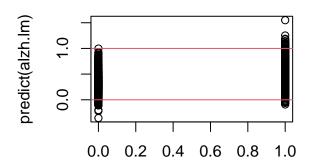
Regression

```
alzheimer=read.csv("alzheimer_data.csv", header = TRUE) pendence
#dim(alzheimer)
#names(alzheimer)
attach(alzheimer)
#length(unique(id))
alzh=(diagnosis>0)*1 #"*1" to create a 0-1 variable
```

A Motivating Example of GLM (continued)

```
alzh.lm = lm(alzh ~ age + female + educ+lhippo + rhippo)als
par(mar = c(4, 4, 0.5, 0.5))
                                                        Review of LM
plot(alzh, predict(alzh.lm)); abline(h=c(0,1), col=2); istic
```

alzh



Regression The Assumption

of Independence

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A Motivating Example of GLM (continued)

- Is alzh.lm a good model for alzh?
- Several assumptions of the LM have been violated, and
- ▶ The predicted values using LM are not between 0 and 1!
- Let $X_i = (x_{i1}, \dots, x_{ip})^T$, i.e., the vector of covariates for the ith subject.
- Let $\pi_i = E(Y_i|X_i)$, the expected probability. We would like to make sure that $\pi_i \in [0,1]$
- How?

Logistic Regression

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▶ Consider the a special transformation of π_i :

$$logit(\pi_i) = log \frac{\pi_i}{1 - \pi_i} \in (-\infty, \infty)$$

- ► This is the so-called "logit" link!
- $\pi_i = E[Y_i|X_i]$: probability of having AD for a subject with covariates X_i .
- $ightharpoonup \frac{\pi_i}{1-\pi_i}$: odds

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We connect π_i and a linear function of the covariates X_i by assuming

$$\log \frac{\pi_i}{1 - \pi_i} = \beta_0 + x_{i1} \times \beta_1 + \ldots + x_{ip} \times \beta_p$$

- Essentially, we model the log-odds.
- ightharpoonup But Y_i is a random variable. We need a distribution. A natural choice is the Bernoulli distribution

$$Y_i|X_i \sim Bernoulli(\pi_i)$$

pmf, mean, variance:

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- Estimation of is typically conducted by maximizing the corresponding likelihood function
- ► How to obtain the likelihood function

$$E(Y_i|X_i) = \pi_i = \frac{\exp\{\beta_0 + x_{i1} \times \beta_1 + \dots + x_{ip} \times \beta_p\}}{1 + \exp\{\beta_0 + x_{i1} \times \beta_1 + \dots + x_{ip} \times \beta_p\}}$$

$$f(Y_i|X_i) = \pi_i^{Y_i} (1-\pi_i)^{1-Y_i}$$
, i.e.,

$$f(Y_i|X_i) = \pi_i \text{ if } Y_i = 1$$

•
$$f(Y_i|X_i) = 1 - \pi_i \text{ if } Y_i = 0$$

▶ independence:
$$f(Y|X) = \prod_{i=1}^n f(Y_i|X_i)$$

$$L(\beta_0, \beta_1, \cdots, \beta_p) = f(Y|X)$$

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- How to obtain the maximum likelihood estimates (MLE) of the parameters $(\beta_0, \dots, \beta_p)$?
 - ► Iteratively re-weighted least squares (IRLS): the default method used by R
 - ► The Newton-Raphson algorithm

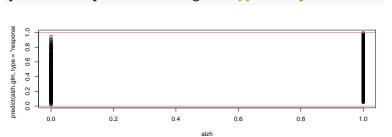
the Motivating Example of Logistic Regression

par(mar = c(4, 4, 0.5, 0.5))

Review of LM amily=bino alzh.glm = glm(alzh ~ age + female + educ+lhippo,



of Independence



#More visualizations

#https://blogs.uoregon.edu/rclub/2016/04/05/plotting-your-log

Interpreting a logistic regression

summary(alzh.glm)\$coefficients[-1,]

Review of LM

Regression

The Assumption

```
Pr ( Sludependence
```

```
z value
##
             Estimate Std. Error
## age
          0.01813761 0.004246088
                                    4.271605 1.940715e-05
## female -1.32020475 0.096534651 -13.675968 1.413151e-42
## educ
          -0.05640342 0.013279326
                                   -4.247461 2.162067e-05
## lhippo
          -1.98502114 0.114028821 -17.408065 7.166544e-68
```

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- ➤ Consider the age variable. The estimated coefficient is 0.018138. What information does it provide?
- ► The estimated log-odds AD for subject *i* is (or add a constant determined by study design, see Day 11 lecture)

$$loigt(\hat{\pi}_i) = \hat{\beta}_0 + \hat{\beta}_{age} age_i + \hat{\beta}_2 female_i + \hat{\beta}_3 educ_i + \hat{\beta}_4 lhippo_i$$

Let $\tilde{\pi}_i$ denote estimated log-odds after one year

$$loigt(\tilde{\pi}_i) = \hat{\beta}_0 + \hat{\beta}_{age}(age_i + 1) + \hat{\beta}_2 female_i + \hat{\beta}_3 educ_i + \hat{\beta}_4 lhippo_i$$

The estimated change in log-odds

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 $logit(\tilde{\pi}_i) - logit(\hat{\pi}_i) = log \frac{\tilde{\pi}_i}{1 - \tilde{\pi}_i} - log \frac{\hat{\pi}_i}{1 - \hat{\pi}_i} = 0.018138$

- The odds of AD in one year later is exp(0.018138) = 1.018303 times of the current odds.
- The estimated increase in odds of AD in a year is $e^{0.018138} 1 = 1.8303\%$
- ► A 95% confidence interval
 - First, obtain a 95% C.I. for the difference in log-odds: (0.018138-1.96*0.004246, 0.018138+1.96*0.004246) = (0.00982, 0.0265)
 - Then, we transform them to increase in odds: $(e^{0.00982} 1, e^{0.0265} 1) = (0.99\%, 2.69\%)$

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- ► What if we are interested in the increase in odds of AD in ten years (everything else is fixed)?
- ▶ The estimated increase in odds of AD in 10 years is

$$e^{10*0.018138} - 1 = 19.89\%$$

► A 95% C.I. for 10-year increase in odds:

Regression

The Assumption of Independence

 $\exp(10*c(0.018138-1.96*0.004246, 0.018138+1.96*0.004246))-1$

[1] 0.1031375 0.3029118

i.e., (10.3%, 30.3%)

```
Very often, we also want to know the significance of a
variable after adjusting for other important covariates?
```

- Does age show a significant effect after adjusting for gender, education, and)hippocampus volume?
- A test for H_0 : $\beta_{age} = 0$ using the Wald test (a type of large-sample test)

```
summary(alzh.glm)$coefficients["age",]
```

```
## Estimate Std. Error z value Pr(>|z|)
## 1.813761e-02 4.246088e-03 4.271605e+00 1.940715e-05
```

 Other tests, such as likelihood ratio test, can also be used

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The Assumption of Independence

 Recall that we used the <u>logit</u> link in the logistic regression

$$g(\pi_i) = logit(\pi_i) = \frac{\pi_i}{1 - \pi_i},$$

where $\pi_i = E(Y_i|X_i)$.

- ► How about LM? $g(\mu_i) = \mu_i$, i.e., LM uses the identity link
- ▶ Poisson $g(\lambda_i) = log(\lambda_i)$. $Y_i | X_i \sim Poisson(\lambda_i)$.

Poisson Regression

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- Poisson regression is often used to model count data
- Why are count data special?
 - Count data are non-negative
 - Count data take integer values
- Count data often violate the assumption of "constant variance"
 - Count data often follow a Poisson distribution
 - ► Consider $K \sim Poisson(\lambda)$. E(K) = ?, Var(K) = , pmf?

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- Neurons may <u>fire</u> selectively for particular types of stimuli
- ➤ To understand whether a neuron is a visual-selective neuron, 20 trials were run for each of the five image categories:
 - ▶ animal, fruit, kids, military, space
- ▶ In each trial, the number of spikes (the number of times that the neuron fired) within a 1-second window was recorded

library(tidyverse)

names(chosen_neuron_data)

```
Review of LM
```

```
#https://www.ics.uci.edu/~zhaoxia/Data/chosen_neuronedata.csuchosen_neuron_data <- read_csv(
```

"https://www.ics.uci.edu/~zhaoxia/Data/chosen_neuron_data.cchosen_neuron_data <- chosen_neuron_data[, c(2:4)]

dim(chosen_neuron_data)

The Assumption of Independence

```
## [1] 100 3
```

table(image categ)

attach(chosen neuron data)

Space

20

20

```
image_categ
##
     Animal
                Fruit
                           Kids Military
         20
                    20
                              20
##
```

sapply(split(n_spikes, image_categ), mean)

Even split of image categories among trials

Animal ## Fruit Kids Military Space ## 0.05 3.60 0.15 0.25 0.05

Poisson Regression: Visualize the count data (by image category)

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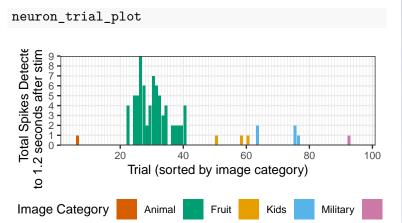
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Fit generalized linear model

```
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```
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```

```
data = chosen neuron data,
                    family = poisson(link="log"))
# Tabulate the coefficient estimates
poisson_neuron_table <- summary(poisson_fit)$coefficients</pre>
```

Poisson GLM with the default log link function poisson_fit <- glm(n_spikes ~ image_categ-1,</pre>

row.names(poisson_neuron_table)=c("Animal", "Fruit", "Kids", "Military", "Space"

Poission Regression: Model Summary

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poisson_neuron_table

```
Pr (> Zudependence
##
             Estimate Std. Error
                                    z value
   Animal
            -2.995732
                        0.9999998 -2.995733 2.737861e-03
## Fruit.
             1.280934
                        0.1178511 10.869084 1.618171e-27
## Kids
            -1.897120
                        0.5773503 -3.285908 1.016541e-03
  Military -1.386294
                        0.4472132 -3.099851 1.936181e-03
            -2.995732
                        0.9999998 -2.995733 2.737861e-03
## Space
```

Poisson Regression: Visualize Observed v.s. Fitted

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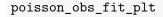
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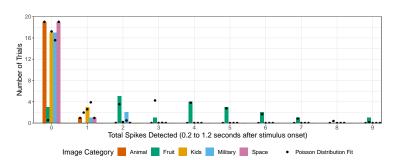
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The Assumption of Independence

- Next, we would like to discuss the significance of the image_categ variable. To do so, we first look at the deviance of a GLM object
- ► The deviance of a GLM object obj is

$$2[log(L_{saturated}) - log(L_{obj})]$$

What is the null hypothesis of no visual-selection?

$$H_0: \beta_{Animal} = \beta_{Fruit} = \beta_{Kids} = \beta_{Military} = \beta_{Space}$$

▶ What is the d.f. in a likelihood ratio test (LRT)?

Poission Regression: The Overall Significance

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Consider two <u>nested</u> models obj1 and obj2, the difference in their deviances is

$$2[log(L_{obj2}) - log(L_{obj1})],$$

which is the LRT statistic.

- What is the saturated model?
 - ▶ Logistic: $\pi_i = y_i$ and $L_{saturated} = 1$
 - Poisson: $\lambda_i = y_i$ and $L_{saturated} = \prod_i \frac{y_i^{y_i} e^{-y_i}}{y_i!}$.

Poission Regression: The Overall Significance

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```
# Test for visual selectivity: Likelihood Ratio Test
poisson_fit0 = glm(n_spikes ~1, data=chosen_neuron_data; fami
```

The Assumption

```
anova(poisson_fit0, poisson_fit, test = "LRT")
## Analysis of Deviance Table
##
## Model 1: n spikes ~ 1
## Model 2: n_spikes ~ image_categ - 1
##
     Resid. Df Resid. Dev Df Deviance Pr(>Chi)
## 1
            99
                  260.985
                            179.77 < 2.2e-16 ***
## 2
            95
                   81.213 4
##
                           0.001 '**' 0.01 '*' 0.05 '.' 0.1 '
## Signif. codes:
```

Poission Regression: The Overall **Significance**

Signif.

codes:

Test for visual selectivity: Rao's score test

anova(poisson_fit0, poisson_fit, test = "Rao")

Review of LM

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'*' 0.05 '.' 0.1

```
## Analysis of Deviance Table
##
## Model 1: n_spikes ~ 1
## Model 2: n_spikes ~ image_categ - 1
                                            Pr(>Chi)
##
    Resid. Df Resid. Dev Df Deviance
                                        Rao
## 1
           99
                 260.985
## 2
           95
                  81.213 4
                              179.77 236.3 < 2.2e-16 ***
##
```

Poission Regression: The Overall Significance

```
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```

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Laulatia

```
# Test for visual selectivity: Wald test. I will explain report poisson_fit_repara <- glm(n_spikes ~ image_categ, data = chosed family = poisson(link="log"))

Wald.stat=poisson_fit_repara$coefficients[2:5] %*%

solve (summary(poisson_fit_repara)$cov.unscaled[2:5,2:5]) %

poisson_fit_repara$coefficients[2:5]

1-pchisq(Wald.stat, df=4)
```

```
## [,1]
## [1,] 0
```

Poission Regression: Model Interpretation

R<gre2e+16

The Assumption of Independence

```
Estimate Std. Error z value \Pr_{\text{Logistic}}^{\text{Review of LM}}(>|z|)
##
                                           1.2809
                                                               0.1179
                                                                            10.869
```

image_categFruit

 $\hat{\beta}_{Fruit} = 1.2809$: What does it tell us?

Recall that we used the log link

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- Note that the model poisson_fit does not include β_0 .
- ► That's why we can estimate β_{Animal} , β_{Fruit} , β_{Kids} , $\beta_{Military}$, β_{space} .
- Question: how should we interprete the estimated coefficients if the intercept term was included?
 - Try poisson_fit_repara <- glm(n_spikes ~ image_categ, data = chosen_neuron_data, family = poisson(link="log"))</p>
 - ► Are the two models equivalent? (Lab activity)

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- ► Re-parameterization
- Parameters:
 - poisson_fit:
 - poisson_fit_repara:
- ► Compare the summary of the two models:
- ► Are they different models?

Parameterization 1: without intercept

```
summary(poisson_fit)
```

```
##
```

```
## Call:
## glm(formula = n_spikes ~ image_categ - 1, family = poisson
```

```
data = chosen neuron data)
```

##

##

```
## Deviance Residuals:
       Min
                  1Q
                        Median
                                      3Q
##
```





```
-2.996
```

```
0.00274 *
< 2e-16 *
```

Review of LM

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```
-2.9957
                                 1.0000
image_categAnimal
                      1.2809
                                 0.1179
                                          10.869
image_categFruit
```

Parameterization 2: with intercept

```
summary(poisson_fit_repara)

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```

Call:

glm(formula = n_spikes ~ image_categ, family = poisson(lin
data = chosen neuron data)

data = chosen_neuron_data)
##
Deviance Residuals:

Min 1Q Median 3Q ## -2.6833 -0.5876 -0.3162 -0.3162

##

Coefficients:
##

ents:

3Q Max

Max 2.3861

361

Estimate Std. Error z value Pr(>|z|) 2.996e+00 1.000e+00 -2.996 0.00274

Regression

The Assumption

(Intercept) -2.996e+00 1.000e+00 -2.996 0.00274 ## image_categFruit 4.277e+00 1.007e+00 4.247 2.16e-05

Regression

The Assumption

- ▶ The Poisson regression we fit provides estimates of β_{Animal} , β_{Fruit} , β_{Kids} , $\beta_{Military}$, β_{space} , which are the log of the Poisson rates
- What if we are interested in difference between specific groups? e.g.,
 - $\begin{array}{c} \beta_{\textit{Fruit}} \beta_{\textit{Animal}} \\ \beta_{\textit{Fruit}} + \beta_{\textit{Animal}} + \beta_{\textit{Kids}} + \beta_{\textit{Military}} + \beta_{\textit{Space}} \end{array}$
 - $\triangleright \beta_{Fruit} \frac{\beta_{Animal} + \beta_{Kids} + \beta_{Military} + \beta_{Space}}{4}$
- They are linear functions of the coefficients, i.e., in the form of $a^T \beta$, where a is a 5-by-1 vector.

Logistic Regression

Regressi

- ► LM/GLM provides not only estimated coefficients but also the variance-covariance of the estimated covariates
 - $\blacktriangleright \text{ Let } \hat{\beta} = c(\hat{\beta}_1, \cdots, \hat{\beta}_p)^T$
 - Let $\hat{\Sigma}$ denote the estimated variance-covariance of $\hat{\beta}$
 - Let a be linear coefficients

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Logistic Regression

Poisson Regressi

The Assumption of Independence

- ► Consider a linear function : $a^T \beta$
- \triangleright Estimate: $a^T \hat{\beta}$
- ► Variance of the estimate: $Var(a^T \hat{\beta}) = a^T \hat{\Sigma} a$
- ► Standard Error (SE): $s.e.(a^T\hat{\beta}) = \sqrt{a^T\hat{\Sigma}a}$
- ► A 95% confidence interval:

$$(a^{T}\hat{\beta} - 1.96 * s.e.(a^{T}\hat{\beta}), a^{T}\hat{\beta} + 1.96 * s.e.(a^{T}\hat{\beta}))$$

ightharpoonup Z-value: $\frac{a^T \hat{\beta} - ?????}{s.e.(a^T \hat{\beta})}$

Inference of Linear Functions of Parameters: Example

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Parameter of interest: $\frac{\beta_{Fruit} + \beta_{Animal} + \beta_{Kids} + \beta_{Military} + \beta_{Space}}{5}$

Study Goals
Review of LM

#extract estimated coefficients and their var-cov
poisson_fit\$coefficients

Logistic Regression

poisson_fit&coefficients

The Assumption

image_categAnimal ## -2.995732

image_categFruit 1.280934

0.0000000

image_categKid -1.89712

image_categSpace ## -2.995732

image categFruit

summary(poisson_fit)\$cov.unscaled #this ia a 5-by5 matrix

```
## image_categAnimal image_categFruit ima
## image categAnimal 0.9999996 0.00000000
```

0.00000000 0.01388889 a%*%poisson fit\$coefficients #estimate

```
Regression
```

The Assumption of Independence

```
[,1]
##
## [1,] -1.598789
```

a=matrix(rep(1/5,5), 1)

```
sqrt(a%*%summary(poisson_fit)$cov.unscaled%*%t(a))
```

```
[,1]
[1.] 0.3192003
```

##

Regression

Regressi

- Linear contrasts are a special family of linear functions
- We say $a^T \beta = \sum_i a_i \beta_i$ is a linear contrast if $\sum a_i = 0$, where $a = (a_1, \dots, a_p)^T$.
- Often, we are interested in whether a linear contrast is zero, i.e., $H_0: a^T \beta = 0$
- ightharpoonup z-value: $\frac{a^T \hat{\beta} 0}{s.e.(a^T \hat{\beta})}$

 $a \leftarrow matrix(c(-1, 1, 0, 0, 0), 1)$

Review of LM

Regression

The Assumption of Independence

```
fruit_animal_var = a%*%summary(poisson_fit)$cov.unscaled%*%t(
```

```
#2 110.1.11.e
print(fruit_animal_est/sqrt(fruit_animal_var))
```

fruit_animal_est = a%*%poisson_fit\$coefficients

```
[,1]
##
## [1,] 4.247274
```

#estimate

#nariance

Regression

The Assumption

```
Simultaneous Tests for General Linear Hypotheses
```

of Independence

Estimate Std. Error z value Pr(>|z|) 1.007 4.247 2.16e-05 ***

Fit: glm(formula = n spikes ~ image categ - 1, family = po data = chosen neuron data)

Linear Hypotheses:

4.277

 $a \leftarrow matrix(c(-1, 1, 0, 0, 0), 1)$

t <- glht(poisson_fit, linfct = a)

library(multcomp)

summary(t)

##

##

##

##

##

1 == 0

```
library(nnet)
```

multinom(diagnosis ~ age + female + educ + lhippo

age

2 671044 0 026772069 _1 227127 _0 04604660 _1

weights: 21 (12 variable)

initial value 2966.253179 ## iter 10 value 2372.326777

final value 2288.461323

converged

Call:

rhippo) ##

Coefficients:

##

(Intercept)

female

educ

multinom(formula = diagnosis ~ age + female + educ + lhipp

Regression

The Assumption of Independence

lhippo

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Other concerns

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Dispersion: under- or over-dispersion

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Zero-inflated Poisson Regression

. . .

► Model selection . . .

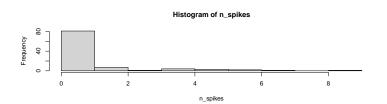
Regression

hist(n_spikes)

Poisson Regressior

Regression

The Assumption of Independence



#Interested inhow to fit a zero-inflated Poisson regression?

#https://www.rdocumentation.org/nackages/nscl/wersions/1.5.5/

The Assumption of Independence

UC Irvine ISI-BUDS Day 12

Zhaoxia \

Study Goals

Review of LM

Logistic Regression

Poisson Regression

Study Goals

Review of LM

Regression

Poisson Regression

The Assumption of Independence

► The common assumption we have made in LM and GLM is that the observations are independent with each other

- This is not always the case
- Examples:

Logistic Regression

Regression

- What is the consequence of ignoring data independence?
 - The damage is probably worse than violations of distributions
 - Fortunately, tools have been developed to account for data dependence
 - Day 13: Linear Mixed-Effects Model (LME)
 - Day 14: Generalized Linear Mixed-Effects Model (GLMM)