

10/17/2024 (Week 8)

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Outline



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Linear Regression

- Single variant regression
- Multivariate regression

Generalized Linear Regression

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• Logistic regression

Study relationship between two variables (X, Y)

where:

- Hypothesis testing : e.g., t-test
- Pearson's correlation coefficient r

$$\rho_{X,Y} = rac{\operatorname{cov}(X,Y)}{\sigma_X \sigma_Y}$$

- Unit free
- Dose not depend on number of samples

- $\bullet\ cov$ is the covariance
- σ_X is the standard deviation of X
- σ_Y is the standard deviation of Y

Pearson's Correlation Test

•
$$H_0: r = 0; \quad H_a: r \neq 0$$

$$r_{xy} = rac{\sum_{i=1}^n (x_i - ar{x})(y_i - ar{y})}{\sqrt{\sum_{i=1}^n (x_i - ar{x})^2} \sqrt{\sum_{i=1}^n (y_i - ar{y})^2}}$$

where:

- n is sample size
- x_i, y_i are the individual sample points indexed with *i*
- $ar{x} = rac{1}{n} \sum_{i=1}^n x_i$ (the sample mean); and analogously for $ar{y}$

Pearson's Correlation Test

- Under H₀: r = 0, with sample size n, the standard error of the correlation coefficient r is given by

$$\sigma_r = \frac{1 - r^2}{\sqrt{n - 2}}$$

• Under H_0 : Test statistic t follows a Student's t-distribution with degrees of freedom n-2

$$t = \frac{r}{\sigma_r} = r \sqrt{\frac{n-2}{1-r^2}}$$

Q1: Why is the Pearson's Correlation Test follows a Student's t-distribution with degrees of freedom of n-2?

Answer

Estimating the population standard deviation has N–1 degrees of freedom because the standard deviation requires an estimate of the mean, so the univariate t-test has N–1 df.

In contrast, in the case of the correlation coefficient, we have to estimate two population standard deviations (one for X and one for Y), plus the covariance.

To do this, we need estimates of both means, so the degrees of freedom for a correlation is N–2.

Q2: What is p-value?

A p-value is a statistical measure that helps you determine the significance of your results in hypothesis testing. Specifically, it represents the probability of obtaining results at least as extreme as those observed in your sample data, assuming that the null hypothesis is true.

Here's a breakdown:

\$

- Null Hypothesis (H0): This is the default assumption that there is no effect or no difference.
- Alternative Hypothesis (H1): This is what you want to prove, indicating that there is an effect or a difference.
- Low p-value (typically ≤ 0.05): Suggests that the observed data is unlikely under the null hypothesis, leading you to reject H0.
- High p-value (> 0.05): Indicates that the observed data is consistent with the null hypothesis, and you fail to reject H0.

It's important to remember that a p-value does not measure the size of an effect or the importance of a result; it merely indicates whether the observed data are statistically significant.

Pearson's Correlation Test: cor.test()

Pearson's product-moment correlation

```
data: age and wholeWeight
t = 41.498, df = 4175, p-value < 2.2e-16
alternative hypothesis: true correlation is not equal to 0
95 percent confidence interval:
    0.5185606 0.5615148
sample estimates:
        cor
    0.5403897
```

Beyond Simple Hypothesis Testing

- Quantify correlation between two variables
- Quantify correlation between one outcome variable and multiple predictor variables
- Account for confounding factors in the test
- Predict one outcome variable by using one or multiple predictor variables

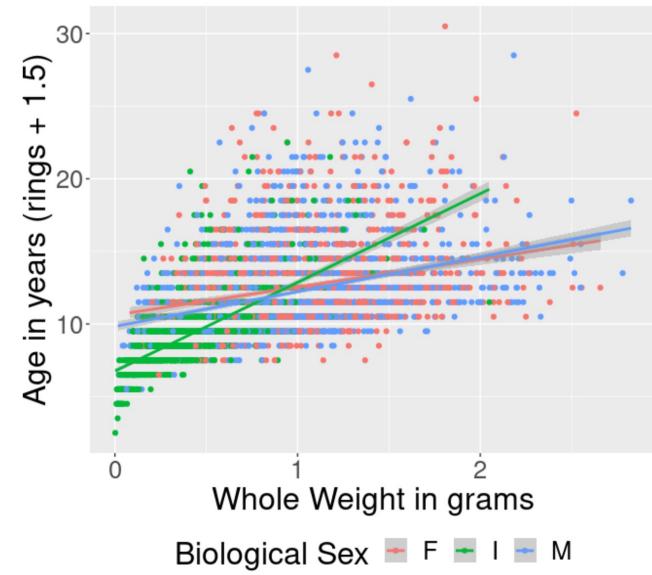
Relationship between one response variable and multiple predictor variables?

Abalones Dataset

Name	Data Type	Measurement Unit	Description
Sex	nominal	_	M, F, and I (infant)
Length	continuous	mm	Longest shell measurement
Diameter	continuous	mm	perpendicular to length
Height	continuous	mm	with meat in shell
Vhole weight	continuous	grams	whole abalone
nucked weight	continuous	grams	weight of meat
scera weight	continuous	grams	gut weight (after bleeding)
hell weight	continuous	grams	after being dried
Rings	integer	_	+1.5 gives the age in years

MUSCLE ATTACHMENT "SCAR" Age of Abalones by Whole Weight Best fit lines shown by sex

Relationship between Abalone age/rings and Whole Weight



Regression



- Technique used for the modeling and analysis of numerical data
- Exploits the relationship between two or more variables so that we can gain information about one of them through knowing values of the other
- Regression can be used for prediction, estimation, hypothesis testing, and modeling causal relationships

Linear Regression

Single variant linear regression model $y_i = \beta_0 + \beta_1 x_i + \varepsilon_i$, i=1,..., n

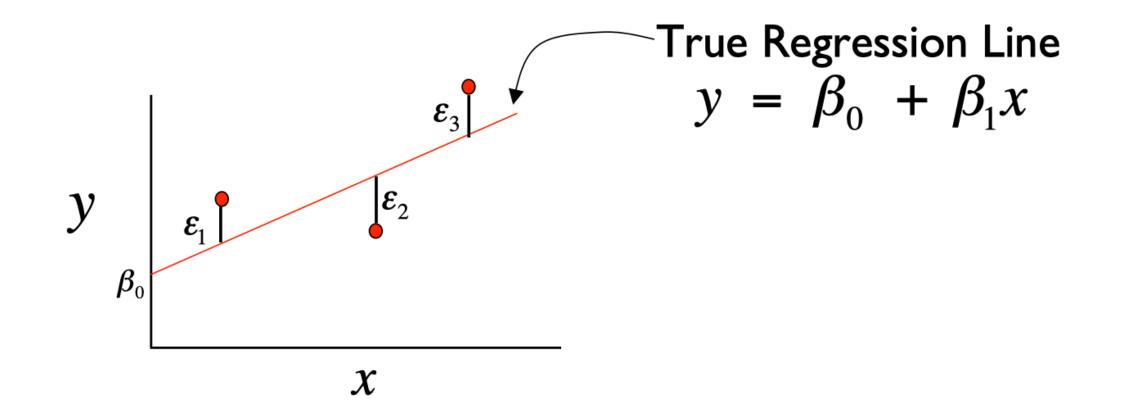
- x_i : Independent (explanatory, predictor, covariate) Variable value for sample i
- y_i: Dependent (response, outcome) Variable value for sample i
- β_0 : Intercept of the fitted linear line
- β_1 : Slope of the fitted linear line, coefficient of X
- $\varepsilon_i \sim N(0, \sigma^2)$: Residual value for sample i

How to fit the model?

• How to find the linear line by estimating the intercept β_0 and slope β_1 ?

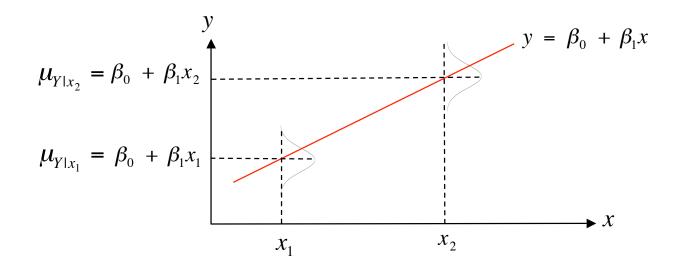
$y = \beta_0 + \beta_1 x$ Δy V Δx β_0 X

Residuals in the linear regression model



The expected value of the outcome variable Y is a linear function of the predictor X

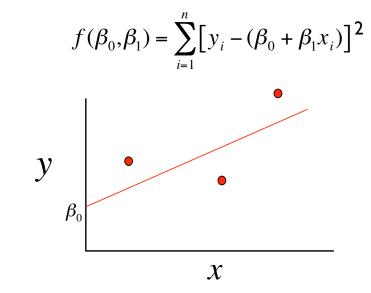
Graphical Interpretation



• For example, if x = height and y = weight then $\mu_{Y|x=60}$ is the average weight for all individuals 60 inches tall in the population

Ordinary Least Square Estimates

- Point estimates of $\hat{\beta}_0$ and $\hat{\beta}_1$ are obtained by the principle of least squares



Calculate the Intercept (β_0): Once you have β_1 , you can calculate the intercept using:

$$eta_0 = ar Y - eta_1 ar X$$

Where:

- \$\overline{X}\$ is the mean of the \$X\$ values

Calculate the Slope (β_1): The slope can be calculated using the formula:

$$eta_1 = rac{n(\sum XY) - (\sum X)(\sum Y)}{n(\sum X^2) - (\sum X)^2}$$

Where:

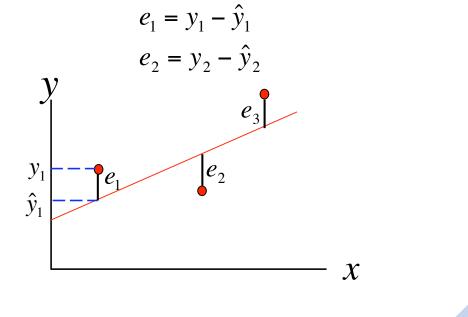
- *n* is the number of observations
- $\sum XY$ is the sum of the product of X and Y
- $\sum X$ is the sum of X values
- $\sum Y$ is the sum of Y values
- $\sum X^2$ is the sum of squared X values

Predicted and Residual Values

• **Predicted**, or fitted, values are values of y predicted by the leastsquares regression line obtained by plugging in $x_1, x_2, ..., x_n$ into the estimated regression line

$$\hat{y}_1 = \hat{\beta}_0 - \hat{\beta}_1 x_1$$
$$\hat{y}_2 = \hat{\beta}_0 - \hat{\beta}_1 x_2$$

• **Residuals** are the deviations of observed and predicted values



```
```{r}
fit1 <- lm(age ~ wholeWeight, data = abalone)
summary(fit1)
```</pre>
```

Linear Regression in R by Im()

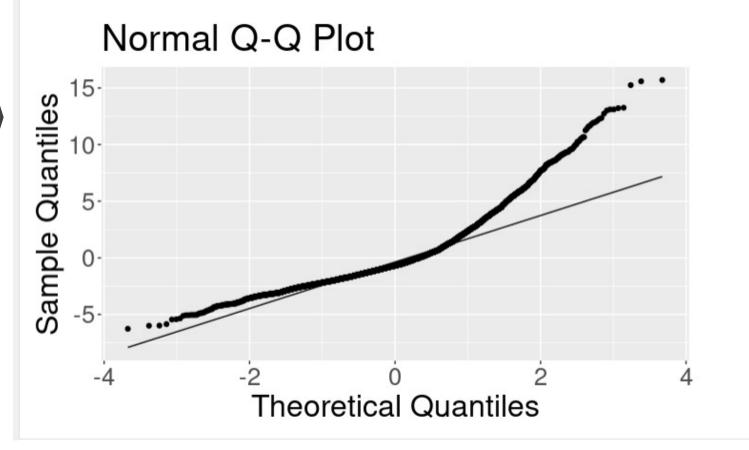
```
Call:
lm(formula = age \sim wholeWeight, data = abalone)
Residuals:
   Min
            10 Median 30
                                 Max
-6.2693 -1.7518 -0.6874 1.0177 15.7029
Coefficients:
           Estimate Std. Error t value Pr(>|t|)
(Intercept) 8.48924 0.08244 103.0 <2e-16 ***
wholeWeight 3.55291 0.08562 41.5 <2e-16 ***
---
Signif. codes: 0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
```

Residual standard error: 2.713 on 4175 degrees of freedom Multiple R-squared: 0.292, Adjusted R-squared: 0.2919 F-statistic: 1722 on 1 and 4175 DF, p-value: < 2.2e-16

Check residuals distribution

```
```{r}
residuals.df <- data.frame(residuals = fit1$residuals)
ggplot(residuals.df, aes(sample = residuals)) +
 stat_qq() + stat_qq_line() +
 labs(x = "Theoretical Quantiles", y = "Sample Quantiles", title = "Normal Q-Q Plot")
```</pre>
```

A X



Relationship between rings/age and whole weight while accounting for Sex? Predict Abalone age/rings by multiple measurements?

Abalones Dataset

Name	Data Type	Measurement Unit	Description
Sex	nominal	_	M, F, and I (infant)
Length	continuous	mm	Longest shell measurement
Diameter	continuous	mm	perpendicular to length
Height	continuous	mm	with meat in shell
Whole weight	continuous	grams	whole abalone
Shucked weight	continuous	grams	weight of meat
Viscera weight	continuous	grams	gut weight (after bleeding)
Shell weight	continuous	grams	after being dried
Rings	integer	_	+1.5 gives the age in years

Multivariate Linear Regression



 Extension of the simple linear regression model to two or more independent/predictor variables

$Y = \beta_0 + \beta_1 X_1 + \beta_2 X_2 + \dots + \beta_p X_p + \epsilon$

- Exercise: fit the following multivariate linear regression model with the Abalone data.
 - Age ~ Sex + length + diameter + height + wholeWeight + shuckedWeight + wisceraWeight + shellWeight

How to quantify categorical independent variable?

Binary variable: coded as 0/1

The sex variable in the abalone dataset has three levels: F, I, M ?

How to quantify categorical independent variable?

- The sex variable in the abalone dataset has three levels: F, M, I?
- Code through (k-1) dummy variables for k levels:

Sex	X1	X2
F	1	0
Μ	0	1
I	0	0

```{r}
fit2 <- lm(age ~ factor(sex) + wholeWeight, data = abalone)
summary(fit2)</pre>

Fit a multivariate linear regression model with sex and wholeWeight

| Call:         |             |             |          |                 |        |
|---------------|-------------|-------------|----------|-----------------|--------|
| lm(formula =  | age ~ facto | or(sex) + ( | wholeWei | ght, data = abo | alone) |
| Residuals:    |             |             |          |                 |        |
|               | 1Q Median   | 30          | Max      |                 |        |
|               | •           |             |          |                 |        |
| -6.0404 -1.74 | 42 -0.5449  | 0.9935 1    | 5.7240   |                 |        |
|               |             |             |          |                 |        |
| Coefficients: |             |             |          |                 |        |
|               | Estimate St | d. Error    | t value  | Pr(>ltl)        |        |
| (Intercept)   | 9.6770      | 0.1290      | 74.987   | < 2e-16 ***     |        |
| factor(sex)I  | -1.5034     | 0.1207      | -12.454  | < 2e-16 ***     |        |
| factor(sex)M  | -0.2684     | 0.1004      | -2.674   | 0.00753 **      |        |
| wholeWeight   | 2.8210      | 0.1013      | 27.849   | < 2e-16 ***     |        |
|               |             |             |          |                 |        |
| Signif. codes | : 0 '***'   | 0.001 '**   | ' 0.01 ' | *' 0.05'.' 0.3  | 1''1   |

Residual standard error: 2.661 on 4173 degrees of freedom Multiple R-squared: 0.3195, Adjusted R-squared: 0.319 F-statistic: 653.2 on 3 and 4173 DF, p-value: < 2.2e-16



### In-Class Exercise : Im()

Question Need to Answer for In-class participation credit.

- What is Regression R-square?
- What dose it mean if you get increased Regression R-square by adding additional predictor variables? The same question is included in Task 5 in Exercise 1. Rmd.

## Generalized Linear Regression

What's the difference between general and generalized linear models?

## General

 $E[Y] = \beta_0 + \beta_1 X_1$ 

 $Y \sim N(\mu, \sigma^2)$ 

## Generalized

 $E[g(Y)] = \beta_0 + \beta_1 X_1$ 

 $Y \sim \begin{cases} Bernoulli, Binomial \\ Poisson \\ Negative binomial \\ etc \end{cases}$ 

 $g \sim$  "link" function to transform Y  $g(Y) \sim N(\mu, \sigma^2)$ 

### Why generalized?



Apply linear regression to outcome variables that are clearly not normally distributed

- Binary : case/control, yes/no, 0/1  $Y \sim Bernoulli(p), \quad 0 \le p \le 1$
- Poisson distributed counts

 $Y \sim Poisson(\lambda), \quad \lambda > 0$ 

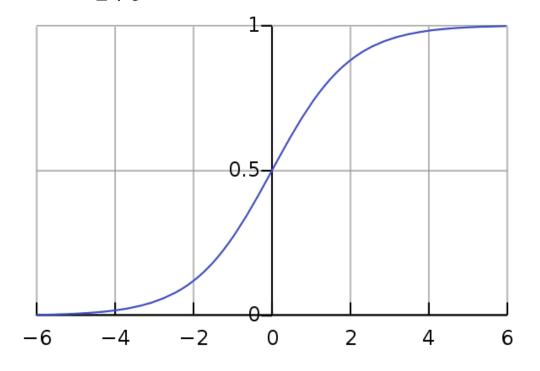
#### Generalized linear regression model

- The mean/expectation function of *Y* can usually be expressed as a function of the distribution parameters
  - Binary outcome: E[*Y*]= *p*
  - Poisson outcome:  $E[Y] = \lambda$
- Model a linear relation ship between E[g(Y)] and explanatory/independent/predictor variables X

## Logistic Regression: *Y*~*Bernoulli* (*p*)

• 
$$l_{\text{LogOdds}} = \log\left(\frac{p}{1-p}\right) = \beta X; \qquad p = Prob(Y = 1)$$

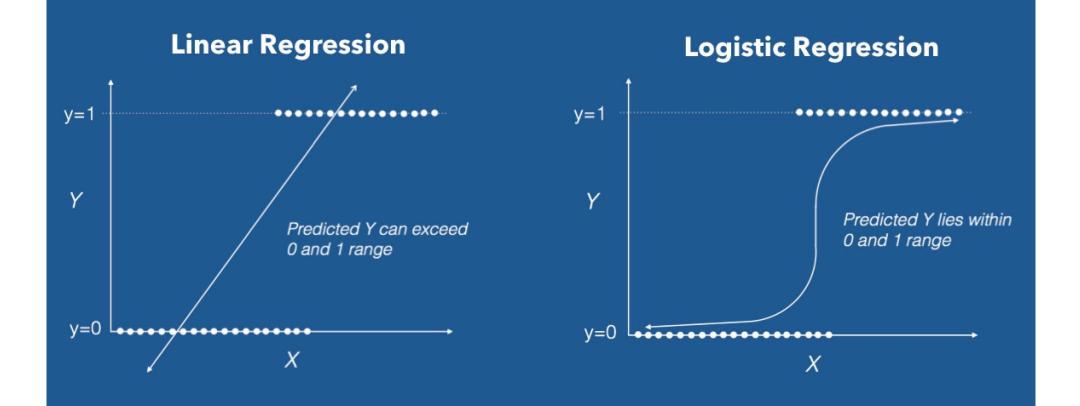
•  $p = \frac{1}{1 + e^{-X\beta}} = \sigma(X\beta)$ , Sigmoid function of  $X\beta$ 



g(E[Y]) is the log odds of success probability or logit

Model will be fitted by maximizing the likelihood function

## Logistic Regression: $Y \sim Bernoulli(p)$



#### Logit link function

Generalized linear model:  $log\left(\frac{l}{1}\right)$ 

$$log\left(\frac{p}{1-p}\right) = \beta_0 + \beta_1 X_1$$

• A one unit change in  $X_1$  leads to a  $\beta_1$  change in the log odds

• In terms of odds:  $odds(Y=1) = e^{b_0 + b_1 X}$ 

• In terms of probability or proportion:  $Pr(Y = 1) = \frac{e^{b_0 + b_1 X}}{1 + e^{b_0 + b_1 X}}$ 

Logit, odds, and probability are different ways of expressing the same thing

#### Logit link function



► Natural log (e) of an odds

➢Often called a *log odds* 

> The logit scale linearizes odds!

 Logits are continuous and are centered on zero (think of as the z-score for the binomial world!)

▶ p = 0.50, odds = 1, then logit = 0

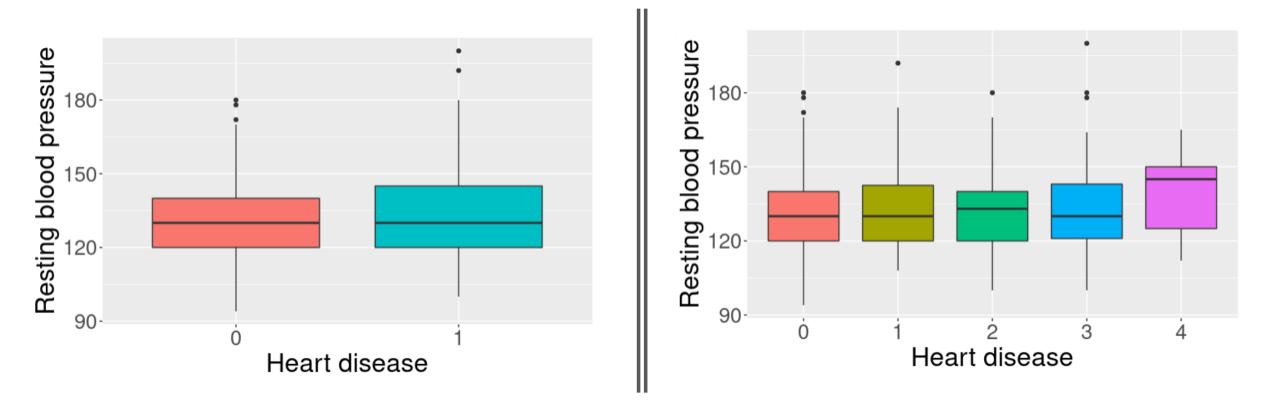
▶ p = 0.70, odds = 2.33, then logit = 0.85

▶ p = 0.30, odds = .43, then logit = -0.85

#### Example dataset : Cleveland heart disease

| Name     | Data Type   | Description                                                                                                                                                                 |
|----------|-------------|-----------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| age      | continuous  | age in years                                                                                                                                                                |
| sex      | binary      | 1 = male; 0 = female                                                                                                                                                        |
| ср       | categorical | chest pain type – 1: typical angina; 2:<br>atypical angina; 3: non-anginal pain; 4:<br>asymptomatic                                                                         |
| trestbps | continuous  | resting blood pressure (in mm Hg on admission to the hospital)                                                                                                              |
| chol     | continuous  | serum cholestoral in mg/dl                                                                                                                                                  |
| fbs      | continuous  | (fasting blood sugar > 120 mg/dl) (1 = true;<br>0 = false)                                                                                                                  |
| restecg  | continuous  | resting electrocardiographic results – 0:<br>normal; 1: having ST-T wave abnormality; 2:<br>showing probable or definite left ventricular<br>hypertrophy by Estes' criteria |
| thalach  | continuous  | maximum heart rate achieved                                                                                                                                                 |
| exang    | binary      | exercise induced angina (1 = yes; 0 = no)                                                                                                                                   |
| oldpeak  | continuous  | ST depression induced by exercise relative to rest                                                                                                                          |
| slope    | categorical | the slope of the peak exercise ST segment-<br>1: upsloping; 2: flat; 3: downsloping                                                                                         |
| са       | continuous  | number of major vessels (0-3) colored by flourosopy                                                                                                                         |
| thal     | categorical | 3 = normal; 6 = fixed defect; 7 = reversable defect                                                                                                                         |
| disease  | categorical | absence (0) vs. presence (1, 2, 3, 4)                                                                                                                                       |

# Study the relationship between resting blood pressure would affect heart disease presence



Study the relationship between resting blood pressure would affect heart disease presence Pearson's product-moment correlation

data: HD and trestbps
t = 2.647, df = 301, p-value = 0.008548
alternative hypothesis: true correlation is not equal to 0
95 percent confidence interval:
 0.03880692 0.25910016
sample estimates:
 cor
 0.1508254

Study the relationship between resting blood pressure would affect heart disease presence

```
Welch Two Sample t-test
```

```
```{r}
fit3 <- glm(HD ~ trestbps, data = cleveland, family = "binomial")
summary(fit3)</pre>
```

Logistic Regression: HD ~ trestbps

Call: $glm(formula = HD \sim trestbps, family = "binomial", data = cleveland)$ Deviance Residuals: 10 Median Min Max 30 -1.4773 -1.0948 -0.9414 1.2394 1.4966 Coefficients: Estimate Std. Error z value Pr(>|z|)(Intercept) -2.483687 0.903634 -2.749 0.00599 ** trestbps 0.017587 0.006796 2.588 0.00966 ** _ _ _ Signif. codes: 0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1

(Dispersion parameter for binomial family taken to be 1)

Null deviance: 417.98 on 302 degrees of freedom Residual deviance: 411.03 on 301 degrees of freedom AIC: 415.03

Number of Fisher Scoring iterations: 4

```
```{r}
fit4 <- glm(HD ~ age + sex + trestbps + factor(thal), data = cleveland, family = "binomial")
summary(fit4)
</pre>
```

```
Call:
glm(formula = HD ~ age + sex + trestbps + factor(thal), family = "binomial",
data = cleveland)
```

Deviance Residuals:

Min	1Q	Median	3Q	Мах
2.0986	-0.7282	-0.4232	0.7656	1.9112

Coefficients:

Estimate Std. Error z value Pr(>|z|)(Intercept) -5.735162 1.360779 -4.215 2.50e-05 \*\*\* 0.052540 0.016683 3.149 0.00164 \*\* age 0.773658 0.339110 2.281 0.02252 \* sex trestbps 0.009081 0.008436 1.076 0.28175 2.691 0.00713 \*\* factor(thal)6 1.511252 0.561693 6.979 2.97e-12 \*\*\* factor(thal)7 2.140144 0.306639 \_ \_ \_ Signif. codes: 0 '\*\*\*' 0.001 '\*\*' 0.01 '\*' 0.05 '.' 0.1 ' ' 1

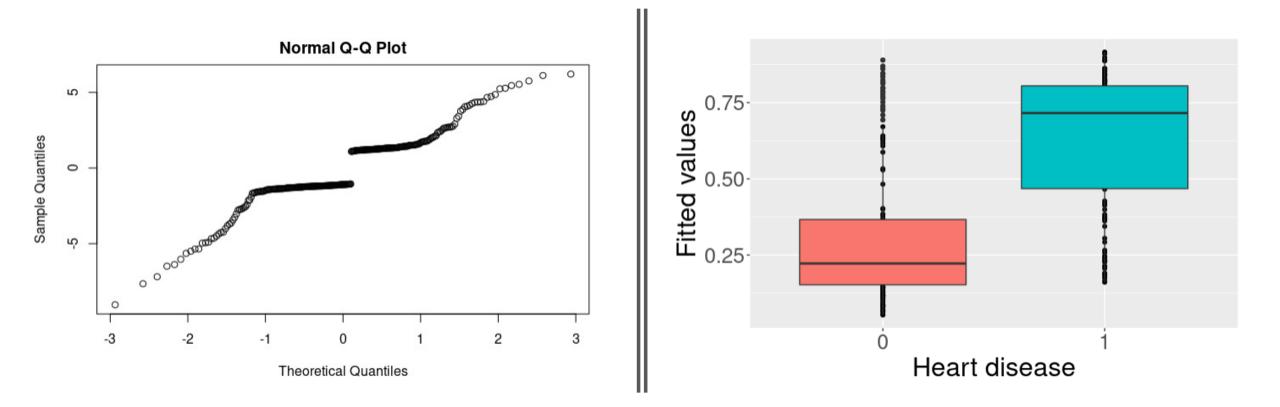
(Dispersion parameter for binomial family taken to be 1)

Null deviance: 415.20 on 300 degrees of freedom Residual deviance: 311.38 on 295 degrees of freedom (2 observations deleted due to missingness) AIC: 323.38

Number of Fisher Scoring iterations: 4

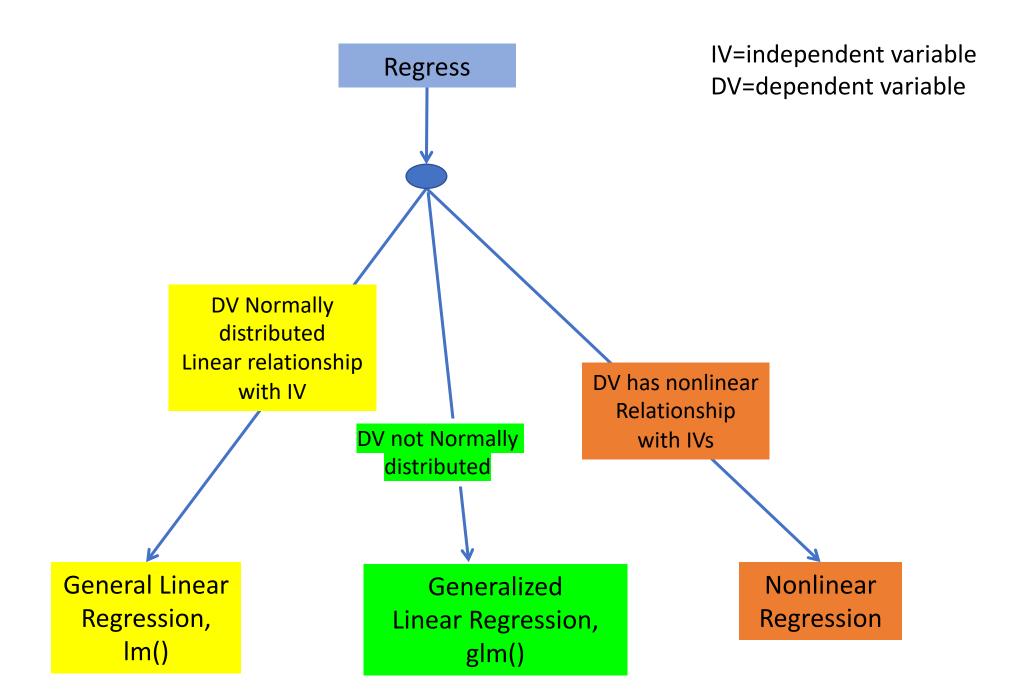
Account for age, sex, and thal

# Logistic regression results



### Generalized linear model families

Normal outcome	• Gaussian	
Binary outcome	• Binomial	
Count outcome	<ul><li>Poisson</li><li>Negative binomial</li></ul>	
Continuous positive outcome	• Gamma • Inverse Gaussian	
Common link functions: identity, logit, log, square-root, inverse, etc.		



## **Checking Assumptions**

- Critically important to examine data and check assumptions underlying the regression model
  - > Outliers
  - > Normality
  - Constant variance
  - Independence among residuals
- Standard diagnostic plots include:
  - > scatter plots of y versus  $x_i$  (outliers)
  - > qq plot of residuals (normality)
  - > residuals versus fitted values (independence, constant variance)
  - $\succ$  residuals versus  $x_i$  (outliers, constant variance)

#### Summary



 Regression offers a single cohesive approach to inference and estimating effect sizes

Response ~ Predictors

- Only reason to stick with t-tests/ANOVA are
  - Mostly just care about "statistical significance"
  - No other confounding covariates
  - Cultural (engrained in biomedical community)

#### Regression or ANOVA/t-tests?

- ANOVA/t-tests thinking emphasize "statistical significance" after experiment
- Regression thinking emphasizes overall weight of an independent variable predictively
- Regression is easy-peasy for "completely randomized" samples
  - Im() –for general linear model
  - glm() –for generalized linear model



#### In-Class Exercise 2 : glm()