MAD – Data Analysis & Biostatistics in R Power - More Regression - Logistic Regression

James R. Hunter, Ph.D.

DIPA, EPM, UNIFESP

9 October 2020



Section 1

Statistical Power

- Statistical Power
- Polynomial Regression

- Statistical Power
- Polynomial Regression
- Section Logistic Regression

- Statistical Power
- Polynomial Regression
- Regression and Analysis of Variance (ANOVA) Kissing Cousins

Section 2

Statistical Power

Conceptual Review

- Null Hypothesis (H_0) : a parameter = some theoretical value
- Alternative Hypothesis (H_1) : \geq , \neq , \leq theoretical value (not H_0)
- \bullet Test statistic calculated from the data assuming H_0 is true
- p-value: probability of observing the theoretical value being tested or a more extreme value
- ullet Small p-vales provide evidence against the H_0
- Level of significance defines the limit of "how small is small enough" for a p-value
 - Notation: α

Power of a Statistical Test

- α provides an idea of the performance of the test across **multiple** samples
- If H_0 is true and $\alpha = 0.01$
 - ▶ In 1% of the **samples**, we would reject H_0 erroneously
 - ▶ In 99% of the **samples**, we would reject H_0 (correct decision)
- We want to reject H_0 when it is really false
- If α is too small, we can reject H_0 even if the correct value of the parameter is close to the value in H_0
- **Power** of a test is the probability of making a correct decision (i.e., reject H_0 when it is really false)
- Higher levels of power in a test mean that the test is more sensitive

Power - Definition

Power is the probability of correctly rejecting a false null hypothesis when the alternative hypothesis is correct

Power Analysis

- Allows us to determine the probability that a statistical significance test will reject a null hypothesis
- Permits the calculation of the number of cases (n) that would be necessary in a sample to achieve a given level of power
- Put more simply: allows us to determine *a priori* the probability that we make a correct decision

2 Types of Statistical Errors

- **Type I** Reject H_0 when it is **true**
 - lacktriangle Occurs with probability lpha
 - Condemning a person falsely of a crime
- **Type II** Not reject H_0 when it is **false**
 - Occurs with probability β
 - $\beta = 1$ power
 - ► A criminal freed in error

Possible Results of a Hypothesis Test

Conclusão do Teste	Estado de Natureza			
		Nula Verdadeira	Nula Falsa	
	Não rejeitar H _o	Correta p = 1 - α	Erro Tipo II p. = B	
	Rejeitar H _o	Erro Tipo L p = α	Correta p = 1 - B	

Factors in Calculating Power - BEAN

- **B**: β Error (Type II)
 - ▶ Probability that significance test will fail to reject H_0 when it is **false**
 - As β error increases, power decreases
- E: Effect Size
 - Magnitude of difference between real population mean and null hypothesized mean relative to standard deviation
 - $\qquad \qquad \frac{(\mu \mu_{H_0})}{\sigma}$
 - ► As effect size increases, power increases
- **A**: α Error (Type I)
 - Probability that statistical test will produce a significant finding when H₀ is true
 - ▶ If $\alpha = 0.05$ and H_0 true, false positive would be found in 5 samples out of 100
- N: Sample size

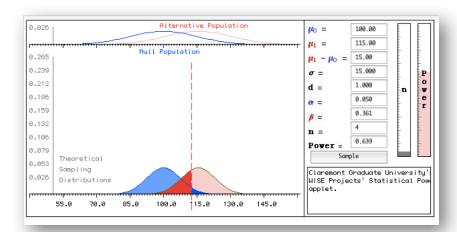
12 / 76

Calculating Power

- If you know three, you can calculate the fourth factor
- n and α are under your direct control

Power – Maximization

- Goal in determining power: maximize the power while maintaining the significance level and minimizing the sample size
- Maximize the probability of finding a true effect while minimizing the chance of finding an effect that does not exist
- Power calculations and sample sized calculated at beginning of project
 during planning
 - Not after already executed experiment. Cannot now try to add new cases



Effect Size

- Of these 4 factors, the most difficult to understand
- Requires some experience with research to apply well
- Table to help beginners use effect size
 - These are general indications
 - With experience you will get a better idea

Table of Effect Size

	Pequeno	Médio	Grande
teste - t	0,2	0,5	0,8
modelo linear	0,02	0,15	0,35
proporções	0,2	0,5	0,8
Qui-quadrado (χ²)	0,1	0,3	0,5

Section 3

How to Calculate Power

Data on Normal Human Temperature

```
temps <- read table(here::here("TempData.txt"), col names = FALSE)
colnames(temps) <- "tempC"</pre>
descr(temps, stats = c("mean", "sd", "min", "q1", "med", "q3", "max", "iqr", "cv"),
     transpose = TRUE)
## Descriptive Statistics
## temps$tempC
## N: 130
                        Std.Dev
                                    Min
                                             01
                                                  Median
                                                              Q3
                                                                            IOR
```

tempC 36.81 0.41 35.72 36.56 36.83 37.06 38.22

Mean

0.50 0.01

Power Calculations with the pwr Package

You can specify 3 of the 4 factors; function calculates the 4th

```
pwr.t.test(n = NULL, d = NULL, sig.level = 0.05, power = NULL,
    type = c("two.sample", "one.sample", "paired"),
    alternative = c("two.sided", "less", "greater"))
```

- \bullet d = effect size
- $sig.level = \alpha$
- You need to specify type and alternative

Execute pwr.t.test() Function

n = 130

d = 0.8

sig.level = 0.05 power = 1

alternative = two.sided

##

##

##

##

##

Conclusion about 130 Person Sample

- Power = 1; Sample larger than necessary
- Test a sample of 10

##

##

##

```
pwr.t.test(n = 10, d = 0.8, sig.level = 0.05,
           type = "one.sample", alternative = "two.sided")
##
##
        One-sample t test power calculation
##
##
                 n = 10
##
                 d = 0.8
         sig.level = 0.05
```

power = 0.6162328

alternative = two.sided

How Many for a Power = 0.95?

n = 22.32455

##

- A sample of 23 would be sufficient to have a very high statistical power
- Tell your colleague that 23 cases would have been enough

Types of Tests Covered in pwr

- pwr.p.test: one-sample proportion test
- pwr.2p.test: two-sample proportion test
- pwr.2p2n.test: two-sample proportion test (unequal sample sizes)
- pwr.t.test: two-sample, one-sample and paired t-tests
- pwr.t2n.test: two-sample t-tests (unequal sample sizes)
- pwr.anova.test: one-way balanced ANOVA
- pwr.r.test: correlation test
- pwr.chisq.test: chi-squared test (goodness of fit and association)
- pwr.f2.test: test for the general linear model (logistic regression, etc.)

Section 4

Final Thoughts on Hypothesis Tests

- Be careful when you interpret a p-value
 - p-value does not tell you if H_0 is true
 - \blacktriangleright What it says: how probable would be the data observed **if** H_0 were true
- Whow you collect the data is the key step in arriving at a correct conclusion
- Always use two-sided tests if you are not absolutely certain that one of the sides is totally uninteresting
- lacktriangle Statistical significance \neq practical significance \neq importance

Effect of Sample Size on p-value

ullet Three samples: flip a fair coin (p = 0.5); always have 60% heads

_				
hea	ads	trials	statistic	p_value
	6	10	6	0.7539063
	60	100	60	0.0568879
6	00	1000	600	0.0000000

- **5** Lack of significance does not mean that H_0 is true
 - Large p-values can occur because of
 - ★ Luck/chance
 - * Problems in data collection
 - ★ H₀ really false
- Specify your hypotheses before collecting data
- Proportion tests and tests of means require
 - Independence of observations
 - Distribution of the estimators approximately normal
- t-test very robust to outliers
 - ▶ Only the most extreme outliers can invalidate t-statistic and p-value

Section 5

Polynomial Regression

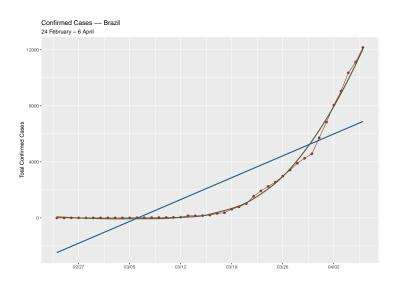
What Is Polynomial Regression

• A way to capture non-linear properties of a variable and retain the linearity of the overall function

Example: Start of COVID-19 Pandemic in Brazil

- Typical of the initial phase of a new infectious disease outbreak
- Deal with period from start of confirmed cases in February through start of April
- Data from Johns Hopkins COVID repository and dashboard

Data

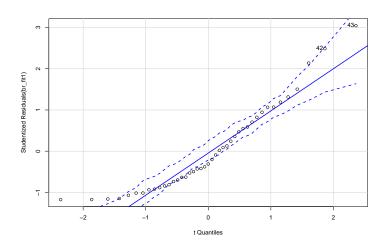


Linear Regression of Cases on Dates

```
br_fit1 <- lm(cases ~ elapsed_date, data = br_cases)</pre>
summary(br fit1)
##
## Call:
## lm(formula = cases ~ elapsed date, data = br cases)
##
## Residuals:
##
      Min
               10 Median
                               30
                                      Max
## -2301.3 -1630.4 -617.5 1254.1 5277.5
##
## Coefficients:
               Estimate Std. Error t value Pr(>|t|)
##
## (Intercept) -2700.86 618.11 -4.370 8.29e-05 ***
## elapsed date 222.89 24.47 9.108 2.11e-11 ***
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 1991 on 41 degrees of freedom
## Multiple R-squared: 0.6693, Adjusted R-squared: 0.6612
## F-statistic: 82.96 on 1 and 41 DF, p-value: 2.113e-11
```

Q-Q Plot of Linear Fit

car::qqPlot(br_fit1)

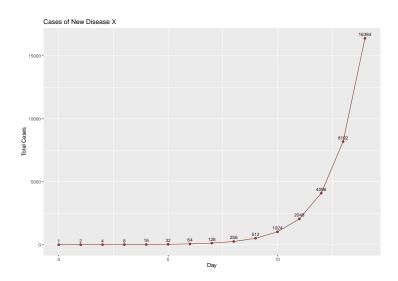


[1] 42 43

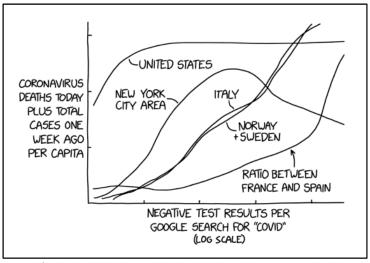
Create a Polynomial Model of the Data

 We know that the initial phases of a new infection follow an exponential curve

```
options(scipen = 1)
r < -2
times <- 14 #14 days = 2 weeks
x <- r^{(0:times)}
Х
    [1]
                               8
                                    16
                                          32
                                                 64
                                                      128
                                                            256
                                                                   512
                                                                        1024
                                                                              2048
   [13] 4096 8192 16384
```



Sort of Like This



I'M A HUGE FAN OF WEIRD GRAPHS, BUT EVEN I ADMIT SOME OF THESE CORONAVIRUS CHARTS ARE LESS THAN HELPFUL.

Create a Quadratic Regression of Elapsed Date

- "Quadratic" means 2nd degree polynomial
 - Will create an equation

$$\hat{Y}_i = \beta_0 + \beta_1 X_i + \beta_2 X_i^2 + \epsilon_i$$

- poly() function (base R) creates regression terms from power 1 up to power in command
 - poly(var, degree = 2, raw = FALSE)
 - degree = put the degree of the polynomial you want to calculate
 - ▶ raw = FALSE use orthogonal polynomials that is polynomials that will have no overlap

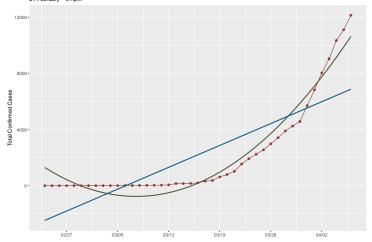
Quadratic Function for elapsed_date

```
br fit2 <- lm(cases~poly(elapsed date, 2, raw = TRUE), data = br cases)
summary(br fit2)
##
## Call:
## lm(formula = cases ~ poly(elapsed_date, 2, raw = TRUE), data = br_cases)
##
## Residuals:
       Min
                 1Q Median
                                  30
                                          Max
## -1297.27 -577.19 22.97 606.22 1502.25
##
## Coefficients:
##
                                     Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                                    1640.0117 352.8630 4.648 3.61e-05 ***
## poly(elapsed_date, 2, raw = TRUE)1 -355.8911 36.9896 -9.621 5.81e-12 ***
## poly(elapsed date, 2, raw = TRUE)2 13.1542 0.8152 16.137 < 2e-16 ***
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 735.7 on 40 degrees of freedom
## Multiple R-squared: 0.956, Adjusted R-squared: 0.9538
## F-statistic: 434.1 on 2 and 40 DF, p-value: < 2.2e-16
```

Plot of Regression

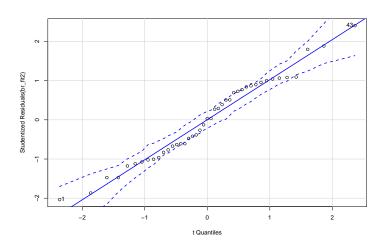
```
br cases %>%
 mutate(fitted_quad = br_fit2$fitted.values) %>%
  ggline(x = "date",
         y = "cases",
         color = "#8F3931FF".
         palette = "uchicago",
         title = "Confirmed Cases -- with Quadratic Regression Line",
         subtitle = "24 February - 6 April".
         xlab = FALSE.
         vlab = "Total Confirmed Cases",
         ggtheme = theme grav()) +
  geom line(aes(y = fitted quad), color = "#58593FFF", size = 1, show.legend = TRUE
  geom_smooth(color = "#155F83FF", method = "lm", se = FALSE) +
  scale x date(date breaks = "7 days", date labels = "%m/%d")
```

Confirmed Cases — with Quadratic Regression Line 24 February – 6 April



Q-Q Plot of Quadratic Fit

car::qqPlot(br_fit2)



[1] 1 43

Cubic Regression of Date and Cases

- Still can be improved
- Try 3rd Degree Cubic Regression
- Can we remove the oscillation in the Q-Q graph?
- Can we reduce the regions of over- and under-estimation?

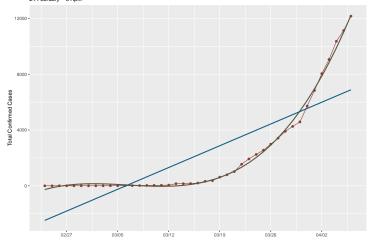
Calculating Cubic Regression

```
br fit3 <- lm(cases~polv(elapsed date, 3, raw = TRUE), data = br cases)
summary(br_fit3)
##
## Call:
## lm(formula = cases ~ poly(elapsed date, 3, raw = TRUE), data = br cases)
##
## Residuals:
##
      Min
               10 Median
                               30
                                      Max
## -781 37 -94 21 22 30 111 31 474 04
##
## Coefficients:
##
                                       Estimate Std. Error t value Pr(>|t|)
## (Intercept)
                                    -429.60935 141.41184 -3.038 0.00423 **
## poly(elapsed date, 3, raw = TRUE)1 178.10207 27.51443 6.473 1.14e-07 ***
## poly(elapsed_date, 3, raw = TRUE)2 -16.84034 1.44413 -11.661 2.77e-14 ***
## poly(elapsed date, 3, raw = TRUE)3 0.45446 0.02159 21.050 < 2e-16 ***
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 211.9 on 39 degrees of freedom
## Multiple R-squared: 0.9964, Adjusted R-squared: 0.9962
## F-statistic: 3636 on 3 and 39 DF, p-value: < 2.2e-16
```

Plot of Cubic Regression

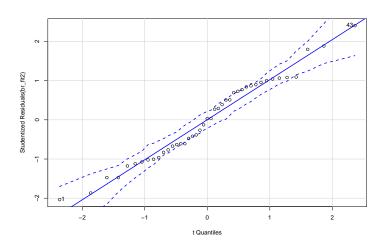
```
br cases %>%
  mutate(fitted cubic = br fit3$fitted.values) %>%
  ggline(x = "date",
         y = "cases",
         color = "#8F3931FF".
         palette = "uchicago",
         title = "Confirmed Cases -- with Cubic Regression Line",
         subtitle = "24 February - 6 April".
         xlab = FALSE.
         vlab = "Total Confirmed Cases",
         ggtheme = theme grav()) +
  geom line(aes(y = fitted cubic), color = "#58593FFF", size = 1, show.legend = TRU
  geom_smooth(color = "#155F83FF", method = "lm", se = FALSE) +
  scale x date(date breaks = "7 days", date labels = "%m/%d")
```

Confirmed Cases — with Cubic Regression Line 24 February – 6 April



Q-Q Plot of Cubic Fit

car::qqPlot(br_fit2)



[1] 1 43

Section 6

One-Way ANOVA - Regression's Kissing Cousin

Analysis of Variance - ANOVA

- Tool developed in 1920's by Sir Ronald Fisher
- Misnomer
 - Technique for comparing differences in means
 - Not variances
- Way to study groups
 - And how some outcome variable can depend on them
- Variety of ANOVA models
- Multiple comparison tests when we want to isolate certain differences
 - Correction for aggressive estimation of p-value
- Clinical trials
 - ► Go-to method of analyzing clinical trial data

Imaginary Clinical Trial of New Drug for COVID-19 Treatment

- New drug "Nontussis"
 - ▶ Directly attacks SARS-CoV-2 virus in respiratory system
- Trial will compare Nontussis to an imaginary version of HCQ and a placebo
- Trial will be double-blinded
- Patients will be divided between
 - ▶ Those on supplemental O_2 (50%)
 - ▶ Those with no supplemental O_2 (50%)
- Outcome measure is a scaled difference (between 1 and 10) of the difference between viral load at enrollment and viral load after 5 days of treatment

Load and View the Data

```
clin_trial <- read_rds(here("covid_clin_trial.rds"))
glimpse(clin_trial)

## Rows: 66

## Columns: 4

## $ pac_id <chr> "1", "2", "3", "4", "5", "6", "7", "8", "9", "10", "11", "12...

## $ drug <chr> "placebo", "placebo", "placebo", "placebo", "placebo", "placebo", "placebo", "placebo", "supp", "supp",
```

Table of the Data

```
clin_trial %>%
  group_by(drug, o2) %>%
  summarise(avg_score = mean(score)) %>%
  knitr::kable()
```

drug	o2	avg_score
hcq	no_supp	4.690909
hcq	supp	4.400000
nontussis	no_supp	6.236364
nontussis	supp	5.981818
placebo	no_supp	4.309091
placebo	supp	4.972727

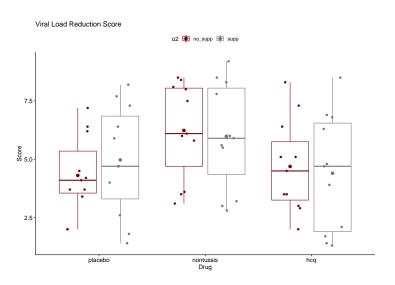
Descriptive Statistics of Data – Supplemental Oxygen

```
clin trial %>%
 filter(o2 == "supp") %>%
 group by(drug, o2) %>%
descr(stats = c("mean", "sd", "min", "med", "max", "igr", "cv"),
     transpose = TRUE)
## Descriptive Statistics
## clin_trial$score
## Group: drug = hcg. o2 = supp
## N: 11
              Mean
                    Std.Dev Min
                                   Median
                                            Max
                       2.53 1.30 4.70 8.50 4.65 0.58
       score 4.40
##
## Group: drug = nontussis, o2 = supp
## N: 11
##
              Mean Std.Dev
                            Min Median Max
       score 5.98
                       2.29 2.80 5.90 9.20 3.70 0.38
## Group: drug = placebo, o2 = supp
##
              Mean
                    Std.Dev
                            Min Median
                                            Max
                                                   IOR
##
       score 4.97 2.36 1.40 4.70 8.20 3.55 0.47
```

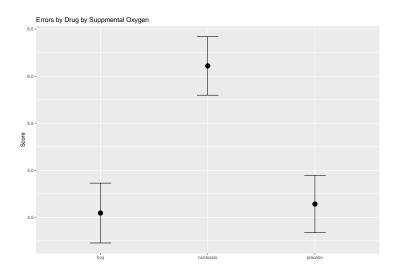
Descriptive Statistics of Data – No Supplemental Oxygen

```
clin trial %>%
 filter(o2 == "no_supp") %>%
 group by(drug, o2) %>%
descr(stats = c("mean", "sd", "min", "med", "max", "igr", "cv"),
     transpose = TRUE)
## Descriptive Statistics
## clin_trial$score
## Group: drug = hcg. o2 = no supp
## N: 11
              Mean
                     Std.Dev
                            Min
                                    Median
                                             Max
                                                   IOR
       score 4.69 1.98 2.00 4.50 8.30 2.50 0.42
##
## Group: drug = nontussis. o2 = no supp
## N: 11
##
              Mean
                     Std.Dev
                            Min Median Max
       score 6.24
                       2.06 3.10 6.10 8.50 3.35 0.33
## Group: drug = placebo, o2 = no supp
##
              Mean
                     Std.Dev
                             Min
                                  Median
                                             Max
                                                   IOR
        score 4.31 1.69 2.00 4.10 7.20 1.80
```

Graph of Data



Graph of Viral Load Reduction Score Errors



Variance of Y (score)

- To understand how this works, need to review variance and sum of squares
- Variance is again based on deviation of individual point from a mean
 - lacktriangle Here, it is Y for each cell compared to the overall mean for Y (\bar{Y})
 - $(Y_{ik} \bar{Y})$ Note subscripts
- Subscripts (and cells) represent
 - ► Each group (k)
 - ► Each **member** within the group (i)
- Formula just assures you pick up all the cells of a cross table

$$Var(Y) = \frac{1}{N} \sum_{k=1}^{K} \sum_{i=1}^{N_k} (Y_{ik} - \bar{Y})^2$$

Variance Looks Like a Mean

- Because it is a mean
- Squared deviations divided by N
- Mean of the squared deviations

Variances and Sum of Squares

- ANOVA works with Total Sum of Squares (SST)
- SS_{tot} is variance not adjusted by N

$$SS_{tot} = \sum_{k=1}^{K} \sum_{i=1}^{N_k} (Y_{ik} - \bar{Y})^2$$

- Divides SST into 2 components
 - \triangleright SS_{betw} Sum of squares between groups
 - ► $SS_{w/in}$ Sum of squares within groups

SS_{betw} - Between groups

ullet Difference between group means $(ar{Y}_k)$ and overall mean $(ar{Y})$

$$SS_{betw} = \sum_{k=1}^K \sum_{i=1}^{N_k} (\bar{Y}_k - \bar{Y})^2$$

$SS_{w/in}$ - Within Groups

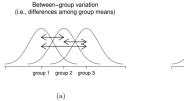
• Difference between the group means (\bar{Y}_k) and the individual values (Y_{ik})

$$SS_{w/in} = \sum_{k=1}^{K} \sum_{i=1}^{N_k} (Y_{ik} - \bar{Y}_k)^2$$

As with regression . . .

$$SS_{tot} = SS_{betw} + SS_{w/in}$$

Graphic Depiction of Variability within and between Groups



(i.e., deviations from group means)

Within-group variation

source: Navarro, Isr v.6, p.430.

Construct Null Hypothesis for the ANOVA

 Null hypothesis: there will be no difference among the drugs in the reduction of viral load

$$H_0$$
: $\mu_{placebo} = \mu_{hcq} = \mu_{nt}$

- Alternative hypothesis: at least one of the means of the drugs will be different than the others

$$H_1: \mu_{\it placebo}
eq \mu_{\it hcq}
eq \mu_{\it nt}$$

Testing the Hypothesis: The F Ratio

- Sums of Squares represent variability: differences of values and group means from other means
- The F ratio is a ratio of variances
- Remember that variances have a degree of freedom
- Here we are working with 2 variances: $SS_{w/in}$ and SS_{betw}
 - We use the sum of squares and not the variances because the N's cancel out
- ... we have two different degrees of freedom
 - $df_{betw} = K 1$ (number of groups less 1)
 - $df_{w/in} = N K$ (number in sample less the number of groups)

Calculate Mean Squares from Sum of Squares

Divide each type of sum of squares by its degrees of freedom

$$MS_{betw} = rac{SS_{betw}}{df_{betw}}$$
 $MS_{w/in} = rac{SS_{w/in}}{df_{w/in}}$

F Ratio is Ratio of Mean Squares

$$F = \frac{MS_{betw}}{MS_{w/in}}$$

- Bigger the F: relatively bigger the separation between group peaks and narrower the curves
- F test statistic has 2 parameters: its 2 different degrees of freedom
- ullet F ratio must be larger than 1 to have any chance of rejecting H_0

Running ANOVA in R

- Basic command aov() with model specified with formula notation
 - As with regression
 - ► Assign to a model name
 - ▶ Use summary() to see full result

```
anova_mod1 <- aov(score~drug, data = clin_trial)
summary(anova_mod1)</pre>
```

```
## Df Sum Sq Mean Sq F value Pr(>F)
## drug 2 33.8 16.90 3.723 0.0296 *
## Residuals 63 286.0 4.54
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
```

Review items in table

Same Information as Shown on Regression Output

lm() is the same operation as aov() behind the curtain

```
lm_mod1 <- lm(score~drug, data = clin_trial)</pre>
summary(lm_mod1)
##
## Call:
## lm(formula = score ~ drug, data = clin_trial)
##
## Residuals:
      Min
               10 Median
                                      Max
## -3.3091 -1.6205 -0.1091 1.8307 3.9545
##
## Coefficients:
                Estimate Std. Error t value Pr(>|t|)
## (Intercept) 4.54545
                            0.45428 10.006 1.22e-14 ***
## drugnontussis 1.56364 0.64245
                                      2.434
                                             0.0178 *
## drugplacebo
                 0.09545 0.64245 0.149 0.8824
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
##
## Residual standard error: 2.131 on 63 degrees of freedom
## Multiple R-squared: 0.1057, Adjusted R-squared: 0.0773
## F-statistic: 3.723 on 2 and 63 DF, p-value: 0.02964
```

Differences in Presentation of Results

- Drugs in lm() appear as dummy variables
- F statistic and df's the same
- p-value of model the same
- lm() estimates of line not useful to ANOVA
 - Interest in ANOVA is in differences in means

Effect Size (η^2) and Coefficient of Determination (R^2)

 Effect size: ratio of between-group sum of squares and total sum of squares

$$\eta^2 = \frac{SS_{betw}}{SS_{tot}}$$

- They measure equivalent ratios
 - ▶ How much of the variance does the model explain
 - Leaving the rest to the residuals
 - lacktriangleright 1sr package has a function that automatically calculates η^2
- $\eta^2 = 1sr::etaSquared(anova_mod1)$ \$eta.sq = 0.1056935, 0.1056935
- R² from lm() function = summary(lm_mod1)\$r.squared = 0.1056935

Multiple Comparison *post hoc* Tests

- If you have an ANOVA with more than two groups and a significant result . . .
 - ▶ Where did the significance come from?
 - ▶ Which comparison of means?
- Null hypothesis really has a 3-way claim:
 - ▶ Nontussis is no better than a placebo
 - ▶ HCQ is no better than a placebo
 - Nontussis is no better than HCQ
- Running multiple t-tests to find where the significance comes from is NOT the answer
- Likely to find significant results simply by chance
 - ▶ The whole hypothesis test system based on controlling Type I error rates

Adjustments (Corrections) When Making Multiple Comparison Tests

- Bonferroni Correction
 - ▶ If I am making m number of tests
 - ightharpoonup Multiply raw p-values times m and then compare to lpha
 - ▶ Very conservative will accept more *H*_Os than other methods
- Benjamini-Hochberg Method (False Discovery Rate)
 - Controls the expected proportion of false discoveries among the rejected H₀s
- Many more

Multiple Comparison in R

- Function pairwise.t.test() conducts pairwise tests and applies correction of your choice
- Feed it the dependent variable (score) and the grouping variable (drug)
- Choose an adjustment method (p.adjust.method)

With Benjamini-Hochberg Method

```
pairwise.t.test(x = clin_trial$score, g = clin_trial$drug, p.adjust.method = "BH")

##
## Pairwise comparisons using t tests with pooled SD
##
## data: clin_trial$score and clin_trial$drug
##
## hcq nontussis
## nontussis 0.039 -
## placebo 0.882 0.039
```

P value adjustment method: BH

##

With Bonferroni Correction

##

```
pairwise.t.test(x = clin_trial$score, g = clin_trial$drug, p.adjust.method = "bonfe")

##

## Pairwise comparisons using t tests with pooled SD

##

## data: clin_trial$score and clin_trial$drug

##

## hcq nontussis

## nontussis 0.053 -

## placebo 1.000 0.077
```

P value adjustment method: bonferroni

More on ANOVA

- clin_trial was a balanced design
 - ▶ All the groups were of the same size
- Unbalanced designs are possible
 - ▶ Calculation of SS_x , df_x more complicated
- Very sophisticated designs possible for multiple dimensional studies
- Non-parametric version using ranks
 - Kruskal-Wallis rank sum test
 - kruskal.test()
 - Use when normality of data and residuals or equality of variances is a problem
- Wide variety of test designs