SIRE516

Basic data analyses

Disclaimer

- This is not a statistics class so I will not cover all statistical aspects behind each analysis.
- I will cover regularly used analyses.
- However, I can provide detailed explanations if time permits.

Exploratory data analysis

Understanding your data

- Before any analysis
 - Read the data dictionary
 - Consult the source (e.g. 'wet lab' people, physicians)
- Type of data
 - Transformation?
 - Type of test and analysis?

Understanding your data

- General data types
 - Categorical
 - Nominal
 - Ordinal
 - Numerical
 - Arbitrary (uniform distribution?)
 - Normal distribution
 - Other: Binomial, Poisson, Beta, etc

Histogram

```
1 ###Histogram
2 library(MASS)
3 ?iris
4
5 #Which parameter is normally distributed?
6 truehist(iris$Sepal.Length)
7 truehist(iris$Sepal.Width)
8 truehist(iris$Petal.Length)
9 truehist(iris$Petal.Width)
```

Quantile-Quantile plot

```
11 ~ #### Q-Q plot ####
12 library(car)
13
14 #Which parameter is normally distributed?
15 qqPlot(iris$Sepal.Length)
16 qqPlot(iris$Sepal.Width)
17 qqPlot(iris$Petal.Length)
18 qqPlot(iris$Petal.Width)
```

Test of normality

```
#### Test of Normality

####### Test of Normality

###### Test of Normali
```

What to do if your data is not normally distributed?

- Transformation
 - Log10: Viral load
- Non-parametric analyses
 - No transformation
 - Arbitrary values

```
#### Data transformation ####

dat <- rnorm(100,mean = 5,sd = 2) # Generate 100 numbers with mean = 5 and sd = 2 (Normal distribution)
truehist(dat)

expdat <- exp(dat) #e^x ~ 2.718282^x
truehist(expdat)

shapiro.test(dat)

tdat <- log10(expdat + 1) # + 1 is important because log10(0) is undefined!
truehist(tdat)
shapiro.test(tdat)

shapiro.test(tdat)

shapiro.test(tdat)</pre>
```

How to present values summarizing your data

- Categorical
 - Count (Percent of count of all category)

```
42 → #### Summarize categorical value ####
    rows <- sample(nrow(iris),91, replace = F) # Sample 91 rows from the iris dataset
    IR <- iris[rows,]</pre>
44
45
    tb <- table(IR$Species)</pre>
46
47
    print(tb)
48
49
    percent <- round(100 * tb/nrow(IR),1)
    print(percent)
50
51
    paste0(tb," (",percent,")")
```

How to present values summarizing your data

- Numerical
 - Mean ± Standard deviation
 - Normal distribution only

```
#### Summarize numerical value (Mean SD) ####

m <- round(mean(IR$Sepal.Length),1)

s <- round(sd(IR$Sepal.Length),1)

paste0(m, " ± ", s)

lm <- round(mean(exp(IR$Sepal.Length)),1)

ls <- round(sd(exp(IR$Sepal.Length)),1)

paste0(lm, " ± ", ls) # SD >= mean implying deviation from normal distribution

llm <- round(mean(log10(1+ exp(IR$Sepal.Length))),1)

lls <- round(sd(log10(1+exp(IR$Sepal.Length))),1)

paste0(llm, " ± ", lls)</pre>
```

How to present values summarizing your data

Numerical

- Median (Interquartile range, IQR, Q25 to Q75)
- Median (Range, Min to Max)
- Almost always suitable for numerical data

```
#### Summarize numerical value (Median) ####

68  q <- round(quantile(IR$Sepal.Length,c(0.5,0.25,0.75)),1)

69  paste0(q[1]," (",q[2],"-",q[3],")")

70  paste0(median(IR$Sepal.Length)," (",min(IR$Sepal.Length),"-",max(IR$Sepal.Length),")")

71

72  lq <- round(quantile(exp(IR$Sepal.Length),c(0.5,0.25,0.75)),1)

73  paste0(lq[1]," (",lq[2],"-",lq[3],")")

74  paste0(round(median(exp(IR$Sepal.Length)),1)," (",round(min(exp(IR$Sepal.Length)),1),

75  "-",round(max(exp(IR$Sepal.Length)),1),")")</pre>
```

Practical 1

Practical 1

- We will create a "baseline" characteristic table of the IRIS data set
- The table will contain four columns one for variable and three for each species
- The first row will be count (%) for each species

```
1 ?iris
2 summary(iris)
```

Hypothesis testing

Why?

- We cannot measure or test all existing data.
- We measure and test a small part (i.e. sample) of the data (i.e. population).
- How can we be sure the small part represents all the data?
 - Sample size → Type 1 & 2 errors
 - Hypothesis testing → Type 1 errors
- We <u>reject</u> a "Null" hypothesis
 - Sufficient evidence

Table of error types		Null hypothesis (H_0) is	
		True	False
Decision about null hypothesis (<i>H</i> ₀)	Don't reject	Correct inference (true negative) (probability = $1-\alpha$)	Type II error (false negative) (probability = β)
	Reject	Type I error (false positive) (probability = α)	Correct inference (true positive) (probability = 1-β)

One-sample testing

- Population mean or proportion was previously estimated
- We test whether the value found in our sample agrees with these values

One-sample test of proportion

- Example: Flipping a normal coin would result in a head in 50% of the trials
- You flip your coin 100 times and get a head 71 times. Is your coin normal or loaded?

```
77 * #### One-sample proportion test ####
78 binom.test(x = 71, n = 100, p = 0.5) # Exact test
79
80 prop.test(x = 71, n = 100, p = 0.5) # Estimation
```

One-sample test of mean

- Example: A survey in 2000 found that Thai male first graders had a mean height of 105 cm with a standard deviation of 5 cm.
- You conduct a similar survey in 100 first graders and found that the mean height is now 107 cm with the same standard deviation. Are the male first graders getting taller?

Test difference in proportions

- Example: Proportion of patients with cancer for two populations
- chisq.test() and prop.test() for large samples (require different inputs)

```
96-#### Difference in proportions ####
 97 smoke <- 510
 98 smoke CA <- 400
 99 non smoke <- 540
100 non smoke CA <- 300
101
102 smoke not CA <- smoke - smoke CA
103 non_smoke_not_CA <- non_smoke - non_smoke_CA
104
105 prop.test(x = c(non smoke CA, smoke CA), n = c(non smoke, smoke))
106
107 m <- matrix(data = c(non smoke CA, non smoke not CA, smoke CA, smoke not CA) , nrow = 2)
108 print(m)
109
110 chisq.test(m)
111 fisher.test(m)
```

Test difference in means

- Example: Heights between male and female first graders.
- Parametric test: t.test
- Non-parametric test: wilcox.test

```
112 - #### Difference in means ####
     male < round(rnorm(n = 100 ,mean = 107, sd = 5),1)
     female \leftarrow round(rnorm(n = 100, mean = 105, sd = 4.5),1)
114
115
     truehist(male)
116
     truehist(female)
117
118
     t.test(male,female, var.equal = F, paired = F)
119
     wilcox.test(male,female, paired = F)
120
121
122 #paired = T for dependent samples (e.g. Before-After mearsurements)
```

Practical 2

Practical 2

- We will perform hypothesis testing with the "Melanoma" dataset.
- We will compare between males and females for any difference in each parameter
- For 'status', regroup to 1 vs 2+3

```
1 library(MASS)
2
3 ?Melanoma
4
5 female <- Melanoma[which(Melanoma$sex == 0),]
6 male <- Melanoma[which(Melanoma$sex == 1),]</pre>
```

Categorical Data Analysis

Contingency table

- Help/visualize for categorical data analysis
- Example:

	Cancer (Case)	No cancer (Control)	
Smoker	400	150	
Non smoker	100	320	
TOTAL	500	470	

Contingency table

Create a table from counts

```
124 = #### Contingency table from counts ####
125  ctable <- matrix(c(400,150,100,320), nrow = 2 , byrow = T) # Matrix fill data by column by default
126  dimnames(ctable) <- list(c("Smoker","Non-smoker"),c("Cancer","No cancer"))
127  print(ctable)
128
129  matrix(c(400,150,100,320), nrow = 2)</pre>
```

Contingency table

Create a table from raw data

```
131 - #### Contingency table from raw data ####
132
133
     #Simulate the data
     rawdat <- data.frame(patientID = 1:(400+150+100+320),
134
135
                            smoking = c(rep("Smoker", 400+150), rep("Non-smoker", 100+320)),
                            status = c(rep("Cancer", 400), rep("No cancer", 150), rep("Cancer", 100), rep("No cancer", 320)))
136
137
     View(rawdat)
      summary(rawdat)
138
139
     ctable2 <- table(rawdat[,c("smoking","status")])</pre>
140
     print(ctable2)
141
```

Test of difference proportions

- chisq.test() → Estimation
 - <u>Caution</u>: if >20% of cells (e.g. 1 in 4 cells of a 2x2 contingency table) is <= 5, Chi-square will not be accurate.
- fisher.test() → Exact test
 - Can be used in most cases. But a large sample size will require intensive computation.

```
143 - #### Test of difference proportions ####
144
     chisq.test(ctable2)
145
146
     fisher.test(ctable2)
147
      small \leftarrow matrix(data = c(1,39,2,20), nrow = 2, byrow = T)
148
      print(small)
149
150
151
     chisq.test(small)
     fisher.test(small)
152
```

- Question: If you decide to start smoking, how much more likely you will get cancer?
- Relative risk (RR)
 - *** Specifically for cohort or cross-sectional study***
 - Not for a case-control study
 - Cases and Controls were specifically recruited
 - Cases from a "rare" disease
 - True prevalence and incidence are lost

$$RR = \frac{\Pr(Disease|Exposed)}{\Pr(Disease|Not exposed)}$$

- Odds ratio (OR)
 - Can be used with cross-sectional, cohort and case-control studies
 - Logistic regression

$$Odds of an event = \frac{\Pr(event \ will \ occur)}{\Pr(event \ will \ not \ occur)}$$

$$OR = \frac{odds \ that \ an \ exposed \ person \ develops \ a \ disease}{odds \ that \ a \ non-exposed \ person \ develops \ a \ disease}$$

	Disease	No disease	Total
Exposed	a	b	a + b
Non-exposed	С	d	c + d
TOTAL	a + c	b + d	a + b + c + d

$${}^{\bullet}RR = \frac{\frac{a}{a+b}}{\frac{c}{c+d}}$$

•
$$OR = \frac{\frac{a}{b}}{\frac{c}{d}}$$

 $= 1 \rightarrow$ No association

< 1 > Negative association (protection)

Package "epitools"

```
#### Measure of association
     library(epitools)
144
145
    #Re-arrange data to match epitools requirement
146
     rawdat$status <- factor(rawdat$status, levels = c("No cancer","Cancer"))
     rawdat$smoking <- factor(rawdat$smoking, levels =c("Non-smoker","Smoker"))</pre>
148
149
150
     ctable3 <- table(rawdat[,c("smoking","status")])</pre>
     print(ctable3)
151
152
     res <- oddsratio(ctable3, method = "wald")
154
     print(res)
155
     res2 <- riskratio(ctable3, method = "wald")
156
157
     print(res2)
158
```

Practical 3

Practical 3

- We will calculate odds ratios for alcohol and tobacco consumption in the "esoph" dataset
- Alcohol consumption → 0-39g/day versus >=40g/day
- Tobacco consumption \rightarrow 0-9g/day versus >=10g/day

```
library(MASS)
library(epitools)

resoph
esoph$slc <- factor(esoph$alcgp, levels(esoph$alcgp), labels = c("0-39g/day",rep(">=40g/day",3)))
esoph$tob <- factor(esoph$tobgp, levels(esoph$tobgp), labels = c("0-9g/day",rep(">=10g/day",3)))
```

Linear Regression

Correlation between two numerical variables

- By knowing X → Estimate Y
- Parametric: Pearson (default)
- Non-parametric: Spearman and Kendall

```
#### Correlation between numerical variables ####
iris
cor.test(iris$Sepal.Length,iris$Petal.Width) #Paired values
cor.test(~Sepal.Length + Petal.Width, iris) #Data.frame

cor.test(iris$Sepal.Length,iris$Petal.Width, method = "s") #Non-paramatric
cor.test(iris$Sepal.Length,iris$Petal.Width, method = "k") #Non-paramatric
```

Correlation between two numerical variables

- P-value:
 - Null hypothesis: the coefficient of correlation = 0
- Interpretation of the coefficient of correlation

Correlation Coefficient Value	Direction and Strength of Correlation
-1.0	Perfectly negative
-0.8	Strongly negative
-0.5	Moderately negative
-0.2	Weakly negative
0.0	No association
+0.2	Weakly positive
+0.5	Moderately positive
+0.8	Strongly positive
+1.0	Perfectly positive

By knowing X \rightarrow Estimate Y

- Simple linear regression: $y = \alpha + \beta x + \varepsilon$
 - α = intercept
 - β = slope
 - ε = error (mean of ε = 0)
 - $\hat{y} = \alpha + \beta x$

```
169 ~ #### Simple linear regression ####
170
171  m <- lm(Sepal.Length ~ Petal.Width , iris)
172  print(m)
173  summary(m)</pre>
```

Interpreting regression result

```
> summary(m)
                        Call:
                        lm(formula = Sepal.Length ~ Petal.Width, data = iris)
                        Residuals:
                             Min
                        -1.38822 -0.29358 -0.04393 0.26429 1.34521
                                                                               P-values:
Intercept (a
                        Coefficients:
                                                                                Test whether any coefficient = 0
                                   Estimate Std. Error t value Pr(>|t|)
                                                                                *Normally, the intercept is kept regardless of p-value
                        (Intercept) 4.77763
                        Petal.Width 0.88858
                                              0.05137
                        Signif. codes: 0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
    Slope (B
                        Residual standard error: 0.478 on 148 degrees of freedom
                        Multiple R-squared: 0.669,
                                                     Adjusted R-squared: 0.6668
                        F-statistic: 299.2 on 1 and 148 DF, p-value: < 2.2e-16
```

 $R^2 \rightarrow \%$ of Y explained (predicted) by your model Used adjusted R^2 for multivariate regression

Multivariate linear regression

- Add more independent variables for prediction
- Categorical variables can be added
 - R does not need recoding of categorical variables for regression

```
175 ▼ #### Multivariate linear regression ####

176

177 mm <- lm(Sepal.Length ~ Sepal.Width + Petal.Length + Petal.Width + Species , iris)

178 print(mm)

179 summary(mm)
```

Multivariate linear regression

-0.723 is added, if and only if, species = versicolor

-1.023 is added, if and only if, species = virginica

```
> summary(mm)
Call:
lm(formula = Sepal.Length ~ Sepal.Width + Petal.Length + Petal.Width +
    Species, data = iris)
Residuals:
    Min
                  Median
-0.79424 -0.21874 0.00899 0.20255 0.73103
Coefficients:
                                                                   These values are not added
                 Estimate Std. Error t value Pr(>|t|)
(Intercept)
                  2.17127
                                     7.760 1.43e-12 ***
                                                                   if species = setosa
Sepal.Width
                  0.49589
                                      5.761 4.87e-08 ***
Petal.Length
                  0.82924
                            0.06853 12.101 < 2e-16
                                                                   (i.e. setosa is set as a reference)
                            0.15120 -2.084 0.03889 *
Petal.Width
                 -0.31516
Speciesversicolor -0.72356
                            0.24017 -3.013 0.00306 **
Speciesvirginica
                 -1.02350
                            0.33373 -3.067 0.00258 **
Signif. codes: 0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1
Residual standard error: 0.3068 on 144 degrees of freedom
Multiple R-squared: 0.8673, Adjusted R-squared: 0.8627
F-statistic: 188.3 on 5 and 144 DF, p-value: < 2.2e-16
```

Inhibit Interpretation/Conversion of Objects

 Function I() in the regression formula allows you to modify/calculate a variable before being fed into the model

```
#### Inhibit Interpretation/Conversion of Objects ####

182  mlog <- lm(Sepal.Length ~ I(log10(1+Petal.Width)), iris)

183  print(mlog)

184  summary(mlog)

185

186  msq <- lm(Sepal.Length ~ I(Petal.Width^2), iris) # Quadratic model

187  print(msq)

188  summary(msq)</pre>
```

Interaction

- Interaction between variables could be added to the model
- For example: male and female might have difference slope of height ~
 weight
- Any interaction must be determined whether it is important or relevant first. Do not rely solely on statistics/statisticians!
- If a interaction is kept in the model, all interacting variables must also be kept regardless of p-values.

```
190 → #### Interaction ####
191 mint <- lm(Sepal.Length ~ Petal.Width*Species, iris)
192 print(mint)
193 summary(mint)</pre>
```

Model selection

- Keep adding variables with limited improvement in performance is not optimal.
 - Simplest model with optimal performance
- Akaike's An Information Criterion (AIC) measures a trade-off between adding variables and performance improvement

```
195 ▼ #### Model selection ####
196    library(MASS)
197
198    sm <- stepAIC(mint)
199    print(sm)
200    summary(sm)</pre>
```

Model selection

- ANOVA could be used to compare two models whether a sufficient improvement is observed in the more complex model.
- $P < 0.05 \rightarrow Improvement$

```
202 - #### Model selection ANOVA ####
203
204 anova(m,mint)
205 anova(m,mm)
```

Prediction with linear regression

New dataset must have the identical structure as the training dataset

```
218 = #### Prediction with linear regression ####
219  newdat <- iris[4:9,]
220  print(newdat)
221
222  predict(mm, newdata = newdat)</pre>
```

Practical 4

Practical 4

- We will perform multivariate regression analysis on the "Davis" dataset
- We will determine how sex, height and interaction between sex and height affect the weight
- Select your best model

```
1 library(car)
2
3 ?Davis
4
5 #Correct errors in the dataset
6 print(Davis[10:13,])
7
8 nDavis <- Davis
9 nDavis[12,2:3] <- Davis[12,3:2]
10
11 #Use nDavis dataset, not Davis</pre>
```

Logistic Regression

Binary outcomes

- Linear regression → numerical outcome
- Logistic regression → binary outcome (e.g. cancer vs no cancer)
- Regression results in (-∞, ∞)
- Probability of binary outcome is [0,1]
- Convert probability to a real number scale
 - Log of odds → Logit

Binary outcomes

- Convert probability to a real number scale
 - Log of odds → Logit

```
207 - #### Logit ####
     library(ggplot2)
208
209
210
     pt <- 10^4 #Try changing number of point
211
     dat <- data.frame(id = 1:pt, prob = seq(0,1,length.out = pt))</pre>
212
213
     ggplot(dat,aes(x = id, y = prob))+
       geom_line()
214
215
     dat$odds <- dat$prob/(1-dat$prob)
216
     ggplot(dat,aes(x = id, y = odds))+
217
218
       geom_line()
219
220
     dat$logit <- log(dat$odds)</pre>
221
     ggplot(dat,aes(x = id, y = logit))+
222 geom_line()
```

Setting logistic regression model

- Setting and selecting logistic regression models are similar to linear regression
- The difference is the interpretation of the result

```
235 - #### Setting a logistic regression model ####
     #From https://towardsdatascience.com/how-to-do-logistic-regression-in-r-456e9cfec7cd
     library(AER)
237
     data(Affairs)
238
     ?Affairs
239
240
241
     Affairs$ynaffair[Affairs$affairs > 0] <- 1
     Affairs$ynaffair[Affairs$affairs == 0] <- 0
     Affairs$ynaffair <- factor(Affairs$ynaffair,levels=c(0,1), labels=c("No","Yes"))
243
244
     table(Affairs$ynaffair)
245
     lgm <- glm(ynaffair ~ gender + age + yearsmarried + children
246
247
                + religiousness + education + occupation +rating,
                data=Affairs, family="binomial")
248
249
     print(lgm)
     summary(lgm)
```

Setting logistic regression model

• The difference is the interpretation of the result

religiousnes

education

rating

occupation

AIC: 627.51

-0.32472

0.02105

0.03092

-0.46845

Number of Fisher Scoring iterations: 4

```
Call:
glm(formula = ynaffair ~ gender + age + yearsmarried + children +
    religiousness + education + occupation + rating, family = "binomial",
    data = Affairs)
Deviance Residuals:
    Min
                  Median
                                        Max
-1.5713 -0.7499 -0.5690 -0.2539
                                    2.5191
Coefficients:
              Estimate Std. Error z value Pr(>|z|)
(Intercept)
              1.37726
                          0.88776
                                    1.551 0.120807
                          0.23909
gendermale
               0.28029
                                    1.172 0.241083
              -0.04426
                          0.01825
                                   -2.425 0.015301 *
               0.09477
                          0.03221
                                    2.942 0.003262 **
yearsmarried
childrenves
               0.39767
                          0.29151
                                    1.364 0.172508
```

-3.618 0.000297 ***

0.417 0.676851

0.431 0.666630

-5.153 2.56e-07

0.08975

0.05051

0.07178

0.09091

(Dispersion parameter for binomial family taken to be 1)

Null deviance: 675.38 on 600 degrees of freedom Residual deviance: 609.51 on 592 degrees of freedom

Signif. codes: 0 '***' 0.001 '**' 0.01 '*' 0.05 '.' 0.1 ' ' 1

For reporting, coefficients must be converted to odds ratio

P-values are still used to determine the significance of each variable

Reporting odds ratio from regression results

```
252 → #### Reporting odds ratio from regression results ####
253
254 exp(cbind(OR = coef(lqm), confint(lqm)))
                     OR
                            2.5 %
                                       97.5 %
(Intercept)
              3.9640180 0.7013467 22.9148790
gendermale
                                   2.1209988
              1.3235091 0.8294178
              0.9567099 0.9223032
                                    0.9908864
age
yearsmarried 1.0994093 1.0326203
                                   1.1718726
childrenyes
             1.4883560 0.8451473
                                    2.6584834
religiousness 0.7227292 0.6049441
                                    0.8605325
education
              1.0212740 0.9254481
                                    1.1284991
occupation
             1.0314027 0.8964089
                                   1.1884105
rating
              0.6259691 0.5227302
                                    0.7470069
```

Prediction with logistic regression

- New dataset must have the identical structure as the training dataset
- Use type = "response" argument to predict a probability of the event

```
256 ~ #### Logistic regression prediction ####
257
258 newdat <- Affairs[c(4,5,9,10),]
259 predict(lgm, newdata = newdat, type = "response")</pre>
```

Practical 5

Practical 5

- We will analyze "Affairs" dataset
- Select the best logistic regression model that predict an affair (e.g. anova, stepAIC)
- If needed, format the type of each variable (e.g. factor, numeric) before fitting a model

```
#From https://towardsdatascience.com/how-to-do-logistic-regression-in-r-456e9cfec7cd
library(AER)
data(Affairs)

?Affairs

Affairs$ynaffair[Affairs$affairs > 0] <- 1
Affairs$ynaffair[Affairs$affairs == 0] <- 0
Affairs$ynaffair <- factor(Affairs$ynaffair,levels=c(0,1), labels=c("No","Yes"))
table(Affairs$ynaffair)</pre>
```

In-class assignment

Early Stage Diabetes Risk Prediction

- Dataset from: https://doi.org/10.24432/C5VG8H
- Create and select the best logistic regression model that predict diabetes
- Report odds ratios from the selected model
- Challenge: Using the selected model to predict the "diabetes probability" (i.e. 0-1) of a 45-year-old male with sudden weight loss, visual blurring and obesity.