Hypothesis testing

Descriptive analysis and basic statistics in biomedical studies using R and Markdown

Juan R Gonzalez juanr.gonzalez@isglobal.org

BRGE - Bioinformatics Research Group in Epidemiology ISGlobal - Barcelona Institute for Global Health http://brge.isglobal.org Hypothesis testing

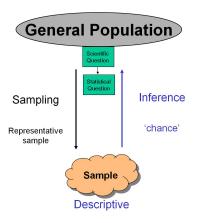


Figure 1: Scheme scientific research

Tests (continuous variables)

One sample test

```
t.test(df$peso, mu=4)
```

```
One Sample t-test

data: df$peso

t = -8.4635, df = 27, p-value = 4.471e-09

alternative hypothesis: true mean is not equal to 4

95 percent confidence interval:

3.016260 3.400169

sample estimates:
mean of x

3.208214
```

► Two independent sample test

```
t.test(peso ~ sexo, data=df)
```

Welch Two Sample t-test

data: peso by sexo

t = 0.39385, df = 25.82, p-value = 0.6969

alternative hypothesis: true difference in means is not equal to 0

95 percent confidence interval:

-0.3024945 0.4458278

sample estimates:

mean in group niño mean in group niña

3.249167 3.177500

Paired t-test

```
t.test(df$horas_an, df$horas_de, paired = TRUE)
```

Paired t-test

► ANOVA (more than 2 groups)

```
Df Sum Sq Mean Sq F value Pr(>F)
naci_ca 2 0.818 0.4092 1.704 0.203
sexo 1 0.035 0.0352 0.147 0.705
Residuals 24 5.763 0.2401
```

Post-hoc

None

P value adjustment method: bonferroni

```
with(df, pairwise.t.test(peso, naci ca , p.adjust="none"))
   Pairwise comparisons using t tests with pooled SD
data: peso and naci_ca
         Española Otras
         0.339
Otras
Sudamérica 0.078 0.461
P value adjustment method: none
  Bonferroni
with(df, pairwise.t.test(peso, naci_ca, p.adjust="bonf"))
   Pairwise comparisons using t tests with pooled SD
data: peso and naci_ca
         Española Otras
         1.00
Otras
Sudamérica 0.23
             1.00
```

Tukey

TukeyHSD (mod)

```
Tukey multiple comparisons of means
   95% family-wise confidence level
Fit: aov(formula = peso ~ naci_ca + sexo, data = df)
$naci ca
                         diff
                                     lwr
                                               upr
                                                       p adj
Otras-Española
                   0.2171429 -0.3493323 0.7836181 0.6102379
Sudamérica-Española 0.4100000 -0.1564752 0.9764752 0.1885514
Sudamérica-Otras
                   0.1928571 -0.4612521 0.8469664 0.7446189
$sexo
                 diff
                            lwr
                                               p adj
                                       upr
niña-niño -0.07166667 -0.4578847 0.3145514 0.7051077
```

Outliers

Analyzing data with outliers may influence the value of a (non-robust) statistic. We can test the null-hypothesis that a variable does not contain **an** outlier. Under the assumption that the data are realizations of one and the same distribution, such a hypothesis can be tested by the Grubbs (1950) test. This test is based on the statistic $g = |suspectvalue - \bar{x}|/s$, where the suspect value is included for the computation of the mean \bar{x} and the standard deviation s.

```
library(outliers)
grubbs.test(df$peso)
```

```
Grubbs test for one outlier

data: df$peso
G = 2.52870, U = 0.75441, p-value = 0.1026
alternative hypothesis: highest value 4.46 is an outlier
```

Since the p-value is not lower than 0.05, the conclusion is that there are no evidences to reject the null- hypothesis of no outliers.

Non-parametric tests

► Wilcoxon test (U Mann Withney)

```
wilcox.test(peso ~ sexo, data=df)
```

Wilcoxon rank sum test with continuity correction

```
data: peso by sexo W = 108.5, p-value = 0.5771 alternative hypothesis: true location shift is not equal to 0
```

Krusdall-Wallis (More than two groups)

```
kruskal.test(peso ~ naci_ca, data=df)
```

```
Kruskal-Wallis rank sum test
data: peso by naci_ca
Kruskal-Wallis chi-squared = 4.6217, df = 2, p-value = 0.09917
```

Two proporions

► Chi-square test

```
freq <- with(df, table(sexo, tip_par))
chisq.test(freq)</pre>
```

```
Pearson's Chi-squared test with Yates' continuity correction
data: freq
X-squared = 1.6106e-31, df = 1, p-value = 1
```

► Fisher test

fisher.test(freq)

sample estimates: odds ratio 0.8710761

```
Fisher's Exact Test for Count Data

data: freq
p-value = 1
alternative hypothesis: true odds ratio is not equal to 1
95 percent confidence interval:
0.06160374 9.21621060
```

freq <- with(df, table(sexo, tip_par))</pre>

Correlation

Pearson correlation test.

Pearson's product-moment correlation

```
cor.test(df$peso, df$edad)
```

Spearman correlation test

Spearman's rank correlation rho

```
cor.test(df$peso, df$edad, method="spearman")
```

```
data: df$peso and df$edad
S = 5678.9, p-value = 0.002215
alternative hypothesis: true rho is not equal to 0
sample estimates:
    rho
-0.5541522
```

Grouping variable	Categorical	Numerical
Categorical	Chi-squared test	t-test
Numerical	Logistic regression	Correlation, Lineal Models, Non-lineal models,

Figure 2: Statistical Tests

# Groups	Parametric (Normal)	Non-parametric
1 ó 2 paired samples	Paired t-test	Wilcoxon (repetead measurements)
2	t-test	U Mann-Whitney (independient)

Figure 3: Testing means

Permutation tests (Advanced)

Some times parametric tests cannot be applied, since there is not known distribution. Therefore, Monte Carlo-based methods can be used instead:

• Carrier of susceptibility allele risk

Cases controls

Figure 4: Permutation testing

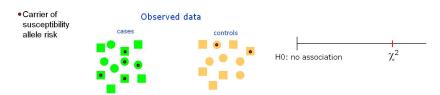


Figure 5: Permutation testing

 Carrier of susceptibility allele risk

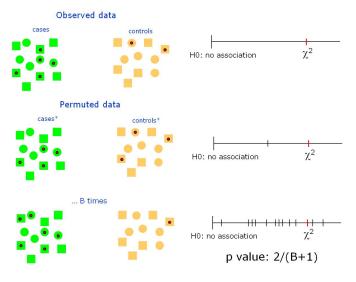


Figure 6: Permutation testing

Let's assume that you are interested in knowing whether there are differences in the *median absolute deviation* (|x - median(x)|) of breastfeeding weeks depending on the type of visits (intensive vs standard). How will you get a p-value?

df\$tx

Regimen visitas asignado

- [1] Intensivo Intensivo Estándar Intensivo Estándar Estándar Intensivo
- [8] Intensivo Estándar Intensivo Estándar Intensivo Estándar Intensivo
- [15] Estándar Estándar Intensivo Intensivo Estándar Intensivo Estándar
- [22] Intensivo Estándar Intensivo Estándar Intensivo Intensivo Estándar Levels: Estándar Intensivo

tx.r

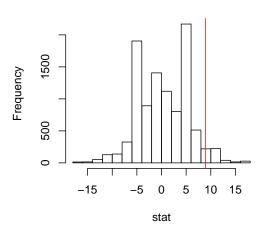
Regimen visitas asignado

- [1] Estándar Intensivo Estándar Intensivo Intensivo Estándar Intensivo
- [8] Intensivo Estándar Intensivo Intensivo Intensivo Estándar Intensivo
- [15] Intensivo Estándar Estándar Intensivo Estándar Estándar Estándar
- [22] Estándar Intensivo Estándar Intensivo Intensivo Intensivo Estándar Levels: Estándar Intensivo

[1] 0.03049695

```
hist(stat)
abline(v=stat.ref, col="red")
```

Histogram of stat

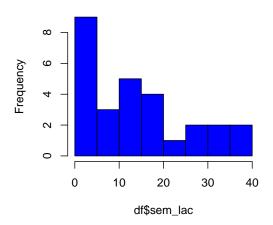


t.test(sem lac ~ tx, data=df)

Welch Two Sample t-test

```
data: sem_lac by tx
t = -3.7597, df = 25.083, p-value = 0.0009121
alternative hypothesis: true difference in means is not equal to 0
95 percent confidence interval:
-21.009082 -6.139636
sample estimates:
mean in group Estándar mean in group Intensivo
6.692308 20.266667
```

Histogram of df\$sem_lac



Session info

sessionInfo()

```
R version 3.4.1 (2017-06-30)
Platform: x86 64-w64-mingw32/x64 (64-bit)
Running under: Windows 10 x64 (build 16299)
Matrix products: default
locale:
[1] LC_COLLATE=Spanish_Spain.1252 LC_CTYPE=Spanish_Spain.1252
[3] LC_MONETARY=Spanish_Spain.1252 LC_NUMERIC=C
[5] LC TIME=Spanish Spain.1252
attached base packages:
              graphics grDevices utils
[1] stats
                                            datasets methods
                                                                 base
other attached packages:
[1] outliers 0.14 Hmisc 4.0-3
                                    ggplot2_2.2.1
                                                   Formula 1.2-2
[5] survival 2.41-3 lattice 0.20-35
loaded via a namespace (and not attached):
 [1] Rcpp 0.12.12
                         compiler_3.4.1
                                             pillar 1.1.0
 [4] RColorBrewer 1.1-2 plvr 1.8.4
                                             base64enc 0.1-3
 [7] tools 3.4.1
                         rpart 4.1-11
                                             digest 0.6.12
[10] evaluate 0.10.1
                         tibble 1.4.2
                                             gtable 0.2.0
[13] htmlTable 1.9
                         checkmate 1.8.3
                                             rlang 0.1.6
[16] Matrix 1.2-10
                         yaml 2.1.16
                                             gridExtra 2.3
[19] stringr 1.3.0
                         knitr 1.20
                                             cluster 2.0.6
[22] htmlwidgets 0.9
                         rprojroot 1.3-2
                                             grid 3.4.1
                                             foreign_0.8-69
[25] nnet 7.3-12
                         data.table_1.10.4
[28] rmarkdown 1.8
                         latticeExtra 0.6-28 magrittr 1.5
```