### Hypotheses tests with continuous variables

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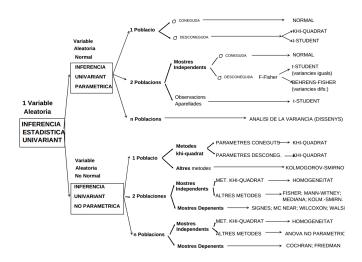
#### **Outline**

- INTRODUCTION
- TYPE OF TESTS
- NORMALITY TESTS
- ONE GROUP COMPARISON
- **5** TWO GROUPS COMPARISON IN INDEPENDENT SAMPLES
- **10** TWO GROUPS COMPARISON IN DEPENDENT SAMPLES
- K GROUPS COMPARISON IN INDEPENDENT SAMPLES
- Multiple comparisons and multiple testing

#### Introduction

- Once the concept of hypothesis testing is established,
- Researchers face the problem of which test should be applied at every possible situation.
- Best solution:
  - understand the problem and the questions addressed
  - know available tests for each problem
  - be aware of applicability assumptions of each test and how to check them.
- Easier to say than to do.
  - Sometimes cheatsheets may be helpful, but be warned against a blind use, that is understand and be critic with the steps.

## Which test is appropriate for which problem



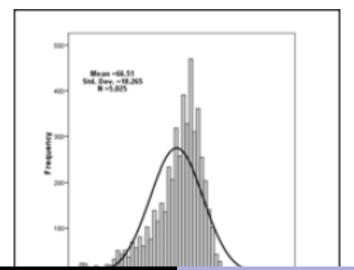
### **Example situation**

- A study was designed to compare two distinct hypertension control programs.
- 60 individuals with HTA were randomly assigned to either one or the other group (30 per group)
- Blood pressure was measured each month during a year

```
with(hta, hist(tas1) )

with(hta, qqnorm(tas1, main="Normal QQplot")); with(hta, qqline(tas1))

par(oldpar) # Vuelve a los parámetros de dibujo originales
```



### **Normality Test**

```
with(hta,shapiro.test(tad1) ) # Shapiro Wilk test

##
## Shapiro-Wilk normality test
##
## data: tad1
## W = 0.96622, p-value = 0.09512
```

### One sample Test

```
with(hta,t.test(tad1,mu=90)) # One sample T.test
##
##
   One Sample t-test
##
## data: tad1
## t = -1.2137, df = 59, p-value = 0.2297
## alternative hypothesis: true mean is not equal to 90
## 95 percent confidence interval:
## 85.80626 91.02707
## sample estimates:
## mean of x
## 88.41667
```

### Homogeneity variance Test

```
library(car)
hta%>%
 group_by(sexo) %>%
 summarise(var = sd(tas1))
## # A tibble: 2 x 2
## sexo
            var
## <chr> <dbl>
## 1 MUJER 17.6
## 2 VARON 22.1
with(hta,leveneTest(tad1~factor(sexo),center="median"))
## Levene's Test for Homogeneity of Variance (center = "median")
        Df F value Pr(>F)
##
## group 1 1.3506 0.2499
        58
##
```

- p value is over 0.05
- We can assume homogeneity of variances

## T test when variances are equal

```
with(hta,t.test(tas1~factor(sexo),var.equal=TRUE ))
##
##
   Two Sample t-test
##
## data: tas1 by factor(sexo)
## t = -0.2471, df = 58, p-value = 0.8057
## alternative hypothesis: true difference in means between grou
## 95 percent confidence interval:
## -11.603461 9.053519
## sample estimates:
## mean in group MUJER mean in group VARON
##
              149.5946
                                  150.8696
```

- Type I Error is over than 0.05
- We cannot reject mean equality

## T test when variances are unequal

```
with(hta,t.test(tas1~factor(sexo),var.equal=FALSE ))
##
##
   Welch Two Sample t-test
##
## data: tas1 by factor(sexo)
## t = -0.23436, df = 39.098, p-value = 0.8159
## alternative hypothesis: true difference in means between grou
## 95 percent confidence interval:
## -12.277927 9.727986
## sample estimates:
## mean in group MUJER mean in group VARON
              149.5946
                                  150.8696
##
```

- Same conclusions as before
- Test is also known as Welch test

## U Mann-Whitney or Sum Rank non parametric test

```
with(hta, wilcox.test(tad1~factor(sexo)
    ,alternative='two.sided',exact=TRUE, correct=FALSE))
##
    Wilcoxon rank sum test
##
##
## data: tad1 by factor(sexo)
## W = 434, p-value = 0.8955
## alternative hypothesis: true location shift is not equal to 0
hta%>%
  group_by(sexo) %>%
  summarise(median = median(tad1))
## # A tibble: 2 x 2
## sexo median
## <chr> <dbl>
## 1 MUJER
               90
## 2 VARON
              90
```

Null Hypothesis cannot be rejected

#### Paired T-test

```
with(hta,t.test(tas1,tas12,paired=TRUE))
##
##
   Paired t-test
##
## data: tas1 and tas12
## t = 6.0672, df = 51, p-value = 1.609e-07
## alternative hypothesis: true mean difference is not equal to
## 95 percent confidence interval:
## 8.518285 16.943253
## sample estimates:
## mean difference
         12.73077
##
summary(hta$tas1)
##
     Min. 1st Qu. Median Mean 3rd Qu. Max.
##
    100.0 140.0 145.0 150.1 160.0 210.0
summary(hta$tas12)
```

## Paired Sign-Rank Wilcoxon Test

#### Read diabetes data

```
library(readx1)
library(dplyr)
library(magrittr)
diabetes <- read_excel("datasets/diabetes.xls")</pre>
sapply(diabetes, class)
##
     numpacie
                     mort tempsviu
                                             edat.
                                                         bmi
                                                                edatdiag
   "numeric" "character" "numeric" "numeric"
                                                  "numeric"
                                                               "numeric"
##
        tabac
                      sbp
                                  dbp
                                              ecg
                                                         chd
                          "numeric" "character" "character"
## "character" "numeric"
diabetes factor <- diabetes %>%
 mutate_if(sapply(diabetes, is.character), as.factor) %>%
 select (-numpacie)
diabetes%>%
 group_by(ecg) %>%
 summarise( n=n().
   mean = mean(edat),
           sd=sd(edat))
## # A tibble: 3 x 4
    ecg
                 n mean
                            sd
   <chr> <int> <dhl> <dhl>
## 1 Anormal 11 64.9 6.76
## 2 Frontera 27 53.8 11.4
## 3 Normal 111 50.5 11.5
```

#### **ANOVA**

anova <- aov (edat~ecg, data=diabetes\_factor)

### Multicomparison

```
library(multcomp)
tuk <- glht(anova, linfct = mcp(ecg = "Tukey"))</pre>
 print(summary(tuk)) # pairwise tests
##
    Simultaneous Tests for General Linear Hypotheses
##
##
## Multiple Comparisons of Means: Tukey Contrasts
##
##
## Fit: aov(formula = edat ~ ecg. data = diabetes factor)
##
## Linear Hypotheses:
##
                          Estimate Std. Error t value Pr(>|t|)
## Frontera - Anormal == 0 -11.094 4.010 -2.767 0.016461 *
## Normal - Anormal == 0 -14.405 3.543 -4.065 0.000217 ***
## Normal - Frontera == 0 -3.310 2.405 -1.376.0.345706
## ---
## Signif. codes: 0 '***' 0.001 '**' 0.05 '.' 0.1 ' ' 1
## (Adjusted p values reported -- single-step method)
```

```
print(confint(tuk, level=0.95)) # confidence intervals
##
##
    Simultaneous Confidence Intervals
##
## Multiple Comparisons of Means: Tukey Contrasts
##
##
## Fit: aov(formula = edat ~ ecg, data = diabetes_factor)
##
## Quantile = 2.3458
## 95% family-wise confidence level
##
##
## Linear Hypotheses:
##
                           Estimate lwr
                                             upr
## Frontera - Anormal == 0 -11.0943 -20.5004 -1.6882
## Normal - Anormal == 0 -14.4046 -22.7168 -6.0923
```

## Normal - Frontera == 0 -3.3103 -8.9531 2.3325

# Multicomparison plot

plot(confint(tuk))

#### Kruskal-Wallis Test

```
diabetes factor%>%
 group_by(ecg) %>%
 summarise(median = median(edat))
## # A tibble: 3 x 2
## ecg median
## <fct> <dbl>
## 1 Anormal 64
## 2 Frontera 53
## 3 Normal 49
kruskal.test(edat~ecg,data=diabetes_factor)
##
##
   Kruskal-Wallis rank sum test
##
## data: edat by ecg
## Kruskal-Wallis chi-squared = 17.483, df = 2, p-value = 0.0001
```

#### **Dunn Test for multiple comparison**

```
library(dunn.test)
with(diabetes factor,dunn.test(edat,ecg,method="bonferroni"))
##
    Kruskal-Wallis rank sum test
##
## data: edat and ecg
  Kruskal-Wallis chi-squared = 17.4826, df = 2, p-value = 0
##
##
##
                              Comparison of edat by ecg
##
                                    (Bonferroni)
## Col Mean-I
## Row Mean | Anormal Frontera
## Frontera | 2.721182
##
                0.0098*
##
##
    Normal |
              4.075469 1.467464
##
                0.0001*
                            0.2134
##
```