Instructions for computer-use in solving the proposed practical activities

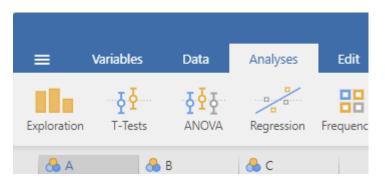
| Instructions for using Jamovi for data analysis | 3 |
|--|----|
| Importing the database | 3 |
| Data type checking | 3 |
| Description of a qualitative variable – frequency table, column graph | 4 |
| Analysis between 2 qualitative variables – Chi square test, Fisher test, contingency table, OR, RR, RA, column graph | |
| Contingency table | .9 |
| Chi Squared and Fisher exact test Results | 9 |
| Medical indicators of association strength1 | 0 |
| Column chart1 | 0 |
| Student test for two independent groups1 | 1 |
| Evaluation of the Student test application conditions – normality1 | 1 |
| Conducting the Student Test for Independent Samples1 | 17 |
| Choosing between the Equal Variance Student Test and the Welch Test – Unequal Variance 1 | 9 |
| Descriptive statistics2 | 20 |
| Numeric | 20 |
| Graph of means2 | 20 |
| Diagnostic Tests2 | 21 |
| ROC analysis2 | 25 |
| Tables of limit values and associated statistics | 28 |
| ROC curve graph2 | 28 |
| Comparison of ROC curves by statistical tests2 | 29 |
| Survival analysis (Kaplan Meyer plot, log-rank test, Cox – HR regresion) | 31 |
| The median survival time3 | 32 |
| The survival probability at different times3 | 33 |

| The Kaplan-Meier survival curve, with the log-rank test | 34 |
|--|----|
| Meta-analysis | 34 |
| Meta-analysis to compare two groups in terms of dichotomous data | 34 |
| Association estimator | |
| Statistical heterogeneity of studies | 36 |
| Forest plot | |
| Publication bias | |
| Installation of additional analysis modules (e.g. for ROC analysis, survival, diagnosis) | |
| Instructions for using EpiInfo and Excel for data analysis | 40 |
| Downloading <i>EpiInfo</i> (for home use) | 40 |
| Starting Epi Info | 40 |
| Analyzing data in Epi Info | 40 |
| Importing an Excel file in Epi Info | 42 |
| Activating the Data Analysis module in Microsoft Excel | 45 |
| Descriptive statistics | 45 |
| Qualitative (categorical) data: | 45 |
| Quantitative data: | 48 |
| Individual description of quantitative variables | 48 |
| Description of a potential relation between two quantitative variables | 50 |
| Survival data: | 51 |
| Data Analysis | 52 |

Instructions for using Jamovi for data analysis

Importing the database

Use the **hamburger** menu icon .



Then select the **Open** option



Afterwards, search for the database folder by pressing the **Browse button**, select the desired file and press the **Open button** to import it.

| 2 | jamovi | ↑ Documents 🖆 Browse |
|------|---------|-------------------------|
| | Jannobr | Q |
| | | .virtualenvs |
| New | | Custom Office Templates |
| Open | | . |

Data type checking

Before starting the analysis, it is necessary to check whether the program has correctly classified the variables in the database. The Activity variable is a qualitative variable, and the other 4 variables are quantitative. To check the type of a variable, **double-click** on **the column title** (variable name). In the case of the Activity variable we ensure that in the **Measure** selection area **type** option is selected **Nominally**. For the other quantitative variables we ensure that in the Measure selection area type option **Continuous** is selected .

| data variable | | | | |
|-------------------|---------------|----------------------|---------------|--------------|
| Activitate | | | | |
| Description | | | |] |
| Measure type Nomi | nal 🔻 🐣 | Levels | | \uparrow |
| Data type Text | ~ | da | | \downarrow |
| Missing values | | nu | | |
| | | | | |
| | | | | ÷ |
| | | Retain unused levels | in analyses 🔵 | |
| | | - | | |
| | data variabli | E | | |
| | CHIT1 | | | |
| | Description | | | |
| | Measure type | Continuous 🗸 🤌 | | |

Description of a qualitative variable – frequency table, column graph

In the Analyses tab, the Exploration button, choose the Descriptives option.

| | Variables |
|-------------|--------------|
| Exploration | ₹ T-Tests |
| Descriptive | es |

The variable of interest (e.g. **disease**, or **risk factor**), should be moved by pressing the arrow next to the **Variables** field.

| Descriptives | | \ni |
|--------------|-----------------------------------|-------|
| ✓ Id Sepsis | Q Variables → Chorioamniotitis | |
| | Split by | |

To display the frequency table, check the **Frequency tables** option.

| Descriptives | Variables across columns \checkmark | 🗸 Frequency tables 🧲 | lla d |
|--------------|---------------------------------------|----------------------|-------|
|--------------|---------------------------------------|----------------------|-------|

A frequency table similar to the one below is obtained:

Frequencies

Frequencies of Chorioamniotitis

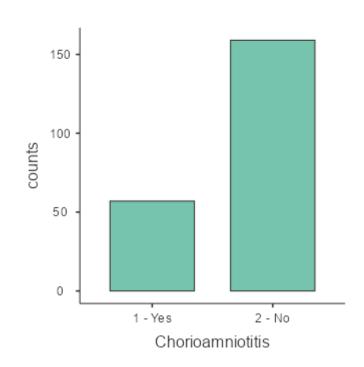
| Chorioamniotitis | Counts | % of Total | Cumulative % |
|------------------|--------|------------|--------------|
| 1 - Yes | 57 | 26.4 % | 26.4 % |
| 2 - No | 159 | 73.6 % | 100.0 % |

To display a column chart, check the **Bar plot** option.

| > Statistics | | |
|----------------|----------------|------------|
| ✓ Plots | | |
| Histograms | Box Plots | Bar Plots |
| Histogram | Box plot | 🗹 Bar plot |
| Density | Label outliers | |
| Q-Q Plots | Violin | |
| Q-Q | 🗌 Data | |
| | Jittered 🗸 | |
| | Mean | |

A column chart is obtained:

Plots



Chorioamniotitis

Analysis between 2 qualitative variables – Chi square test, Fisher test, contingency table, OR, RR, RA, column graph

In the Analyses tab, the Frequencies module one can choos the option Independent Samples (X² test of association).

| | ≡ Variable | s Data | Analyses | Edit | |
|----|-----------------|--------|------------|-----------------|---|
| Ex | ploration T-Tes | ITI | Regression | Frequencies | Factor Surv |
| | NrCrt. | 🔒 Sex | 🔒 Aspirina | One Sample Pro | oportion Tests |
| 1 | 1 | m | nu | 2 Outcome | |
| 2 | 2 | m | nu | | Binomial test |
| 3 | 3 | m | da | N Outcom | |
| 4 | 4 | f | nu | | χ² Goodness of fit |
| 5 | 5 | m | nu | Contingency Ta | bles |
| 6 | 6 | f | da | | |
| 7 | 7 | m | da | Independe | ent Samples x ² test of association |
| 8 | 8 | m | nu | Deline of Court | |
| 9 | 9 | f | nu | Paired Sam | McNemar test |
| 10 | 10 | f | nu | | |
| 11 | 11 | m | nu | Log Linear | Regression |
| 12 | 12 | m | nu | Log-Linear | Regression |

Grouping variable (e.g. **risk factor**, or **treatment**), should be moved by pressing the arrow next to the **Rows field**.

The variable that is of interest (e.g. **disease**, or **complication**, or **treatment result**), should be moved by pressing the arrow next to the **Columns field**.

As options we recommend the following choices: Tests: **X2**, **Fisher's exact test**; Comparative measures (2x2 only): **Odds ratio**, **Relative risk**, Attributable risk/absolute risk reduction (**Difference in proportion**), **Confidence intervals**; Counts: **Observed counts**, **Expected counts**; Percentages: **Row**; Plots: column (**Bar Plot**), **Y-Axis**: percentages, **within rows**; Bar **Type**, Stacked; **X-Axis**: **Rows**.

| Contingency Tables | \ominus |
|--|---|
| Id Gender D1Ulceration (mm) D2Ulceration (mm) D1-D2 (mm) | Rows → Aspirin Aspirin Columns → Pa Recovery Counts (optional) → Layers → |
| ✓ Statistics | |
| Tests | Comparative Measures (2x2 only) |
| ✓ χ ² | 🗸 Odds ratio |
| χ^{z} continuity correction | Log odds ratio |
| Likelihood ratio | Relative risk |
| ✓ Fisher's exact test | Difference in proportions |
| z test for difference in 2 proportions | s 🔽 Confidence intervals |
| Hypothesis | Interval 95 % |
| Oroup 1 ≠ Group 2 | Compare rows 🗸 |
| Group 1 > Group 2 | |
| Group 1 < Group 2 | |

| ✓ Cells | |
|-----------------------------|-------------------------|
| Counts | Percentages |
| Observed counts | Row |
| Expected counts | Column |
| | Total |
| ✓ Plots | |
| Plots | Y-Axis |
| 🗸 Bar Plot | Ocounts |
| Bar Type | Percentages within rows |
| O Side by side | X-Axis |
| Stacked | - |
| | Rows |
| | Columns |

Contingency table

| Contingency Tables | | | | | |
|--------------------|--------------|--------|-----------|---------|--|
| | | Vinde | Vindecare | | |
| Aspirina | | da | nu | Total | |
| da | Observed | 182 | 18 | 200 | |
| | Expected | 162 | 38.0 | 200 | |
| | % within row | 91.0 % | 9.0 % | 100.0 % | |
| nu | Observed | 142 | 58 | 200 | |
| | Expected | 162 | 38.0 | 200 | |
| | % within row | 71.0 % | 29.0 % | 100.0 % | |
| Total | Observed | 324 | 76 | 400 | |
| | Expected | 324 | 76.0 | 400 | |
| | % within row | 81.0 % | 19.0 % | 100.0 % | |

Chi Squared and Fisher exact test Results

The value of p, corresponding to the tests, is in the column p.

| - | | | |
|-------|----|----|----|
| - × - | Τ- | | - |
| γ | 12 | 51 | ς. |
| Λ. | | | - |

| | Value | df | р |
|---------------------|-------|----|--------|
| χ² | 26.0 | 1 | < .001 |
| Fisher's exact test | | | < .001 |
| N | 400 | | |

Medical indicators of association strength

Medical indicator results are presented in the **Comparative Measures** table, where the point estimator is in the **Value** column, and the ends of the 95% confidence interval are in the **Lower, Upper** (95% Confidence Intervals) columns.

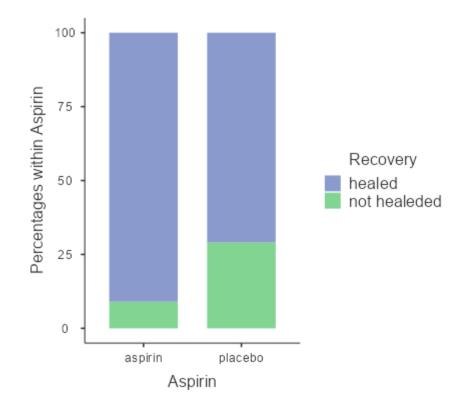
Comparative Measures

| | | 95% Confidence Interva | |
|-----------------------------|--------|------------------------|-------|
| | Value | Lower | Upper |
| Difference in 2 proportions | 0.200ª | 0.126 | 0.274 |
| Odds ratio | 4.13 | 2.33 | 7.32 |
| Relative risk | 1.28 ° | 1.16 | 1.41 |

Rows compared

Column chart

Finally, we have the graphical representation of the association between the two qualitative variables.



Student test for two independent groups

Evaluation of the Student test application conditions – normality

The Student Test can only be applied if the data of both compared groups follow a normal distribution. Various methods can be used to assess normality: data distribution indicators: skewness - asymmetry coefficient, excess kurtosis; tests for normality; graphs (histogram, QQ plot).

Data distribution indicators can be used to learn about the normality of the data. If **both** the skewness **coefficient and** the kurtosis coefficient (**both coefficients**) are **within the range (-1; 1)** it is a suggestion that the data follow a **normal distribution**; if the skewness coefficient **or** the excess kurtosis (**any of the coefficients**) is **outside the range (-1; 1)** we have a suggestion that the data **does not follow a normal distribution**. Example in the table: for variable D2 ulceration (mm) – suggests that data do not follow a normal distribution (skewness: yes=4.30, no=2.54; kurtosis: yes=20.2; no=6.58); for variable D1-D2 (mm) – suggests that data follows a normal distribution (skewness: yes=-0.0718, no=0.119; kurtosis: yes=-0.480; no=-0.403).

One option for assessing the normality of data is with the help of statistical tests to assess normality. If the sample has less than 50 observations we look at the result of the Shapiro-Wilk test, if

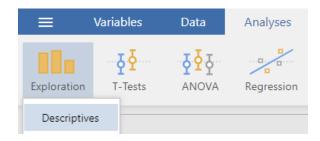
it has more than 50 observations we look at the result of the Kolmogorov-Smirnov test (in case it is available). The outcome of interest is **the p-value** of the test (Shapiro-Wilk p). If the p-value is **less than 0.05**, it is a suggestion that the data does not follow a normal distribution, otherwise if the p-value is greater than 0.05, it is a suggestion that the data follows a normal distribution. Example in the table: for variable D2 ulceration (mm) – suggests that data does not follow a normal distribution (yes: p<0.001, no: p<0.001); for variable D1-D2 (mm) – suggests that data follows a normal distribution (yes: p=0.547, no: p=0.512).

Another **way** to **evaluate the normality of the** data is with the help of graphs, such as the histogram, or the Q-Q-plot graph.

Histogram chart interpretation. If the graph follows an **approximately, relatively symmetrical bell shape**, there is a suggestion that the data follows a **normal distribution**, **otherwise** we have a suggestion that the data **does not follow a normal distribution**. Example in the image: for variable D2 Ulceration (mm) – suggests that data does not follow a normal distribution; for variable D1-D2 (mm) – suggests that data follows a normal distribution.

Interpretation of the Q-Q plot graph. If the tendency of the observations is to be arranged far from the solid line (which represents a normal distribution), then it is a suggestion that the data does not follow a normal distribution, otherwise, if the tendency of the observations is to be close to the solid line, it is a suggestion that the data follow a normal distribution.

The normality evaluation will be done in the **Analysis** tab, the **Exploration** menu, the **Descriptives** command.



In the **Variables** section, the quantitative variables for which normality is to be evaluated (e.g. D2Ulceration (mm), D1-D2 (mm)) are moved. In the **Split by** section, the qualitative variable is moved, which identifies the groups to be compared (e.g. Aspirin).

| Descriptives | | \ni |
|---|--------|---|
| Id Q a Sex a Recovery D1Ulceration (mm) | → | Variables D2Ulceration (mm) D1-D2 (mm) |
| Descriptives Variables across columns 🗸 | → ` | Split by Split by Split by Image: Split by |

In the Statistics section, check Skewness, Kurtosis, Shapiro-Wilk.

| ✓ Statistics | |
|-----------------------------------|-----------------------|
| Sample Size | Central Tendency |
| 🔽 N 🔽 Missing | 🗸 Mean |
| Percentile Values | 🗹 Median |
| Cut points for 4 equal groups | Mode |
| Percentiles 25,50,75 | Sum |
| Dispersion | Distribution |
| 🗸 Std. deviation 🖌 Minimum | 🗸 Skewness |
| 🗌 Variance 🗹 Maximum | 🗸 Kurtosis |
| Range IQR | Normality |
| Mean Dispersion | 🗹 Shapiro-Wilk |
| Std. error of Mean | Outliers |
| Confidence interval for Mean 95 % | Most extreme 5 values |

In the Plots section, check Histogram and Q-Q.

| ✓ Plots | | | |
|-------------|----------------|-----------|--|
| Histograms | Box Plots | Bar Plots | |
| 🗸 Histogram | Box plot | Bar plot | |
| Density | Label outliers | | |
| Q-Q Plots | Violin | | |
| 🗸 Q-Q | Data | | |
| - | Jittered 🗸 | | |
| | Mean | | |

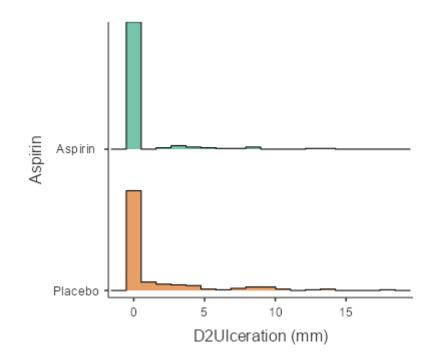
A table with statistics on each variable is obtained, for each group:

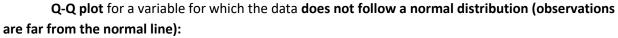
| | Aspirin | D2Ulceration (mm) | D1-D2 (mm) |
|---------------------|---------|-------------------|------------|
| Ν | Aspirin | 200 | 200 |
| | Placebo | 200 | 200 |
| Missing | Aspirin | 0 | 0 |
| | Placebo | 0 | 0 |
| Mean | Aspirin | 0.547 | 23.7 |
| | Placebo | 1.49 | 22.3 |
| Median | Aspirin | 0.00 | 23.6 |
| | Placebo | 0.00 | 21.9 |
| Standard deviation | Aspirin | 1.97 | 3.85 |
| | Placebo | 3.12 | 4.18 |
| Minimum | Aspirin | 0.00 | 13.8 |
| | Placebo | 0.00 | 12.6 |
| Maximum | Aspirin | 13.6 | 32.9 |
| | Placebo | 17.9 | 33.2 |
| Skewness | Aspirin | 4.30 | -0.0718 |
| | Placebo | 2.54 | 0.119 |
| Std. error skewness | Aspirin | 0.172 | 0.172 |
| | Placebo | 0.172 | 0.172 |
| Kurtosis | Aspirin | 20.2 | -0.480 |
| | Placebo | 6.58 | -0.403 |
| Std. error kurtosis | Aspirin | 0.342 | 0.342 |
| | Placebo | 0.342 | 0.342 |
| Shapiro-Wilk W | Aspirin | 0.313 | 0.994 |
| | Placebo | 0.554 | 0.993 |
| Shapiro-Wilk p | Aspirin | < .001 | 0.547 |
| | Placebo | < .001 | 0.512 |

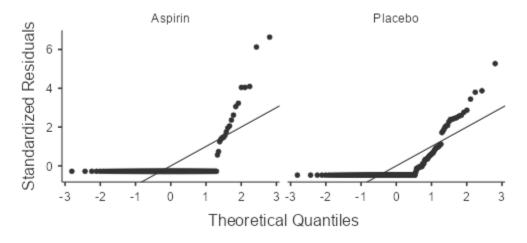
Descriptives

The **histogram** for a variable for which the data **does not follow a normal distribution (marked** right- skewness, marked kurtosis):

D2Ulceration (mm)

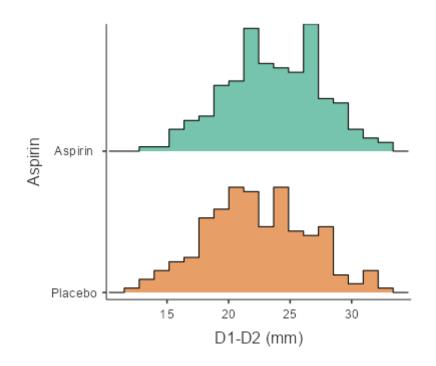




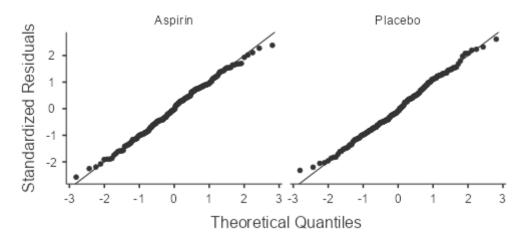


The **histogram** chart for a variable for which the data follows an approximately normal distribution (bell-shaped, relatively symmetrical chart):

D1-D2 (mm)



Q-Q plot for a variable for which the data follows a normal distribution (the observations are close to the normal line):



In the Analyses tab, choose the Independent Samples T-Test option.

Conducting the Student Test for Independent Samples

In the Analyses tab, choose the Independent Samples T-Test option.

| = ' | Variables | Data | Analyses | Edit |
|-------------|--------------|-----------------------|------------|-------------|
| Exploration | ₹ T-Tests | ₽ ₽ ANOVA | Regression | Frequencies |
| | Independe | nt Samples T-1 | Test | |
| Indep | Paired Sam | Paired Samples T-Test | | |
| | One Sampl | e T-Test | _ | |

The qualitative variable that identifies the compared groups should be moved by pressing the arrow next to the Grouping Variable field.

The quantitative variable(s) that are of interest should be moved by pressing the arrow next to the **Dependent Variable field**.

| Independent Samples T-Test | (- | € |
|--|--|---|
| ✓ Id Q Sex Q Recovery ✓ D1Ulceration (mm) ✓ D2Ulceration (mm) | → Dependent Variables → D1-D2 (mm) Grouping Variable → Sa Aspirin | ◆ |

Check the following options:

- 1. in the Tests section: **Student's**, **Welch's section**
- 2. in section Additional statistics: Mean difference, Confidence interval, Descriptives, Descriptive plots.
- 3. In section Assumptions checks: Homogeneity test.

| Tests | Additional Statistics | | |
|------------------------------------|-----------------------|------|---|
| ✓ Student's | 🗸 Mean difference | | |
| Bayes factor | ✓ Confidence interval | 95 % | 6 |
| Prior 0.707 | Effect size | | |
| ✓ Welch's | Confidence interval | 95 % | 6 |
| Mann-Whitney U | Descriptives | | |
| Hypothesis | Descriptives plots | | |
| Oroup 1 ≠ Group 2 | Assumption Checks | | |
| O Group 1 > Group 2 | ✓ Homogeneity test | | |
| O Group 1 < Group 2 | Normality test | | |
| Missing values | Q-Q plot | | |
| Exclude cases analysis by analysis | | | |
| O Exclude cases listwise | | | |

Choosing between the Equal Variance Student Test and the Welch Test - Unequal Variance

In the case of the Student test for independent groups, there are two variants, for the situation where equal variances or unequal variances are assumed. To choose between the two, tests can be used to compare variances. In the Homogeneity of Variances Tests table, we look at the p-value obtained with the Leven test. If the p-value is less than 0.05, it is a suggestion that groups have unequal variances, otherwise it is a suggestion that groups have equal variances.

Assumptions

| Homogeneity of Variances Test (Levene's) | | | | |
|--|------|----|-----|-------|
| | F | df | df2 | р |
| D1-D2 (mm) | 1.17 | 1 | 398 | 0.280 |

Note. A low p-value suggests a violation of the assumption of equal variances

[3]

If **the variances** are assumed to be **equal**, we choose the corresponding results for **the Student test**, otherwise we choose the corresponding ones for the **Welch test** in the **Independent Samples T**-

Test table. In column P we have the p value of the test, in the column Mean difference, we have the difference between the averages of the two groups, and in Lower, Upper (95% Confidence Interval), there is the 95% confidence interval associated with the difference between the averages of the groups.

| Independent Samples T-Test | | | | | | | | |
|----------------------------|-------------|-----------|-----|--------|-----------------|---------------|------------|---------------|
| | | | | | | | 95% Confid | ence Interval |
| | | Statistic | df | р | Mean difference | SE difference | Lower | Upper |
| D1-D2 (mm) | Student's t | 3.44 | 398 | < .001 | 1.38 | 0.402 | 0.591 | 2.17 |

Independent Samples T-Test

Note. H_a µ_{Aspirin} ≠ µ_{Placebo}

Descriptive statistics

Numeric

Descriptive statistics can be found numerically in the **Group Descriptives table**. **The number of subjects** is in column **N**, **mean** in column **Mean**, **standard deviation** in column **SD**.

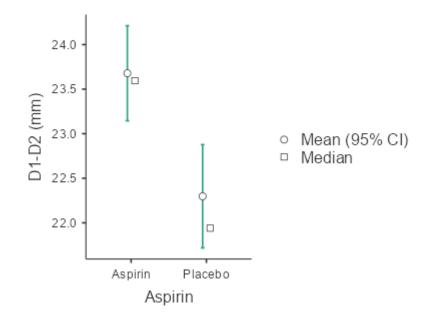
| Group Descriptives | | | | | | |
|--------------------|---------|-----|------|--------|------|-------|
| | Group | Ν | Mean | Median | SD | SE |
| D1-D2 (mm) | Aspirin | 200 | 23.7 | 23.6 | 3.85 | 0.272 |
| | Placebo | 200 | 22.3 | 21.9 | 4.18 | 0.296 |

Graph of means

A graph of means **is also provided**, where the circle represents the mean, error bars represent the 95% confidence interval, and square represents the median.

Plots

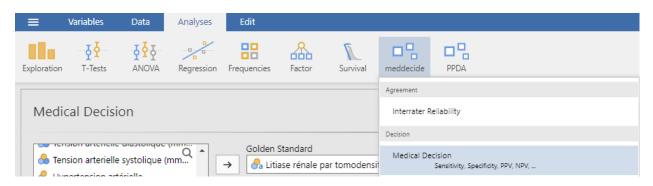




Diagnostic Tests

Make sure that the **Meddecide – Functions for medical decision in ClinicoPath** module is installed. See the chapter **Installing an additional module**.

In the **Analyses** tab, press the **Meddecide** button, from the menu, select the Medical Decision option (Sensitivity, Specificity, PPV, NPV).



Select the variable that represents the standard test (for example, BCC) and press the arrow button next to the Golden Standard field. Specify which category of the variable represents the presence of the disease, as indicated by the standard test (here With), in the Positive Level list. Select the variable that represents the new test (for example, Confocal microscopy) and press the arrow button next to the New Test field. Specify which category of the variable represents a positive test result (here Positive), in the Positive Level list.

| Medical Decision | | | | \ni |
|------------------|---|---|---|----------|
| Id | Q | → | Golden Standard BCC Positive Level With New Test Confocal microscopy Positive Level Positive | <u>م</u> |

Select the following options: in the Table section: Original Data, Footnotes, 95% CI; in the Plots section: Fagan Nomogram.

| Table |
|--|
| 🗸 Original Data |
| Footnotes |
| ✓ 95% CI |
| Prior Probability |
| Prior Probability (prevalence) |
| 0.3 |
| Plots |
| 🗸 Fagan Nomogram |
| Original Data |
| Confocal microscopy With Without Negative 6 20 Positive 57 5 |

| Co | nfocal microscopy | BCC | n |
|----|-------------------|--------|------|
| 1 | Negative | With | 6 |
| 2 | Negative | Withou | t 20 |
| 3 | Positive | With | 57 |
| 4 | Positive | Withou | t 5 |

Check if the data is correctly placed in the table below, compared to the table at the above! If there are any problems, change the Positive Level option.

Recoded Data for Decision Test Statistics

| | Gold Positive | Gold Negative | Total |
|---------------|---------------|---------------|-------|
| Test Positive | 57 | 5 | 62 |
| Test Negative | б | 20 | 26 |
| Total | 63 | 25 | 88 |

Below there are statistics of the totals of the participants, by different categories:

| | n |
|-------------------------|-----------------|
| | |
| Totalundefined | 88ª |
| Diseased undefined | 63 ^b |
| Healthyundefined | 25ª |
| Positive Testsundefined | 62° |
| Negative Testsundefined | 26 ^f |
| True Testundefined | 779 |
| Wrong Testundefined | 11 ^h |

^a Total Number of Subjects

^b Total Number of Subjects with Disease

^d Total Number of Healthy Subjects

^e Total Number of Positive Tests

^f Total Number of Negative Tests

⁹ Total Number of True Test Results

^h Total Number of Wrong Test Results

Below there are the classic statistics of diagnostic tests, with their 95% confidence interval.

| | n |
|-------------------|-----------------|
| Totalundefined | 88ª |
| Diseasedundefined | 63 ^b |
| Healthyundefined | 25₫ |

| Positive Testsundefined | 62 ^e |
|-------------------------|-----------------|
| Negative Testsundefined | 26 ^f |
| True Testundefined | 77 ⁹ |
| Wrong Testundefined | 11 ^h |

^a Total Number of Subjects

- ^b Total Number of Subjects with Disease
- ^d Total Number of Healthy Subjects
- ^e Total Number of Positive Tests
- ^f Total Number of Negative Tests
- ⁹ Total Number of True Test Results
- ^h Total Number of Wrong Test Results

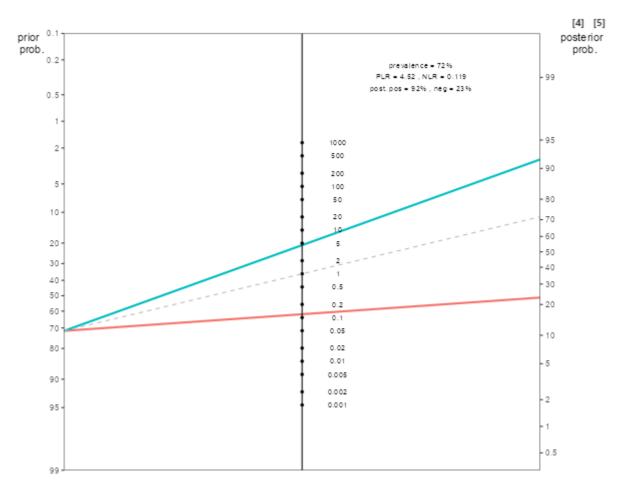
| | | 95% Confidence Interv | |
|-------------------------------|----------|-----------------------|-------|
| Decision Statistics | Estimate | Lower | Upper |
| Apparent prevalence | 70.5% | 59.8% | 79.7% |
| True prevalence | 71.6% | 61.0% | 80.7% |
| Test sensitivity | 90.5% | 80.4% | 96.4% |
| Test specificity | 80.0% | 59.3% | 93.2% |
| Diagnostic accuracy | 87.5% | 78.7% | 93.6% |
| Positive predictive value | 91.9% | 82.2% | 97.3% |
| Negative predictive value | 76.9% | 56.4% | 91.0% |
| Proportion of false positives | 20.0% | 6.8% | 40.7% |
| Proportion of false negative | 9.5% | 3.6% | 19.6% |
| False Discovery Rate | 8.1% | 2.7% | 17.8% |
| False Omission Rate | 23.1% | 9.0% | 43.6% |

| 95% | Confidence | Interval |
|------------|------------|----------|
|------------|------------|----------|

| Decision Statistics | Estimate | Lower | Upper |
|-------------------------------------|----------|---------|---------|
| Diagnostic odds ratio | 38.000 | 10.4445 | 138.255 |
| Number needed to diagnose | 1.419 | 1.1162 | 2.518 |
| Youden's index | 0.705 | 0.3971 | 0.896 |
| Likelihood ratio of a positive test | 4.524 | 2.0571 | 9.949 |
| Likelihood ratio of a negative test | 0.119 | 0.0542 | 0.261 |

Below there is the **Fagan nomogram**, based on the prevalence observed in the sample, imagining a positive test result, and another negative.

Fagan nomogram



ROC analysis

In the table Analyzes , press the PPDA button .

| ≡ | Variables | Data | Analyses | Edit | | | | |
|-------------|--------------|----------------------|----------|-------------|--------|----------|-----------|--|
| Exploration | ₹ T-Tests | ₽ ₽₽ ANOVA | | Frequencies | Factor | Survival | meddecide | |

you can't find it, it may not be displayed. In the table **Analyzes**, on the right, tap on + **Modules** and check if it is not present in the list of installed modules.

| 25 | Factor | Survival | | _ | meddecide | | CHR SiMpbyAlges | ee |
|-------|-------------|-------------|----|----|------------------------------------|--------------------|--------------------|----|
| | | | | ja | movi library | | | |
| | | | (- | М | anage installed | | | |
| | | | Ir | | lled Modules scatr | | | |
| den | t Variable | | | | | | | ٠ |
| DAI | | | | | blandr Bland-Altman Metho | d Comparison | | |
| CR | | | | | | | | |
| alpro | otectina | | | | deathwatch Death Watch | | | |
| HIT1 | | | 4 | | flexplot Graphically Based Da | ata Analysis | | l |
| /aria | ble | | | | gamlj | | | |
| ctivi | tate | | ~ | | General Analyses for | Linear Models i | n jamovi | |
| Vari | able | | | | jeva Evidential Analyses f | or Common Stat | istical Te | l |
| | | | - | | jjstatsplot Wrapper for qqstats | plot | | l |
| | | | | | MAJOR Meta-Analysis for JA | MOVI | | l |
| | | | - | | meddecide Functions for Medica | al Decision in Cli | nicoPath | |
| M | lodule - ps | - | | | psychoPDA Psychometrics & Pos | st-Data Analysis | | |
| | Show i | n main menu | | | SimplyAgree | | | |

You can check that module to be visible in the table **Analyzes** by clicking the **Show in main option menus** .

If it is not installed, follow the steps indicated in the chapter **Installing the additional analysis module** .

Clicking on the PPDA module, select the ROC Test option from the menu

| | PPDA |
|---|-------------------------------|
| t | Differential Item Functioning |
| 4 | Binary LogR |
| 1 | Measure Diagnostics |
| | Test ROC |

Select the variable that represents **the standard test** (eg Activity), and press the **arrow button** next to the **Class field variables**. Select the quantitative **variables** that represent **the tests of interest** and press the arrow button next to the **Dependent variable field**.

| 🔶 nr_crt | Q | Dependent Variable |
|-----------------|---|--------------------|
| 😪 Activitate | | \rightarrow |
| 🔶 CDAI | | |
| 🔶 PCR | | |
| 🔶 Calprotectina | | A |
| 🔶 сніт1 | | Class Variable |
| | | → |
| | | Group Variable |
| | | → |

In the end you will get something similar to the image below:

| 🤌 nr_crt 🛛 🔍 | Dependent Variable |
|--------------|---|
| - | → 🤣 CDAI |
| | |
| | Calprotectina |
| | СНІТІ |
| | / · · · · · · · · · · · · · · · · · · · |
| | Class Variable |
| | → 🔗 Activitate 🔗 |
| | Group Variable |
| | → |

Tables of limit values and associated statistics

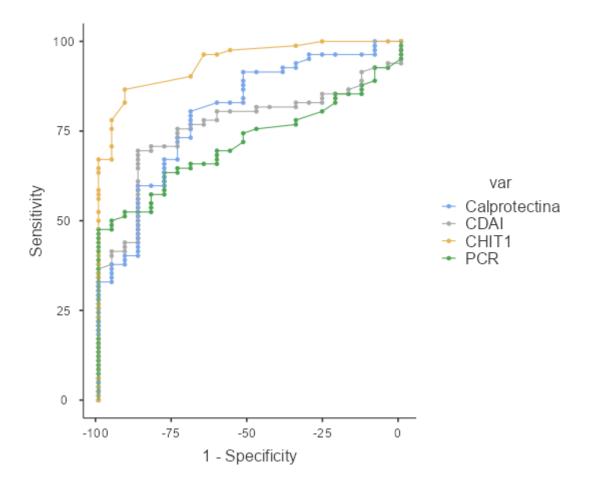
The program has already done the ROC analysis and presents you on the right side tables for different cutoff values (**Cutpoint**), with sensitivity (**Se**), specificity (**Sp**), positive predictive value (**PPV**) and negative (**NPV**), Youden 's index (**Youden's index**), the area under the ROC curve (**AUC**).

Thus **the best limit value**, which has the Youden index has the highest value, is 210, having the associated sensitivity and specificity of 80.49% and 69.57%, respectively.

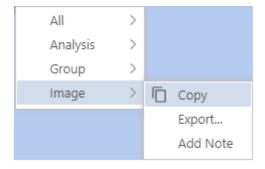
| Cutpoint | Sensitivity (%) | Specificity (%) | PPV (%) | NPV (%) | Youden's index | AUC | Metric Score |
|----------|-----------------|-----------------|---------|---------|----------------|-------|--------------|
| 210 | 80.49% | 69.57% | 90.41% | 50% | 0.501 | 0.801 | 1.50 |
| 215 | 79.27% | 69.57% | 90.28% | 48.48% | 0.488 | 0.801 | 1.49 |
| 220 | 78.05% | 69.57% | 90.14% | 47.06% | 0.476 | 0.801 | 1.48 |
| 221.6 | 76.83% | 69.57% | 90% | 45.71% | 0.464 | 0.801 | 1.46 |
| 240 | 75.61% | 69.57% | 89.86% | 44.44% | 0.452 | 0.801 | 1.45 |
| 280 | 73.17% | 73.91% | 90.91% | 43.59% | 0.471 | 0.801 | 1.47 |
| 300 | 71.95% | 73.91% | 90.77% | 42.5% | 0.459 | 0.801 | 1.46 |
| 400 | 67.07% | 78.26% | 91.67% | 40% | 0.453 | 0.801 | 1.45 |
| 596 | 59.76% | 86.96% | 94.23% | 37.74% | 0.467 | 0.801 | 1.47 |
| 600 | 58.54% | 86.96% | 94.12% | 37.04% | 0.455 | 0.801 | 1.45 |

ROC curve graph

Below is the graph with the ROC curve (Receiver operating characteristic - the operating characteristic of the receiver).



To copy the image press the right mouse button and select Image Copy



Comparison of ROC curves by statistical tests

To perform **statistical tests that compare the ROC curves** with each other, press the **> button** on the right **Advanced**, and select the **De Long's test option**.

| > Visualization | |
|-------------------|--|
| > Advanced | |

| ✓ Advanced | |
|----------------|-------------------|
| ✓ DeLong's tes | t |
| Method | Maximize metric 🗸 |
| Metric | Sum: Sens/Spec 🗸 |

The results look like the following:

DeLong Test of Difference between AUCs

| Estimated A | AUC's: | | | | |
|-------------|-------------|-------------|-----------|----------|-------------|
| AUC SD(| Hanley) P(H | 0: AUC=0.5) | SD(DeLong | g) P(H0: | AUC=0.5) |
| 1 0.762 | 0.050 | 0.000 | 0.04 | 18 | 0.000 |
| 2 0.717 | 0.055 | 0.000 | 0.04 | 19 | 0.000 |
| 3 0.801 | 0.045 | 0.000 | 0.05 | 50 | 0.000 |
| 4 0.942 | 0.022 | 0.000 | 0.02 | 22 | 0.000 |
| | | | | | |
| Pairwise o | omparisons: | | | | |
| AUC | Difference | CI(lower) | CI(upper) | P.Value | Correlation |
| 1 vs. 2 | 0.045 | -0.059 | 0.149 | 0.400 | 0.400 |
| 1 vs. 3 | -0.039 | -0.145 | 0.066 | 0.465 | 0.409 |
| 1 vs. 4 | -0.179 | -0.266 | -0.092 | 0.000 | 0.406 |
| 2 vs. 3 | -0.084 | -0.187 | 0.019 | 0.110 | 0.437 |
| 2 vs. 4 | -0.224 | -0.318 | -0.131 | 0.000 | 0.268 |
| 3 vs. 4 | -0.140 | -0.228 | -0.052 | 0.002 | 0.455 |
| | | | | | |
| Overall te | st: | | | | |
| p-value = | 1.28e-06 | | | | |

The first table lists the estimates of **the areas under the ROC curve** for each diagnostic test, the standard deviation, as well as a **statistical test of significance for one ROC curve**, for each selected variable in the order of their selection. Thus for the CDAI clinical activity score, the area under the ROC curve is 0.717, and the result is statistically significant P (H0: AUC=0.5) being less than 0.05. (valerur du p<0.001). **The null hypothesis** of the test for an **ROC curve** is that the area under the ROC curve for CDAI, compared to histopathological examination, is not statistically significantly different from 0.5. **The alternative hypothesis** of the test is that the area under the ROC curve for CDAI compared to histopathological examination, is not statistically significantly different from 0.5. **The alternative hypothesis** of the test is that the area under the ROC curve for CDAI compared to histopathological examination, is not statistically significantly different from 0.5.

In the second table are comparisons between tests taken two by two, presenting the difference between the surfaces of the ROC curves (AUC Difference), with 95% confidence interval, statistical test value (**P.Value**). For example, comparing CDAI clinical activity score (1) with C-reactive protein (2), the result is not statistically significant (p=0.400). **Null hypothesis** of **the two-ROC curve comparison** test: There is no statistically significant difference between the diagnostic accuracy of the CDAI clinical activity score and C-reactive protein, as measured by AUC, with histopathological examination as standard. **Alternative hypothesis** of the two-ROC curve comparison test: There is a statistically significant difference between the diagnostic accuracy of the CDAI clinical activity score and C-reactive protein, as measured by AUC, with histopathological examination as standard. **Alternative hypothesis** of the two-ROC curve comparison test: There is a statistically significant difference between the diagnostic accuracy of the CDAI clinical activity score and C-reactive protein, measured by AUC, with histopathological examination as standard.

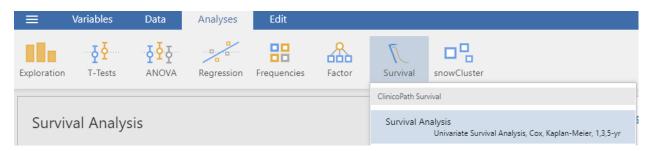
Finally, the result of a global statistical test comparing all ROC curves with each other is presented.

Survival analysis (Kaplan Meyer plot, log-rank test, Cox – HR regresion)

Make sure that the **jsurvival - Survival Module of ClinicoPath** for Jamovi 0.0.2 is installed (e.g. by verifying if the Survival button is present in the Analyses tab). See the chapter **Installation of additional analysis module (e.g. for ROC analysis, survival, diagnosis)**.

Check that the type of variables is well chosen in the **Data** tab, and that the order of the categories for each variable is as you prefer.

In the Analyses tab, clik Survival button, and choose in the menu the option Survival Analysis (Univariate Survival Analysis, Cox, Kaplan-Meier, 1,3,5-yr).



Move the variable that indicates the **survival time** with a click on the arrow next to the **Time Elapsed** text box. Move the variable that indicates the **censorship status/event** with a click on the arrow next to the **Outcome** text box. Make sure that the **category** that indicates **the event** is selected from the **Event Level** list (e.g. 1). Move the grouping variable with a click on the arrow next to the **Explanatory Variable** field (e.g. Type, Treatment, Risk Factor).

| Survival Analysis | | | \ni |
|--|---------------|--|---------------|
| Patient no. Patient no. | Q → | Time Elapsed Duration_months Outcome | * |
| | \rightarrow | Censorship_status | \$8 h. |
| | | Event Level 1 | ~ |
| | → | Explanatory Variable | -1.0 |
| | | 🐣 Histological_type | |

In the Plots section, check the Survival Plot option.

You can change the options of the graph: to show the p-value, the table with the number of patients at risk, the cases with censored observations. Please check **risktable** to make sure the legend is visible.

| ✓ Plots |
|----------------------|
| Plots |
| ✓ Survival Plot |
| KMunicate-Style Plot |
| Cumulative Events |
| Cumulative Hazard |
| Plot Arguments |
| Plot End Time 60 |
| Time Interval 12 |
| 95% CI |
| ✓ risktable |
| censored |

To find out the probability of survival at different times, you can enter these times in the **Survival table** section, in the **Cutpoints** field.

| ✓ Survival Tables | | | | | | |
|----------------------|--|--|--|--|--|--|
| Survival table | | | | | | |
| Cutpoints 12, 36, 60 | | | | | | |

The median survival time

In the **Median Survival Summary and Table** section, the medians of survival time with 95% confidence intervals can be found in the text before the table. Also, in the table, in the **Median** column we find the median by group, and the 95% confidence interval in the columns **Lower** and **Upper 95% Confidence interval.** For example, for the group with small cell histology, the median survival is 2 months.

Median Survival Summary and Table - Histological_type

When Histological_type is small cell, median survival is 2 [1 - 2, 95% CI] months. When Histological_type is squamous cell, median survival is 4 [3 - 10, 95% CI] months.

| | | | | | | 95% Confidence Interval | |
|---------------------------------|---------|--------|-------|----------|--------|-------------------------|-------|
| Levels | Records | Events | rmean | se_rmean | Median | Lower | Upper |
| Histological_type=small cell | 48 | 45 | 2.72 | 0.497 | 2.00 | 1.00 | 2.00 |
| Histological_type=squamous cell | 35 | 31 | 7.76 | 1.601 | 4.00 | 3.00 | 10.00 |

Median Survival Table: Levels for Histological_type

Cox Regression-hazard ratio

In the table with the results of the Cox regression (**Cox Regression Summary and Table**), you can find the hazard ratio with the 95% confidence interval, in the **HR (Univariable) column**. To find out which group is **the reference group**, in the **HR (Univariable)** column you look for the group that shows the - **sign**. In the example below, it can be seen that the reference group is the histopathological small cell type. So, HR is obtained by dividing the hazard of participants with cancer of squamous histology, to the hazard of participants with cancer of small cell histology.

Cox Regression Summary and Table - Histological_type

When Histological_type is squamous cell, there is 0.43 (0.26-0.72, p=0.001) times risk than when Histological_type is small cell.

Cox Table- Histological_type

| Explanatory | Levels | all | HR (Univariable) |
|-------------------|---------------|-----------|---------------------------|
| Histological_type | small cell | 48 (57.8) | - |
| | squamous cell | 35 (42.2) | 0.43 (0.26-0.72, p=0.001) |

The survival probability at different times

The probability of survival at different points in time (e.g. in the **Time column**, at 12 months) can be found in the **Survival Summary and Table**, in the **Survival column**, and the 95% confidence interval in the **Lower, and Upper 95% confidence interval columns**), but also in text format, before the table. For example, the group with small cell histology has a probability of survival at 12 months of 5.9%.

1, 3, 5-yr Survival Summary and Table - Histological_type

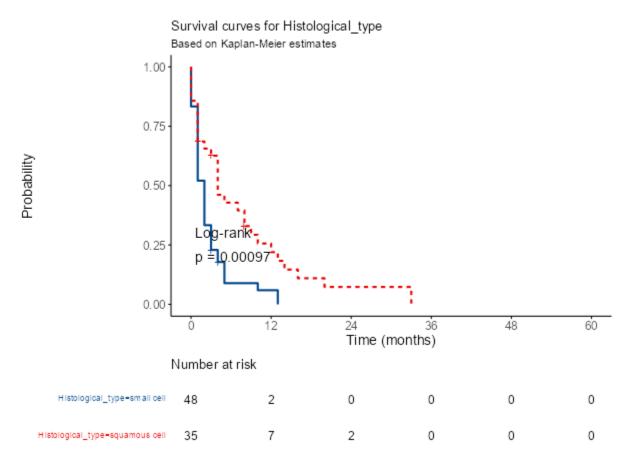
When Histological_type=small cell, 12 month survival is 6% [1.6%-22%, 95% CI]. When Histological_type=squamous cell, 12 month survival is 22% [11.1%-43%, 95% CI].

| | | | | | 95% Confidence Interval | |
|---------------------------------|------|----------------|------------------|----------|-------------------------|--------|
| Levels | time | Number at Risk | Number of Events | Survival | Lower | Upper |
| Histological_type=small cell | 12 | 2 | 43 | 5.9 % | 1.6 % | 21.6 % |
| Histological_type=squamous cell | 12 | 7 | 25 | 22.0 % | 11.1 % | 43.3 % |

1, 3, 5 year Survival - Histological_type

The Kaplan-Meier survival curve, with the log-rank test

The Kaplan-Meier survival curve, with the log-rank p-value, and the table with the number of subjects at risk, can be found in the Survival **Plot** section. In the table with the number of subjects at risk, at the beginning of the study there are 48 living subjects (at risk of having the event – death) in the group with small cell histology, and 35 subjects in the group with squamous histology. At 12 months there are only 2 living subjects (at risk of the event – death) left in the group with small cell histology, and 7 subjects in the group with squamous histology.



Meta-analysis

Meta-analysis to compare two groups in terms of dichotomous data

We make sure that **the MAJOR module** is **installed**. If it is not installed, follow the instructions in the **Installation of additional analysis modules** chapter.

We make sure that **the type of variables** representing the number of events, respectively the total number of subjects, is of the **Continuous type**.

From the MAJOR module, we choose the Dichotomous Models option.

| | meddecide | PPDA | SimplyAgree | | | |
|--|----------------------|------|-------------|--|--|--|
| Meta Analysis | | | | | | |
| Correlation | n Coefficients (r, I | N) | | | | |
| Dichotomous Models | | | | | | |
| Effect Sizes and (Sampling Variances or Standard Errors) | | | | | | |
| Mean Differences (n, M, SD) | | | | | | |
| Proportions | | | | | | |
| Reliability Generalization | | | | | | |

Move the variable indicating the number of events in the experimental/intervention/exposed group to the **Number of Incidents in Experimental Group** field ; the variable indicating the number of events in the control/non-exposed group to the **Number of Incidents in Control Group** field; the variable indicating the total number of participants in the experimental/intervention/exposed group in the **Total Sample Size for Experimental Group** field; the variable indicating the total number of participants in the **Total Sample Size for Experimental Group** field; the variable indicating the total number of participants in the study in the **Total Sample Size for Control Group** field; the variable indicating the name of the study in the **Study Label** field.

In the **Model Options** section, choose for **Model Estimator**, **Restricted Maximum-Likelihood**; for **Model Measures**, Log odds ratio (for case-control studies), or risk difference (for cohort studies); check **Back-Transform Log Odds Ratio to Odds Ratio**.

The rest of the options can be left as they are. If desired, it can be reduced in case of reduced heterogeneity to the Fail-Safe N Rosenberg method.

| Dichotomous Models | | \ominus |
|--------------------|---|---|
| 🔗a Study Type | | Number of Incidents in Control Group |
| | - | Total Sample Size for Experimental Group Total Exposed |
| | - | Total Sample Size for Control Group Image: Control Size for Con |
| | - | Study Label |
| | | Moderator (optional) |

| ✓ Model Options | |
|--------------------------------|--------------------|
| Model estimator Restricted Max | ximum-Likelihood 🗸 |
| Model measures Log odds ratio | ~ |
| Moderator type No Moderator | ~ |
| ✓ Back-Transform Log Odds Rat | tio to Odds Ratio |

Association estimator

If Log odds ratio has been selected as the estimator of the association, as well as if **Back-Transform Log Odds Ratio to Odds Ratio** has been checked, then in the table with the same name, the value of the OR point estimator of the can be found, as well as the 95% confidence interval.

| Back-Transform Log Odds Ratio to Odds Ratio | | | | | |
|---|-------|-------|--|--|--|
| Odds Ratio CI Lower Bound CI Upper Bound | | | | | |
| 1.542 | 1.135 | 2.096 | | | |

In **the Random-Effects Model table**, the estimator value of the log odds ratio/log risk ratio/risk difference association is reported in the **Estimate** column with the confidence interval in the last two columns. The **p-value** for its statistical significance is in the **p**-column.

Random-Effects Model (k = 10)

| | Estimate | se | Z | р | CI Lower Bound | CI Upper Bound |
|-----------|----------|-------|------|-------|----------------|----------------|
| Intercept | 0.433 | 0.157 | 2.77 | 0.006 | 0.126 | 0.740 |
| | | | | | | |

Note. Tau² Estimator: Restricted Maximum-Likelihood

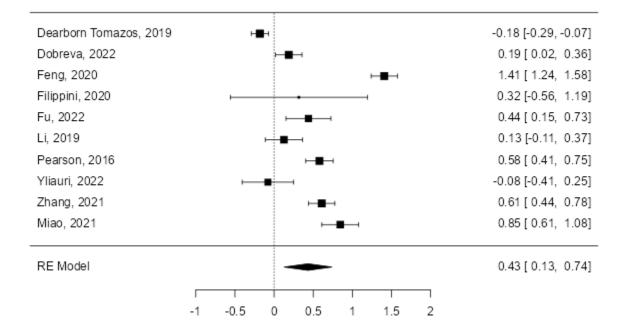
Statistical heterogeneity of studies

In the **Heterogeneity Statistics** table, in the column I^2 there is the inconsistency index, and in column **p**, the p-value of the associated test.

Heterogeneity Statistics Tau² |² Η² R² Tau df Q р 0.471 0.222 (SE= 0.1147) 95.71% 23.304 9.000 288.885 < .001 .

Forest plot

In the **Forest plot** section, you have the forest plot. Unfortunately, for the estimators of the log odds ratio, log risk ratio, the logarithm values of the OR or RR are written in the graph. The advantage is the symmetry of the confidence intervals. The downside is that it is more difficult to interpret. The value of 0 is equivalent to an OR or RR of 1. Values on the graph greater than 0 for OR/RR are greater than 1, and values less than 0 are less than 1.



Publication bias

The publication bias can be assessed with formal statistical tests and with plots, like the funnel plot.

Statistics

To evaluate the publication bias, in the **Publication Bias Assessment** section, one can use the result p-value (column p) of the statistical test for funnel plot asymmetry (**Regression Test for Funnel Plot Asymetry**).

Publication Bias Assessment

| Fail-Safe N Analysis (File Drawer Analysis) | Analysis (File Drawer Ar | nalysis) | |
|---|--------------------------|----------|--|
|---|--------------------------|----------|--|

| Fail-safe N | р |
|-------------|--------|
| 593.000 | < .001 |

Note. Fail-safe N Calculation Using the Rosenthal Approach

Rank Correlation Test for Funnel Plot Asymmetry

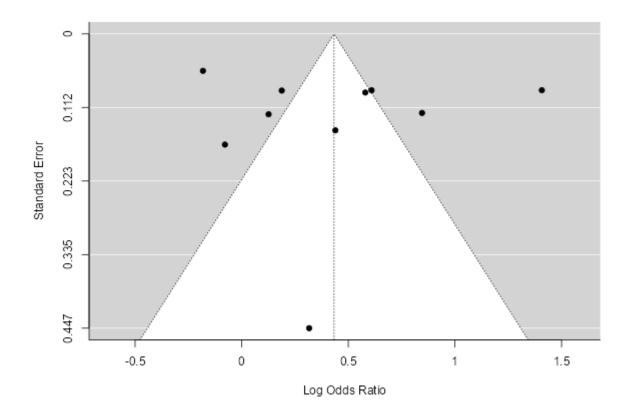
| Kendall's Tau | | р |
|---------------|--------|-------|
| | -0.156 | 0.601 |

| Regression | Test for | Funnel | Plot Asymmetry |
|------------|----------|--------|----------------|
|------------|----------|--------|----------------|

| Z | р |
|--------|-------|
| -0.306 | 0.760 |

Funnel plot

The funnel plot can be found in the **Funnel plot section**.



Installation of additional analysis modules (e.g. for ROC analysis, survival, diagnosis)

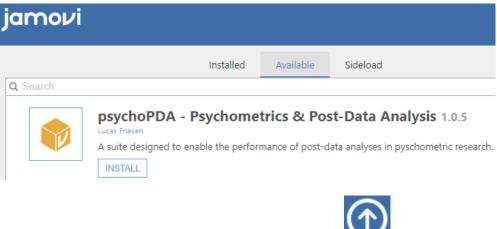
In the table Analyzes , on the right, tap on + Modules and check if it is not present in the list of installed modules.



To install analysis modules in addition to the default ones, select tab **Analyzes**, and on the right side press the + **Modules button**, then select **Manage installed**.



Select tab **Available**, scroll until you find the desired module (eg psychoPDA) and press the **INSTALL button**.



Return to the analysis window by pressing the arrow button:

Instructions for using EpiInfo and Excel for data analysis

Downloading EpiInfo (for home use)

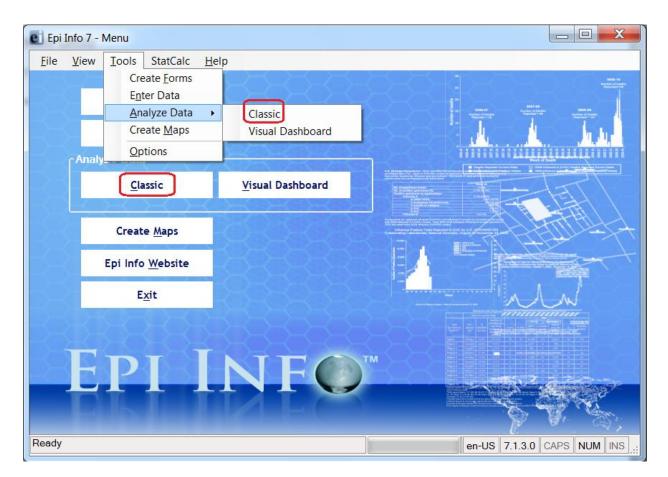
Use an Internet browser to navigate to: <u>http://www.cdc.gov/epiinfo/installation.htm</u> and follow the installation instructions.

Starting Epi Info

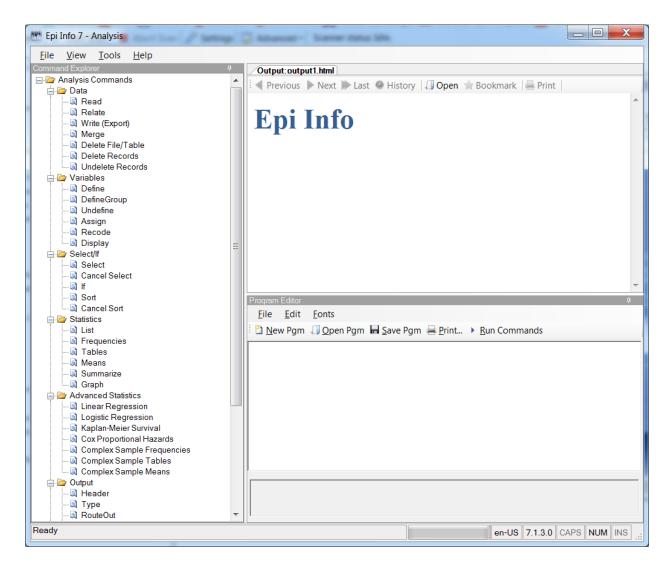
Start button, shortcut Epi InfoTM 7 or Start button, All Programs, CDC, Epi Info 7, Epi InfoTM 7

Analyzing data in Epi Info

Click on **Classic** in the **Analyze Data section** or using the menu.



The data analysis window:



Importing an Excel file in Epi Info

In the commands section - **Command Explorer** – left hand side of the window, **Analysis Commands**, in the section : **Data** – click on **Read**. The window for importing data will open. Select the type of file to import.

| Read | ? X |
|---------------------------|--------------|
| Recent Data Sources | |
| Database <u>T</u> ype | • |
| Epi Info 7 Project | |
| Data Source | |
| (none) | |
| Show | |
| Data Source Explorer | |
| | |
| | |
| | |
| | |
| | |
| | |
| | |
| | |
| | |
| | |
| OK Cancel Clear Save Only | <u>H</u> elp |

If the Excel file has the .xlsx, extension we choose Microsoft Excel 2007 Workbook (.xlsx), if the file has the .xls extension, we choose Microsoft Excel 97-2003 Workbook (.xls).

| Database <u>Type</u> |
|--|
| Epi Info 7 Project |
| Epi Info 7 Project |
| Microsoft Access 2002-2003 (.mdb) |
| Microsoft Access 2007 (.accdb) |
| Microsoft Excel 97-2003 Workbook (xls) |
| Microsoft Excel 2007 Workbook (xlsx) |
| Microsoft SQL Server Database |
| Flat ASCII File |
| MySQL Database |
| PostgreSQL Database |

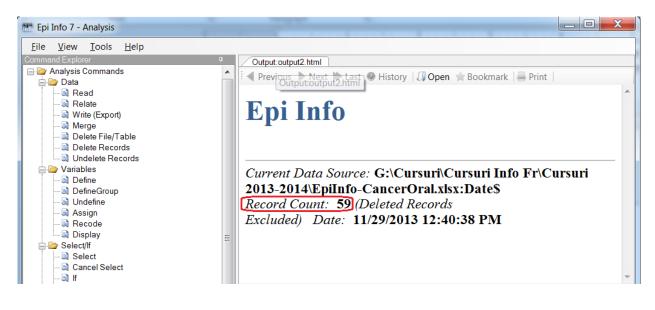
After this we search for the file to import after clicking on the button corresponding to **Data Source**. In the window that will open we click the button of the **Location** field, and we browse for the Excel file. We select the file and click **OK**, letting checked the option : **First row contains header information**.

| Open Existing File | X |
|---|----------------|
| Please enter the filename and path to the existing Exce | workbook below |
| Location: | |
| | |
| First row contains header information | |
| | OK Cancel |

After this EpiInfo shows all the worksheets in the file, in the **Data Source Explorer** section. Select the worksheet that contains the data and click the OK button to import it.

| Database <u>T</u> ype | | | |
|---------------------------------------|--------------------------------------|--|--|
| Microsoft Excel 2007 Workbook (.xlsx) | | | |
| Data Source | | | |
| G:\Cursuri\Cursuri Info Fr\Cursur | ri 2013-2014\Epilnfo-CancerOral.xlsx | | |
| Show | | | |
| ✓ <u>F</u> orms | <u>√</u> <u>T</u> ables | | |
| Data Source Explorer | | | |
| Date\$ | | | |
| Sheet2\$ | | | |
| Sheet3\$ | | | |

After importing Epi Info shows in the results section (**Output**), the imported file name and the number of records in the file (**Record Count**).



Activating the Data Analysis module in Microsoft Excel

Click on a void cell, then click on *Add-Ins* in the *Tools* menu. Check the box next to *Analysis ToolPack* and then click OK. Select another empty cell, then search for *Data Analysis* in the *Tools* menu.

If *Data Analysis* does not appear in the *Tools* menu, despite being checked in *Add-Ins*, uncheck the box in *Add-Ins* and repeat the above procedure.

Descriptive statistics

Qualitative (categorical) data:

Frequency tables

Use the **COUNTIF** function in **Microsoft Excel** to count how many times each value taken by a variable appears in the database (its absolute frequency).

E.g. to find out how many female persons (coded with F in the file) are in the sample, a void cell at the future location of the frequency table should contain a similar formula to =COUNTIF(A2:A58, "F"), if data regarding gender was recorded in cells A2 to A58. A correct frequency table should look like this:

| Gender | Number of subjects |
|--------|--------------------|
| Male | 20 |
| Female | 37 |
| Total | 57 |

Table 1. Gender distribution in the studied sample

Note that any table has to be labeled on top of it, using a clear and precise title. Select the table, right click on this selection and choose *Caption* in order to label the table. Row and column labels should be visible and easily understandable by the reader, with no need to search for further explanations in order to understand the content of the table.

Pie charts

Follow the instructions above to create a frequency table using **COUNTIF**.

Select only the cells containing the absolute frequencies and their labels (do not select the total or column labels). Use *Insert - Graph* and select *Pie*. Click *Next*. In the *Chart Options* window click on the tab *Data Labels* and tick *Percentage*. Continue and finish the chart wizard. A correct pie chart should look like this:

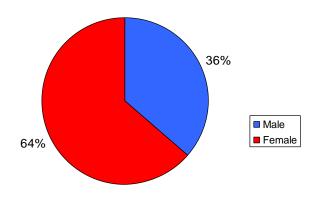


Figure 1. Gender distribution in the studied sample

Note that pie charts have to be labeled using visible percentages.

If you plan to use the chart in a *PowerPoint* presentation, make sure to label it on top, using a clear and precise chart title (what, how and for which subjects has been represented?).

If you plan to use the chart as a figure in a *Word* document, erase the chart title in *Excel* but remember to label the chart in *Microsoft Word*: select the figure, right click on this selection and use *Caption*.

All labels and legend entries should be visible and easily understandable by the reader, with no need to search for further explanations in order to understand the content of the figure.

Frequency tables in Epi Info

In Command Explorer section Statistics we choose Frequencies .

In the new window we select the variable of interest in the list of **Frequency of** and we click the boutton **OK**.

| Frequencies | | 2 X |
|--|---|----------------------|
| ILL Freq % + 20 35% - 37 65% Total 57 100% | Erequency of * Bain de bouche avec d alcool Fumer ou tabac Hauteur (cm) ID Poids (kg) Sexe | St <u>r</u> atify by |
| • O <u>u</u> tput to Table | OK Cancel Clear | Save Only Help |

In the Output we get the frequency table and the corresponding confidence intervals.

FREQ Gen

| GEN | Frequency | Percent | Cum. Percent | |
|-------|-----------|---------|--------------|--|
| F | 14 | 23.73% | 23.73% | |
| M | 45 | 76.27% | 100.00% | |
| Total | 59 | 100.00% | 100.00% | |

95% Conf Limits

F 13.62% 36.59% M 63.41% 86.38%

Contingency tables

In *Microsoft Excel*, select any cell containing data. Then, click in the menu bar *Data* – *Pivot Table* – *Pivot Chart Report*. Work your way through the wizard and obtain a new worksheet containing an empty pivot table and a field list.

Drag and drop the field representing a prognostic factor (the risk factor, the new diagnostic test or the new treatment, depending on the given research scenario) to the area labeled **Drop Row Fields Here**. Drag and drop the field representing an outcome (the disease, the reference diagnostic test or the treatment response, depending on the given research scenario) to the area labeled **Drop Column Fields Here**. Finally, drag and drop any of the formerly used fields to the area labeled **Drop Data Fields Here**.

Rename row and column labels so that they are easily understandable by the reader, with no need to search for further explanations in order to understand the content of the table (*e.g. If male gender was coded as m, rename the corresponding row label: male*)

Right click on a row label and select order, to correct the row order in your contingency table. Right click on a column label and select order, to correct the column order in your contingency table.

After inserting the contingency table into your *Word* document, remember to label it using *Caption* and a correct title.

The column chart associated to a contingency table

After creating a pivot contingency table, select *Insert - Chart* from the menu bar.

To hide the chart buttons right click the button *Count of* and select *Hide Pivot Chart Field Buttons*.

To show frequency labels, right click the empty chart area towards the upper left corner, select *Chart Options* and, in the *Data Labels* tab, tick *Percentage* or *Value*.

Then, switch to the *Titles* tab and define clear and precise titles for your chart axes, including the units of measurement between brackets, where necessary.

After inserting the chart into your *Word* document, remember to label it using *Caption* and a correct title.

Quantitative data:

Individual description of quantitative variables

Mean, median, standard deviation, 95% confidence interval for means

In *Microsoft Excel*, use *Tools* – *Data Analysis* – *Descriptive Statistics* to simultaneously compute the most important descriptive parameters for selected quantitative variables.

In the *Descriptive Statistics* window tick options *Summary Statistics* and *Confidence Level for Mean*.

To find the lower limit of the 95% confidence interval, compute *Mean* minus *Confidence Level (95%)*.

To find the upper limit of the 95% confidence interval, compute *Mean* plus *Confidence Level (95%)*.

Frequency table and Histogram

In *Microsoft Excel*, use *Tools* – *Data Analysis* – *Descriptive Statistics* to compute minimum, maximum and range for the desired quantitative variable.

Choose a convenient bin size for the variable of interest (a round-figure for which 7-10 non-overlapping intervals of that similar size will cover the whole variable range).

Label a void column as "*Bin Variable name (units of measurement)*" on the same worksheet as the variable of interest.

Below this label, insert the value for minimum+ the chosen bin size.

Use *Edit – Fill – Series* (select options: in Columns, Step value=bin size, Stop value=maximum-bin size) to complete the column containing the bin values for your variable.

Now use Tools – Data Analysis – Histogram:

For *Input Range* select the range of cells containing the quantitative variable for which you want to plot a frequency table and histogram. For *Bin Range* select the newly created column. In both cases, include the column labels in your selection and tick *Labels*.

In *New Worksheet Ply* write a suggestive name for the worksheet that will contain the frequency table and histogram for your variable.

In order to display the histogram you need to select *Chart Output*.

After pressing the OK button, both frequency table and histogram will appear in a raw, unfinished form.

In order to be comprehensible, both the frequency table and the histogram need adjustments:

- 1. replace the upper limit of each bin shown in the brute table with the corresponding bin interval
- 2. delete the chart legend since its information is redundant and only takes up space
- 3. delete the title *Histogram*, since you will label the figure in *Microsoft Word*, using *Caption*
- 4. resize the chart area as needed, in order to have a clear view over your histogram
- eliminate spaces between columns by right clicking any of the columns and using *Format Data Series – Options* and adjusting the *Gap Width*
- 6. verify the content and font size of all labels, to make sure that your histogram is easily understandable by anyone who reads your work.

A correct frequency table and a correct histogram should look like this:

Table 2. Weight distribution in the studied sample

| Weight intervals (kg) | No. of subjects |
|-----------------------|-----------------|
| <=40 | 21 |
| (40-50] | 144 |
| (50-60] | 297 |
| (60-70] | 240 |
| (70-80] | 145 |
| (80-90] | 79 |
| (90-100] | 45 |
| >100 | 29 |

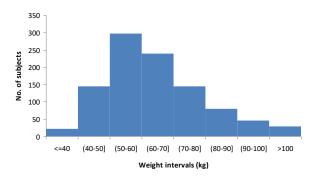


Figure 2. Histogram of weight in the studied sample

Description of a potential relation between two quantitative variables

Scatter chart

In *Microsoft Excel*, select the columns containing the two quantitative variables, including their labels. Then select *Insert – Chart* and choose *XY (Scatter)*.

Advance to step 3 of the chart wizard and define correct titles for both X and Y axes. Do not forget to specify after the title of each axis the corresponding units of measurement, between round brackets.

Then click on the *Legend* tab and uncheck the *Show legend* box, since no useful information derives from a legend when investigating only two variables at once.

In the *Titles* tab write the precise title of each axis, including the units of measurement in parentheses.

If you plan to use the chart in a *PowerPoint* presentation, make sure to label it on top, using a clear and precise chart title (what, how and for which subjects has been represented?).

If you plan to use the chart as a figure in a *Word* document, erase the chart title in *Excel* but remember to label the chart in *Microsoft Word*: select the figure, right click on this selection and use *Caption*.

All labels should be visible and easily understandable by the reader, with no need to search for further explanations in order to understand the content of the figure.

After finishing the chart wizard, right-click on any point from the data cloud and select *Add Trendline*. The most common trend of data clouds is a linear one. In the *Options* tab check *Display equation on chart* and *Display R-squared value on chart*.

To highlight the trendline using a contrasting color, right-click on the trendline and use *Format Trendline*.

To highlight the trendline labels using a contrasting color, right-click on the label box and use *Format Data Labels*.

A correct scatter chart should look like this:

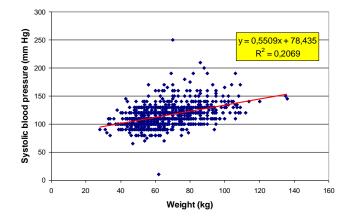


Figure 3. Relation between weight and systolic blood pressure for all subjects included in the studied sample

Survival data:

Median of survival time

In *Microsoft Excel*, use *Tools – Data Analysis – Descriptive Statistics*. In the *Descriptive Statistics* window check *Summary Statistics*.

Survival probability chart

In the Analysis module of EpiInfo click Kaplan-Meyer Survival, from the left panel.

Complete the dialog box as seen in the image below:

| Kaplan-Meier Survival | | | | X |
|---|--------------|--|-----------------------|---|
| Censored Variable CensorVariable Time Variable Time_month_ Group Variable Gender Weight | • | Value for U "complete Time Unit Months Graph Type Survival P Output to T | y" e robability | • |
| | <u>S</u> ave | Only | <u>O</u> K | |
| Clear | <u>H</u> elp | | <u>C</u> ancel | |

Change the *Group Variable* as needed for your comparison.

Data Analysis

Performing a Student test (t-test) in Excel

Before performing the test you need to sort your data according to the groups that you wish to compare. (e.g. if you wish to compare cholesterol values of males with cholesterol values of females, you need to sort your data by gender).

To sort your data, click on any cell inside your data range, then use **Data – Sort**.

If the groups that you wish to compare are independent (e.g. comparing cholesterol values of women with those of men), use *Tools – Data Analysis – t-Test: Two-Sample Assuming Unequal Variances.*

If the groups that you wish to compare are paired (e.g. comparing cholesterol values of the same subjects, before and after treatment), use **Tools – Data Analysis – t-Test: Paired Two Sample for Means.**

In the test window, select for *Variable 1 Range* the cells containing the quantitative variable corresponding to the first group (e.g. initial cholesterol values for women) and for *Variable 2 Range* the cells containing the quantitative variable corresponding to the second group (e.g. initial cholesterol values for men), without selecting the column label. Pay attention not to select the grouping variable (e.g. *Gender*) instead of the corresponding quantitative variable that you wish to compare.

Since the null hypothesis (H₀) of your reasoning states the absence of difference between mean values of the compared variable, introduce 0 in the *Hypothesized mean difference* box.

Give a title to the future worksheet that will contain the test results, by entering a suggestive name in the *New Worksheet Ply* box.

Immediatly after pressing OK, rename the generic lables *Variable 1* and *Variable 2* using suggestive labels: include information regarding both the quantitative variable you have compared and the grouping variable that sets them apart. This will allow you to easily interpret your test results later on.

The two-tailed p-value rendered by the test shows the statistical significance of the investigated difference between the mean values of the compared groups.

If the rendered two-tailed p-value includes the letter E followed by a negative figure this means in fact a very low (i.e. significant) p-value (e.g. $p = 3,22342E-6 = 3,22342 \times 10^{-6} = 0,0000032234$).

The test results also include the mean values of the compared variables. By subtracting them, you will be able to evaluate the difference between mean values, thus appraising the clinical significance of this difference.

Computing the contingency table, Risk Ratio (RR) and Odds Ratio (OR) in EpiInfo

Performing a Chi-square (X^2) test in EpiInfo

In the *Analysis* module of *EpiInfo* click *Tables*, from the left panel.

In the dialog window that opens, select from the drop-down lists the *Exposure Variable* (e.g. the risk factor, treatment, etc.) and the *Outcome Variable* (e.g. the disease suspected to be an outcome of the risk factor, the improvement of health suspected to be an outcome of the treatment, etc.), then press OK.

| Tables | | ? 💌 |
|---|---|------------------------|
| | Outcome Variable Colesterol LDL crescut | Stratify by |
| Exposure Variable Diabet zaharat ▼ Weight | ILL WATER + + 20 - 37 | |
| O <u>u</u> tput to Table | | Optional Page Settings |
| | | Columns Per Page |
| | OK Cancel Clear | Save Only Help |

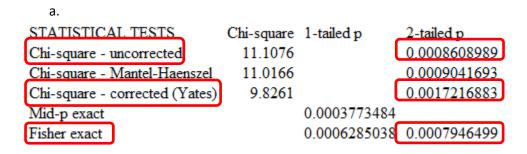
| | Colesterol I | Colesterol LDL crescut | | |
|----------------|--------------|------------------------|---------|--|
| DIABET ZAHARAT | da | nu | Total | |
| da | 40 | 42 | 82 | |
| Row% | 48.78% | 51.22% | 100.00% | |
| Col% | 85.11% | 56.00% | 67.21% | |
| nu | 7 | 33 | 40 | |
| Row% | 17.50% | 82.50% | 100.00% | |
| Col% | 14.89% | 44.00% | 32.79% | |
| TOTAL | 47 | 75 | 122 | |
| Row% | 38.52% | 61.48% | 100.00% | |
| Col% | 100.00% | 100.00% | 100.00% | |

Depending on the type of data collection used in your research scenario, interpret only the appropriate indicator (RR / OR) and its **95% Confidence Interval** displayed to the right of the **Point Estimate** of each indicator.

| Single | Table | Ana | lysis |
|--------|-------|-----|-------|
| - | | | |

| | Point | 95% Confide | nce Interval |
|----------------------------|----------|-------------|--------------|
| | Estimate | Lower | Upper |
| PARAMETERS: Odds-based | | | |
| Odds Ratio (cross product) | 4.4898 | 1.7831 | 11.3049 (T) |
| Odds Ratio (MLE) | 4.4367 | 1.8082 | 11.9542 (M) |
| | | 1.6804 | 13.2787 (F) |
| PARAMETERS: Risk-based | | | |
| Risk Ratio (RR) | 2.7875 | 1.3725 | 5.6611 (T) |
| Risk Difference (RD%) | 31.2805 | 15.2896 | 47.2714 (т) |

In most cases, the two-tailed p-value rendered by the *Chi-square - uncorrected* test shows the statistical significance of the investigated difference between the frequency distribution in the compared groups. Yet, sometimes, when one or more expected frequencies are lower than 5, a message will be displayed below the test results, telling you to interpret the p-value rendered by the *Fisher exact* test.



Comparing quantitative data (Test student/ANOVA/ ...) in Epi Info

In **Command Explorer, Statistics** section we choose the command **Means**.

| Means | | 8 2 |
|---|--|--|
| Means of DZ slab controlat vs DZ contro | Cross-tabulate by Value of Colesterol HDL (mg/dl) | Stratify by |
| Weight Output to Table | ILL HEIGHT + - 62.0 20 40 62.5 37 59 | Optional Page Settings Columns Per Page No Line Wrap |
| | OK Cancel Clear | Save Only Help |

In the opened window choose the quantitative variable from the list from **Means of** and the grouping variable in the list from **Cross-tabulate by Value of**, then press the OK button.

In the window with the results (output), we have:

1. Descriptive statistics for groups:

| Descriptive Statistics for Each Value of Crosstab Variable | | | | | | iable |
|--|-----------|----------|-------|----------|----------|-------------|
| | Oł | os To | otal | Mean | Variance | Std Dev |
| DZ controla | t 45.0 | 000 2502 | .0000 | 55.6000 | 112.4727 | 10.6053 |
| DZ slab contro | olat 37.0 | 000 1932 | .0000 | 52.2162 | 77.8408 | 8.8227 |
| | Minimum | 25% | Med | ian 75 | % Maxin | num Mode |
| DZ controlat | 39.0000 | 48.0000 | 55.00 | 000 62.0 | 000 80.0 | 000 54.0000 |
| DZ slab controlat | 36.0000 | 46.0000 | 52.00 | 000 58.5 | 000 76.0 | 000 41.0000 |

1. The result of the t test (Student) to compare the means of two independent samples with equal variances (Pooled) or unequal (Unequal) variances

T-Test

| | Method | Mean | 95% CL Mean | Std Dev |
|--------------------------|---------------|--------|----------------|---------|
| Diff (Group 1 - Group 2) | Pooled | 3.3838 | -0.9633 7.7309 | 9.8432 |
| Diff (Group 1 - Group 2) | Satterthwaite | 3.3838 | -0.8867 7.6543 | |
| | | | | |
| Method | Variances | DF t | Value Pr > t | |
| Pooled | Equal | 80 | 1.55 0.1253 | |
| Satterthwai | te Unequal | 79.98 | 1.58 0.1188 | |

2. **The result of ANOVA test** to compare the averages of three or more independent samples with equal variances:

ANOVA, a Parametric Test for Inequality of Population Means

(For normally distributed data only)

| Variation | SS | ďf | MS | F statistic |
|-----------|------------|----|-----------|-------------|
| Between | 232.49071 | 1 | 232.49071 | 2.39957 |
| Within | 7751.07027 | 80 | 96.88838 | |
| Total | 7983.56098 | 81 | | |
| | | | | |

P-value = 0.12532

3. The result of the Bartlett test to compare the variances of two independent samples:

Bartlett's Test for Inequality of Population Variances

Bartlett's chi square= 1.30103 df=1 P value=0.25403

A small p-value (e.g., less than 0.05 suggests that the variances are not homogeneous and that the ANOVA may not be appropriate.

4. The results of nonparametric tests for comparing two independent samples (Mann-Whitney test / Wilcoxon Two-Sample) or more than two independent samples (Kruskal-Wallis test):

Mann-Whitney/Wilcoxon Two-Sample Test (Kruskal-Wallis test for two groups)

Kruskal-Wallis H (equivalent to Chi square) = 1.4703 Degrees of freedom = 1 P value = 0.2253

Performing a Log-rank test for survival analysis in EpiInfo

In the Analysis module of EpiInfo click Kaplan-Meyer Survival, from the left panel.

| Censored Variable | | Value for U | ncensored |
|------------------------|--------------|-------------------|------------|
| CensorVariable | • | "complete | |
| Ti <u>m</u> e Variable | | Time <u>U</u> nit | |
| Time_month_ | • | Months | |
| Group Variable | | Graph Type | |
| Gender | • | Survival P | robability |
| Weight | | Output to Ta | able |
| | • | | |
| | | | |
| | <u>S</u> ave | Only | <u>O</u> K |
| | | | |

Complete the dialog box as seen in the image below:

Change the Group Variable as needed for your comparison.

Below the survival chart you will find the result of the Log-rank test.

Performing a Cox Regression and computing the Hazard Ratio in EpiInfo

In the *Analysis* module of *EpiInfo* click *Cox proportional hazard*, from the left panel.

Complete the dialog box as seen in the image below:

| Cox Proportional Hazards | | |
|---------------------------------------|----------------------|-----------------------------|
| Censored Variable | Value for Uncensored | St <u>r</u> atify By |
| Ti <u>m</u> e Variable Time_months | Ti <u>m</u> e Unit | - <u> </u> |
| Group Variable Gender | Other Variables | - |
| Weight | Make Dummy | |
| Confidence Limits 95% | 1 | Make E <u>x</u> tended Term |
| Output to Table | | |
| ()=Make Dummy Variables | | |
| | | |
| | Graph Options | Save Only OK |
| | Clear | Help Cancel |

Change the *Group Variable* as needed for your comparison.

The Hazard Ratio (HR), its 95% confidence interval and the statistical significance of the Cox regression model will be listed below the regression model chart.

Performing a Simple Linear Regression in Microsoft Excel

Use Data Analysis - Regression from the Tools menu in Microsoft Excel.

For *Input Y Range* select the cell range that contains the dependent variable (**y**) in your sample, the one you want to predict using a simple linear regression.

For *Input X Range* select the cell range that contains the independent variable (**x**) in your sample, the one you want to use in order to predict the dependent variable (**y**), using a simple linear regression.

Make sure to include in your selection the cells containing labels for both the dependent and the independent variable and check the *Labels* box. Also check the *Confidence Level* box for a 95% CI and enter a suggestive name for the new worksheet where your simple linear regression will be saved.

Performing a Multiple Linear Regression in Microsoft Excel

Use Data Analysis - Regression from the Tools menu in Microsoft Excel.

For *Input Y Range* select the cell range that contains the dependent variable (**y**) in your sample, the one you want to predict using a multiple linear regression.

For *Input X Range* select the contiguous cell range that contains the independent variables $(x_1, x_2, ..., x_n)$ in your sample, the ones you want to use in order to predict the dependent variable (y), using a multiple linear regression. If the independent variables do not form a contiguous cell range, cut isolated variables before using *Regression*, and insert them into adjacent columns in order to form a contiguous cell range.

Make sure to include in your selection the cells containing labels, for all dependent and independent variables and check the *Labels* box. Also check the *Confidence Level* box for a 95% CI and enter a suggestive name for the new worksheet where your multiple linear regression will be saved.