

## Exercise 1

A group of researchers wanted to predict the death from ventilator associated pneumonia in patients admitted to an intensive care unit. The authors measures the blood concentration of proANP on day 0 (D0) and day 4 (D4), and recorded the age and gender of the patients. They used the data to predict survival, in the group of 71 patients in which 26 died.

The percent of males and females distributed between survivors and non-survivors are shown below:

| Gender (%) | survivors (n = 45) | non-survivors (n = 26) | Total (n = 71) |
|------------|--------------------|------------------------|----------------|
| Male       | 66.7               | 46.2                   | 59.2           |
| Female     | 33.3               | 53.8                   | 40.8           |
| Total      | 100                | 100                    | 100            |

Use the  $\chi^2$ -test to test for association between gender and survival. Is there a significant association?

First we need to convert the percentages to original numbers.

| Gender (%) | survivors (n = 45) | non-survivors (n = 26) | Total (n = 71) |
|------------|--------------------|------------------------|----------------|
| Male       | 30                 | 12                     | 42             |
| Female     | 15                 | 14                     | 29             |
| Total      | 45                 | 26                     | 71             |

Then the expected numbers are calculated:

| Gender (%) | survivors (n = 45) | non-survivors (n = 26) | Total (n = 71) |
|------------|--------------------|------------------------|----------------|
| Male       | 26.6               | 15.4                   | 42             |
| Female     | 18.4               | 10.6                   | 29             |
| Total      | 45                 | 26                     | 71             |

Then the normalized differences between the observed and expected values are calculated and summed:

| Gender (%) | survivors (n = 45) | non-survivors (n = 26) | Total (n = 71) |
|------------|--------------------|------------------------|----------------|
| Male       | 0.429242           | 0.742919               | 1.172161       |
| Female     | 0.621661           | 1.075952               | 1.697613       |
| Total      | 1.050903           | 1.818871               | 2.869774       |

Therefore  $\chi^2 = 2.87$ . From the table in the appendix the p value is between 0.10 and 0.05. This is usually not considered statistically significant.

Is the  $\chi^2$ -test valid in this case?

Yes. All expected values are above 5.

What is the odds ratio of survival for genders?

$$or = \frac{ad}{bc} = \frac{30 \cdot 14}{12 \cdot 15} = 2.33$$

What is the 95% confidence interval of the odds ratio?

First the standard error of the natural logarithm transformed odds ratio is calculated:

$$SE(\ln(or)) = \sqrt{\frac{1}{a} + \frac{1}{b} + \frac{1}{c} + \frac{1}{d}} = \sqrt{\frac{1}{30} + \frac{1}{12} + \frac{1}{15} + \frac{1}{14}} = 0.5$$

Then the odds ratio is log transformed

## Solutions for contingency exercises

$$\ln(or) = 0.847$$

The 95% CI of the log transformed odds ratio is then

$$95\% \text{ CI} = 0.847 - 1.96 * 0.50 \text{ to } 0.847 + 1.96 * 0.50 = -0.142 \text{ to } 1.837$$

The 95% CI of the odds ratio is then:

$$95\% \text{ CI} = e^{-0.142} \text{ to } e^{1.837} = 0.87 \text{ to } 6.28$$

Is the odds ratio in agreement with the  $\chi^2$ -test and does it provide the same information.

The odds ratio is in agreement with the  $\chi^2$ -test because 1 is included in the 95% confidence interval we did not find a significant difference. If we believe that there is a difference between genders the odds ratio give an indication of the strength of the association between gender and survival. That is that the odds for males to survive is 2.33 times as high as the odds for males to survive.

## Exercise 2

Kan forskerne bruge Chi-kvadrat testen?

Det kan de ikke fordi 2 af 4 forventede værdier er under 5:

|               | Treatment A | Treatment B |    |
|---------------|-------------|-------------|----|
| <b>2 arms</b> | 7           | 7           | 14 |
| <b>3 arms</b> | 3           | 3           | 6  |
|               | 10          | 10          | 20 |

Hvilken test bør de bruge?

De skulle have brugt Fisher's exact test

Er der en signifikant forskel på antallet af patienter med bivirkninger i de to behandlingsgrupper?

For et beregne Fisher's exact test skal man beregne sandsynligheden for at finde netop de observerede værdier fra Tabel 2:

|               | Treatment A | Treatment B | Total |
|---------------|-------------|-------------|-------|
| <b>2 arms</b> | 5           | 9           | 14    |
| <b>3 arms</b> | 5           | 1           | 6     |
| <b>Total</b>  | 10          | 10          | 20    |

$$p = \frac{14! 6! 10! 10!}{20! 5! 5! 9! 1!} = 0,065$$

Ligeledes skal sandsynligheden for at finde mere ekstreme observationer beregnes. I dette tilfælde er det bare en mere ekstrem situation, som give de samme totaler:

|               | Treatment A | Treatment B | Total |
|---------------|-------------|-------------|-------|
| <b>2 arms</b> | 4           | 10          | 14    |
| <b>3 arms</b> | 6           | 0           | 6     |
| <b>Total</b>  | 10          | 10          | 20    |

$$p = \frac{14! 6! 10! 10!}{20! 4! 10! 6! 0!} = 0,0054$$

Sandsynligheden for at forskerne har opnået deres observationer ved et tilfælde er altså  $p=0,07$ .

## Solutions for contingency exercises

Hvad er odds ratio mellem de to behandlingsgrupper for at gro en tredje arm?

$$or = \frac{5 \cdot 9}{5 \cdot 1} = 9$$

Hvad er konfidensintervallet for odds ratioen?

$$\ln(or) = 2,20$$
$$SE(\ln(or)) = \sqrt{\frac{1}{5} + \frac{1}{9} + \frac{1}{5} + \frac{1}{1}} = 1,23$$

$$2,20 - 1,96 \cdot 1,23 \text{ to } 2,20 + 1,96 \cdot 1,23 = -0,211 \text{ to } 4,61$$

The confidenceinterval is thus

$$e^{-0,211} \text{ to } e^{4,61} \Rightarrow 0,81 \text{ to } 100$$

Kan odds ratioen vise en statistisk signifikant forskel mellem de to behandlinger?

Da konfidensintervallet overlapper 1 kan men heller ikke vha odds ratio vise en signifikant forskel på bivirkninger mellem de to behandlingsgrupper.

### Opgave 3

Først skal man lave apgarscore om til en variabel som indikerer om scoren er over 7. Derefter laves en krydstabel:

smoking \* apgarOver7 Crosstabulation

|         |     |                | apgarOver7 |      | Total |
|---------|-----|----------------|------------|------|-------|
|         |     |                | ,00        | 1,00 |       |
| smoking | no  | Count          | 10         | 10   | 20    |
|         |     | Expected Count | 12,0       | 8,0  | 20,0  |
|         | yes | Count          | 8          | 2    | 10    |
|         |     | Expected Count | 6,0        | 4,0  | 10,0  |
| Total   |     | Count          | 18         | 12   | 30    |
|         |     | Expected Count | 18,0       | 12,0 | 30,0  |

Man kan se at en af cellerne er under 4, så man bør lave en Fishers exact test i stedet for en Chi-kvadrat test. Her er resultatet for dem begge:

Solutions for contingency exercises

Chi-Square Tests

|                                    | Value              | df | Asymp. Sig. (2-sided) | Exact Sig. (2-sided) | Exact Sig. (1-sided) | Point Probability |
|------------------------------------|--------------------|----|-----------------------|----------------------|----------------------|-------------------|
| Pearson Chi-Square                 | 2,500 <sup>a</sup> | 1  | ,114                  | ,235                 | ,117                 |                   |
| Continuity Correction <sup>b</sup> | 1,406              | 1  | ,236                  |                      |                      |                   |
| Likelihood Ratio                   | 2,647              | 1  | ,104                  | ,141                 | ,117                 |                   |
| Fisher's Exact Test                |                    |    |                       | ,235                 | ,117                 |                   |
| Linear-by-Linear Association       | 2,417 <sup>c</sup> | 1  | ,120                  | ,235                 | ,117                 | ,096              |
| N of Valid Cases                   | 30                 |    |                       |                      |                      |                   |

a. 1 cells (25,0%) have expected count less than 5. The minimum expected count is 4,00.

b. Computed only for a 2x2 table

c. The standardized statistic is -1,555.

Vi har altså ikke kunnet vise at der er en sammenhæng mellem rygning og Apgar scoren, men det er IKKE det samme som at påstå at en sådan sammenhæng ikke er til stede.