

Unit 4 – Probabilities in Epidemiology

Solutions

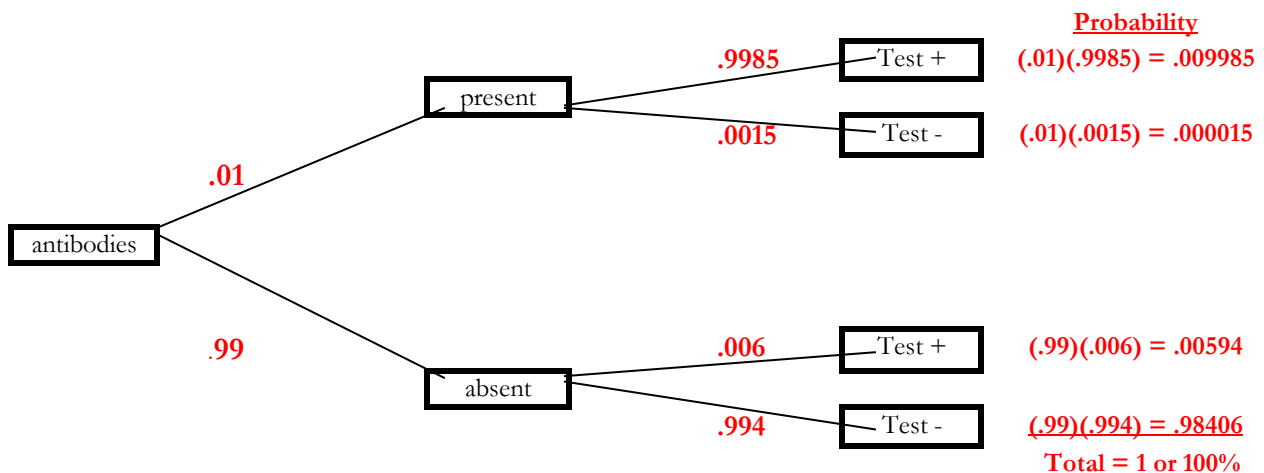
Question #1

Enzyme immunoassay tests are used to screen blood specimens for the presence of antibodies to HIV, the virus that causes AIDS. The presence of antibodies indicates the presence of the HIV virus. The test is quite accurate but is not always correct. The following table gives the probabilities of positive and negative test results when the blood tested does and does not actually contain antibodies to HIV.

| | Test Result | |
|--------------------|--------------|--------------|
| | Positive (+) | Negative (-) |
| Antibodies present | 0.9985 | 0.0015 |
| Antibodies absent | 0.0060 | 0.9940 |

Suppose that 1% of a large population carries antibodies to HIV in their blood.

- (a) Draw a tree diagram for selecting a person from this population (outcomes: antibodies present or absent) and for testing their blood (outcomes: test positive or negative).



- (b) What is the probability that the test is positive for a randomly chosen person in this population?

Answer: .015925

This tree shows 4 mutually exclusive outcomes for a person who either has or does not have the antibody and who either tests positive or negative.

Thus, the answer is obtained by summing the probability of the mutually exclusive outcomes that satisfy the event of a positive test.

$$\begin{aligned}\Pr[\text{test positive}] &= \Pr[\text{antibody and positive test}] + \Pr[\text{NO antibody and positive test}] \\ &= .009985 + .00594 \\ &= .015925\end{aligned}$$

- (c) What is the probability that a person in this population has the HIV virus, given that they test negative?

Answer: 0.0000152, representing a 0.0015% chance, approximately.

Using Bayes Rule

$$\begin{aligned}\Pr[\text{antibody}|\text{test-}] &= \frac{\Pr[\text{antibody and test-}]}{\Pr[\text{test-}]} \\ &= \frac{\Pr[\text{antibody}] \cdot \Pr[\text{test-}|\text{antibody}]}{\Pr[\text{antibody}] \cdot \Pr[\text{test-}|\text{antibody}] + \Pr[\text{NOantibody}] \cdot \Pr[\text{test-}|\text{Noantibody}]} \\ &= \frac{(.01)(.0015)}{(.01)(.0015) + (.99)(.994)} \\ &= .0000152\end{aligned}$$

Question #2

In introductory epidemiology, one of the study designs that is introduced is the (prospective) **cohort study**. In this type of study involving two groups, the investigator enrolls a pre-set number (set by design) of participants into each of the two groups that are generically described as “exposed” and “not exposed” and follows them forward to a designated end of the observation period, at which point one or more outcomes are measured.

The following table is from a **cohort study** of Danish men and women that investigated two outcomes, alcohol intake and mortality, in relationship to a number of possible influences: sex, age, body mass index, and smoking. Shown in this table is a cross-tabulation of alcohol intake and death, by sex and level of alcohol intake.

Table 8.2 The distribution of alcohol intake and deaths by sex and level of alcohol intake. Reproduced from *BMJ*, 308, 302–6, courtesy of BMJ Publishing Group

| Alcohol intake (beverages a week)* | Men | | Women | |
|---------------------------------------|-------------------|---------------------|-------------------|---------------------|
| | No of subjects | No (%) of deaths | No of subjects | No (%) of deaths |
| <1 | 625 | 195 (31.2) | 2472 | 394 (15.9) |
| 1–6 | 1183 | 252 (21.3) | 3079 | 283 (9.2) |
| 7–13 | 1825 | 383 (21.0) | 1019 | 96 (9.4) |
| 14–27 | 1234 | 285 (23.1) | 543 | 46 (8.5) |
| 28–41 | 585 | 118 (20.2) | 72 | 6 (8.3) |
| 42–69 | 388 | 99 (25.5) | 29 | 5 (17.2) |
| > 69 | 211 | 66 (31.3) | 20 | 1 (5.0) |
| Total | 6051 | 1398 (23.1) | 7234 | 831 (11.5) |

*One beverage contains 9–13 g alcohol.

- (a) From the information in the table, construct a table with 2 rows and 2 columns. Define your rows by sex and your columns by mortality. What you will have constructed is called a **contingency table**, and specifically, a **2x2 table**.

Answer:

2x2 table

| | Dead | Alive | |
|-------|------|-------|-------|
| Men | 1398 | 4653 | 6051 |
| Women | 831 | 6403 | 7234 |
| | 2229 | 11056 | 13285 |

Some preliminary calculations to get the numbers

| Men | dead | alive | row total |
|--------------|------|-------|-----------|
| | 195 | 430 | 625 |
| | 252 | 931 | 1183 |
| | 383 | 1442 | 1825 |
| | 285 | 949 | 1234 |
| | 118 | 467 | 585 |
| | 99 | 289 | 388 |
| | 66 | 145 | 211 |
| Column total | 1398 | 4653 | 6051 |

| Women | dead | alive | row total |
|--------------|------|-------|-----------|
| | 394 | 2078 | 2472 |
| | 283 | 2796 | 3079 |
| | 96 | 923 | 1019 |
| | 46 | 497 | 543 |
| | 6 | 66 | 72 |
| | 5 | 24 | 29 |
| | 1 | 19 | 20 |
| Column total | 831 | 6403 | 7234 |

Yielding:

2x2 table

| | Dead | Alive | |
|-------|------|-------|-------|
| Men | 1398 | 4653 | 6051 |
| Women | 831 | 6403 | 7234 |
| | 2229 | 11056 | 13285 |

- (b) Next, construct the following contingency table, again with 2 rows and 2 columns.
 Define your first row to be persons who consume less than one beverage per week.
 Define your second row to be persons who consume more than 69 beverages per week.
 Define your columns by mortality.

Answer:

| | Dead | Alive | |
|--------------------------|------|-------|------|
| Less than 1 drink/week | 589 | 2508 | 3097 |
| More than 69 drinks/week | 67 | 164 | 231 |
| | 656 | 2672 | 3328 |

Some preliminary calculations

Men

| | Dead | Alive | Row Total |
|--------------------------|------|-------|-----------|
| Less than 1 drink/week | 195 | 430 | 625 |
| More than 69 drinks/week | 66 | 145 | 211 |
| | 261 | 575 | 836 |

Women

| | Dead | Alive | Row Total |
|--------------------------|------|-------|-----------|
| Less than 1 drink/week | 394 | 2078 | 2472 |
| More than 69 drinks/week | 1 | 19 | 20 |
| | 395 | 2097 | 2492 |

Answer is the sum of the two tables. For example, in row 1 & column 1,
 $589 = 195 + 394$:

| | Dead | Alive | Row Total |
|--------------------------|------|-------|-----------|
| Less than 1 drink/week | 589 | 2508 | 3097 |
| More than 69 drinks/week | 67 | 164 | 231 |
| | 656 | 2672 | 3328 |

- (c) Using the information in your 2x2 table that you constructed in Question #2b, calculate the risk of death among persons who consume less than one beverage per week. Then calculate the risk of death among persons who consume more than 69 beverages per week.

| | Dead | Alive | | Risk of Death = |
|--------------------------|------|-------|------|--------------------------|
| Less than 1 drink/week | 589 | 2508 | 3097 | $589/3097 = 0.190184049$ |
| More than 69 drinks/week | 67 | 164 | 231 | $67/231 = 0.29004329$ |
| | 656 | 2672 | 3328 | |

- (d) In 1-2 sentences, compare the two risk estimates you obtained in Question #2c.

The estimated risk of death is approximately 1.5 times greater for persons who drink more than 69 drinks/week (29% risk) relative to those who drink less than 1 drink/week (19% risk).

Question #3

Another study design that is introduced in introductory epidemiology is the **case-control study**. This study design also calls for the comparison of two groups. Here, however, the investigator enrolls set (again, set by design) numbers of participants, defined by their disease status at the start of the study. **“Cases”** are the enrollees with disease. **“Controls”** are the enrollees who do not have the disease under investigation. The investigation involves looking back in time (“retrospective review”) at the histories of all study participants. The goal of this “back in time” look is to see if the cases are different from the controls with respect to their history of some exposure of interest.

The table below is from a **case-control study** that investigated the relationship of occurrences of Down Syndrome (**cases**) to history of exposure to maternal smoking during pregnancy. Shown in the table are some characteristics of the mothers, together with their status with respect to their history of smoking during pregnancy.

Table 8.3 Basic characteristics of mothers in a case-control study of maternal smoking and Down syndrome. Reproduced from *Amer. J. Epid.*, 149, 442–6, courtesy of Oxford University Press

Selected characteristics of Down syndrome cases and birth-matched controls. Washington State, 1984–1994

| | Cases (n = 775) | | Controls (n = 7750) | |
|---------------------------------|-----------------|------|---------------------|------|
| | No. | % | No. | % |
| Smoking during pregnancy | | | | |
| Age < 35 years | | | | |
| Yes | 112 | 20.0 | 1411 | 20.2 |
| No | 421 | 75.0 | 5214 | 74.6 |
| Unknown | 28 | 5.0 | 363 | 5.2 |
| Aged ≥ 35 years | | | | |
| Yes | 15 | 7.0 | 108 | 14.2 |
| No | 186 | 86.9 | 611 | 80.2 |
| Unknown | 13 | 6.1 | 43 | 5.6 |

- (a) Using the information in the table, construct separate 2x2 contingency tables, one for mothers aged < 35 years and the other for mothers aged ≥ 35 years. Define rows by exposure (smoked during pregnancy versus not). Define columns by case status (cases versus controls).

Age < 35

| | Case | Control | |
|--------------------------------|------|---------|------|
| Hx smoking during pregnancy | 112 | 1411 | 1523 |
| Did not smoke during pregnancy | 421 | 5214 | 5635 |
| | 533 | 6625 | 7158 |

Age ≥ 35

| | Case | Control | |
|--------------------------------|------|---------|-----|
| Hx smoking during pregnancy | 15 | 108 | 123 |
| Did not smoke during pregnancy | 186 | 611 | 797 |
| | 201 | 719 | 920 |

- (b) For each of the 2x2 tables you constructed in Question #3a, calculate two odds:
- Odds of smoking during pregnancy among cases
 - Odds of smoking during pregnancy among controls

Age < 35

| | Case | Control | |
|--------------------------------|------|---------|------|
| Hx smoking during pregnancy | 112 | 1411 | 1523 |
| Did not smoke during pregnancy | 421 | 5214 | 5635 |
| | 533 | 6625 | 7158 |

| | Cases | Controls |
|----------------------|-------------|-------------|
| Odds of hx smoking = | 112/421= | 1411/5214= |
| | 0.266033254 | 0.270617568 |

Age ≥ 35

| | Case | Control | |
|--------------------------------|------|---------|-----|
| Hx smoking during pregnancy | 15 | 108 | 123 |
| Did not smoke during pregnancy | 186 | 611 | 797 |
| | 201 | 719 | 920 |

| | Cases | Controls |
|----------------------|-------------|-------------|
| Odds of hx smoking = | 15/186= | 108/611= |
| | 0.080645161 | 0.176759411 |

- (c) Using the calculations of odds that you obtained in Question #3b, calculate two odds ratios:

- Odds Ratio for history of maternal smoking among mothers age < 35
- Odds Ratio for history of maternal smoking among mothers age ≥ 35

Age < 35

| | Case | Control | |
|--------------------------------|------|---------|------|
| Hx smoking during pregnancy | 112 | 1411 | 1523 |
| Did not smoke during pregnancy | 421 | 5214 | 5635 |
| | 533 | 6625 | 7158 |

| | Cases | Controls | OR=odds of hx(cases)/odds of hx(controls) |
|----------------------|-------------|-------------|---|
| Odds of hx smoking = | 112/421= | 1411/5214= | 0.26603/0.2706= |
| | 0.266033254 | 0.270617568 | 0.983059807 |

Age ≥ 35

| | Case | Control | |
|--------------------------------|------|---------|-----|
| Hx smoking during pregnancy | 15 | 108 | 123 |
| Did not smoke during pregnancy | 186 | 611 | 797 |
| | 201 | 719 | 920 |

| | Cases | Controls | OR=odds of hx (cases)/odds of hx (controls) |
|----------------------|-------------|-------------|---|
| Odds of hx smoking = | 15/186= | 108/611= | 0.0806/0.1768= |
| | 0.080645161 | 0.176759411 | 0.456242533 |

(d) In 1-2 sentences, interpret your results in Question #3c.

This case-control study provides no evidence of an adverse association of maternal smoking during pregnancy and Down Syndrome births. Among mothers < 35 years of age, the estimated odds ratio (OR = 0.98) is nearly equal to the null value of 1. Among mothers ≥ 35 years of age, the estimated odds ratio (OR = 0.46) is substantially less than 1.

Question #4

This question is intended to re-enforce your appreciation of the distinction between the two study designs: prospective cohort versus case-control.

In 1-2 sentences, why can't you calculate risk in a case-control study?

In a case-control study, participants are not selected on the basis of their exposure to the predictor of interest and then followed for the occurrences of the outcome, which would then permit the estimation of risk. Instead, participants are selected on the basis of their already having the outcome or not; indeed, these might even be equal sample sizes. The column totals in your 2x2 table therefore cannot be used to estimate risk of outcome.