Statistics 305/605: Introduction to Biostatistical Methods for Health Sciences

Chapter 15, part 3: McNemar's Test

Jinko Graham

2018-10-22

Paired Data and McNemar's Test

- The chi-square test is not appropriate when the study is designed to collect paired data.
- Example: Study of smoking habits over time.
 - ▶ A random sample of 2110 people were questioned about smoking status in 1980 and again in 1982.
 - Are smoking status and year associated? i.e, does the population proportion of smokers differ by year?
 - ▶ Test $H_0: p_{1980} p_{1982} = 0$ vs. $H_a: p_{1980} p_{1982} \neq 0$, where p_{1980} and p_{1982} are the population proportions of smokers in 1980 and 1982, respectively.
- ► The data might look as follows:

| | Smoking | |
|--------|---------|------|
| person | 1980 | 1982 |
| 1 | no | yes |
| 2 | no | no |
| 3 | yes | no |
| : | : | : |
| 2110 | yes | yes |

▶ The data could be arranged in a table as:

| | | Year | | |
|-------|-----|------|------|------|
| | | 1980 | 1982 | |
| Smoke | Yes | 717 | 696 | 1413 |
| | No | 1393 | 1414 | 2807 |
| | | 2110 | 2110 | 4220 |

- ▶ But this table is misleading: the 4220 observations that it reports are from 2110 double-counted people.
- ► The 4220 observations counted in the table are not independent, but rather paired observations from just 2110 people.

► To clarify the paired nature of the data, they are typically arranged as:

| | | 1982 (after) | | |
|---------------|-----------|--------------|-----------|------|
| | | Smoker | Nonsmoker | |
| 1980 (before) | Smoker | 620 | 97 | 717 |
| | Nonsmoker | 76 | 1317 | 1393 |
| | | 696 | 1414 | 2110 |

- ▶ The observations that are counted in the cells of this table are people and are independent.
- ► The cells with the same before- and after-status of the subject are called *concordant*.
- ► The cells with different before- and after-status are called discordant.
- McNemar's test is a contrast between the discordant cells, and ignores the concordant cells.

| | 1982 (after) | | |
|-----------|--------------|--|--|
| | Smoker | Nonsmoker | |
| Smoker | 620 | r = 97 | 717 |
| Nonsmoker | s = 76 | 1317 | 1393 |
| | 696 | 1414 | n = 2110 |
| | | $\begin{array}{c} & \text{Smoker} \\ \text{Smoker} & 620 \\ \text{Nonsmoker} & s = 76 \end{array}$ | $\begin{array}{c cccc} & Smoker & Nonsmoker \\ Smoker & 620 & r = 97 \\ Nonsmoker & s = 76 & 1317 \end{array}$ |

- Our sample of 2110 individuals has 620 continuing smokers, r = 97 quitters and s = 76 starters in 1982.
- In 1980 and 1982, the sample proportions of smokers are, respectively,

$$\hat{p}_{1980} = (620 + r)/n = (620 + 97)/2110$$

 $\hat{p}_{1982} = (620 + s)/n = (620 + 76)/2110$

- ▶ So, $\hat{p}_{1980} \neq \hat{p}_{1982}$ when $r \neq s$; or when $\hat{p}_{start} \neq \hat{p}_{quit}$, where
 - $\hat{p}_{start} = s/n$ is the proportion starting in 1982
 - $\hat{p}_{quit} = r/n$ is the proportion quitting in 1982.
- ► Re-express $H_0: p_{1980} p_{1982} = 0$ vs. $H_a: p_{1980} p_{1982} \neq 0$ as

•
$$H'_0$$
: $p_{quit} = p_{start}$ vs. H'_a : $p_{quit} \neq p_{start}$.

McNemar's Test

- As always, base the hypothesis test on the difference in observed proportions $\hat{p}_{start} \hat{p}_{quit} = (r s)/n$ and its standard error.
- We'll skip the derivation but the test statistic ends up being:

$$X^2 = \frac{(r-s)^2}{r+s} \sim \chi_1^2.$$

- ▶ If the number of quitters, r, is very different from the number of starters, s, the statistic X^2 is **big** and we reject H_0 in favour of H_a .
- ► An alternate form that uses a continuity correction for small samples (text, page 351) is

$$X^2 = \frac{(|r-s|-1)^2}{r+s} \sim \chi_1^2.$$

McNemar's Test for the Smoking Data

▶ We have r = 97 and s = 76. The test statistic with continuity correction is

$$\frac{(|97-76|-1)^2}{97+76} = 2.31$$

and the corresponding p-value is 0.128 (see R demo).

▶ Taking $\alpha = .05$, there is insufficient statistical evidence to conclude that smoking status is associated with year (the pvalue 0.128 is > 0.05).

Other Examples of Paired Data

- Scoring individuals from the same matched pair.
 - e.g. case-control pairs in which the control has been matched to the case on a number of characteristics.
- Scoring the same experimental unit with two different techniques
- ▶ Ratings of the same experimental unit by two different raters
- Genetic variants that were transmitted/not transmitted from a parent (experimental unit) to its offspring

Example: Transmission/Disequilibrium Test (TDT)

- ▶ Spielman *et al.*, 1993 click developed a test for genetic linkage that is an application of McNemar's test
- Genetic linkage DNA segments that are physically close together on a chromosome tend to be co-transmitted from parent to offspring.
 - ▶ A genetic marker that is close to, or linked, to a disease-causing mutation should tend to be co-transmitted with the disease.
- ► Application to type 1 (autoimmune) diabetes (T1D):
 - ▶ A genetic marker called 5'FP located near the insulin gene was associated with T1D in the population. Is 5'FP linked to a disease-causing mutation?
 - ▶ If so, we should see certain variants of 5'FP over-represented in transmissions from parents to children affected by T1D.

Diabetes Data from Spielman et al.

- The marker had two variants denoted 1 and X.
- ► Study of 124 heterozygous parents of children with T1D; heterozygous means that they carry a 1 and an X.
- The dataset has
 - one row for each parent, and
 - two columns, one for the variant that was transmitted from the parent to the affected child, and one for the variant that was not transmitted.

| ## | | transmitted | untransmitted |
|----|---|-------------|---------------|
| ## | 1 | 1 | X |
| ## | 2 | 1 | X |
| ## | 3 | 1 | X |
| ## | 4 | 1 | X |
| ## | 5 | 1 | X |
| ## | 6 | 1 | X |
| ## | 7 | 1 | X |
| ## | 8 | 1 | X |

McNemar's test on T1D Data

Cross tabulating transmitted and untransmitted variables for the heterozygous parents yields the following table.

```
## untransmitted
## transmitted 1 X
## 1 0 78
## X 46 0
```

McNemar's test applied to the T1D data (see R demo):

```
##
## McNemar's Chi-squared test
##
## data: tt
## McNemar's chi-squared = 8.2581, df = 1, p-value = 0.004057
```

- ► There is strong evidence that the 1 variant is preferentially transmitted over the X variant to the affected child.
 - ▶ The marker near the insulin gene is genetically linked to T1D.