Brian Caffo

Table of contents

Independer group t intervals

Likelihoo method

Unequal variances

# Mathematical Biostatistics Bootcamp: Lecture 10, T Confidence Intervals

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Table of contents

group *t* intervals

Likelihoo method

Unequal variances

- 1 Table of contents
- 2 Independent group t intervals
- 3 Likelihood method
- 4 Unequal variances

Unequal variance

## Independent group t confidence intervals

- Suppose that we want to compare the mean blood pressure between two groups in a randomized trial; those who received the treatment to those who received a placebo
- We cannot use the paired t test because the groups are independent and may have different sample sizes
- We now present methods for comparing independent groups

Unequal variances

- Let  $X_1, \ldots, X_{n_x}$  be iid  $N(\mu_x, \sigma^2)$
- Let  $Y_1, \ldots, Y_{n_y}$  be iid  $N(\mu_y, \sigma^2)$
- Let  $\bar{X}$ ,  $\bar{Y}$ ,  $S_x$ ,  $S_y$  be the means and standard deviations
- Using the fact that linear combinations of normals are again normal, we know that  $\bar{Y} \bar{X}$  is also normal with mean  $\mu_y \mu_x$  and variance  $\sigma^2(\frac{1}{p_x} + \frac{1}{p_y})$
- The pooled variance estimator

$$S_p^2 = \{(n_x - 1)S_x^2 + (n_y - 1)S_y^2\}/(n_x + n_y - 2)$$

is a good estimator of  $\sigma^2$ 

Unequal variance

#### Note

- The pooled estimator is a mixture of the group variances, placing greater weight on whichever has a larger sample size
- If the sample sizes are the same the pooled variance estimate is the average of the group variances
- The pooled estimator is unbiased

$$E[S_p^2] = \frac{(n_x - 1)E[S_x^2] + (n_y - 1)E[S_y^2]}{n_x + n_y - 2}$$
$$= \frac{(n_x - 1)\sigma^2 + (n_y - 1)\sigma^2}{n_x + n_y - 2}$$

• The pooled variance estimate is independent of  $\bar{Y} - \bar{X}$  since  $S_x$  is independent of  $\bar{X}$  and  $S_y$  is independent of  $\bar{Y}$  and the groups are independent

Table of

Independent group t intervals

Likelihoo method

Unequal variances

 The sum of two independent Chi-squared random variables is Chi-squared with degrees of freedom equal to the sum of the degrees of freedom of the summands

Therefore

$$(n_x + n_y - 2)S_p^2/\sigma^2 = (n_x - 1)S_x^2/\sigma^2 + (n_y - 1)S_y^2/\sigma^2$$

$$= \chi_{n_x - 1}^2 + \chi_{n_y - 1}^2$$

$$= \chi_{n_x + n_y - 2}^2$$

Unequal variance

## Putting this all together

• The statistic

$$\frac{\frac{\bar{Y} - \bar{X} - (\mu_y - \mu_x)}{\sigma \left(\frac{1}{n_x} + \frac{1}{n_y}\right)^{1/2}}}{\sqrt{\frac{(n_x + n_y - 2)S_\rho^2}{(n_x + n_y - 2)\sigma^2}}} = \frac{\bar{Y} - \bar{X} - (\mu_y - \mu_x)}{S_\rho \left(\frac{1}{n_x} + \frac{1}{n_y}\right)^{1/2}}$$

is a standard normal divided by the square root of an independent Chi-squared divided by its degrees of freedom

- Therefore this statistic follows Gosset's t distribution with  $n_x + n_y 2$  degrees of freedom
- Notice the form is (estimator true value) / SE

Unequal variances

• Therefore a  $(1-\alpha) \times 100\%$  confidence interval for  $\mu_V - \mu_X$  is

$$\bar{Y} - \bar{X} \pm t_{n_x + n_y - 2, 1 - \alpha/2} S_p \left( \frac{1}{n_x} + \frac{1}{n_y} \right)^{1/2}$$

- Remember this interval is assuming a constant variance across the two groups
- If there is some doubt, assume a different variance per group, which we will discuss later

#### Likelihood method

• Exactly as before,

$$\frac{\bar{Y} - \bar{X}}{S_{\rho} \left(\frac{1}{n_{x}} + \frac{1}{n_{y}}\right)^{1/2}}$$

follows a non-central t distribution with non-centrality parameter  $\frac{\mu_{y}-\mu_{x}}{\sigma\left(\frac{1}{n_{x}}+\frac{1}{n_{y}}\right)^{1/2}}$ 

• Therefore, we can use this statistic to create a likelihood for  $(\mu_y - \mu_x)/\sigma$ , a standardized measure of the change in group means

Brian Caffo

Table of contents

Independent group *t* intervals

Likelihood method

Unequal variance: Example from Rosner Fundamentals of Biostatistics, Page 304

- Comparing SBP for 8 oral contraceptive users versus 21 controls
- $\bar{X}_{OC}=132.86$  mmHg with  $s_{OC}=15.34$  mmHg
- $\bar{X}_C=127.44$  mmHg with  $s_C=18.23$  mmHg
- Pooled variance estimate

$$s_p^2 = \frac{7(15.34)^2 + 20(18.23)^2}{8 + 21 - 2} = 307.8$$

- $t_{27,.975} = 2.052$  (in R, qt(.975, df = 27))
- Interval

$$132.86 - 127.44 \pm 2.052 \left\{ 307.8 \left( \frac{1}{8} + \frac{1}{21} \right)^{1/2} \right\} = [-9.52, 20.36]$$

Independen group t intervals

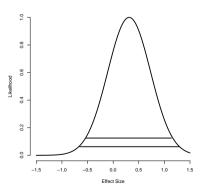
Likelihood method

Unequal variances

# Likelihood plot for the effect size

Reasonable values for the effect size from the confidence interval

$$[-9.52, 20.36]/sp = [-.54, 1.16]$$



Independent group t intervals

Likelihoo method

Unequal variances

#### Unequal variances

Note that under unequal variances

$$\bar{Y} - \bar{X} \sim N\left(\mu_y - \mu_x, \frac{\sigma_x^2}{n_x} + \frac{\sigma_y^2}{n_y}\right)$$

The statistic

$$\frac{\bar{Y} - \bar{X} - (\mu_y - \mu_x)}{\left(\frac{\sigma_x^2}{n_x} + \frac{\sigma_y^2}{n_y}\right)^{1/2}}$$

approximately follows Gosset's t distribution with degrees of freedom equal to

$$\frac{\left(S_{x}^{2}/n_{x}+S_{y}^{2}/n_{y}\right)^{2}}{\left(\frac{S_{x}^{2}}{n_{x}}\right)^{2}/(n_{x}-1)+\left(\frac{S_{y}^{2}}{n_{y}}\right)^{2}/(n_{y}-1)}$$

Unequal variances • Comparing SBP for 8 oral contraceptive users versus 21 controls

- $ar{X}_{OC}=$  132.86 mmHg with  $s_{OC}=$  15.34 mmHg
- $ar{X}_C=$  127.44 mmHg with  $s_C=$  18.23 mmHg
- df = 15.04,  $t_{15.04,.975} = 2.13$
- Interval

$$132.86 - 127.44 \pm 2.13 \left(\frac{15.34^2}{8} + \frac{18.23^2}{21}\right)^{1/2} = [-8.91, 19.75]$$