Brian Caffo

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# Mathematical Biostatistics Bootcamp: Lecture 10,

#### Brian Caffo

Department of Biostatistics

Johns Hopkins Bloomberg School of Public Health

Johns Hopkins University

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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  $\sigma^2(\frac{1}{n_X} + \frac{1}{n_Y})$
- 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 variances

### 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

Unequal variances

#### Result

- 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 variances

## Putting this all together

The statistic

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

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

## Confidence interval

• Therefore a  $(1-\alpha) \times 100\%$  confidence interval for  $\mu_{\rm Y} - \mu_{\rm X}$  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

Unequal variance

### Likelihood method

Exactly as before,

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

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

• 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

Unequal variances

## Example

Unequal variances

## Unequal variances

Note that under unequal variances

$$ar{Y} - ar{X} \sim N\left(\mu_y - \mu_x, rac{\sigma_x^2}{n_x} + rac{\sigma_y^2}{n_y}
ight)$$

• 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

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