# Statistics 305: Introduction to Biostatistical Methods for Health Sciences

Chapter 11: Inference for Two Means

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# Comparison of Two Means (Chapter 11)

#### Context

- We have measurements sampled from two populations.
- ► Want to make inference about the difference between distributions.
- ▶ In particular, we're interested in the difference between the two population means, denoted  $\mu_1$  and  $\mu_2$ .

#### Notation

- ▶ Let  $x_{11},...,x_{1n_1}$  denote a sample from the first population and  $x_{21},...,x_{2n_2}$  denote a sample from the second.
- ▶ The sample averages  $\bar{x}_1$  and  $\bar{x}_2$  estimate the population means  $\mu_1$  and  $\mu_2$ , respectively; so  $\bar{x}_1 \bar{x}_2$  estimates  $\mu_1 \mu_2$ .
- ▶ We're interested in confidence intervals for  $\mu_1 \mu_2$  and tests of  $H_0: \mu_1 \mu_2 = 0$  vs.  $H_a: \mu_1 \mu_2 \neq 0$ .

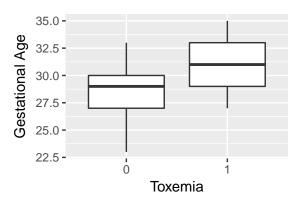
#### Example: Low Birthweight Infants

- Data on 100 infants with birth weight less than 1500g.
  - ▶ Variables are: head circumference (cm), birth length (cm), gestational age (wks), birth weight (g), mother's age (yrs), and mother's status for toxemia (1=high blood pressure during pregnancy, 0=not)
- Compare the distribution of variables such as age and birth weight in moms with to mom's without toxemia
- ▶ The first few rows of the data set are as follows:

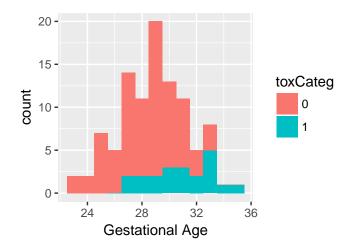
##		headcirc	length	gestage	birthwt	momage	toxemia
##	1	27	41	29	1360	37	0
##	2	29	40	31	1490	34	0
##	3	30	38	33	1490	32	0
##	4	28	38	31	1180	37	0
##	5	29	38	30	1200	29	1
##	6	23	32	25	680	19	0

## Gestational Age by Toxemia

- Question: Does the distribution of gestational age differ in moms with toxemia vs. moms without toxemia?
- Explore differences by toxemia status graphically, using boxplots (below) and histograms (next slide) in the sample.



# Gestational Age by Toxemia: Histograms



# Gestational Age by Toxemia: Summary Statistics

► The sample means and SDs of gestational age for each toxemia category are summarized below.

```
## # A tibble: 2 x 3
## toxCateg mean sd
## <fctr> <dbl> <dbl> *dbl>
## 1 0 28.35443 2.320687
## 2 1 30.90476 2.321740
```

- ► The sample means of the gestational ages differ between the toxemia groups, but the sample SDs look the same.
- Could the difference in the sample means be due to chance?

## Outline of Inference Approach

- ▶ Same basic approach to inference as in the one-sample problem:
  - Inference is based on the sampling distribution of the statistic  $\bar{X}_1 \bar{X}_2$
- ▶ Transform  $\bar{X}_1 \bar{X}_2$  to a *pivotal quantity*, Z, if population SDs  $\sigma_1$  and  $\sigma_2$  are known.
- ▶ When  $\sigma$ 's are unknown, as is typically the case, we substitute estimates to obtain a *pivotal quantity* T.
- Confidence intervals and hypothesis tests follow from the sampling distribution of T.
- Note: The following topics from the text are omitted:
  - ▶ Paired-samples *t*-test (Section 11.1 of text)
  - ► Two-sample *t*-test assuming equal SDs (Section 11.2.1)

# Sampling Distribution of $\bar{X}_1 - \bar{X}_2$

- ▶ We have simple random samples (SRSs) of size  $n_1$  for group 1 and  $n_2$  for group 2.
- ▶ These samples are independent.
- ▶ Then the distribution of  $\bar{X}_1 \bar{X}_2$  has
  - mean  $\mu_1 \mu_2$  and
  - ▶ SD  $\sqrt{\sigma_1^2/n_1 + \sigma_2^2/n_2}$ , where  $\sigma_1$  and  $\sigma_2$  are the population SDs for group 1 and group 2, respectively.
- ▶ If the sample sizes are large (will give a rule-of-thumb later), the CLT tells us that the shape of this distribution is approximately normal.

#### **Z** Transformation

- "Standardizing" a random variable by subtracting its population mean and dividing by its population SD gives a random variable with mean 0 & SD 1.
- For normal random variables, the transformation does not change the distribution; the standardized random variable is still normally distributed.
- ▶ Conclude that if  $\bar{X}_1 \bar{X}_2$  is approximately normal with mean  $\mu_1 \mu_2$  and SD  $\sqrt{\sigma_1^2/n_1 + \sigma_2^2/n_2}$ , then

$$Z = rac{(ar{X}_1 - ar{X}_2) - (\mu_1 - \mu_2)}{\sqrt{\sigma_1^2/n_1 + \sigma_2^2/n_2}} \sim N(0, 1),$$

where  $\sim$  means "is distributed as".

#### T Transformation

▶ Inserting sample SDs  $s_1$  and  $s_2$  for the parameters  $\sigma_1$  and  $\sigma_2$  in Z gives

$$T = \frac{(\bar{X}_1 - \bar{X}_2) - (\mu_1 - \mu_2)}{\sqrt{s_1^2/n_1 + s_2^2/n_2}}$$

- What is the distribution of T?
  - Unfortunately not a t-distribution, but we can approximate it by a t-distribution with ν df.
  - We won't ever use the formula for  $\nu$  but if you're curious it is given on page 270 of the text.
- ▶ Instead, computer software such as R automatically calculates  $\nu$  for us, as shown next.

### Illustration with Gestational Age and Toxemia

► The following is the output of the t.test() function for these data (see the R demo for details):

```
##
## Welch Two Sample t-test
##
## data: gestage by toxCateg
## t = -4.4745, df = 31.465, p-value = 9.365e-05
## alternative hypothesis: true difference in means is not equal to 0
## 95 percent confidence interval:
## -3.712089 -1.388574
## sample estimates:
## mean in group 0 mean in group 1
## 28.35443 30.90476
```

- ▶ The df is  $\nu = 31.465$ .
- ▶ We can also see that the software computed a 95% confidence interval and p-value. More on these in a few slides.

#### Confidence Intervals for $\mu_1 - \mu_2$

▶ The level-*C* CI for  $\mu_1 - \mu_2$  is of the form

estimate  $\pm$  margin of error

- ▶ The estimate is  $\bar{x}_1 \bar{x}_2$
- ▶ The margin of error is  $t^* \times SE$  where
  - $t^*$  is the upper (1-C)/2 critical value of the t-distribution with  $\nu$  df, and
  - ► SE is the **estimated SD** of  $\bar{x}_1 \bar{x}_2$ :  $\sqrt{s_1^2/n_1 + s_2^2/n_2}$

#### Application to Gestational Age and Toxemia

- Calculate a 90% CI for the difference between mean gestational age in the toxemia and non-toxemia groups.
- ▶ The relevant sample statistics to four digits are as follows:

group	sample mean $(\bar{x})$	sample SD $(s)$	sample size (n)
1: non-toxemia	28.35	2.321	79
2: toxemia	30.90	2.322	21

- ► Estimate is  $\bar{x}_1 \bar{x}_2 = 28.35 30.90 = -2.55$
- ▶ Margin of error is  $t^* \times SE$  where
  - computer software calculates a critical value of  $t^* = 1.695$  (see the R demo).
  - ► the SE is  $\sqrt{s_1^2/n_1 + s_2^2/n_2} = \sqrt{2.321^2/79 + 2.322^2/21} = 0.570.$
  - ▶ Hence the margin of error is  $1.696 \times 0.570 = 0.966$ .
- ► CI is (-2.55 0.966, -2.55 + 0.966) = (-3.516, -1.584).

#### Interpretation

- ▶ The 90% CI is approximately (-3.5, -1.6).
- The text suggests an interpretation such as:
  - "90% of intervals constructed in this way cover the true difference between mean gestational age in the non-toxemia and toxemia groups."
- Another common style of interpretation is:
  - We are 90% confident that the true difference between mean gestational age in the non-toxemia and toxemia groups is between −3.5 and −1.6."
- ▶ Or, we might find it more natural to switch the order of the groups, which would switch the sign of the difference:
  - "We are 90% confident that the true difference between mean gestational age in the toxemia and non-toxemia groups is between 1.6 and 3.5"

#### Hypothesis Test

For the null hypothesis  $H_0: \mu_1 - \mu_2 = 0$  of no difference between the groups, the formula for the observed t-statistic is

$$t = \frac{(\bar{x}_1 - \bar{x}_2) - 0}{\sqrt{s_1^2/n_1 + s_2^2/n_2}}$$

- ▶ Observed values of t that are extreme in the sense of being more compatible with  $H_a$  are taken as evidence against  $H_0: \mu_1 \mu_2 = 0$ .
- ▶ The *p*-value is the chance of a value that is as or more extreme than what we observed, under the null hypothesis.
- Using the same logic as in the one-sample problem (Ch 10), and taking T to be a random variable with a t distribution on  $\nu$  df, we have:
  - for  $H_a: \mu_1 \mu_2 \neq 0$ , the *p*-value is  $p = 2P(T \geq |t|)$ ,
  - for  $H_a: \mu_1 \mu_2 > 0$ , the *p*-value is  $p = P(T \ge t)$  and
  - ▶ for  $H_a$ :  $\mu_1 \mu_2 < 0$ , the *p*-value is  $p = P(T \le t)$ .

# Application to Gestational Age and Toxemia

We can re-use the calculations from the CI example (page 13), for the difference in sample means  $(\bar{x}_1 - \bar{x}_2 = -2.55)$  and the SE of the difference  $(\sqrt{s_1^2/n_1 + s_2^2/n_2} = 0.570)$ :

$$t = \frac{(\bar{x}_1 - \bar{x}_2) - 0}{\sqrt{s_1^2/n_1 + s_2^2/n_2}} = \frac{-2.55}{0.570} = -4.474.$$

- We use computer software (R demo) to calculate that the p-value for a 2-sided alternative hypothesis is equal to  $9.4 \times 10^{-5}$ , or 0.000094.
  - ▶ This *p*-value is twice the probability that a random variable, *T*, with a t distribution on 31.465 degrees of freedom, is greater than or equal to the absolute value of the observed t statistic, |*t*|.

#### Interpretation

- ► Small *p*-values (e.g. < .05) are taken as evidence against the null hypothesis.
- ▶ Our *p*-value of  $9.4 \times 10^{-5}$  is very small.
- If we had set a level of  $\alpha=10\%$  for the test, we'd declare that: "We reject the null hypothesis that the mean gestational age is the same in the toxemia and non-toxemia groups at the 10% level."
- ▶ If we hadn't set a level of the test in advance, we might report our results as:

"There is strong evidence that the mean gestational age is different in the toxemia and non-toxemia groups (p < 0.001)."

#### Cause and Effect

- Our two-sample t test has revealed that toxemia and gestational age are associated.
  - The distribution of gestational age is different in the two toxemia groups (different means, lower in toxemia group).
- But, an association does not mean that toxemia has a causal effect on gestational age.
  - It could be that gestational age affects toxemia.
  - Or, there could be a hidden confounding variable that affects both toxemia and gestational age, that accounts for their association.

(More on confounding later, when we study multiple regression.)

### Relationship Between Confidence Intervals and Tests

- ▶ In the low birth weight example, the 90% CI does not cover zero, and the hypothesis test of  $H_0: \mu_1 \mu_2 = 0$  vs  $H_a: \mu_1 \mu_2 \neq 0$  rejects the null hypothesis at the 10% level.
- Conversely, when a 90% CI does cover zero, the corresponding test against a two-sided alternative will retain the null hypothesis at the 10% level.
- ▶ This is a general property of tests of a population parameter  $\theta$ .
  - ▶ A level- $\alpha$  test of  $H_0$ :  $\theta = \theta_0$  versus  $H_a$ :  $\theta \neq \theta_0$  retains the null hypothesis if and only if the level  $(1 \alpha) \times 100\%$  CI covers  $\theta_0$ .

#### **Assumptions**

- ▶ The assumptions for inference are that:
  - ► The data are two SRSs from the two parent populations (e.g. mothers with toxemia and mothers without toxemia).
  - Either
    - 1. the data measurements in the parent populations are normally distributed with mean  $\mu_i$  and sd  $\sigma_i$ , written  $N(\mu_i, \sigma_i)$ , or
    - 2. the sample size  $n=n_1+n_2$  is "large" enough to rely on the CLT for the sample means  $\bar{X}_1$  and  $\bar{X}_2$ .
- Guidelines for n (from the text Basic Practice of Statistics by D. Moore):
  - ► For *n* < 15, use the *t*-based CI and hypothesis test if the data look to be approximately normally distributed.
  - For  $15 \le n < 40$ , use the *t*-based CI and hypothesis test, *except* in the presence of outliers or strong skewness in the data distribution.
  - ▶ For large samples ( $n \ge 40$ ), you can use the t-based CI and hypothesis test, even for clearly skewed distributions (because of the CLT).

# Checking the assumptions for the low-birth-weight example

- ▶ There were n = 100 babies in this data set.
- According to the rules-of-thumb on the previous slide, we can use the t-based CI and hypothesis test even if the population distributions are skewed.

## Summary

- ▶ Inference for the difference between two population means is based on either Z (SDs known) or T (SDs unknown).
- lacktriangle Confidence intervals are of the form estimate  $\pm$  margin of error
  - the margin of error is a critical value times SE
- ▶ To test the null hypothesis  $H_0: \mu_1 \mu_2 = 0$  against an alternative  $H_a$  we compute a test statistic t (or z if SDs known) and p-value
  - ightharpoonup can compare p-value to a significance level lpha to obtain a test
- Inference is considered reliable when the parent populations are normal, or when rules of thumb for sample sizes are satisfied.