



### BIOSTATISTICS

#### Lecture 7

## Hypothesis about Means and Proportions of Two Populations

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- Independent and dependent samples
- ◆ Comparison of means: t-test
- Paired t-test
- Comparison of two proportions

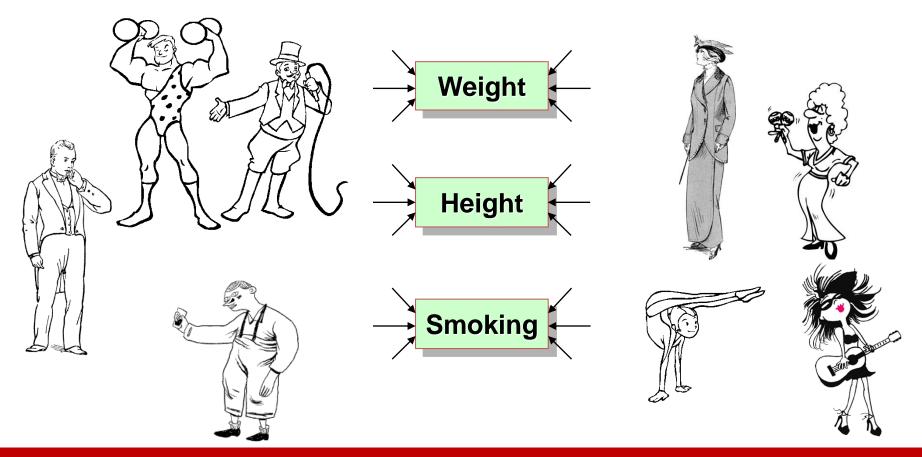


#### TWO POPULATIONS

**Independent Samples** 

#### **Independent samples**

Samples selected from two populations in such a way that the elements making up one sample are chosen independently of the elements making up the other sample.





#### TWO POPULATIONS

#### **Dependent Samples**

#### **Matched samples**

Samples in which each data value of one sample is matched with a corresponding data value of the other sample.

## Before treatment After treatment **Analysis** Prepare Target mRNAs RT/PCR Generate Microarray Hybridise target mixture to microarray



#### **MEANS OF TWO POPULATIONS**

#### **Example**

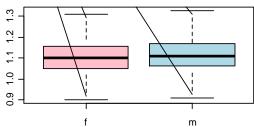
mice.xls

outliers are removed from boxplots

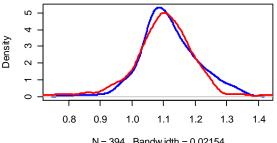
Final body weights (g) 20 m

Body weight distributions Density 10 30 50 20 N = 394 Bandw idth = 1.499

Weights change (g)



Distributions of weight change



N = 394 Bandwidth = 0.02154

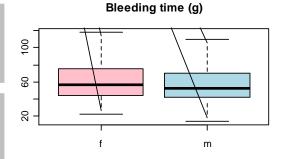
Distributions of bleeding times

N = 381 Bandw idth = 5.729

Q1: Is body weight for male and female significantly different?

Q2: Is weight change for male and female significantly different?

Q3: Is bleeding time for male and female significantly different?



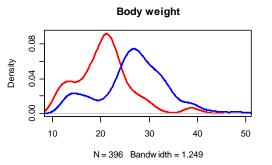
Density 0.000 50 150 100 200

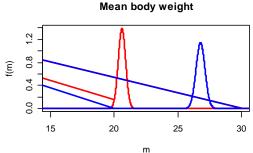


#### **MEANS OF TWO POPULATIONS**

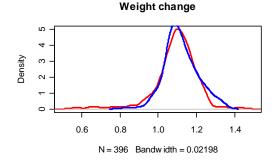
**Example** 

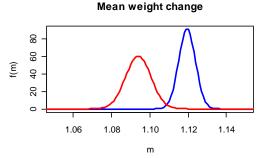
Q1: Is body weight for male and female significantly different?



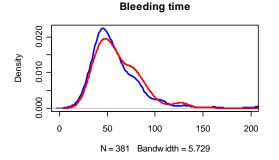


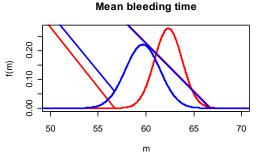
Q2: Is weight change for male and female significantly different?





Q3: Is bleeding time for male and female significantly different?







#### **HYPOTHESES**

**Theory** 

#### Two tail hypothesis

$$H_0$$
:  $\mu_1 = \mu_2$ 

$$H_a: \mu_1 \neq \mu_2$$

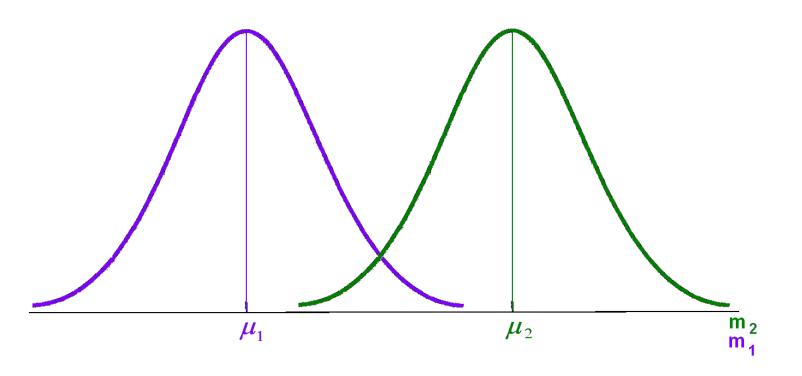
#### One tail hypothesis

$$H_0: \mu_1 \geq \mu_2$$

$$H_a$$
:  $\mu_1 < \mu_2$ 

$$H_0$$
:  $\mu_1 \le \mu_2$ 

$$H_a: \mu_1 > \mu_2$$





#### COMPARING MEANS

Theory

As we know how to work with standard hypotheses (comparison with constant  $\mu_0$ ), let us transform our hypothesis:

$$H_0$$
:  $\mu = \mu_0$   
 $H_a$ :  $\mu \neq \mu_0$ 

$$H_0: \mu_1 = \mu_2$$
 $H_a: \mu_1 \neq \mu_2$ 

$$H_0$$
:  $\mu_2 - \mu_1 = 0$   
 $H_a$ :  $\mu_2 - \mu_1 \neq 0$ 

$$H_a$$
:  $\mu_2 - \mu_1 \neq 0$ 

To use it, we need to know what is the distribution of  $D = m_2 - m_1$ 

#### Distribution of sum or difference of 2 normal random variables

The sum/difference of 2 (or more) normal random variables is a normal random variable with mean equal to sum/difference of the means and variance equal to SUM of the variances of the compounds.

Variables	$m_1$	$m_2$	$m_2-m_1$
Means	$\mu_1$	$\mu_2$	$\mu_2 - \mu_1$
Variances	$\sigma_1^{\ 2}$	$\sigma_2^2$	$\sigma_1^2 + \sigma_2^2$



#### COMPARING MEANS

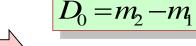
Theory

$$H_0$$
:  $\mu_2 - \mu_1 = D_0$ 

$$H_0$$
:  $\mu_2 - \mu_1 = D_0$   
 $H_a$ :  $\mu_2 - \mu_1 \neq D_0$ 

$$D_0 = \mu_2 - \mu_1$$

$$\sigma_{m_2-m_1} = \sqrt{\frac{\sigma_1^2}{n_1} + \frac{\sigma_2^2}{n_2}}$$



$$s_{m_2 - m_1} = \sqrt{\frac{s_1^2}{n_1} + \frac{s_2^2}{n_2}}$$

Statistics to be used for hypothesis testing:

if  $\sigma$  is known: z-statistics

$$z = \frac{m_2 - m_1 - D_0}{\sqrt{\frac{\sigma_1^2}{n_1} + \frac{\sigma_2^2}{n_2}}}$$

if  $\sigma$  is unknown: t-statistics

$$t = \frac{m_2 - m_1 - D_0}{\sqrt{\frac{s_1^2}{n_1} + \frac{s_2^2}{n_2}}}$$

This is what we call a t-test !!!



#### COMPARING MEANS

**Unpaired t-test: Algorithm** 

$$H_0$$
:  $\mu_2 - \mu_1 = D_0$ 

$$H_a$$
:  $\mu_2 - \mu_1 \neq D_0$ 

$$D_0 = m_2 - m_1$$

$$H_0: \mu_2 - \mu_1 = D_0$$
 $H_a: \mu_2 - \mu_1 \neq D_0$ 
Usually  $D_0 = 0$ 

$$s_{m_2 - m_1} = \sqrt{\frac{s_1^2 + s_2^2}{n_1 + n_2}}$$

#### 1. Build the statistics to be used for hypothesis testing:

$$t = \frac{m_2 - m_1 - D_0}{\sqrt{\frac{s_1^2}{n_1} + \frac{s_2^2}{n_2}}}$$

#### t-distribution has following degrees of freedom:

$$t = \frac{m_2 - m_1 - D_0}{\sqrt{\frac{s_1^2 + s_2^2}{n_1 + n_2}}}$$

$$df = \frac{\left(\frac{s_1^2 + s_2^2}{n_1 + n_2}\right)^2}{\frac{1}{n_1 - 1}\left(\frac{s_1^2}{n_1}\right)^2 + \frac{1}{n_2 - 1}\left(\frac{s_2^2}{n_2}\right)^2}{\left(\frac{s_1^2 + s_2^2}{n_2}\right)^2}$$

$$(n_1 + n_2)/2 < df < n_1 + n_2$$

$$df = (n-1)\frac{(s_1^2 + s_2^2)^2}{(s_1^4 + s_2^4)}$$

#### 2. Calculate the p-value

$$\Rightarrow$$
 = TDIST(ABS(t), df, 2)

**②. Or simply do:** 

 $\Rightarrow$  = T.TEST (array1, array2, 2, 3)

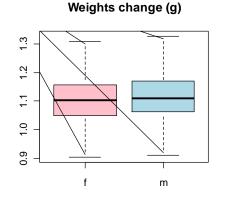


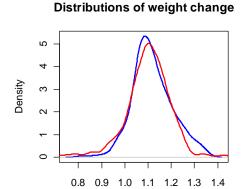
#### **UNPAIRED T-TEST**

#### **Example**

#### mice.xls

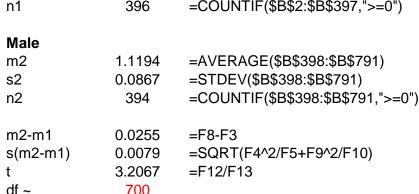
Q2: Is the mean of weight change for male and female significantly different?



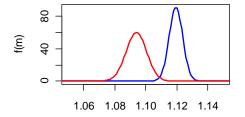


N = 394 Bandwidth = 0.02154

Parameter	Value	Command
Female		
m1	1.0939	=AVERAGE(\$B\$2:\$B\$397)
s1	0.1320	=STDEV(\$B\$2:\$B\$397)
n1	396	=COUNTIF(\$B\$2:\$B\$397,">=0")



p-value (1) 0.001403856 =TDIST(F14,700,2) p-value (2) 0.00140541 =TTEST(\$B\$2:\$B\$397,\$B\$398:\$B\$791,2,3) 





#### PAIRED T-TEST

#### **Theory and Example**

#### Paired t-test

In a paired t-test, instead of testing  $H_0$ :  $\mu_2 - \mu_1 = 0$ , use following steps:

- 1. Build a new random value  $y = x_1 x_2$  (subtract matched values).
- 2. Test whether one-sample mean  $\mu_v$  = 0 (see Lecture 6)

#### bloodpressure.xls

#### Systolic blood pressure (mmHg)

Subject	BP before	BP after
1	122	127
2	126	128
3	132	140
4	120	119
5	142	145
6	130	130
7	142	148
8	137	135
9	128	129
10	132	137
11	128	128
12	129	133

The systolic blood pressures of n=12 women between the ages of 20 and 35 were measured before and after usage of a newly developed oral contraceptive.

Q: Does the treatment affect the systolic blood pressure?

#### **Unpaired test**

**Paired test** 

Test	p-value	
unpaired	0.414662	
paired	0.014506	



#### COMPARING PROPORTIONS

**Theory** 

$$H_0$$
:  $\pi_1 = \pi_2$ 

$$H_a$$
:  $\pi_1 \neq \pi_2$ 

$$H_0$$
:  $\pi_1 - \pi_2 = 0$   
 $H_a$ :  $\pi_1 - \pi_2 \neq 0$ 

$$H_0: \pi_1 = \pi_2$$

$$H_a: \pi_1 \neq \pi_2$$

$$H_a: \pi_1 - \pi_2 \neq 0$$

$$\sigma_{p_1 - p_2} = \sqrt{\frac{p_1(1 - p_1)}{n_1} + \frac{p_2(1 - p_2)}{n_2}}$$

#### Pooled estimator of $\pi$

An estimator of a population proportion obtained by computing a weighted average of the point estimators obtained from two independent samples.

$$p = \frac{n_1 p_1 + n_2 p_2}{n_1 + n_2}$$

$$z = \frac{p_1 - p_2}{\sqrt{p(1-p)\left(\frac{1}{n_1} + \frac{1}{n_2}\right)}}$$

$$\sigma_{p_1-p_2} = \sqrt{p(1-p)\left(\frac{1}{n_1} + \frac{1}{n_2}\right)}$$



= 2\*NORM.S.DIST(-ABS(z),TRUE)



#### **COMPARING PROPORTIONS**

**Example** 

SWR/J	MA/MyJ	
f	f	
f	f	
f	f	
f	f	
f	f	
f	f	
f	f	
f	f	
f	m	
f	m	
m	m	
m	m	
m	m	
m	m	
m	m	
m	m	
m	m	
m	m	
m	m	
	m	
	m	
	m	
	m	

mice.xls

Q: Is the male proportion significantly different in these mouse strains (0.47 and 0.65)?

$$p = \frac{n_1 p_1 + n_2 p_2}{n_1 + n_2}$$

$$z = \frac{p_1 - p_2}{\sqrt{p(1-p)\left(\frac{1}{n_1} + \frac{1}{n_2}\right)}}$$

Gives less precise estimation OR

$$z = \frac{p_1 - p_2}{\sqrt{\frac{p_1(1-p_1)}{n_1} + \frac{p_2(1-p_2)}{n_2}}}$$

$$\Rightarrow$$
 = 2\*NORM.S.DIST(-ABS(z),TRUE)

	SWR/J	MA/MyJ	pooled
count male	9	15	24
n	19	23	42
р	0.474	0.652	0.571
z	-1.16		
p-val	0.244658997	p-value = 0.24	



#### **QUESTIONS?**

# Thank you for your attention

