# PG Dip in High Intensity Psychological Interventions

# Inference about means

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# Methods for inference about means

#### Large samples

- Single mean: Normal method (z method)
- Paired data: Normal method (z method)
- > Two samples: Normal method (z method)

# Small samples

- > Single mean: One sample t method
- ➤ Paired data: Paired t method
- > Two samples: Two sample t method (independent samples t method, two group t method)

#### Single mean, large sample method

Confidence interval using the Normal distribution.

Method can be used for any large sample.

Data may be from any distribution.

Distribution of birthweight in 1749 singleton pregnancies to Caucasian mothers in South London.



standard deviation = 563.2 g.

the mean = 13.5 g  $= 563.2 / \sqrt{1769}$ 



Confidence interval using the Normal distribution.

mean = 3296.0 g, standard deviation = 563.2 g, standard error of the mean = 13.5 g.

Large sample  $\rightarrow$  mean from Normal distribution with SD = 13.5 well estimated from data.



95% of observations from a Normal distribution are within 1.96 standard deviations from the mean.

96% confidence interval = 3296.0 - 1.96 × 13.5 g to 3296.0 + 1.96 × 13.5 g

= 3270 to 3322 g.

#### Single mean, large sample method

#### Assumptions:

- The observations are independent. We should not have, for example, a group of 100 observations where there are 10 subjects with 10 observations on each.
- The sample is large enough for the standard errors to be well estimated. My rule of thumb is 100 for one group.

# Paired data, large sample method

Confidence interval for mean difference using the Normal distribution.

Example: interventions for depression delivered using the internet.

Recruited 525 people with symptoms of depression identified in a survey.

Randomly allocated to websites:

- BluePages, information about depression (n = 166),
- MoodGYM, cognitive behaviour therapy (n = 182),
- attention placebo (n = 178).

Christensen H, Griffiths KM, Jorm AF. (2004) Delivering interventions for depression by using the internet: randomised controlled trial. *British Medical Journal* **328**, 265-268.

#### Paired data, large sample method

Baseline depression score (0-60) and fall after six weeks by treatment group for 525 patients with depression (Christensen *et al.*, 2004)

	number	Baseline mean	scores SD	Fall mea	in an	scores SD	
BluePages	165	21.1	10.4	3	. 9	9.1	
MoodGYM	182	21.8	10.5	4	2	9.1	
Controls	178	21.6	11.1	1	. 0	8.4	

90 (17%) of subjects did not return post-intervention questionnaires and the authors assumed that their scores were unchanged.

Differences must have a large spike of at least 90 observations at zero.

They could not have a Normal distribution.

#### Paired data, large sample method

BluePages: 165 subjects, mean fall in depression score = 3.9, standard deviation = 9.1.

Standard error of the mean difference = 0.71.

95% confidence interval for the mean fall is  $3.9 - 1.96 \times 0.71$  to  $3.9 + 1.96 \times 0.71$ = 2.5 to 5.3 points on the depression scale.

This is an interval estimate for the mean fall in depression score assuming non-responders do not change.

# Paired data, large sample method

BluePages: 165 subjects, mean fall in depression score = 3.9, standard deviation = 9.1, standard error of the mean = 0.71.

Test of significance

- null hypothesis: mean change in population is zero,
- alternative hypothesis: there is a change, in either direction.

Large sample  $\rightarrow$  mean will be from a Normal distribution with standard deviation equal to the standard error of the mean.

Observed sample mean minus the unknown population mean divided by the standard error will be an observation from the Standard Normal distribution.

#### Paired data, large sample method

BluePages: 165 subjects, mean fall in depression score = 3.9, standard deviation = 9.1, standard error of the mean = 0.71. Test of significance

• null hypothesis: mean change in population is zero.

If true then sample mean over standard error will be from a Standard Normal distribution.

z = 3.9/0.71 = 5.49.

P = 0.0000004.

Usually quote this as P<0.0001.

This is the large sample Normal test for a single mean, also called the z test for a single mean.

#### Paired data, large sample method

#### Assumptions:

- The observations are independent. We should not have, for example, a group of 100 observations where there are 10 subjects with 10 observations on each.
- The sample is large enough for the standard errors to be well estimated. My rule of thumb is 100 for one group.
- The mean and standard deviation of differences are constant, i.e. not related to the size of the variable.

Check by plotting difference against the average of the two measurements for the subject.

See later under paired t test.

#### Two means, large sample method

		Fall	in s	cores
	number	mean	SD	SE of mean
BluePages	165	3.9	9.1	0.71
MoodGYM	182	4.2	9.1	0.67
Controls	178	1.0	8.4	0.63

Confidence interval for difference between two means, BluePages minus MoodGYM.

Difference = -0.3

Standard error for the difference =  $\sqrt{0.71^2 + 0.67^2} = 0.98$ 

N.B. This only works when the groups are independent.

95% CI =  $-0.3 - 1.96 \times 0.98$  to  $-0.3 + 1.96 \times 0.98$ = -2.2 to +1.6

#### Two means, large sample method

		Fall	in s	cores
	number	mean	SD	SE of mean
BluePages	165	3.9	9.1	0.71
MoodGYM	182	4.2	9.1	0.67
Controls	178	1.0	8.4	0.63

Test of null hypothesis means are equal, BluePages versus MoodGYM.

Difference = -0.3, standard error for the difference = 0.98.

If null hypothesis true, difference/standard error will be from a Standard Normal distribution.

difference/standard error = -0.3/0.98 = -0.31.

From Normal distribution P = 0.8.

Large sample Normal or z test for two means.

# Two means, large sample method

#### Assumptions:

- The observations and groups are independent. We should not have links between observations in the two groups, such as a matched study where each subject in one group is matched, e.g. by age and sex, with a subject in the other group.
- The samples are large enough for the standard errors to be well estimated. My rule of thumb is at least 50 in each group.

# One sample t method for small samples

Example: nine patients with chronic non-healing wounds (Shukla *et al.*, 2004).

Biopsies were assessed using the microscopic angiogenesis grading system (MAGS) score, which provides an index of how well small blood vessels are developing and hence of epithelial regeneration.

High scores are good.

Observations: 20, 31, 34, 39, 43, 45, 49, 51, and 63.

mean = 41.7, standard deviation = 12.5, standard error of the mean = 4.2.

Shukla VK, Rasheed MA, Kumar M, Gupta SK, Pandey SS. (2004) A trial to determine the role of placental extract in the treatment of chronic non-healing wounds. *Journal of Wound Care* **13**, 177-9,

# One sample t method for small samples

Observations: 20, 31, 34, 39, 43, 45, 49, 51, and 63.

mean = 41.7, standard deviation = 12.5, standard error of the mean = 4.2.

95% confidence interval for the mean:

If the sample were large we could use

mean - 1.96 standard errors to mean + 1.96 standard errors.

We cannot use the large sample Normal method because the samples are too small.

The standard error will not be sufficiently well estimated.

#### One sample t method for small samples

Observations: 20, 31, 34, 39, 43, 45, 49, 51, and 63.

mean = 41.7, standard deviation = 12.5, standard error of the mean = 4.2.

95% confidence interval for the mean:

mean - ? standard errors to mean + ? standard errors.

We use Student's t distribution.

For a small sample, we must assume that the differences themselves follow a Normal distribution.

95% confidence interval:

mean  $- t_{0.05}$  standard errors to mean  $+ t_{0.05}$  standard errors.

#### One sample t method for small samples

Observations: 20, 31, 34, 39, 43, 45, 49, 51, and 63.

mean = 41.7, standard deviation = 12.5, standard error of the mean = 4.2.

95% confidence interval for the mean:

mean –  $t_{0.05}$  standard errors to mean +  $t_{0.05}$  standard errors.

What is *t*<sub>0.05</sub>?

This comes from Student's t distribution, defined as the distribution followed by a sample mean minus a population mean all divided by the standard error, when the observations follow a Normal distribution.

where  $t_{0.05}$  is the two-sided 5% point of the t distribution with degrees of freedom = number of observations minus one.









	Τw	<i>i</i> o taile	d prob	bability po	oints of	the t	Distrib	ution	
D.f.		Proba	ability	1	D.f.		Proba	ability	
	0.10	0.05	0.01	0.001		0.10	0.05	0.01	0.001
	(10%)	(5%)	(1%)	(0.1%)		(10%)	(5%)	(1%)	(0.1%)
1	6.31	12.70	63.66	636.62	16	1.75	2.12	2.92	4.02
2	2.92	4.30	9.93	31.60	17	1.74	2.11	2.90	3.97
3	2.35	3.18	5.84	12.92	18	1.73	2.10	2.88	3.92
4	2.13	2.78	4.60	8.61	19	1.73	2.09	2.86	3.88
5	2.02	2.57	4.03	6.87	20	1.73	2.09	2.85	3.85
6	1.94	2.45	3.71	5.96	21	1.72	2.08	2.83	3.82
7	1.90	2.36	3.50	5.41	22	1.72	2.07	2.82	3.79
8	1.86	2.31	3.36	5.04	23	1.71	2.07	2.81	3.77
9	1.83	2.26	3.25	4.78	24	1.71	2.06	2.80	3.75
10	1.81	2.23	3.17	4.59	25	1.71	2.06	2.79	3.73
11	1.80	2.20	3.11	4.44	30	1.70	2.04	2.75	3.65
12	1.78	2.18	3.06	4.32	40	1.68	2.02	2.70	3.55
13	1.77	2.16	3.01	4.22	60	1.67	2.00	2.66	3.46
14	1.76	2.15	2.98	4.14	120	1.66	1.98	2.62	3.37
15	1.75	2.13	2.95	4.07	00	1.65	1.96	2.58	3.29
D.f.	= Degi	rees of	f free	dom					
<b>oo</b> = ;	infini	ty, sa	me as	the Stand	ard No:	rmal D:	istrib	ution	



# One sample t method for small samples

Observations: 20, 31, 34, 39, 43, 45, 49, 51, and 63.

mean = 41.7, standard deviation = 12.5, standard error of the mean = 4.2.

95% confidence interval for the mean:

mean  $- t_{0.05}$  standard errors to mean  $+ t_{0.05}$  standard errors.

What is *t*<sub>0.05</sub>?

 $t_{0.05}$  is the two-sided 5% point of the t distribution with degrees of freedom = number of observations minus one.

This is the degrees of freedom for the sample variance.

We have 9 observations so 9 - 1 = 8 degrees of freedom.

	Τw	<i>i</i> o taile	d prob	pability p	oints of	the t	Distrib	ution	
D.f.		Proba	ability	7	D.f.		Proba	bility	,
	0.10	0.05	0.01	0.001		0.10	0.05	0.01	0.001
	(10%)	(5%)	(1%)	(0.1%)		(10%)	(5%)	(1%)	(0.1%)
1	6.31	12.70	63.66	636.62	16	1.75	2.12	2.92	4.02
2	2.92	4.30	9.93	31.60	17	1.74	2.11	2.90	3.97
3	2.35	3.18	5.84	12.92	18	1.73	2.10	2.88	3.92
4	2.13	2.78	4.60	8.61	19	1.73	2.09	2.86	3.88
5	2.02	2.57	4.03	6.87	20	1.73	2.09	2.85	3.85
6	1.94	2.45	3.71	5.96	21	1.72	2.08	2.83	3.82
7	1.90	2.36	3.50	5.41	22	1.72	2.07	2.82	3.79
8	1.86	2.31	3.36	5.04	23	1.71	2.07	2.81	3.77
9	1.83	2.26	3.25	4.78	24	1.71	2.06	2.80	3.75
10	1.81	2.23	3.17	4.59	25	1.71	2.06	2.79	3.73
11	1.80	2.20	3.11	4.44	30	1.70	2.04	2.75	3.65
12	1.78	2.18	3.06	4.32	40	1.68	2.02	2.70	3.55
13	1.77	2.16	3.01	4.22	60	1.67	2.00	2.66	3.46
14	1.76	2.15	2.98	4.14	120	1.66	1.98	2.62	3.37
15	1.75	2.13	2.95	4.07	00	1.65	1.96	2.58	3.29
D.f.	= Deg	rees of	f free	dom					
oo = ;	infini	ty, sa	me as	the Stand	lard No:	rmal D:	istribu	ution	

#### One sample t method for small samples

Observations: 20, 31, 34, 39, 43, 45, 49, 51, and 63.

mean = 41.7, standard deviation = 12.5, standard error of the mean = 4.2.

95% confidence interval for the mean:

mean –  $t_{0.05}$  standard errors to mean +  $t_{0.05}$  standard errors.

 $t_{0.05} = 2.31$ 

95% confidence interval for the mean:

41.7 - 2.31 × 4.2 to 41.7 + 2.31 × 4.2

= 32.0 to 51.4 MAGS units.

# One sample t method for small samples Assumptions:

- $\succ$  The observations are independent.
- $\succ$  The observations are from a Normal distribution.







MAGS scor placenta wounds (S	re before a l extract : Shukla <i>et a</i>	and after in 9 pati al., 2004	treatment with to ents with non-heal )	pi in
Subject	Before	After	Difference	
1	20	32	12	
2	31	47	16	
3	34	43	9	
4	39	43	4	
5	43	55	12	
6	45	52	7	
7	49	61	12	
8	51	55	4	
9	63	71	8	
Mean			9.33	
Standard	deviation		4.03	

			-
			-
			-
			-
			-

# Paired t method

For a small sample, we must assume that the differences themselves follow a Normal distribution.

95% confidence interval:

mean difference –  $t_{0.05}$  standard errors to mean difference +  $t_{0.05}$  standard errors.

where  $t_{0.05}$  is the two-sided 5% point of the t distribution with degrees of freedom = number of observations minus one.

Test of significance: refer mean difference / standard error to the t distribution with degrees of freedom = number of observations minus one.

# Paired t method

Example: Increase in MAGS score

Mean difference = 9.33, SE = 1.34 litres/min.

9 differences, hence 9 - 1 = 8 degrees of freedom.

	Τw	<i>i</i> o taile	d prob	pability po	oints o	f the t	Distrib	ution	
D.f.		Proba	ability	7	D.f.		Proba	bility	,
	0.10	0.05	0.01	0.001		0.10	0.05	0.01	0.001
	(10%)	(5%)	(1%)	(0.1%)		(10%)	(5%)	(1%)	(0.1%)
1	6.31	12.70	63.66	636.62	16	1.75	2.12	2.92	4.02
2	2.92	4.30	9.93	31.60	17	1.74	2.11	2.90	3.97
3	2.35	3.18	5.84	12.92	18	1.73	2.10	2.88	3.92
4	2.13	2.78	4.60	8.61	19	1.73	2.09	2.86	3.88
5	2.02	2.57	4.03	6.87	20	1.73	2.09	2.85	3.85
6	1.94	2.45	3.71	5.96	21	1.72	2.08	2.83	3.82
7	1.90	2.36	3.50	5.41	22	1.72	2.07	2.82	3.79
8	1.86	2.31	3.36	5.04	23	1.71	2.07	2.81	3.77
9	1.83	2.26	3.25	4.78	24	1.71	2.06	2.80	3.75
10	1.81	2.23	3.17	4.59	25	1.71	2.06	2.79	3.73
11	1.80	2.20	3.11	4.44	30	1.70	2.04	2.75	3.65
12	1.78	2.18	3.06	4.32	40	1.68	2.02	2.70	3.55
13	1.77	2.16	3.01	4.22	60	1.67	2.00	2.66	3.46
14	1.76	2.15	2.98	4.14	120	1.66	1.98	2.62	3.37
15	1.75	2.13	2.95	4.07	00	1.65	1.96	2.58	3.29
D.f.	= Degi	rees of	ffreed	dom					
oo = ;	infini	ty, sa	ne as	the Stand	ard No	rmal D:	istribu	ution	



# Paired t method

# Example: Increase in MAGS score

Mean difference = 9.33, SE = 1.34 litres/min.

9 differences, hence 9 - 1 = 8 degrees of freedom.

Using the 8 d.f. row, we get  $t_{0.05} = 2.31$ .

The 95% confidence interval:

9.33 – 2.31 × 1.34 to 9.33 + 2.31 × 1.34 = 6.2 to 12.4.

# Paired t method

# Example: Increase in MAGS score

Mean difference = 9.33, SE = 1.34 litres/min.

9 differences, hence 9 - 1 = 8 degrees of freedom.

Using the 8 d.f. row, we get  $t_{0.05} = 2.31$ .

The 95% confidence interval:

9.33 - 2.31 × 1.34 to 9.33 + 2.31 × 1.34 = 6.2 to 12.4.

#### Test of significance:

Mean/SE = 9.33/1.34 = 6.96

	Τw	<i>i</i> o taile	d prot	pability po	pints o	f the t	Distrib	ution	
D.f.		Proba	ability	7	D.f.		Proba	ability	7
	0.10	0.05	0.01	0.001		0.10	0.05	0.01	0.001
	(10%)	(5%)	(1%)	(0.1%)		(10%)	(5%)	(1%)	(0.1%)
1	6.31	12.70	63.66	636.62	16	1.75	2.12	2.92	4.02
2	2.92	4.30	9.93	31.60	17	1.74	2.11	2.90	3.97
3	2.35	3.18	5.84	12.92	18	1.73	2.10	2.88	3.92
4	2.13	2.78	4.60	8.61	19	1.73	2.09	2.86	3.88
5	2.02	2.57	4.03	6.87	20	1.73	2.09	2.85	3.85
6	1.94	2.45	3.71	5.96	21	1.72	2.08	2.83	3.82
7	1.90	2.36	3.50	5.41	22	1.72	2.07	2.82	3.79
8	1.86	2.31	3.36	5.04	23	1.71	2.07	2.81	3.77
9	1.83	2.26	3.25	4.78	24	1.71	2.06	2.80	3.75
10	1.81	2.23	3.17	4.59	25	1.71	2.06	2.79	3.73
11	1.80	2.20	3.11	4.44	30	1.70	2.04	2.75	3.65
12	1.78	2.18	3.06	4.32	40	1.68	2.02	2.70	3.55
13	1.77	2.16	3.01	4.22	60	1.67	2.00	2.66	3.46
14	1.76	2.15	2.98	4.14	120	1.66	1.98	2.62	3.37
15	1.75	2.13	2.95	4.07	00	1.65	1.96	2.58	3.29
D.f.	= Degi	rees of	ffreed	dom					
<b>oo</b> = ;	infini	ty, sa	ne as	the Stand	ard No	rmal D:	istribu	ution	



# Paired t method

# Example: Increase in MAGS score

Mean difference = 9.33, SE = 1.34 litres/min.

9 differences, hence 9 - 1 = 8 degrees of freedom.

Using the 8 d.f. row, we get  $t_{0.05} = 2.31$ .

The 95% confidence interval:

9.33 – 2.31 × 1.34 to 9.33 + 2.31 × 1.34 = 6.2 to 12.4.

#### Test of significance:

Mean/SE = 9.33/1.34 = 6.96

From t table, P<0.001. From computer program, P=0.0001.

#### Paired t method

# Assumptions:

- > The observations are independent.
- > The differences follow a Normal distribution.
- The mean and standard deviation of differences are constant, i.e. not related to the size of the variable.





This is also called the unpaired t method or test and the two group t method, Student's two sample t test.

Example: Capillary density (per mm<sup>2</sup>) in the feet of ulcerated patients and a healthy control group (data supplied by Marc Lamah)

		Cont	rols		Ulce	erated	d pati	ients
-	17.5	34.5	36.5	52.0	9.0	18.5	23.0	27.5
	27.5	31.0	38.0		11.0	20.0	23.0	28.0
	27.0	35.5	40.0		12.5	20.0	24.0	28.5
	29.5	33.5	39.5		18.0	22.0	26.5	29.0
	27.0	35.5	40.0		18.0	22.5	26.5	44.5
	29.0	34.0	40.0		18.0	22.5	27.0	
nber		19				2	23	
an		34.0	80			2	22.59	
		7.3	29				7.31	

-				
_		Controls		Ulcerated patients
-	17.5	34.5 36.5	52.0	9.0 18.5 23.0 27.5
	27.5	31.0 38.0		11.0 20.0 23.0 28.0
	27.0	35.5 40.0		12.5 20.0 24.0 28.5
	29.5	33.5 39.5		18.0 22.0 26.5 29.0
	27.0	35.5 40.0		18.0 22.5 26.5 44.5
	29.0	34.0 40.0		18.0 22.5 27.0
Number		19		23
lean		34.08		22.59
SD		7.29		7.31



Capillary density (per mm2) in the feet of ulcerated patients and a healthy control group (data supplied by Marc Lamah)

	Control	s Ul	Ulcerated patients				
	17.5 34.5 36 27.5 31.0 38 27.0 35.5 40 29.5 33.5 39 27.0 35.5 40 29.0 34.0 40	.5 52.0 9.   .0 11.   .0 12.   .5 18.   .0 18.   .0 18.	0 18.5 23.0 27.5 0 20.0 23.0 28.0 5 20.0 24.0 28.5 0 22.0 26.5 29.0 0 22.5 26.5 44.5 0 22.5 27.0				
Number	19		23				
Mean	34.08		22.59				
SD	7.29		7.31				
The standard error will not be sufficiently well estimated.							

#### Two sample t method

We cannot use the large sample Normal method because the samples are too small.

The standard error will not be sufficiently well estimated.

The distribution of the standard error estimate depends on the distribution of the observations themselves.

We must make two assumptions about the data:

- 1. the observations come from Normal distributions,
- the distributions in the two populations have the same variance. (N.B. The populations, not the samples from them, have the same variance.)

# Two sample t method

If the distributions in the two populations have the same variance, we need only one estimate of variance. We call this the common or pooled variance estimate.

The degrees of freedom are number of observations minus 2.

We use this common estimate of variance to estimate the standard error of the difference between the means.

Capillary density example:

Common variance = 53.31, SD = 7.30 capillaries/mm<sup>2</sup>, df = 19 + 23 - 2 = 40.

SE of difference = 2.26 capillaries/mm<sup>2</sup>.

Difference (control - ulcer) = 34.08 - 22.59 = 11.49capillaries/mm<sup>2</sup>.

Capillary density example:

Common variance = 53.31, SD = 7.30 capillaries/mm<sup>2</sup>, df = 19 + 23 - 2 = 40.

SE of difference = 2.26 capillaries/mm<sup>2</sup>.

Difference (control – ulcer) = 34.08 – 22.59 = 11.49 capillaries/mm<sup>2</sup>.

95% confidence interval for difference:

11.49 - ? × 2.26 to 11.49 + ? × 2.26.

? comes not from the Normal distribution but the t distribution with 40 degrees of freedom.

Two tailed probability points of the t Distribution									
D.f.	Probability			D.f.		Probability			
	0.10	0.05	0.01	0.001		0.10	0.05	0.01	0.001
	(10%)	(5%)	(1%)	(0.1%)		(10%)	(5%)	(1%)	(0.1%)
1	6.31	12.70	63.66	636.62	16	1.75	2.12	2.92	4.02
2	2.92	4.30	9.93	31.60	17	1.74	2.11	2.90	3.97
3	2.35	3.18	5.84	12.92	18	1.73	2.10	2.88	3.92
4	2.13	2.78	4.60	8.61	19	1.73	2.09	2.86	3.88
5	2.02	2.57	4.03	6.87	20	1.73	2.09	2.85	3.85
6	1.94	2.45	3.71	5.96	21	1.72	2.08	2.83	3.82
7	1.90	2.36	3.50	5.41	22	1.72	2.07	2.82	3.79
8	1.86	2.31	3.36	5.04	23	1.71	2.07	2.81	3.77
9	1.83	2.26	3.25	4.78	24	1.71	2.06	2.80	3.75
10	1.81	2.23	3.17	4.59	25	1.71	2.06	2.79	3.73
11	1.80	2.20	3.11	4.44	30	1.70	2.04	2.75	3.65
12	1.78	2.18	3.06	4.32	40	1.68	2.02	2.70	3.55
13	1.77	2.16	3.01	4.22	60	1.67	2.00	2.66	3.46
14	1.76	2.15	2.98	4.14	120	1.66	1.98	2.62	3.37
15	1.75	2.13	2.95	4.07	00	1.65	1.96	2.58	3.29
D.f. = Degrees of freedom									
$\infty$ = infinity, same as the Standard Normal Distribution									

#### Two sample t method

Capillary density example:

Common variance = 53.31, SD = 7.30 capillaries/mm<sup>2</sup>, df = 19 + 23 - 2 = 40.

SE of difference = 2.26 capillaries/mm<sup>2</sup>.

Difference (control - ulcer) = 34.08 - 22.59 = 11.49

capillaries/mm<sup>2</sup>.

95% confidence interval for difference:

 $11.49 - ? \times 2.26$  to  $11.49 + ? \times 2.26$ .

? comes not from the Normal distribution but the t distribution with 40 degrees of freedom.

 $\begin{array}{l} 11.49 - 2.02 \times 2.26 \text{ to } 11.49 + 2.02 \times 2.26 \\ = 6.92 \text{ to } 16.07 \text{ capillaries/mm}^2. \end{array}$ 

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Difference (control – ulcer) = 34.08 – 22.59 = 11.49 capillaries/mm<sup>2</sup>.

Test of significance, null hypothesis that in the population the difference between means = 0:

(difference - 0)/SE = 11.49/2.26 = 5.08.

If the null hypothesis were true, this would be an observation from the t distribution with 40 degrees of freedom.

Two tailed probability points of the t Distribution									
D.f.	Probability			D.f.		Probability			
	0.10	0.05	0.01	0.001		0.10	0.05	0.01	0.001
	(10%)	(5%)	(1%)	(0.1%)		(10%)	(5%)	(1%)	(0.1%)
1	6.31	12.70	63.66	636.62	16	1.75	2.12	2.92	4.02
2	2.92	4.30	9.93	31.60	17	1.74	2.11	2.90	3.97
3	2.35	3.18	5.84	12.92	18	1.73	2.10	2.88	3.92
4	2.13	2.78	4.60	8.61	19	1.73	2.09	2.86	3.88
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#### Two sample t method

Capillary density example:

Test of significance, null hypothesis that in the population the difference between means = 0:

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If the null hypothesis were true, this would be an observation from the t distribution with 40 degrees of freedom.

From the table, the probability of such an extreme value is less than 0.001.

Using a computer program:  $\mathsf{P}$  = 0.0000, which we write  $\mathsf{P}{<}0.0001.$ 

# Assumptions of two sample t method

- > Observations are independent.
- Distribution of capillary density follows a Normal distribution in each population.
- $\succ$  Variances are the same in each population.

# Two sample t method

#### Assumptions of two sample t method

- Distribution of capillary density follows a Normal distribution in each population.
- > Variances are the same in each population.



# Two sample t method

# Effect of deviations from assumptions

Methods using the t distribution depend on some strong assumptions about the distributions from which the data come.

In general for two equal sized samples the t method is very resistant to deviations from Normality, though as the samples become less equal in size the approximation becomes less good.

The most likely effect of skewness is that P values are too large and confidence intervals too wide.