## Chapter 583

# Studentized Range Tests for Equivalence

## Introduction

This procedure computes power and sample size of equivalence tests of the means of two or more groups which are analyzed using a studentized range tests. It turns out that the question of equivalence among a set of means is often more meaningful than the question of equality.

Methodology for testing the equivalence of the means of two groups has received much attention. However, testing equivalence among three or more groups has not received much attention. The article by Shieh (2016) gives results for two competing test procedures: The F-test and the studentized range test. Results for the F-test are available in **PASS** in another procedure. This procedure provides power and sample size results the studentized range test.

While the F-test is by far the most commonly used method for testing the equality of two or more means, Shieh (2016) showed that when testing for mean equivalence, neither test is always optimal. In fact, the studentized range test is more powerful when the actual range is close to the equivalence boundary.

# **Technical Details for the Studentized Range Test**

Suppose G groups each have a normal distribution and with means  $\mu_1, \mu_2, ..., \mu_G$  and common variance  $\sigma^2$ . Let  $N_1, N_2, ..., N_G = N_i$  denote the common sample size of all groups and let N denote the total sample size. In this case of equal group sizes,  $N = GN_i$ . The multigroup equivalence problem requires one to show that the means are sufficiently close to each other. Shieh (2016) considering whether the difference between the minimum and maximum means (the range of the means) is sufficiently small so that the differences among the means can be regarded as of no practical importance.

## The One-Way Model

Consider the usual one-way fixed-effects model

$$Y_{gj} = \mu_g + \varepsilon_{gj}$$

where  $Y_{gj}$  is response,  $\mu_g$  are the treatment means, and  $\varepsilon_{gj}$  are the independent, normally distributed error with zero mean and common variance  $\sigma^2$ . Here the subscript g indexes the G groups, and the subscript j indexes the  $N_i$  subjects in each group.

Cohen (1988) showed that hypotheses about the G means may be obtained using either the variance of the means in terms of the F-test or their range in terms of the studentized range.

## **Equivalence Hypothesis**

The hypothesis of mean equivalence is

$$H_0: \frac{\delta}{\sigma} \ge \frac{\delta_0}{\sigma}$$
 versus  $H_1: \frac{\delta}{\sigma} < \frac{\delta_0}{\sigma}$ 

where  $\delta = \mu_{Max} - \mu_{Min}$  represents the range and  $\delta_0$  is the equivalence bound.

## **Studentized Range Statistic**

The studentized range statistic is defined as follows

$$Q = \frac{\left[\max_{g=1 \text{ to } G} (\bar{Y}_g) - \min_{g=1 \text{ to } G} (\bar{Y}_g)\right] \sqrt{N_i}}{S}$$

where  $\overline{Y}_q$  are the sample means, and S is the sample variance.

It turns out that the distribution of Q is a function of the pairwise mean differences  $\mu_g - \mu_h$ , not just the range (the maximum of these differences).

The cumulative distribution function, from which the power can be computed, is given by

$$\Theta(q) = P\{Q \le q\} = E_K \left\{ \sum_{g=1}^{G} E_{Z_g} \left[ \prod_{\substack{h=1 \\ h \ne g}}^{G} \left( \Phi\{Z_g + \delta_{gh} \sqrt{N_i}\} - \Phi\{Z_g + \delta_{gh} \sqrt{N_i} - q\sqrt{K/(N-G)}\} \right) \right] \right\}$$

where  $\delta_{gh}=\mu_g-\mu_h$ , K is a chi-square random variable with N-G degrees of freedom,  $\Phi\{z\}$  is the CDF of a standard normal distribution,  $Z_g$  are independent standard normal random variables,  $E_K\{x\}$  is the expectation with respect to K, and  $E_{Z_g}\{x\}$  is the expectation with respect to  $Z_g$ .

Note that the critical value is based on the set of group means. It cannot be determined from just  $\delta_0$ . When only  $\delta_0$  is specified, the least favorable configuration (LFC) of the means is used. This is given by

$$\{\mu_1, \dots, \mu_G\} = \left\{ -\frac{\delta_0}{2}, \frac{\delta_0}{2}, 0, \dots, 0 \right\}$$

If a sample size is desired, it can be determined using a standard binary search algorithm.

## **Example 1 – Finding Power**

An experiment is being designed to assess the equivalence of the means of four groups using a studentized range test with a significance level of 0.05. Previous studies have shown a standard deviation of 2. The maximum range of the four means allowed in equivalent means is 2. Power calculations assume that the actual range is 1. To better understand the relationship between power and sample size, the researcher wants to compute the power for several group sample sizes between 20 and 100. The sample sizes will be equal across all groups.

## Setup

If the procedure window is not already open, use the PASS Home window to open it. The parameters for this example are listed below and are stored in the **Example 1** settings file. To load these settings to the procedure window, click **Open Example Settings File** in the Help Center or File menu.

Solve For	Power
Alpha	0.05
G (Number of Groups)	4
Ni (Sample Size Per Group)	20 40 60 80 100
μ1i's Input Type	Enter Range of Means H1
δ1 (Range of Means H1)	1
μ0i's Input Type	Enter Range of Means H0
δ0 (Range of Means H0 or Equiv Limit)	2
σ (Standard Deviation)	2

## **Output**

Click the Calculate button to perform the calculations and generate the following output.

#### **Numeric Reports**

#### **Numeric Results**

Solve For: Power Number of Groups: 4

	Samp	ole Size	Range of Means		Ctondord	
Power	Total N	Group Ni	H1 δ1	H0 (Equivalence) δ0	Standard Deviation σ	Alpha
0.3921	80	20	1	2	2	0.05
0.6847	160	40	1	2	2	0.05
0.8463	240	60	1	2	2	0.05
0.9270	320	80	1	2	2	0.05
0.9659	400	100	1	2	2	0.05

Power The probability of rejecting a false null hypothesis of non-equivalence in favor of the alternative hypothesis of equivalence.

N The total number of subjects in the study.

Ni The number of items sampled per group.

δ1 The range of the group means assumed by the alternative hypothesis. It is the value at which the power is computed. Note that you must have δ1 < δ0.

δ0 The range of the group means assumed by the null hypothesis. This value is the equivalence limit (bound). Note that you must have  $\delta 1 < \delta 0$ .

σ The standard deviation of the responses for all groups.

Alpha The significance level of the test: the probability of rejecting the null hypothesis of non-equivalent means when it is actually true.

#### **Summary Statements**

A one-way design with 4 groups will be used to test whether the 4 means are equivalent, using an equivalence bound (for the range) of 2. The comparison will be made using a studentized range equivalence test with a Type I error rate ( $\alpha$ ) of 0.05. The common within-group standard deviation of responses for all groups is assumed to be 2. To detect a range of group means of 1, with group sample sizes of 20 subjects per group (for a total of 80 subjects), the power is 0.3921.

#### **Dropout-Inflated Sample Size**

Dropout Rate	Sample Size N	Dropout- Inflated Enrollment Sample Size N'	Expected Number of Dropouts D
20%	80	100	20
20%	160	200	40
20%	240	300	60
20%	320	400	80
20%	400	500	100

Dropout Rate The percentage of subjects (or items) that are expected to be lost at random during the course of the study and for whom no response data will be collected (i.e., will be treated as "missing"). Abbreviated as DR. Ν The evaluable sample size at which power is computed (as entered by the user). If N subjects are evaluated out of the N' subjects that are enrolled in the study, the design will achieve the stated power. N' The total number of subjects that should be enrolled in the study in order to obtain N evaluable subjects, based on the assumed dropout rate. N' is calculated by inflating N using the formula N' = N / (1 - DR), with N' always rounded up. (See Julious, S.A. (2010) pages 52-53, or Chow, S.C., Shao, J., Wang, H., and Lokhnygina, Y. (2018) pages 32-33.)

D The expected number of dropouts. D = N' - N.

#### **Dropout Summary Statements**

Anticipating a 20% dropout rate, 100 subjects should be enrolled to obtain a final sample size of 80 subjects.

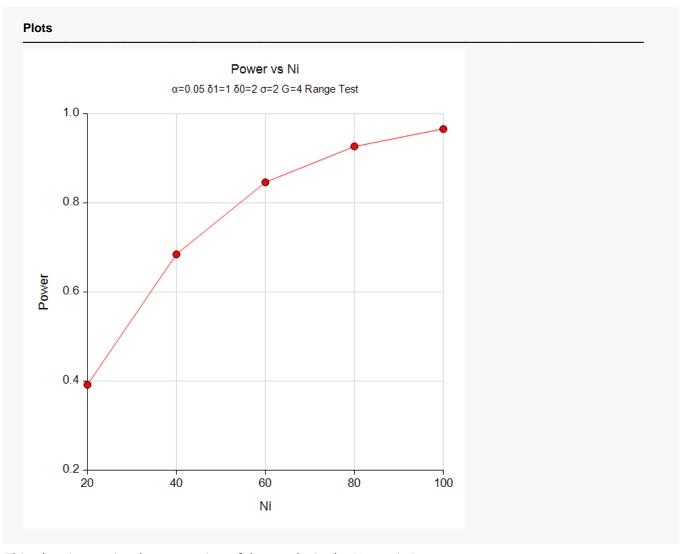
#### References

Shieh, G. 2016. 'A comparative appraisal of two equivalence tests for multiple standardized effects'. Computer Methods and Programs in Biomedicine, Vol 126, Pages 110-117. http://dx.doi.org/10.1016/j.cmpb.2015.12.004 Wellek, Stefan. 2010. Testing Statistical Hypotheses of Equivalence and Noninferiority, 2nd Edition. CRC Press.

Cohen, Jacob. 1988. Statistical Power Analysis for the Behavioral Sciences. Lawrence Erlbaum Associates. Hillsdale, New Jersey.

This report shows the numeric results of this power study.

### **Plots Section**



This plot gives a visual presentation of the results in the Numeric Report.

# Example 2 - Finding the Sample Size Necessary to Reject

Continuing with the last example, we will determine how large the sample size would need to have been for alpha = 0.05 and power = 0.80 or 0.9.

## Setup

If the procedure window is not already open, use the PASS Home window to open it. The parameters for this example are listed below and are stored in the **Example 2** settings file. To load these settings to the procedure window, click **Open Example Settings File** in the Help Center or File menu.

Solve For	Sample Size
Power	0.8 0.9
Alpha	0.05
G (Number of Groups)	4
μ1i's Input Type	Enter Range of Means H1
δ1 (Range of Means H1)	1
μ0i's Input Type	Enter Range of Means H0
δ0 (Range of Means H0 or Equiv Limit)	2
σ (Standard Deviation)	2

## **Output**

Click the Calculate button to perform the calculations and generate the following output.

Solve Fo Number	r: of Groups:	Sample S 4	ize			
	Sample Size			Range of Means	01	
Power	Total N	Group Ni	H1 δ1	H0 (Equivalence) δ0	Standard Deviation σ	Alpha
0.8014	212	53	1	2	2	0.05
0.9013	288	72	1	2	2	0.05

This report shows the necessary sample sizes for achieving powers of 0.8 and 0.9.

# Example 3 - Validation using Shieh (2016)

Shieh (2016) page 115 presents an example in which alpha = 0.05, G = 4,  $\sigma = 7.47583$ ,  $\delta 1 = 0.1821(7.47583) = 1.36135$ ,  $\delta 0 = 7.47583$ , and power = 0.8. The resulting sample size is 27 per group for a total of 108.

## Setup

If the procedure window is not already open, use the PASS Home window to open it. The parameters for this example are listed below and are stored in the **Example 3** settings file. To load these settings to the procedure window, click **Open Example Settings File** in the Help Center or File menu.

Solve For	Sample Size
Power	0.8
Alpha	0.05
G (Number of Groups)	4
μ1i's Input Type	Enter Range of Means H1
δ1 (Range of Means H1)	1.36135
μ0i's Input Type	Enter Range of Means H0
δ0 (Range of Means H0 or Equiv Limit)	7.47583
σ (Standard Deviation)	7.47583

## Output

Click the Calculate button to perform the calculations and generate the following output.

Solve For: Number of Groups:			Sample Size 4					
	Sample Size		R	ange of Means	01			
Power	Total N	Group Ni	H1 δ1	H0 (Equivalence) δ0	Standard Deviation σ	Alpha		
0.811	108	27	1.361	7.476	7.476	0.05		

**PASS** also found Ni = 27 and N = 108.

# Example 4 – Comparing the Impact of Mean Configuration on Power and Sample Size

This example will look at the impact the configuration of the means assumed by the null hypothesis on the required sample size. The three configurations suggested by Cohen (1988) will be studied.

Let alpha = 0.05, G = 5,  $\sigma$  = 1,  $\delta$ 1 = 0.4, and power = 0.8. The following three configurations will be entered in the spreadsheet and used to specify  $\delta$ 0. Note that all three configurations result in  $\delta$ 0 = 0.8.

Column	Means H0	Comments
C1	-0.4, -0.4, 0.4, 0.4, 0.4	Minimum mean variation
C2	-0.4, -0.2, 0, 0.2, 0.4	Intermediate mean variation
C3	-0.4, 0, 0, 0, 0.4	Maximum mean variation

## Setup

If the procedure window is not already open, use the PASS Home window to open it. The parameters for this example are listed below and are stored in the **Example 4** settings file. To load these settings to the procedure window, click **Open Example Settings File** in the Help Center or File menu.

Solve For	Sample Size
Power	0.8
Alpha	0.05
G (Number of Groups)	5
μ1i's Input Type	Enter Range of Means H1
δ1 (Range of Means H1)	0.4
μ0i's Input Type	Enter Columns Containing Sets of µ0i's
Columns Containing Sets of µ0i's	1-3
σ (Standard Deviation)	1

#### **Input Spreadsheet Data**

Row	C1	C2	C3
1	-0.4	-0.4	-0.4
2	-0.4	-0.2	0.0
3	0.4	0.0	0.0
4	0.4	0.2	0.0
5	0.4	0.4	0.4

## **Output**

Click the Calculate button to perform the calculations and generate the following output.

#### **Numeric Results**

Solve For: Sample Size

Number of Groups: 5

			(	Group Mean	ıs		
	Sample Size			H0 (Equivalenc		Standard	
Power	Total N	Group Ni	Range δ1	Means μ0i	Range δ0	Deviation σ	Alpha
0.8106	145	29	0.4	C1(1)	0.8	1	0.05
0.8032	350	70	0.4	C2(2)	0.8	1	0.05
0.8028	415	83	0.4	C3(3)	8.0	1	0.05

Item	Values
C1(1) C2(2) C3(3)	-0.4, -0.4, 0.4, 0.4, 0.4 -0.4, -0.2, 0, 0.2, 0.4 -0.4, 0, 0, 0, 0.4

These results show that the sample size required varies from 145 to 415 depending on the configuration.