## USING STATISTICS IN RESEARCH

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## SECTION 5.

## HYPOTHESIS TESTING EXTENSIONS

In this section, we examine extensions to the hypothesis testing methods described above: although the methods differ slightly, the general approach remains the unchanged. We proceed using the same strategy, outlined below, For a data sample  $x_1, ..., x_n$ , with corresponding random variables  $X_1, ..., X_n$ , we

- 1. consider a pair of competing hypotheses,  $H_0$  and  $H_1$
- 2. define a suitable test statistic  $T = T(X_1, ..., X_n)$  (that is, some function of the original random variables; this will define the **test statis**-**tic**), and a related **pivotal random variable**  $T_{PQ} = T_{PQ}(X)$
- 3. assume that  $H_0$  is true, and compute the sampling distribution of T,  $f_T$  or  $F_T$ ; this is the **null distribution**

- 4. compute the **observed** value of T,  $t = T(x_1, ..., x_n)$ ; this is the **test** statistic
- 5. assess whether t is a surprising observation from the distribution  $f_T$ . If it is surprising, we have evidence to reject  $H_0$ ; if it is not surprising, we cannot reject  $H_0$

This strategy can be applied to more complicated normal examples, and also non-normal and non-parametric testing situations. It is a general strategy for assessing the statistical evidence for or against a hypothesis.

## 5.1 ANALYSIS OF VARIANCE

The first extension we consider still presumes a normality assumption for the data, but extends the ideas from Z and T tests, which compare at most two samples, to allow for the analysis of any number of samples. **Analysis of variance** or **ANOVA** is used to display the sources of variability in a collection of data groups. The ANOVA F-test compares variability **between** groups with the variability **within** groups.

### 5.1.1 ONE-WAY ANOVA

The T-test can be extended to allow a test for differences between more than two data samples. Suppose there are K groups of sizes  $n_1, ..., n_K$  (let  $n = n_1 + ... + n_K$ ) from different populations. The model can be represented as follows: let  $y_{kj}$  be the jth observation in the kth group,

then

$$y_{kj} = \mu_k + \varepsilon_{kj}$$

for k = 1, ..., K, and  $\varepsilon_{kj} \sim N(0, \sigma^2)$ . This model assumes that

$$Y_{kj} \sim N\left(\mu_k, \sigma^2\right)$$

and that the expectations for the different groups are different. We can view the data as a table comprising K columns, with each column corresponding to a sample. The groups are commonly referred to as **FAC-TORS**. To test the hypothesis that each population has the same mean, that is, the hypotheses

 $H_0$  :  $\mu_1 = \mu_2 = \dots = \mu_K$ 

 $H_1 : not H_0$ 

an **Analysis of Variance (ANOVA)** F-test may be carried out. To carry out a test of the hypothesis, the following **ANOVA table** should be completed;

Source	D.F.	Sum of squares	Mean square	F
Between Samples	K-1	FSS	FSS/(K-1)	$\frac{FSS/(K-1)}{RSS/(n-K)}$
Within Samples	n-K	RSS	RSS/(n-K)	
Total	n-1	TSS		

where, and

$$TSS = \sum_{k=1}^{K} \sum_{j=1}^{n_k} (y_{kj} - \overline{y}_{..})^2 \quad RSS = \sum_{k=1}^{K} \sum_{j=1}^{n_k} (y_{kj} - \overline{y}_k)^2 \quad FSS = \sum_{k=1}^{K} n_k (\overline{y}_k - \overline{y}_k)^2$$

where TSS is the **total** sum-of-squares (i.e. total deviation from the overall data mean  $\overline{y}$ .) RSS is the **residual** sum-of-squares (i.e. sum of deviations from individual group means  $\overline{y}_k$ , k = 1, ..., K) and FSS is the **fitted** sum-of-squares (i.e. weighted sum of deviations of group means from the overall

data mean, with weights equal to number of data points in the individual samples) Note:that

$$TSS = FSS + RSS$$

The definitions of these three sums of squares quantities gives insight into how ANOVA works by decomposing the total variation in the observed data

- $\bullet$  TSS is the **overall** variation
- FSS is the variation caused by the **systematic** component (that is, the differences in group means)
- $\bullet$  RSS is the **random** variation

If the F statistic is calculated in this way, and compared with an F distribution with parameters K-1, n-K, the hypothesis that all the individual samples have the same mean can be tested.

**EXAMPLE** Three genomic segments were used to studied in order to discover whether the distances (in kB) between successive occurrences of a particular motif were substantially different. Several measurements were taken using for each segment;

	Method				
	SEGMENT A	SEGMENT B	SEGMENT C		
	42.7	44.9	41.9		
	45.6	48.3	44.2		
	43.1	46.2	40.5		
	41.6		43.7		
			41.0		
Mean	43.25	46.47	42.26		
Variance	2.86	2.94	2.66		

For these data, the ANOVA table is as follows;

Source	D.F.	Sum of squares	Mean square	$\overline{F}$
SEGMENTS	2	34.1005	17.0503	6.11
Residual	9	25.1087	2.7899	
Total	11	59.2092		

and the F statistic must be compared with an  $F_{2,9}$  distribution. For a significance test at the 0.05 level, F must be compared with the 95th percentile (in a **one-sided** test) of the  $F_{2,9}$  distribution. This value is 4.26. Therefore, the F statistic **is** surprising, given the hypothesized model, and therefore there is evidence to reject the hypothesis that the segments are identical.

## 5.1.2 TWO-WAY ANOVA

One-way ANOVA can be used to test whether the underlying means of several groups of observations are equal. Now consider the following data collection situation. Suppose there are K treatments, and L groups of observations that are believed to have different responses, that all treatments are administered to all groups, and measurement samples of size n are made for each of the  $K \times L$  combinations of treatments  $\times$  groups. The experiment can be represented as follows: let  $y_{klj}$  be the jth observation in the kth treatment on the lth group, then

$$y_{klj} = \mu_k + \delta_l + \varepsilon_{klj}$$

for k = 1, ..., K, l = 1, ..., L, and again  $\varepsilon_{klj} \sim N(0, \sigma^2)$ .

This model assumes that  $Y_{kj} \sim N\left(\mu_k + \delta_l, \sigma^2\right)$  and that the expectations for the different samples are different. We can view the data as a 3 dimensional-table comprising K columns and L rows, with n observations for each column  $\times$  row combination, corresponding to a sample.

It is possible to test the hypothesis that each **treatment**, and/or that each **group** has the same mean, that is, the two null hypotheses

$$H_0$$
 :  $\mu_1 = \mu_2 = \dots = \mu_K$ 

$$H_0$$
 :  $\delta_1 = \delta_2 = \dots = \delta_K$ 

against the alternative  $H_1$ :not  $H_0$  in each case. For these tests, a **Two-way Analysis of Variance (ANOVA)** F-test may be carried out. The

Two-Way ANOVA table is computed as follows

Source	D.F.	Sum of squares	Mean square	F
TREATMENTS	K-1	$FSS_1$	$FSS_1/(K-1)$	$\frac{FSS_1/(K-1)}{RSS/(R+1)}$
GROUPS	L-1	$FSS_2$	$FSS_2/(L-1)$	$\frac{FSS_2/(L-1)}{RSS/(R+1)}$
Residual	R+1	RSS	RSS/(R+1)	
Total	N-1	TSS		

where  $N = K \times L \times n$ , R = N - L - K. and again

$$TSS = FSS_1 + FSS_2 + RSS.$$

In the table below, there are K=6 Treatments, and L=3 Groups, and n=1

	I	II	III	GROUP totals
1	0.96	0.94	0.98	2.88
2	0.96	0.98	1.01	2.95
3	0.85	0.87	0.86	2.58
4	0.86	0.84	0.90	2.60
5	0.86	0.87	0.89	2.62
6	0.89	0.93	0.92	2.74
TREATMENT totals	5.38	5.43	5.56	16.37

There are two natural hypotheses to test; first, do the TREATMENTS differ, and second, do the GROUPS differ?

Two-way analysis of variance can be used to analyze such data. Given two sources of variation the data can be thought of as a table with the rows and columns representing these two sources . Two-way analysis of variance studies the variability due to the GROUPS effect (here, variability in the

specimens) and the variability due to the "column" effect (variability in the TREATMENTS), and calibrates them against the average level of variability in the data overall. Having performed the appropriate calculations, the results are displayed in an ANOVA table. For example, for the data above

Source	D.F.	Sum of squares	Mean square	$\overline{F}$
TREATMENT	5	0.040828	0.0081656	31.54
GROUP	2	0.002878	0.001439	5.57
Residual	10	0.002589	0.0002589	
Total	17	0.046295		

The two F statistics can be interpreted as follows; the first (F = 31.54) is the test statistic for the test of equal means in the **rows**, that is, that there is no difference between **TREATMENTS**. This statistic must be compared with an  $F_{5,10}$  distribution (the two degrees of freedom being the

entries in the degrees of freedom column in the specimens and residual rows of the ANOVA table). The 95th percentile of the  $F_{5,10}$  distribution is 3.33, and thus the test statistic is **more extreme** than this critical value, and thus the hypothesis that each specimen has the same mean can be **rejected**.

The second F statistic, (F = 5.57), is the test statistic for the test of equal means in the **columns**, that is, that there is no difference between **GROUPS**. This statistic must be compared with an  $F_{2,10}$  distribution (the two degrees of freedom being the entries in the degrees of freedom column in the methods and residual rows of the ANOVA table). The 95th percentile of the  $F_{2,10}$  distribution is 4.10, and thus the test statistic is **more extreme** than this critical value, and thus the hypothesis that each method has the same mean can be **rejected**.

#### 5.1.3 ANOVA: KEY ASSUMPTIONS

In ANOVA, there are three key assumptions

- (i) all data are **independent**
- (ii) the data are **normally** distributed
- (iii) the data subgroups (defined by the cross classification by factors) have **equal variances**.

Of these three points, (i) can be assessed by consideration of the study design, (ii) can be tested formally using methods that will be described in later sections, and (iii) can be tested using statistical hypothesis testing in the following way using **Levene's Test** 

#### LEVENE'S TEST FOR HOMOGENEITY OF VARIANCE

The Levene test is defined for the two hypotheses as follows: suppose that the data Y of size n is partitioned into K subgroups of sizes  $n_1, ..., n_K$  where  $n = n_1 + ... + n_K$ . It is of interest to test whether the subgroups have the same variance, that is the hypothesis

$$H_0$$
:  $\sigma_1 = \sigma_2 = \dots = \sigma_K$ 

 $H_1$ :  $\sigma_i \neq \sigma_j$  for at least one pair (i,j).

Test Statistic

$$W = \frac{(n-K)\sum_{i=1}^{K} n_i (\overline{Z}_i - \overline{Z})^2}{(K-1)\sum_{i=1}^{K} \sum_{j=1}^{n_k} n_i (Z_{ij} - \overline{Z}_i)^2}$$

where  $Z_{ij}$  can have one of the following three definitions:

- 1.  $Z_{ij} = |Y_{ij} \overline{Y_i}|$ , where  $\overline{Y_i}$  is the mean of the *i*th subgroup.
- 2.  $Z_{ij} = |Y_{ij} Y_i^{(MEDIAN)}|$  where  $Y_i^{(MEDIAN)}$  is the median of the *i*th subgroup.
- 3.  $Z_{ij} = |Y_{ij} Y_i^{(TRIMMED)}|$  where  $Y_i^{(TRIMMED)}$  is the 10% trimmed mean of the *i*th subgroup.

The three choices for defining  $Z_{ij}$  determine the robustness (ability of the test to not falsely detect unequal variances when the underlying data are not normally distributed) and power (ability of the test to detect unequal variances when the variances are in fact unequal) of Levene's test. The Levene test rejects the hypothesis that the variances are equal at significance level  $\alpha$  (typically,  $\alpha = 0.05$ ) if

$$W > F_{K-1,n-K}(1-\alpha)$$

where  $F_{K-1,n-K}(1-\alpha)$  is the  $(1-\alpha)$ % quantile of the Fisher F distribution with K-1 and n-K degrees of freedom.

# 5.2 NON-NORMAL DATA: COUNTS AND PROPORTIONS

The one and two sample tests described in earlier sections can also be applied to non-normal data. A common form of non-normal data arise when the counts of numbers of "successes" or "failures" that arise in a fixed number of trials. In this case, the Binomial distribution model is appropriate; in a one sample testing, we model the number of successes, X, by assuming  $X \sim Binomial(n, \theta)$  and test hypotheses about  $\theta$ , whereas in the two sample case, we assume that the number of successes in the two samples are random variables  $X_1$  and  $X_2$ , where  $X_1 \sim Binomial(n_1, \theta_1)$  and  $X_2 \sim Binomial(n_2, \theta_2)$ , and perhaps test the null hypothesis

$$H_0: \theta_1 = \theta_2$$

against some alternative hypothesis  $(\theta_1 \neq \theta_2, \theta_1 > \theta_2 \text{ or } \theta_1 < \theta_2)$ 

#### 5.2.1 ONE-SAMPLE TESTING

In the one sample case, two alternative approaches can be adopted. The first is a so-called **exact** test, where the distribution of the chosen test statistic under  $H_0: \theta = c$  is computed exactly, giving exact critical values and p-values. The second is a approximate test based on a Normal approximation to the binomial distribution.

For the exact test, we note that, **if**  $H_0$  **is true**, and  $\theta = c$ , then  $X \sim Binomial(n,c)$  so the critical values in a two-sided test can be computed directly by inspection of the Binomial(n,c) c.d.f; that is

$$F_{BIN}(C_{R_1}; n, \theta = c) = 0.025$$
  $C_{R_2} = F_{BIN}(0.975; n, \theta = c)$ 

where  $F_{BIN}(-; n, \theta)$  is the c.d.f. of the  $Binomial(n, \theta)$  distribution

$$F_{BIN}(x; n, \theta) = \sum_{i=0}^{\lfloor x \rfloor} \binom{n}{i} \theta^{i} (1 - \theta)^{n-i}$$

and  $\lfloor x \rfloor$  is the smallest whole number not greater than x. For the approximate test, we use the fact that

$$X \sim Binomial(n, \theta) \approx Normal(n\theta, n\theta(1 - \theta))$$

and hence random variable Z

$$Z = \frac{X - n\theta}{\sqrt{n\theta(1 - \theta)}}$$

is approximately distributed as Normal(0,1). For the approximate test of  $H_0: \theta = c$ , we therefore use the test statistic

$$z = \frac{x - nc}{\sqrt{nc(1 - c)}}$$

(x is the actual, observed count) and compare this with the standard normal c.d.f.. This test is virtually equivalent to the one-sample t-test.

#### 5.2.2 TWO SAMPLE TESTING

For a two sample test of  $H_0: \theta_1 = \theta_2$ , we use a similar normal approximation to the one-sample case. If  $H_0$  is true, then there is a common probability  $\theta$  determining the success frequency in both samples, and the maximum likelihood estimate of  $\theta$  is

$$\widehat{\theta} = \frac{x_1 + x_2}{n_1 + n_2} = \frac{x}{n}, \text{ say}$$

and thus it can be shown that the test statistic.

$$z = \frac{\frac{x_1}{n_1} - \frac{x_2}{n_2}}{\sqrt{\frac{(n_1 + n_2)}{n_1 n_2} \left(\frac{x_1 + x_2}{n_1 + n_2}\right) \left(1 - \frac{x_1 + x_2}{n_1 + n_2}\right)}}$$

has an approximate standard Normal distribution.

## 5.3 CONTINGENCY TABLES

Contingency tables are constructed when a sample of data of size n are classified according to D factors, with each factor having  $k_d$  levels or categories, for d = 1, ..., D. When the classification is complete, the result can be represented by a D-way table of  $k_1 \times k_2 \times ... \times k_D$  "cells", with each cell containing a fraction of the original data. For example, if D = 2, the table consists of  $k_1$  rows and  $k_2$  columns, and the number data in cell (i, j) is denoted  $n_{ij}$  for  $i = 1, ..., k_1$  and  $j = 1, ..., k_2$ , where

$$\sum_{i=1}^{k_1} \sum_{j=1}^{k_2} n_{ij} = n$$

Such a table when D=2,  $k_1=4$  and  $k_2=6$  is illustrated below

			COLUMN					
		1	2	3	4	5	6	Total
	1	$n_{11}$	$n_{12}$	$n_{13}$	$n_{14}$	$n_{15}$	$n_{16}$	$n_{1.}$
	2	$n_{21}$	$n_{22}$	$n_{13} \\ n_{23} \\ n_{33}$	$n_{14} \\ n_{24}$	$n_{25}$	$n_{26}$	$n_{2.}$
ROW	3	$n_{31}$	$n_{32}$	$n_{33}$	$n_{34}$	$n_{35}$	$n_{36}$	$n_{3.}$
	4	$n_{41}$	$n_{42}$	$n_{43}$	$n_{44}$	$n_{45}$	$n_{46}$	$n_4$ .
	Total	$n_{.1}$	$\overline{n}_{.2}$	$n_{.3}$	$\overline{n}_{.4}$	$n_{.5}$	$n_{.6}$	n

## 5.3.1 CHI-SQUARED GOODNESS-OF-FIT TEST

It is often of interest to test whether row classification is **independent** of column classification, as this would indicate **independence** between row and column factors. An approximate test can be carried out using a **Chi-Squared Goodness-of-Fit** statistic; if the independence model is correct, the expected cell frequencies  $\hat{n}_{ij}$  can be calculated as

$$\hat{n}_{ij} = \frac{n_i \cdot n_{.j}}{n}$$
  $i = 1, ..., k_1, \ j = 1, ..., k_2$ 

where  $n_i$  is the *total* of cell counts in row i and  $n_{.j}$  is the *total* of cell counts in column j, and that, under independence, the  $\chi^2$  test statistic

$$\chi^2 = \sum_{i=1}^{k_1} \sum_{j=1}^{k_2} \frac{(n_{ij} - \hat{n}_{ij})^2}{\hat{n}_{ij}}$$

has an approximate chi-squared distribution with  $(k_1 - 1)(k_2 - 1)$  degrees of freedom.

## 5.3.2 LIKELIHOOD RATIO TEST

Another approximate test is based on a Likelihood Ratio (LR) statistic

$$LR = 2\sum_{i=1}^{k_1} \sum_{j=1}^{k_2} n_{ij} \log \frac{n_{ij}}{\hat{n}_{ij}}$$

This statistic also has an approximate Chi-squared distribution

$$\chi^2_{(k_1-1)(k_2-1)}$$

again given that  $H_0$  is true.

We will see further analysis of count data in section (5.3) below.

## $5.3.3 \quad 2 \times 2 \text{ TABLES}$

When  $k_1 = k_2 = 2$ , the contingency table reduces to a two-way binary classification

		COLUMN			
		1	2	Total	
	1	$n_{11}$	$n_{12}$	$n_{1.}$	
ROW	2	$n_{21}$	$n_{22}$	$n_{2.}$	
	Total	$n_{.1}$	$n_{.2}$	$\overline{n}$	

In this case we can obtain some more explicit tests: one is again an **exact test**, the other is based on a normal approximation. The chi-squared test described above is feasible, but other tests may also be constructed:

#### • FISHER'S EXACT TEST FOR INDEPENDENCE

Suppose we wish to test for **independence** between the row and column variables of a contingency table. When the data consist of two categorical variables, a contingency table can be constructed reflecting the number of occurrences of each factor combination. **Fisher's exact test** assesses whether the classification according to one factor is independent of the classification according to the other, that is the test is of the null hypothesis  $H_0$  that the factors are independent, against the general alternative, **under the assumption that the row and column totals are fixed**.

The data for such a table comprises the row and column totals  $(n_1, n_2, n_{.1}, n_{$ 

is

$$p(n_{11}) = \frac{\binom{n_{1.}}{n_{11}}\binom{n_{2.}}{n_{21}}}{\binom{n}{n_{.1}}} = \frac{n_{1.}!n_{.1}!n_{2.}!n_{.2}!}{n!n_{11}!n_{12}!n_{21}!n_{22}!}$$

where  $n! = 1 \times 2 \times 3 \times ... \times (n-1) \times n$ .

For the p-value, we need to assess the whether or not the observed table is surprising under this null distribution; suppose we observe  $n_{11} = x$ , then we can compare p(x) with all p(y) for all feasible y, that is y in the range  $\max\{0, n_1 - (n - n_1)\} \le y \le \min\{n, n_1\}$ . We are thus calculating the null distribution **exactly** given the null distribution assumptions and the row and column totals; if the observed test statistic lies in the tail of the distribution, we can reject the null hypothesis of independent factors.

#### • MANTEL-HAENSZEL TEST FOR INDEPENDENCE

This test allows you to test for independence between two factors in the presence of a third, and possibly related variable. It extends the two-way Chi-squared test of independence described above; the test statistic is a chi-squared type statistic, and the null distribution under independence is a Chi-squared distribution.

## • McNEMAR'S TEST FOR SYMMETRY IN PAIRED SAM-PLES

In a  $2 \times 2$  table representing paired data (where observations are, for example, matched in terms of medical history or genotype, or phenotype) the usual chi-squared test is not appropriate, and **McNemar's** test can instead be used. Consider the following table for a total of n matched pairs of observations, in which each individual in the pair has been classified (or randomized to class) A or B, with one A one B in each pair, and then the outcome (disease status, survival status)

recorded.

		ig  A		
		YES	NO	Total
	YES	$n_{11}$	$n_{12}$	$n_{1.}$
В	NO	$n_{21}$	$n_{22}$	$n_{2.}$
	Total	$n_{.1}$	$n_{.2}$	n

that is,  $n_{11}$  pairs were observed for which both A and B classified individuals had disease/survival status YES, whereas  $n_{12}$  pairs were observed for which the A individual had status NO, but the B individual had status YES, and so on.

An appropriate test statistic here for a test of symmetry or "discordancy" in these results (that is, whether the two classifications are significantly different in terms of outcome) is

$$\chi^2 = \frac{(n_{12} - n_{21})^2}{n_{12} + n_{21}}$$

which effectively measures how different the off-diagonal entries in the table are. This statistic is an adjusted Chi-squared statistic, and has a  $\chi_1^2$  distribution under the null hypothesis that there is no asymmetry. Again a one-tailed test is carried out: "surprising" values of the test statistic are large.

## 5.4 NON-PARAMETRIC TESTS

The standard test for the equality of expectations of two samples is the two-sample T-test. This test is predicated on the assumption of normality of the underlying distributions. In many cases, such an assumption is inappropriate, possible due to distributional asymmetry or the presence of outliers, and thus other tests of the hypothesis of equality of population locations must be developed.

Some of the standard non-parametric tests used in statistical analysis are described below: we concentrate on two-sample tests for the most part. All of these tests can be found in good statistics packages.

#### 5.4.1 THE MANN-WHITNEY-WILCOXON TEST

Consider two samples  $x_1, ..., x_{n_1}$  and  $y_1, ..., y_{n_2}$ . The **Mann-Whitney-Wilcoxon** test proceeds as follows; first, sort the pooled sample into ascending order. Add up the ranks of the data from sample one to get  $u_1$  say. Repeat for sample two to get  $u_2$ . Note that

$$u_1 + u_2 = \frac{(n_1 + n_2)(n_1 + n_2 + 1)}{2}$$

The Mann-Whitney-Wilcoxon statistic is  $u_1$ . It can be shown that, under the hypothesis that the data are from populations with the equal medians, then  $u_1$  has an approximate normal distribution with mean and variance

$$\frac{n_1(n_1+n_2+1)}{2} \qquad \frac{n_1n_2(n_1+n_2+1)}{12}$$

This is the non-parametric alternative to the two sample t-test.

## 5.4.2 THE KOLMOGOROV-SMIRNOV TEST

The two-sample Kolmogorov-Smirnov test is a non-parametric test for comparing two samples via their **empirical cumulative distribution function**. For data  $x_1, ..., x_n$ , the empirical c.d.f. is the function  $\widehat{F}$  defined at x by

$$\widehat{F}(x) = \frac{c(x)}{n}$$
  $c(x) = \text{"Number of data } \le x$ "

Thus, for two samples, we have two empirical c.d.f., and the (two-sided) **Kolmogorov-Smirnov** test that the two samples come from the same underlying distribution is based on the statistic

$$T = \max_{x} \left| \widehat{F}_{1}(x) - \widehat{F}_{2}(x) \right|.$$

It is easy to show that  $0 \le T \le 1$ , but the null distribution of T is not available in closed form. Fortunately, the p-value probability in the test for test statistic t, p = P[T > t] can be obtained for various different sample sizes using statistical tables or packages.

## 5.4.3 THE CHI-SQUARED GOODNESS-OF-FIT TEST

It is often required to test whether a sample can be well modelled using a particular distribution; the **chi-squared goodness-of-fit test** is the most commonly used test. It is a non-parametric test for which the null distribution can be well approximated by a Chi-squared distribution. It is studied in more detail in the *Model Validation* section below.

## 5.4.4 THE SHAPIRO-WILK NORMALITY TEST

It is often required to test whether a sample of data are normally distributed; if they are, then many of the tests described above can be utilized. The **Shapiro-Wilk** test can be used to test this hypothesis; the test statistic is commonly denoted W, and critical and p- values from its null distribution are available from tables or statistics packages. Not that also the **Kolmogorov-Smirnov** test can be used in a one-sample problem to test any distributional assumptions, including normality.

### 5.4.5 THE KRUSKAL-WALLIS TEST

The Kruskal-Wallis rank test is a nonparametric alternative to a one-way analysis of variance. The null hypothesis is that the true location parameter is the same in each of the samples. The alternative hypothesis is that at least one of the samples has a different location. Unlike one-way ANOVA, this test does not require normality

### 5.4.6 THE FRIEDMAN RANK SUM TEST

The **Friedman rank sum test** is a nonparametric alternative to a two-way analysis of variance. It is appropriate for data arising from an experiment in which exactly one observation was collected from each experimental unit, or group, under each treatment. The elements of the samples are assumed to consist of a treatment effect, plus a group effect, plus independent and identically distributed residual errors

## 5.5 EXACT TESTS

We have seen Chi-squared tests being used to test hypotheses about data. These tests involved the construction of a chi-squared statistic, of the form

$$\chi^2 = \sum_{i=1}^{k} \frac{(O_i - E_i)^2}{E_i}$$

where  $O_i$  and  $E_i$  are the observed and expected/predicted counts in each of k (cross) classifications or categories.

The distribution of the test-statistic is typically approximated by a Chisquared distribution with an appropriately chosen degrees of freedom. This approximation is good when the sample size is large, but not good when the table is "sparse", with some low (expected) cell entries (under the null hypothesis). The approximation breaks down for small sample sizes due to the inappropriateness of the Normal approximation referred to in section (5.2) We have also seen two examples of **Exact Tests:** the exact binomial test in section (5.2) and Fisher's Exact Test in section (5.3.3). For these tests, we proceeded as follows, mimicking the general hypothesis strategy outlined at the start of the section.

- 1. Write down a null hypothesis  $H_0$  and a suitable alternative hypothesis  $H_1$
- 2. Construct a test statistic T deemed appropriate for the hypothesis under study
- 3. Compute the null distribution of T, that is the sampling distribution of T if  $H_0$  is true,  $f_T$
- 4. Compare the observed value of T, t = T(x) for sample data  $x = (x_1, ..., x_n)$  with the null distribution and assess whether the observed test statistic is a surprising observation from  $f_T$ ; if it is reject  $H_0$

Step 3 is crucial: for some tests (for example, one and two sample tests based on the Normal distribution assumption), it is possible to find  $f_T$  analytically for appropriate choices of T in Step 2. For others, such as the chi-squared goodness of fit and related tests,  $f_T$  is only available approximately. However, the null distribution (and hence the critical regions and p-value) can, in theory, **always** be found: it is the probability distribution of the statistic T under the model restriction imposed by the null hypothesis.

# 5.6 PERMUTATION TESTS AND RESAM-PLING METHODS

For most of the hypothesis tests above, we start with the assumptions and work forward to derive the sampling distribution of the test statistic under the null hypothesis. For **permutation tests**, we will reverse the procedure, since the sampling distribution involves the permutations which give the procedure its name and are the key theoretical issue in understanding the test. For **resampling** or **bootstrap** methods, we will resample the original data uniformly and randomly so as to explore the variability of a test statistic.

## 5.6.1 PERMUTATION TESTS

A permutation is a reordering of the numbers 1, ..., n. For example, (1, 2, 3, 4, 5, 6), (1, 3, 2, 4, 5, 6), (4, 5, 2, 6, 1, 3) (3, 2, 1, 6, 4, 5) are all permutations of the numbers 1 through 6 (note that this includes the standard order in first line). There are  $n! = 1 \times 2 \times 3 \times ... \times n$  permutations of n objects.

The central idea of permutation tests refers to rearrangements of the data. The null hypothesis of the test specifies that **the permutations** are all equally likely. The sampling distribution of the test statistic under the null hypothesis is computed by forming all (or many) of the permutations, calculating the test statistic for each and considering these values all equally likely.

Consider the following two group example, where we want to test for any significant difference between the groups.

Group 1 : 55, 58, 60

Group 2 : 12, 22, 34

Here are the steps we will follow to use a permutation test to analyze the differences between the two groups. For the original order the sum for Group 1 is 173. In this example, if the groups were truly equal (and the null hypothesis was true) then randomly moving the observations among the groups would make no difference in the sum for Group 1. Some of the sums would be a little larger than the original sum and some would be a bit smaller. For the six observations there are 720 permutations of which there are 20 distinct combinations for which we can compute the sum of Group 1.

	GROUP 1	GROUP 2	SUM		GROUP 1	GROUP 2	SUM
1	55, 58, 60	12, 22, 34	173	11	12, 22, 60	55, 58, 34	94
2	55, 58, 12	60, 22, 34	125	12	12, 58, 22	55, 60, 34	92
3	55, 58, 22	12, 60, 34	135	13	55, 12, 22	12, 55, 58	89
4	55, 58, 34	12, 22, 34	148	14	12, 34, 60	55, 58, 34	106
5	55, 12, 60	58, 22, 34	127	15	12, 58, 34	55, 22, 60	104
6	55, 22, 60	12, 58, 34	137	16	55, 12, 34	12, 58, 60	101
7	55, 34, 60	12, 22, 58	149	17	22, 34, 60	55, 58, 34	116
8	12, 58, 60	55, 22, 34	130	18	22, 58, 34	55, 22, 60	114
9	22, 58, 60	12, 55, 34	140	19	55, 22, 34	12, 58, 60	111
10	34, 58, 60	12,22,55	152	20	12, 22, 34	55, 58, 60	68

Of these 20 different orderings only **one** has a Group 1 sum that greater than or equal to the Group 1 sum from our original ordering. Therefore the probability that a sum this large or larger would occur by chance alone is 1/20 = 0.05 and can be considered to be statistically significant.

#### 5.6.2 MONTE CARLO METHODS

Above, the permutation yielded an **exact test** because we were able to enumerate all of the possible combinations. In larger examples it will not be possible, so we will have to take a large number of random orderings, sampled uniformly from the permutation distribution. A general **Monte Carlo** strategy for two sample testing is outlined below:

- 1. For two sample tests for samples of size  $n_1$  and  $n_2$ , compute the value of the test statistic for the observed sample  $t^*$
- 2. Randomly select one of the  $(n_1 + n_2)!$  permutations, re-arrange the data according to this permutation, allocate the first  $n_1$  to pseudosample 1 and the remaining  $n_2$  to pseudo-sample 2, and then compute the test statistic  $t_1$

- 3. Repeat 2. N times to obtain a random sample of  $t_1, t_2, ..., t_N$  of test statistics from the TRUE null distribution.
- 4. Compute the *p*-value by reporting

$$\frac{\text{Number of } t_1, t_2, ..., t_N \text{ more extreme than } t^*}{N}$$

this value will be a good approximation to the true p-value if the Monte Carlo sample size N is large enough.

# 5.6.3 RESAMPLING METHODS AND THE BOOT-STRAP

In statistical analysis, we usually interested in obtaining estimates of a parameter via some statistic, and also an estimate of the variability or uncertainty attached to this point estimate, and a confidence interval for the true value of the parameter.

Traditionally, researchers have relied on normal approximations to obtain standard errors and confidence intervals. These techniques are valid only if the statistic, or some known transformation of it, is asymptotically normally distributed. If the normality assumption does not hold, then the traditional methods should not be used to obtain confidence intervals. A major motivation for the traditional reliance on normal-theory methods has been computational tractability, computational methods remove the reliance on asymptotic theory to estimate the distribution of a statistic.

Resampling techniques such as the bootstrap and jackknife provide estimates of the standard error, confidence intervals, and distributions for any statistic. The fundamental assumption of bootstrapping is that the observed data are representative of the underlying population. By resampling observations from the observed data, the process of sampling observations from the population is mimicked. The key techniques are

• THE BOOTSTRAP: In bootstrap resampling, B new samples, each of the same size as the observed data, are drawn with replacement from the observed data. The statistic is first calculated using the observed data and then recalculated using each of the new samples, yielding a bootstrap distribution. The resulting replicates are used to calculate the bootstrap estimates of bias, mean, and standard error for the statistic.

• THE JACKKNIFE: In jackknife resampling, a statistic is calculated for the n possible samples of size n-1, each with one observation left out. The default sample size is n-1, but more than one observation may be removed. Jackknife estimates of bias, mean, and standard error are available and are calculated differently than the equivalent bootstrap statistics.

Using the bootstrap and jackknife procedures, all informative summaries (mean, variance, quantiles etc) for the sample-based estimates' sampling distribution can be approximated.