How to read a paper

Statistics for the non-statistician. I: Different types of data need different statistical tests

This is the fourth in a series of 10 articles introducing non-experts to finding medical articles and assessing their value

Unit for Evidence-Based Practice and Policy, Department of Primary Care and Population Sciences, University College London Medical School/ Royal Free Hospital School of Medicine, Whittington Hospital, London N19 5NF Trisha Greenhalgh, *senior lecturer*

p.greenhalgh@ucl. ac.uk

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As medicine leans increasingly on mathematics no clinician can afford to leave the statistical aspects of a paper to the "experts." If you are numerate, try the "Basic Statistics for Clinicians" series in the *Canadian Medical Association Journal*,¹⁴ or a more mainstream statistical textbook.⁵ If, on the other hand, you find statistics impossibly difficult, this article and the next in this series give a checklist of preliminary questions to help you appraise the statistical validity of a paper.

Have the authors set the scene correctly?

Have they determined whether their groups are comparable, and, if necessary, adjusted for baseline differences?

Most comparative clinical trials include either a table or a paragraph in the text showing the baseline characteristics of the groups being studied. Such a table should show that the intervention and control groups are similar in terms of age and sex distribution and key prognostic variables (such as the average size of a cancerous lump). Important differences in these characteristics, even if due to chance, can pose a challenge to your interpretation of results. In this situation, adjustments can be made to allow for these differences and hence strengthen the argument.⁶

What sort of data have they got, and have they used appropriate statistical tests?

Numbers are often used to label the properties of things. We can assign a number to represent our height, weight, and so on. For properties like these, the measurements can be treated as actual numbers. We can, for example, calculate the average weight and height of a group of people by averaging the measurements. But consider an example in which we use numbers to label the property "city of origin," where 1 = London, 2 = Manchester, 3 = Birmingham, and so on. We could still calculate the average of these



Summary points

In assessing the choice of statistical tests in a paper, first consider whether groups were analysed for their comparability at baseline

Does the test chosen reflect the type of data analysed (parametric or non-parametric, paired or unpaired)?

Has a two tailed test been performed whenever the effect of an intervention could conceivably be a negative one?

Have the data been analysed according to the original study protocol?

If obscure tests have been used, do the authors justify their choice and provide a reference?

numbers for a particular sample of cases, but we would be completely unable to interpret the result. The same would apply if we labelled the property "liking for x" with 1=not at all, 2=a bit, and 3=a lot. Again, we could calculate the "average liking," but the numerical result would be uninterpretable unless we knew that the difference between "not at all" and "a bit" was exactly the same as the difference between "a bit" and "a lot."

All statistical tests are either parametric (that is, they assume that the data were sampled from a particular form of distribution, such as a normal distribution) or non-parametric (they make no such assumption). In general, parametric tests are more powerful than nonparametric ones and so should be used if possible.

Non-parametric tests look at the rank order of the values (which one is the smallest, which one comes next, and so on) and ignore the absolute differences between them. As you might imagine, statistical significance is more difficult to show with non-parametric tests, and this tempts researchers to use statistics such as the r value inappropriately. Not only is the r value (parametric) easier to calculate than its non-parametric equivalent but it is also much more likely to give (apparently) significant results. Unfortunately, it will give a spurious estimate of the significance of the result, unless the data are appropriate to the test being used. More examples of parametric tests and their non-parametric equivalents are given in table 1.

Another consideration is the shape of the distribution from which the data were sampled. When I was at school, my class plotted the amount of pocket money received against the number of children receiving that amount. The results formed a histogram the same shape as figure 1—a "normal" distribution. (The term "normal" refers to the shape of the graph and is used

Table 1 Some commonly used statistical tests

Parametric test	Example of equivalent non-parametric test	Purpose of test	Example
Two sample (unpaired) t test	Mann-Whitney U test	Compares two independent samples drawn from the same population	To compare girls' heights with boys' heights
One sample (paired) <i>t</i> test	Wilcoxon matched pairs test	Compares two sets of observations on a single sample	To compare weight of infants before and after a feed
One way analysis of variance (<i>F</i> test) using total sum of squares	Kruskall-Wallis analysis of variance by ranks	Effectively, a generalisation of the paired <i>t</i> or Wilcoxon matched pairs test where three or more sets of observations are made on a single sample	To determine whether plasma glucose level is higher one hour, two hours, or three hours after a meal
Two way analysis of variance	Two way analysis of variance by ranks	As above, but tests the influence (and interaction) of two different covariates	In the above example, to determine if the results differ in male and female subjects
χ^2 test	Fisher's exact test	Tests the null hypothesis that the distribution of a discontinuous variable is the same in two (or more) independent samples	To assess whether acceptance into medical school is more likely if the applicant was born in Britain
Product moment correlation coefficient (Pearson's <i>r</i>)	Spearman's rank correlation coefficient (\mathbf{r}_{σ})	Assesses the strength of the straight line association between two continuous variables.	To assess whether and to what extent plasma HbA ₁ concentration is related to plasma triglyceride concentration in diabetic patients
Regression by least squares method	Non-parametric regression (various tests)	Describes the numerical relation between two quantitative variables, allowing one value to be predicted from the other	To see how peak expiratory flow rate varies with height
Multiple regression by least squares method	Non-parametric regression (various tests)	Describes the numerical relation between a dependent variable and several predictor variables (covariates)	To determine whether and to what extent a person's age, body fat, and sodium intake determine their blood pressure

because many biological phenomena show this pattern of distribution). Some biological variables such as body weight show "skew normal" distribution, as shown in figure 2. (Figure 2 shows a negative skew, whereas body weight would be positively skewed. The average adult male body weight is 70 kg, and people exist who weigh 140 kg, but nobody weighs less than nothing, so the graph cannot possibly be symmetrical.)

Non-normal (skewed) data can sometimes be transformed to give a graph of normal shape by performing some mathematical transformation (such as using the variable's logarithm, square root, or reciprocal). Some data, however, cannot be transformed into a smooth pattern. For a very readable discussion of the normal distribution see chapter 7 of Martin Bland's *Introduction to Medical Statistics.*⁵

Deciding whether data are normally distributed is not an academic exercise, since it will determine what type of statistical tests to use. For example, linear regression will give misleading results unless the points on the scatter graph form a particular distribution about the regression line—that is, the residuals (the perpendicular distance from each point to the line) should themselves be normally distributed. Transforming data to achieve a normal distribution (if this is indeed achievable) is not cheating: it simply ensures that data values are given appropriate emphasis in assessing the overall effect. Using tests based on the normal distribution to analyse non-normally distributed data, however, is definitely cheating.

If the authors have used obscure statistical tests, why have they done so and have they referenced them?

The number of possible statistical tests sometimes seems infinite. In fact, most statisticians could survive with a formulary of about a dozen. The rest should generally be reserved for special indications. If the paper you are reading seems to describe a standard set of data which have been collected in a standard way, but the test used has an unpronounceable name and is not listed in a basic statistics textbook, you should smell a rat. The authors should, in such circumstances, state why they have used this test, and give a reference (with page numbers) for a definitive description of it. Are the data analysed according to the original protocol? If you play coin toss with someone, no matter how far you fall behind, there will come a time when you are one ahead. Most people would agree that to stop the game then would not be a fair way to play. So it is with research. If you make it inevitable that you will (eventually) get an apparently positive result you will also make it inevitable that you will be misleading yourself about the justice of your case.⁷ (Terminating an intervention trial prematurely for ethical reasons when subjects in one arm are faring particularly badly is a different matter and is discussed elsewhere.⁷)

Raking over your data for "interesting results" (retrospective subgroup analysis) can lead to false conclu-



sions.⁸ In an early study on the use of aspirin in preventing stroke, the results showed a significant effect in both sexes combined, and a retrospective subgroup analysis seemed to show that the effect was confined to men.⁹ This conclusion led to aspirin being withheld from women for many years, until the results of other studies¹⁰ showed that this subgroup effect was spurious.

This and other examples are included in Oxman and Guyatt's, "A consumer's guide to subgroup analysis," which reproduces a useful checklist for deciding whether apparent subgroup differences are real.¹¹

Paired data, tails, and outliers

Were paired tests performed on paired data?

Students often find it difficult to decide whether to use a paired or unpaired statistical test to analyse their data. There is no great mystery about this. If you measure something twice on each subject—for example, blood pressure measured when the subject is lying and when standing—you will probably be interested not just in the average difference of lying versus standing blood pressure in the entire sample, but in how much each individual's blood pressure changes with position. In this situation, you have what is called "paired" data, because each measurement beforehand is paired with a measurement afterwards.

In this example, it is using the same person on both occasions which makes the pairings, but there are other possibilities (for example, any two measurements of bed occupancy made of the same hospital ward). In these situations, it is likely that the two sets of values will be significantly correlated (for example, my blood pressure next week is likely to be closer to my own blood pressure last week than to the blood pressure of a randomly selected adult last week). In other words, we would expect two randomly selected paired values to be closer to each other than two randomly selected unpaired values. Unless we allow for this, by carrying out the appropriate paired sample tests, we can end up with a biased estimate of the significance of our results.

Was a two tailed test performed whenever the effect of an intervention could conceivably be a negative one?

The term "tail" refers to the extremes of the distribution—the areas at the outer edges of the bell in figure 1. Let's say that the graph represents the diastolic blood pressures of a group of people of which a random sample are about to be put on a low sodium diet. If a low sodium diet has a significant lowering effect on blood pressure, subsequent blood pressure measurements on these subjects would be more likely to lie within the left tail of the graph. Hence we would analyse the data with statistical tests designed to show whether unusually low readings in this patient sample were likely to have arisen by chance.

But on what grounds may we assume that a low sodium diet could only conceivably put blood pressure down, but could never do the reverse, put it up? Even if there are valid physiological reasons in this particular example, it is certainly not good science always to assume that you know the direction of the effect which your intervention will have. A new drug intended to relieve nausea might actually exacerbate it, or an educational leaflet intended to reduce anxiety might increase it. Hence, your statistical analysis should, in general, test the hypothesis that either high or low values in your dataset have arisen by chance. In the language of the statisticians, this means you need a two tailed test, unless you have very convincing evidence that the difference can only be in one direction.

Were "outliers" analysed with both common sense and appropriate statistical adjustments?

Unexpected results may reflect idiosyncrasies in the subject (for example, unusual metabolism), errors in measurement (faulty equipment), errors in interpretation (misreading a meter reading), or errors in calculation (misplaced decimal points). Only the first of these is a "real" result which deserves to be included in the analysis. A result which is many orders of magnitude away from the others is less likely to be genuine, but it may be so. A few years ago, while doing a research project, I measured several different hormones in about 30 subjects. One subject's growth hormone levels came back about 100 times higher than everyone else's. I assumed this was a transcription error, so I moved the decimal point two places to the left. Some weeks later, I met the technician who had analysed the specimens and he asked, "Whatever happened to that chap with acromegaly?"

Statistically correcting for outliers (for example, to modify their effect on the overall result) requires sophisticated analysis and is covered elsewhere.⁶

The articles in this series are excerpts from *How to read a paper: the basics of evidence based medicine.* The book includes chapters on searching the literature and implementing evidence based findings. It can be ordered from the BMJ Bookshop: tel 0171 383 6185/6245; fax 0171 383 6662. Price £13.95 UK members, £14.95 non-members.

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