

Comparing methods of measurement: why plotting difference against standard method is misleading

J Martin Bland, Douglas G Altman

Summary

When comparing a new method of measurement with a standard method, one of the things we want to know is whether the difference between the measurements by the two methods is related to the magnitude of the measurement. A plot of the difference against the standard measurement is sometimes suggested, but this will always appear to show a relationship between difference and magnitude when there is none. A plot of the difference against the average of the standard and new measurements is unlikely to mislead in this way. We show this theoretically and illustrated by a practical example.

Lancet 1995; **346**: 1085-87.

Introduction

In earlier papers [1,2] we discussed the analysis of agreement between methods of clinical measurement. We had two issues in mind: to demonstrate that the analyses then in general use were incorrect and misleading, and to recommend a more appropriate method. We saw the aim of such studies as to determine whether two methods agreed sufficiently well for them to be used interchangeably. This led us to suggest that the analysis should be based on the differences between measurements on the same subject by the two methods. The mean difference would be the estimated bias, the systematic difference between methods, and the SD of the differences would measure random fluctuations around this mean. We recommended 95% limits of agreement, mean difference plus or minus 2 (more precisely, 1.96) SDs, which would tell us how far apart measurements by the two methods were likely to be for most individuals.

Figure 1 shows a typical data set, the measurement of systolic blood pressure by a test method, finger pressure, and a standard method, arm blood pressure. This is a random subsample of 200 observations from a larger data set. [3, 4] The sub-sample was drawn to avoid cluttered graphs. The mean difference, finger minus arm, is 4.3 mm Hg and the SD is 14.6 mm Hg. Hence the lower 95% limit is $4.3 - 1.96 \times 14.6 = -24$ mm Hg and the upper 95% limit is $4.3 + 1.96 \times 14.6 = 33$ mm Hg. Thus we estimate that for 95% of subjects the finger measurement will be between 24 mm Hg below the arm measurement and 33 mm Hg above it.

For the mean and SD of the differences to be meaningful estimates we must assume that they are reasonably constant throughout the range of measurement. We suggested checking this assumption graphically. [1, 2] The usual plot, method one versus method two, is inefficient because the

points tend to be clustered along the line of equality (figure 1), especially if the two methods give closely related measurements. We therefore proposed that the difference be plotted against the average of the measurements by the two methods (figure 2).

In figure 2 we have also added the 95% limits of agreement and the regression line of difference on average. The main departure from assumptions we were expecting was an increase in variability, shown by an increase in the scatter of the differences, as the magnitude of the measurement increased. There may also be a trend in the bias, a tendency for the mean difference to rise or fall with increasing magnitude. Either would show that the methods did not agree equally through the range. In figure 2, for example, there is an increase in bias with magnitude, shown by the positive slope of the regression line. Such deviations from assumptions can often be dealt with by a suitable transformation, usually logarithmic. [2] In particular, this approach will be effective if the differences are proportional to the magnitude of the measurement.

The 95% limits of agreement approach has been widely adopted and the *Lancet* paper [2] widely cited. [5] However, it is sometimes argued that when one method may be regarded as a 'gold standard', it is presumably more accurate than the other method and so we should plot the difference against the gold standard. [6,7] We think that this idea is misguided and is likely to lead to misinterpretation. Here we will show why, and that the plot of difference against average is almost always preferable.

Plotting difference against average

We denote our standard measurement by S , the new or test measurement by T , their variances by σ_S^2 and σ_T^2 , and their correlation by ρ . If the study includes a wide range of measurements, and unless the two methods of

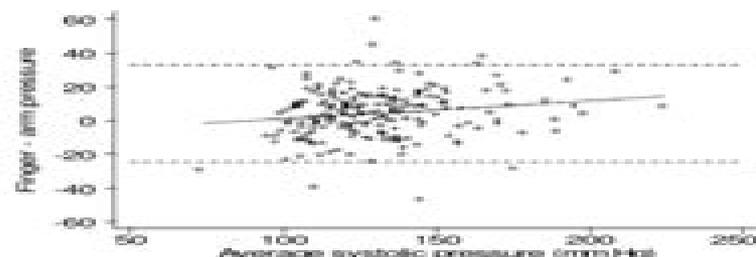
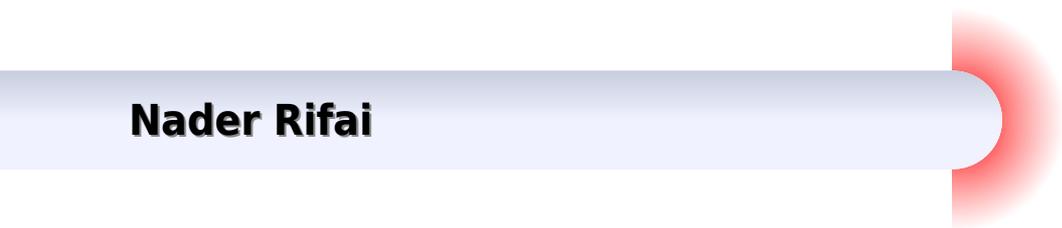


Figure 1. Test and standard measurements, with line of equality

Department of Public Health Sciences, St George's Hospital Medical School, Cranmer Terrace, London SW17 0RE (J M Bland, Ph.D) and Medical Statistics Laboratory, Imperial Cancer Research Fund, London WC2A 3PX (D G Altman, B.Sc.)

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cohorts and outdoor workers Controversial topics such as environmental effects of sunscreen products and the risks and benefits of UV radiation in the context of skin cancer vitamin D and cardiovascular and metabolic health are also covered

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Table of Contents Comparing Methods Of Measurement Why Plotting Difference

1. Understanding the eBook Comparing Methods Of Measurement Why Plotting Difference
 - The Rise of Digital Reading Comparing Methods Of Measurement Why Plotting Difference
 - Advantages of eBooks Over Traditional Books
2. Identifying Comparing Methods Of Measurement Why Plotting Difference
 - Exploring Different Genres
 - Considering Fiction vs. Non-Fiction
 - Determining Your Reading Goals
3. Choosing the Right eBook Platform
 - Popular eBook Platforms
 - Features to Look for in an Comparing Methods Of Measurement Why Plotting Difference
 - User-Friendly Interface
4. Exploring eBook Recommendations from Comparing Methods Of Measurement Why Plotting Difference
 - Personalized Recommendations

- Comparing Methods Of Measurement Why Plotting Difference User Reviews and Ratings
- Comparing Methods Of Measurement Why Plotting Difference and Bestseller Lists
- 5. Accessing Comparing Methods Of Measurement Why Plotting Difference Free and Paid eBooks
 - Comparing Methods Of Measurement Why Plotting Difference Public Domain eBooks
 - Comparing Methods Of Measurement Why Plotting Difference eBook Subscription Services
 - Comparing Methods Of Measurement Why Plotting Difference Budget-Friendly Options
- 6. Navigating Comparing Methods Of Measurement Why Plotting Difference eBook Formats
 - ePub, PDF, MOBI, and More
 - Comparing Methods Of Measurement Why Plotting Difference Compatibility with Devices
 - Comparing Methods Of Measurement Why Plotting Difference Enhanced eBook Features
- 7. Enhancing Your Reading Experience
 - Adjustable Fonts and Text Sizes of Comparing Methods Of Measurement Why Plotting Difference
 - Highlighting and Note-Taking Comparing Methods Of Measurement Why Plotting Difference
 - Interactive Elements Comparing Methods Of Measurement Why Plotting Difference
- 8. Staying Engaged with Comparing Methods Of Measurement Why Plotting Difference
 - Joining Online Reading Communities
 - Participating in Virtual Book Clubs
 - Following Authors and Publishers Comparing Methods Of Measurement Why Plotting Difference
- 9. Balancing eBooks and Physical Books Comparing Methods Of Measurement Why Plotting Difference
 - Benefits of a Digital Library
 - Creating a Diverse Reading Collection Comparing Methods Of Measurement Why Plotting Difference
- 10. Overcoming Reading Challenges
 - Dealing with Digital Eye Strain
 - Minimizing Distractions
 - Managing Screen Time
- 11. Cultivating a Reading Routine Comparing Methods Of Measurement Why Plotting Difference
 - Setting Reading Goals Comparing Methods Of Measurement Why Plotting Difference
 - Carving Out Dedicated Reading Time
- 12. Sourcing Reliable Information of Comparing Methods Of Measurement Why Plotting Difference
 - Fact-Checking eBook Content of Comparing Methods Of Measurement Why Plotting Difference

- Distinguishing Credible Sources
13. Promoting Lifelong Learning
- Utilizing eBooks for Skill Development
 - Exploring Educational eBooks
14. Embracing eBook Trends
- Integration of Multimedia Elements
 - Interactive and Gamified eBooks

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