Introduction

Most health professionals are aware that estimates based on a random sample of a population are subject to error due to sampling variability. Fewer people are aware that rates and percentages based on a full population count are also estimates subject to error. Random error may be substantial when the measure, such as a rate or percentage, has a small number of events in the numerator (e.g., less than 20). A rate observed in a single year can be considered as a sample or estimate of the true or underlying rate. This idea of an “underlying” rate is an abstract concept, since the rate observed in one year did actually occur. However, since annual observed rates may fluctuate dramatically, it is the underlying rate that health policies should seek to address. The larger the numerator of the observed rate, the better the observed rate will estimate the underlying rate.

Many publications of the State Center for Health Statistics contain rates or percentages with a small numerator. This is a problem with a measure such as the infant mortality rate. In a single year many counties may have only one or two infant deaths and such rates in a small population may fluctuate dramatically from year to year. One means of addressing this problem is to look at five-year rates where the numerator will be larger. Even with five-year rates, however, many counties will have few events and therefore unstable rates. Many cause-specific death rates for individual counties will have small numerators. This statistical problem is compounded when age-adjusted rates are produced because, in the process of calculating an age-adjusted rate by the direct method, the deaths and population are broken up into smaller groups. Rates are calculated for a number of specific age groups and numerators for each rate are often small.

Some customers of the State Center for Health Statistics may treat our published rates and percentages as completely accurate. Unfortunately, there is the danger of making unwarranted comparisons between geographic areas or comparisons over time when the rates or percentages have small numerators. We do not consider it feasible to completely ignore all rates based on small numbers. In one sense, the rates do describe what actually happened in a year, but you must use caution and interpret any comparisons critically. The following section provides some methods for quantifying random errors in rates as a basis for making decisions about when changes or differences in rates are meaningful.
Calculation of Errors in Rates

The formulas presented here provide a means of estimating the confidence interval around a single rate and for determining whether the difference between two rates is statistically significant.

*Definition: A confidence interval is a range above and below an observed rate within which we would expect the “true” rate to lie a certain percentage of the time (usually 95%).

Calculation of a confidence interval recognizes that an observed rate is not a precise estimate of the underlying rate because the observed rate is influenced by random error. The formulas below are exactly the same as the ones used for a random sample from a larger population. The population rate for a given year based on a complete count can be considered a sample of one of a large number of possible measurements, all of which cluster in a normal distribution (bell curve) around the “true” (unknown) rate of the population. The larger the numerator of the measured rate, the better the rate will estimate the true or underlying rate of the population. The confidence interval accounts for only random measurement error. Systematic errors or biases in measurement may still be present and cannot be assessed by these formulas.

These formulas apply to any proportion or simple (crude) rate. Random errors may also be estimated for adjusted rates and other more complex measures, but a description of this is beyond the scope of the present Primer.

Proportions vs. Percentages vs. Rates

The formulas below are expressed in terms of \( p \), or the proportion or fraction of a population that has a certain characteristic (e.g., death, low birthweight, early prenatal care). In this context, the terms proportion, percentage, and rate are used interchangeably. For example, in 1995 Wake County had a resident population of approximately 518,000 out of which approximately 2,900 died during the year. The proportion who died is \( \frac{2,900}{518,000} = .005598 \). For the percentage who died, multiply by 100; the result is .5598%. A percentage is simply a rate per 100. For a rate per 1,000, multiply the proportion by 1,000; the result is 5.598 deaths per 1,000 population. The number of deaths per 100,000 is 559.8. So the multiplier is completely arbitrary, though for rare events we usually use 1,000 or higher so that the rate is not a decimal fraction.

The formulas presented below use \( p \), or the proportion, so a percentage or rate has been converted back to the proportion (by dividing by the multiplier) in these examples.

**Infant Death Rates**

The infant death rates (expressed per 1,000 live births) reported in State Center for Health Statistics publications are not strictly proportions since the deaths and births occurred during a particular calendar year. Though approximately one-half of infant deaths occur on the first day of life, some of the infant deaths that occur in a given year are to babies born in the previous calendar year. Technically, the more correct way to compute the proportion of babies who before their first birthday would be to use a linked birth/infant death file to track a population of births (also called a birth cohort) through the first year of life. But in practicality this difference is small. We suggest that the formulas below may reasonably be used for infant deaths rates reported as usual based on year of occurrence and expressed as the proportion of babies who die.

\[ \text{Infant death rate} = \frac{\# \text{deaths under 1 year of age}}{\# \text{of live births}} \times 1000 \]

**Confidence Intervals**

We can compute a confidence interval around a proportion or rate. The confidence interval is the interval within which we would expect the “true” rate to fall a certain percentage of the time. A 95% confidence interval is frequently used, which means using a multiplier (“Z” value) of 1.96. For a 99% confidence interval, one would use the multiplier 2.57. Let us say that in hypothetical Rocky County there are 20 infant deaths (d) out of a population of 1,900 live births (n) in a single year. The proportion dying (p) is \( \frac{20}{1,900} = .0105 \). You can also say that 1.05 percent died or that the infant death rate is 10.5 per 1,000 births for that year.

**Formula:**

\[ 95\% \text{ Confidence Interval} = p \pm 1.96 \sqrt{\frac{p \cdot q}{n}} \]
where \( q = 1-p \). This formula works for any value of \( p \), though for small values of \( p \) (.01 or less), the value of \( q \) is very close to 1 and may therefore be ignored. In the current example this calculates out to:

\[
q = 1-.0105 = .9895
\]

95% Confidence Interval =

\[
.0105 \pm 1.96 \sqrt{.0105 \times .9895 / 1900} = .0105 \pm .0046.
\]

Expressed in the traditional way in terms of infant deaths per 1,000 live births, we can say that we are 95% sure that the true infant death rate for this population is between 5.9 and 15.1. These limits are quite large. A useful rule of thumb is that any rate with fewer than 20 events in the numerator will have a confidence interval that is wider than the rate itself. In the current example of a rate of 10.5 per 1,000 with a numerator of 20, the width of the confidence interval is 9.2.

\[\text{Formula:}\]

Width of the confidence interval = higher limit - lower limit

In the current example this is: 15.1 - 5.9 = 9.2

**Combining Data for Greater Precision**

One way to reduce the error of a rate is to combine several years of data. Another way is to combine geographic areas; for example, look at regional rather than county-level rates. In the example above, let us assume that over a five year period in Rocky County we observed five times as many infant deaths and live births (100 and 9,500 respectively) as in the example above.

The five-year infant death rate would still be 10.5, but with the larger numerator, the range of the 95% confidence interval would be much smaller (8.5 to 12.5). Try the calculations so you can verify this result. In general, you have to quadruple the sample size (\( n \)) to cut the random error in half.

**Differences Between Rates**

When comparing rates, you might want to assess the statistical significance of a change in a rate over time, or of the difference between two rates in one period of time (for example between two geographic areas or population groups).

The **standard error of the difference between two rates** is computed as:

\[\text{Formula:}\]

\[
SE = \sqrt{\frac{p_1 q_1}{n_1} + \frac{p_2 q_2}{n_2}}
\]

where \( p_1 \) and \( p_2 \) are the two rates to be compared expressed as proportions. The difference between the two proportions can be considered statistically significant at the 95% confidence level if the difference is greater than 1.96 multiplied by the standard error (computed above).

As an example, take a county where the percentage of women who smoked during pregnancy (from the birth certificates) declined from 21.4% in 2000 to 16.7% in 2005. We want to know if this change is statistically significant at the 95% confidence level. In 2000, the mother smoked for 150 births (\( d_1 \)) out of 700 total births (\( n_1 \)). In 2005, the mother smoked for 125 births (\( d_2 \)) out of 750 total births (\( n_2 \)). The proportions are

\[
p_1 = \frac{d_1}{n_1} = .214 \quad \text{and} \quad p_2 = \frac{d_2}{n_2} = .167 \quad \text{(or 21.4\% and 16.7\%).}
\]

Therefore, the calculation of 1.96 times the standard error of the difference is as follows:

\[
1.96 \times SE = 1.96 \sqrt{\frac{.214 \times .786}{700} + \frac{.167 \times .833}{750}} = .0404
\]

Since the difference between the two proportions of .047 (i.e., .214 - .167) exceeds 1.96 times the standard error of the difference (i.e., .0404), we can say that the decline in the smoking percentage in this county is statistically significant at the 95% confidence level. Or stated another way, the probability is less than .05 (or 5\%) that the observed decline in smoking was due to chance.

The formula for the standard error of the difference can be used to solve for any unknown in the equation. For example, if you want to know what the exact level of statistical significance of an observed difference between two proportions is, solve for the multiplier (“Z”) by dividing the observed difference by the standard error of the difference and look up the probability value for Z in a table of areas under the normal curve. In the smoking example presented above, the probability that the observed decline would occur just due to random variation in the percentages is .02. Please verify this result by consulting a table of
areas under the normal curve in your statistical text or online. For assistance with this or for other questions, contact the State Center for Health Statistics.

**Other Issues**

These formulas are based on parameters of the normal curve and in some cases will be only an approximation. If \( n \) (sample or population size, also denominator of the proportion or rate) is less than 30, or if the number of events (numerator of the proportion) is less than 5, these formulas become less reliable and readers should contact the State Center for Health Statistics for more appropriate alternatives.

Another important consideration is the issue of practical versus statistical significance. If \( n \) is large enough, almost any difference will be statistically significant. However, the same difference may be of very little practical or clinical significance. It is the responsibility of the user of statistics to evaluate whether observed differences, which may be statistically significant, are of real public health importance.

Finally, the issue of using rates versus actual counts should be mentioned. Rates or proportions allow more standardized comparisons between populations of different size, but there may be substantial random measurement error involved. In many cases just looking at the number of events is appropriate; do not always rush to calculate a proportion or rate. If the number of infant deaths in a county increased from one in 2007 to two in 2008 and the number of births remained about the same, looking at the infant mortality rate would erroneously suggest that the problem had become twice as great. In this case, each infant death could be investigated as unique sentinel health event. Examining the numbers behind the rates is always a good idea, and in some cases just looking at the numbers makes more sense.

This section on calculation of errors in rates demonstrates that an observed rate or proportion should not be taken as an exact measure of the true value in a population. Even measures based on complete reporting from a population may have a substantial random error component.

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**Key Points to Remember**

- If the number of events (numerator) is less than 20, your statistic may be unreliable due to random error. Interpret it with caution and look at the raw numbers too!
- You can calculate a confidence interval around your statistic to get an idea of the precision of your estimate. Narrower confidence intervals indicate more precise estimates.
- To get greater precision and increase your sample size, combine data from several years, or several places during one year.
- You can check if the difference between two rates is statistically significant at a certain level by comparing the difference between the rates to the standard error of the difference multiplied by “\( Z \)” (usually \( Z = 1.96 \)).
- It is up to you, the health professional, to decide whether the difference between two rates is *clinically* important, no matter whether it is *statistically* significant or not!
Statistical Guidelines

To address the problems of rates based on small numbers, the State Center for Health Statistics has adopted the following statistical guidelines:

♦ All publications of the State Center for Health Statistics that contain rates or percentages should contain a caution about interpreting rates or percentages based on small numbers. This caution should be featured prominently in the introductory material, and then discussed in more detail in the methods or technical notes section. See the 2006 North Carolina Vital Statistics, Volume 1 and Volume 2, for examples of this.

♦ Such a caution should accompany any information that is sent out to a customer as a special data request, if the information contains rates or percentages based on small numbers.

♦ When rates or percentages are published or distributed, the numerators should also be shown if possible.

♦ When maps of rates are produced, where possible there should be a legend warning the reader to “interpret with caution” for rates or percentages based on a very small numerator, e.g., less than 20 events.

♦ At every opportunity, customers of the State Center for Health Statistics should be educated about statistical issues, and especially about the potential for misinterpretation when comparisons are made using rates or percentages based on small numbers.

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