

Sampling & Confidence Intervals

Mark Lunt

Centre for Epidemiology Versus Arthritis
University of Manchester



14/11/2023

Principles of Sampling

- Often, it is not practical to measure every subject in a population.
- A reduced number of subjects, a sample, is measured instead.
 - Cheaper
 - Quicker
 - More thorough
- Sample needs to be chosen in such a way as to be representative of the population

Types of Sample

- Simple Random
- Stratified
- Cluster

- Quota
- Convenience
- Systematic

Simple Random Sample

- Every subject has the same probability of being selected.
- This probability is independent of who else is in the sample.
- Need a list of every subject in the population (*sampling frame*).
- Statistical methods depend on randomness of sampling.
- Refusals mean the sample is no longer random.

- Divide population into distinct sub-populations.
 - E.g. into age-bands, by gender
- Randomly sample from each sub-population.
 - sampling probability is same for everyone in a sub-population
 - sampling probability differs between sub-populations
- More efficient than a simple random sample if variable of interest varies more *between* sub-populations than *within* sub-populations.

- Randomly sample groups of subjects rather than subjects
- Why ?
 - List of subjects not available, list of groups is
 - Cheaper and easier to recruit a number of subjects at the same time.
 - In intervention studies, may be easier to treat groups: randomise hospitals rather than patients.
- Need a reasonable number of clusters to assure representativeness.
- The more similar clusters are, the better cluster sampling works.
- Cluster samples need special methods for analysis

- Deliberate attempt to ensure proportions of subjects in each category in a sample match the proportion in the population.
- Often used in market research: quotas by age, gender, social status.
- Variables not used to define the quotas may be very different in the sample and population.
- Proportion of men and of elderly may be correct, not proportions of elderly men.
- Probability of inclusion is unknown, may vary greatly between categories
- Cannot assume sample is representative.

Systematic & Convenience Samples

Systematic Take every n^{th} subject.

- If there is clustering (or periodicity) in the sampling frame, may not be representative.
- Shared surnames can cause problems.
- Randomly order and take every n^{th} subject: random.

Convenience Take a random sample of easily accessible subjects

- May not be representative of entire population.
- E.g. people going to G.P. with sore throat easy to identify, not representative of people with sore throat.

Estimating from Random Samples

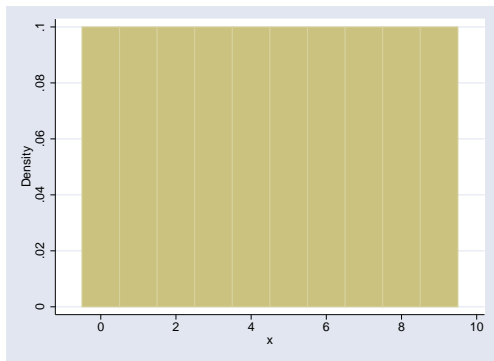
- We are interested in what our sample tells us about the population
- We use sample statistics to estimate population values
- Need to keep clear whether we are talking about sample or population
- Values in the population are given Greek letters $\mu, \pi \dots$, whilst values in the sample are given equivalent Roman letters $m, p \dots$
- Suppose we have a population, in which a variable x has a mean μ and standard deviation σ . We take a **random** sample of size n . Then
 - Sample mean \bar{x} should be close to the population mean μ .
 - However, if several samples are taken, \bar{x} in each sample will differ slightly.

Variation of \bar{x} around μ

- How much the means of different samples differ depends on
 - Sample Size** The mean of a small sample will vary more than the mean of a large sample.
 - Variance in the Population** If the variable measured varies little, the sample mean can only vary little.
- I.e. variance of \bar{x} depends on variance of x and on sample size n .

Example

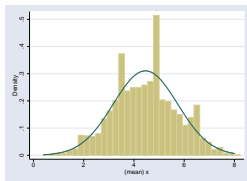
Consider consider a population consisting of 1000 copies of each of the digits 0, 1, ..., 9. The distribution of the values in this population is



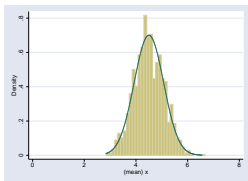
Example: Samples

- Samples of size 5, 25 and 100
- 2000 samples of each size were randomly generated
- Mean of x (\bar{x}) was calculated for each sample
- Histograms created for each sample size separately

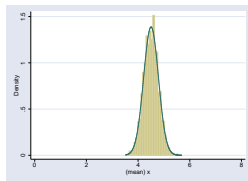
Example: Distributions of \bar{x}



Size 5



Size 25



Size 100

Properties of \bar{x}

$E(\bar{x}) = \mu$ i.e. on average, the sample mean is the same as the population mean.

Standard Deviation of $\bar{x} = \frac{\sigma}{\sqrt{n}}$ i.e the uncertainty in \bar{x} increases with σ , decreases with n . The standard deviation of the sampling distribution of the mean is also called the **Standard Error**

\bar{x} is normally distributed This is true whether or not x is normally distributed, provided n is sufficiently large. Thanks to the *Central Limit Theorem*.

- Standard deviation of the *sampling distribution* of a statistic
- Sampling distribution: the distribution of a statistic as sampling is repeated
- All statistics have sampling distributions
- Statistical inference is based on the standard error

Example: Sampling Distribution of \bar{x}

$$\mu = 4.5 \quad \sigma = 2.87$$

Size of samples	Mean \bar{x}		S.D. \bar{x}	
	Predicted	Observed	Predicted	Observed
5	4.5	4.47	1.29	1.26
25	4.5	4.51	0.57	0.57
100	4.5	4.50	0.29	0.30

Estimating the Variance

In a population of size N , the variance of x is given by

$$\sigma^2 = \frac{\sum(x_i - \mu)^2}{N} \quad (1)$$

This is the *Population Variance*

In a sample of size n , the variance of x is given by

$$s^2 = \frac{\sum(x_i - \bar{x})^2}{n - 1} \quad (2)$$

This is the *Sample Variance*

Why $n - 1$ rather than N

Population $\sigma^2 = \frac{\sum(x_i - \mu)^2}{N}$

Sample $s^2 = \frac{\sum(x_i - \bar{x})^2}{n-1}$

- Use $n - 1$ rather than n because we don't know μ , only an imperfect estimate \bar{x} .
- Since \bar{x} is calculated from the sample (i.e. from the x_i), x_i will tend to be closer to \bar{x} than it is to μ .
- Dividing by n would underestimate the variance
- With a reasonable sample size, makes little difference.

Suppose that you want to estimate π , the proportion of subjects in the population with a given characteristic. You take a random sample of size n , of whom r have the characteristic.

- $p = \frac{r}{n}$ is a good estimator for π .
- If you create a variable x which is 1 for subjects which have the characteristic and 0 for those who do not, then $p = \bar{x}$
- If the sample is large, p will be normally distributed, even though x isn't

Confidence Intervals

- The distribution of \bar{x} approaches normality as n gets bigger.
- \bar{x} can be thought of as a random draw from a distribution with mean μ and standard deviation $\frac{\sigma}{\sqrt{n}}$.
- If samples could be taken repeatedly, 95% of the time, the \bar{x} would lie between $\mu - 1.96 \frac{\sigma}{\sqrt{n}}$ and $\mu + 1.96 \frac{\sigma}{\sqrt{n}}$.
- As a consequence, 95% of the time, μ would lie between $\bar{x} - 1.96 \frac{\sigma}{\sqrt{n}}$ and $\bar{x} + 1.96 \frac{\sigma}{\sqrt{n}}$.
- This is a 95% confidence interval for the population mean.
- If, as is usually the case, σ is unknown, can use its estimate s .

Confidence Interval Example

In 216 patients with primary biliary cirrhosis, serum albumin had a mean value of 34.46 g/l and a standard deviation of 5.84 g/l.

$$\begin{aligned} \text{Standard deviation of } x &= 5.84 \\ \Rightarrow \text{Standard error of } \bar{x} &= \frac{5.84}{\sqrt{216}} \\ &= 0.397 \\ \Rightarrow \text{95\% Confidence Interval} &= 34.46 \pm 1.96 \times 0.397 \\ &= (33.68, 35.24) \end{aligned}$$

So, the mean value of serum albumin in the *population* of patients with primary biliary cirrhosis is probably between 33.68 g/l and 35.24 g/l.

Confidence Intervals for Proportions

- p is normally distributed with standard error $\sqrt{\frac{p(1-p)}{n}}$ **provided n is large enough.**
- This can be used to calculate a confidence interval for a proportion.
- Exact confidence intervals can be calculated for small n (less than 20, say) from tables of the binomial distribution.
- A reference range for a proportion is meaningless: a subject either has the characteristic or they do not.

Confidence Interval around a Proportion: Example

100 subjects each receive two analgesics, X and Y, for one week each in a randomly determined order. They then state a preference for one drug. 65 prefer X, 35 prefer Y. Calculate a 95% confidence interval for the proportion preferring X.

$$\begin{aligned}\text{Standard Error of } p &= \sqrt{\frac{0.65 \times 0.35}{100}} \\ &= 0.0477\end{aligned}$$

$$\begin{aligned}\Rightarrow \text{95\% Confidence Interval} &= 0.65 \pm 1.96 \times 0.0477 \\ &= (0.56, 0.74)\end{aligned}$$

So, in the general population, it is likely that between 56% and 74% of people would prefer X.

- The `ci` command produces confidence intervals
- For proportions, you use the `binomial` option

Confidence Intervals and Big Data

- 95% Confidence Interval holds population value 95% of the time *provided sample is random*
- In practice, selection bias will have some effect
- Will tend to move confidence interval up or down
- Problem with big data is that confidence intervals are narrow: slight movement may be enough to move population value out of confidence interval
- *Coverage* may be lower than nominal 95% level if bias is large compared to sampling error
- See “Uncertainty beyond sampling error”

Sample Size Calculations

- Primary outcome of a study is a statistic (mean, proportion, relative risk, incidence rate, hazard ratio etc)
- The larger the study, the more precisely we can estimate our statistic
- We can calculate how many subjects we need to achieve adequate precision if we
 - know how the distribution of the statistic changes with increasing numbers of subjects
 - Have a definition of “adequate”
- Power-based calculations are more complicated (for next week).

Sample size for precision of mean

Suppose that we want to know μ to a certain level of precision.

- We can be 95% certain that μ lies within

$$\bar{x} \pm \frac{1.96\sigma}{\sqrt{n}}$$

- The width of this interval depends on n , which we control.
- Therefore, we can select n to give our chosen width.
- Need to use an estimate for σ , for which we can use s .

Sample Size Formula

Suppose we want to fix the width of the 95% confidence interval to $2W$, i.e. 95% CI = $\bar{x} \pm W$. Then

$$\begin{aligned}W &= 1.96 \times \text{Standard Error} \\ &= 1.96 \times \frac{\sigma}{\sqrt{n}} \\ \Rightarrow W^2 &= \frac{1.96^2 \sigma^2}{n} \\ \Rightarrow n &= \left(\frac{1.96\sigma}{W} \right)^2\end{aligned}$$

Sample Size Example

In the primary biliary cirrhosis example, suppose that we wish to know the mean serum albumin in cirrhosis patients to within 0.5 g/l. How many patients would we need to study (assuming a standard deviation of 5.84 g/l) ?

$$\begin{aligned}W &= 0.5 \\ \sigma &= 5.84 \\ \Rightarrow n &= \left(\frac{1.96\sigma}{W} \right)^2 \\ &= \left(\frac{1.96 \times 5.84}{0.5} \right)^2 \\ &\approx 524\end{aligned}$$