



BIOSTATISTICS



Biostatistics index

STATISTICS

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EPIDEMIOLOGY

Analytical Epidemiology
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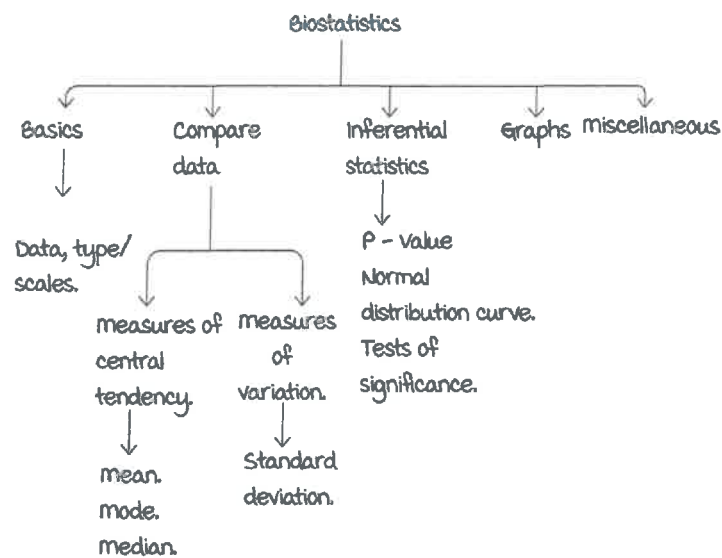
SCREENING

Advanced concepts in screening of disease

INTRODUCTION TO DATA IN BIOSTATISTICS

Uses :

- Define cut-offs.
- Understand variation.
- To present data.
- To make inference (provide evidence).



Data

00:07:54

Quantitative	Qualitative
• Continuous. • measurable. • E.g. weight, height, AST, ALT levels. • mean of data can be calculated.	• Discrete. • Countable. • E.g. No. of people who are sick/ healthy, alive/dead. Gender. • Proportions/ percentages can be calculated.

Pulse rate is a data which is discrete and countable, however it is quantitative as we calculate its mean.

BP is quantitative data.

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Scales of data

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Nominal	Ordinal	Interval	Ratio
• Named data. • No sequence • E.g. Gender, religion, blood groups.	• Inherent order. • Has a sequence. • E.g. Stage, grade, the severity of the disease.	• Interval between two values is present. • No start point/no absolute zero. • E.g. °C, dB.	• Ratio can be calculated. • There is zero point/absolute zero. • E.g. Na, K, FEV levels.

Interval type of data :

Example : 20 °C is not half as hot as 40 °C, but colder compared to 40 °C. Here the intensity of data is measured. Also, the temperature can go below 0 °C (in minus °C), which means there is no absolute zero.

Ratios :

Example : A weak fragile child weighs 20 kg when the ideal weight should have been 40 kg in the same age group. The ideal weight is 2 x child's age, which means the values can be expressed in multiples (double, triple) of each other i.e calculation of ratios is possible. Also, there is absolute zero/ no value below zero.

MEASURES OF CENTRAL TENDENCY AND VARIATION

Measures of Central tendency

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Mean :

1. Arithmetic mean :

$$\text{Average} = \frac{\Sigma(\text{summation})}{n}$$

2. Geometric mean :

- Calculated in case of : Exponential data.

Extreme values.

- Example : Human development index

(India = 0.647, ranked at 129 in 2019)

3. Harmonic mean :

- Calculated in case of : Inverse data.

Fractional values.

Advantages :

- Best measure of central tendency.
- Easiest to calculate.

Disadvantages :

- most affected by extreme values.

Median :

Central value after arranging in ascending or descending order.

Advantages :

- Least affected by extreme values.

Mode :

The most frequently occurring value.

Mode = 3 median - 2 mean.

Advantages :

- The most robust measure of central tendency.
- The last to be affected by extreme values.

Data with extreme values : Preferred measure is median.

Preferred mean is geometric mean.

Measures of variation

00:14:11

1. Range :

Range = maximum to minimum.

2. Standard deviation :

Gives the mean deviation of every value from the mean.

Formula : The root of the mean of squared deviation.

$$SD = \sqrt{\frac{\sum (x - \bar{x})^2}{n}}$$

In case of a small sample,

$$SD = \sqrt{\frac{\sum (x - \bar{x})^2}{n - 1}}$$

 $n - 1$ is the correction for the small sample ($n < 30$).

3. Variance :

Variance (V) = SD^2

$$V = \frac{\sum (x - \bar{x})^2}{n}$$

4. Coefficient of variation (CV) :

Absolute variation between 2 different populations.

$$CV = \frac{SD}{\text{mean}} \times 100$$

5. Standard error :

Gives the error in different studies in terms of standard deviation.

Alternatively, gives the variation between values when different researches are done.

a. Standard error for mean :

- For quantitative data.

$$SE_m = \frac{SD}{\sqrt{n}}$$

**Measures of
Central Tendency
and Variation**

b. Standard error for proportions :

- For qualitative data.

$$SE_p = \sqrt{\frac{PQ}{N}}$$

P : Prevalence.

Q : 100 - prevalence.

n : Sample size.

If p- value or Confidence interval is provided as input,
Standard error has to be calculated and **not** the
Standard deviation.

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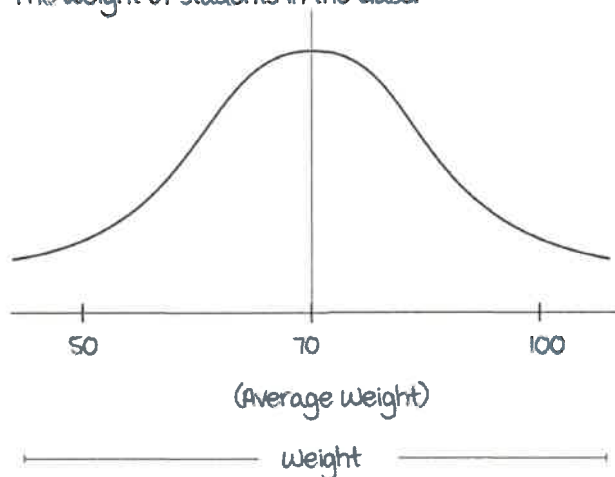
NORMAL DISTRIBUTION CURVE

Normal distribution curve

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It represents the distribution of data in a bell-shaped curve, in a large sample.

Eg : The weight of students in the class.



Features of Normal distribution curve :

It is also known as the Gaussian distribution curve.

It is a bilaterally symmetrical bell-shaped curve.

The ends never touch the baseline.

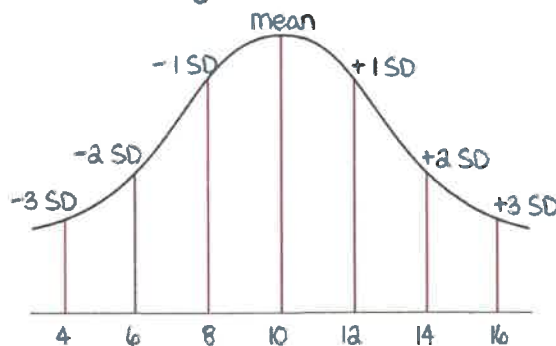
mean = median = mode $\rightarrow$ coincide at 0 or the centrepoint.

SD = 1.

AUC = 1 (Area Under Curve), means the whole population is accounted for.

Eg : mean Hb ($\bar{x}$ Hb) at a place = 10 gm% $\pm$ 2 g%.

where 1 SD = 2 g%



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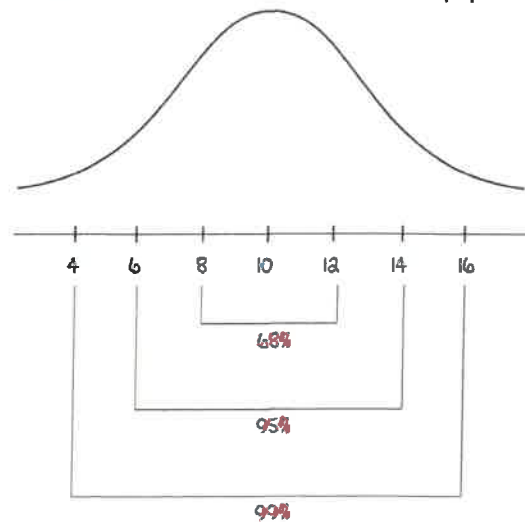
Assumptions in normal distribution curve :**First assumption**

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Between the -1 SD and $+1$ SD : 68% of the population lies.

Between the -2 SD and $+2$ SD : 95% of the population lies.

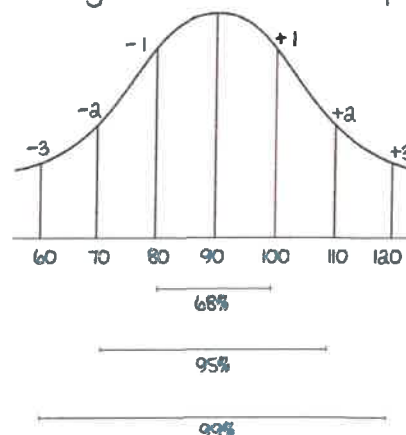
Between the -3 SD and $+3$ SD : 99% of the population lies.



Eg : mean blood glucose = 90 ± 10 SD.

How much of the population will be expected to fall between :

- 80 to 100 mg/dl = 68% population.
- 70 to 110 mg/dl = 95% population.
- 70 to 100 mg/dl = 68% + 13.5% population $[(95-68)/2=13.5]$
- more than 70 mg/dl = $100 - 2.5\% = 97.5\%$ population.
- Less than 100 mg/dl = 84% population $(100-13.5+2+0.5)$.
- more than 100 mg/dl = 16% population.
- Less than 60 mg/dl = $100 - 99 = 1/2 = 0.5\%$ population.
- Less than 120 mg/dl = $100 - 0.5\% = 99.5\%$ population.



Normal Distribution Curve

Q. The mean blood glucose from 5929 ANC females in the state of Maharashtra was found to be 130 ± 5 mg/dl. The cut off for diagnosing GDM was kept as higher than 140 mg/dl. How many pregnant females are expected to be GDM diagnosed?

- A. < 50. C. 100 to 200.
B. 50 to 100. D. 200 to 500.

mean = 130, +1 SD = 135, +2 SD = 140, +3 SD = 145

-1 SD = 125, -2 SD = 120, -3 SD = 115.

To be GDM diagnosed, they must belong to above +2 SD of population.

Above +2 SD = 100 - 95% (between +2 and -2 SD) - 2.5% (less than -2 SD) = 2.5%

2.5% of 5929 ~ 150 females, which falls under range of 100-200.

Second assumption : Zone of Normalcy

00:20:51

Zone of normalcy/normal zone :

Between the -2 SD and +2 SD = 95% of population.

Z score :

It is also called standard deviate.

It gives the location of the value in terms of the standard deviation (SD).

The cut off for Z score : ± 2 SD / ± 1.96 SD.

If the Z score is > 2 : Abnormal Z score.

It is calculated by = $\frac{\text{Observed value} - \text{Expected value}}{\text{SD}}$

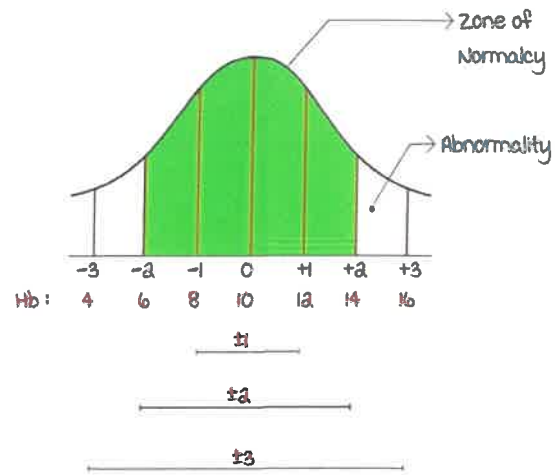
Eg : Observed value of Hb = 15 gm /dl.

Expected value (always the mean value) = 10

SD = 2

Z score = $\frac{15 - 10}{2} = 2.5$

Z score 2.5 : It lies 2.5 SD away from the mean.



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CONCEPT OF PROBABILITY VALUE

P value

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P value :

Probability value (chance of events expressed in decimals).

Normal value ranges from 0 to 1.

0 : Lowest probability.

1 : maximum probability.

Standard errors (SE) :

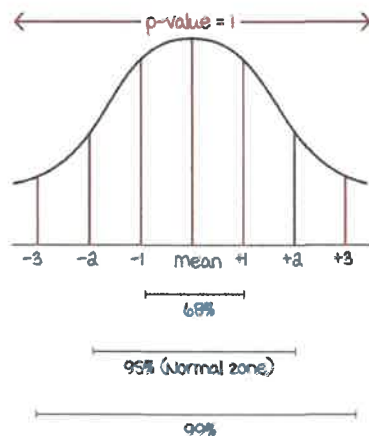
$\pm 1, \pm 2, \pm 3, \dots$

Confidence limit/interval :

± 1 to -1 = 68% confidence interval.

± 2 to -2 = 95% confidence interval.

± 3 to -3 = 99% confidence interval.



In the normal distribution curve :

The highest probability is towards the centre : 1.

The lowest probability lies on either side of the curve.

At ± 2 to -2 standard deviation the P value is : 0.05 -

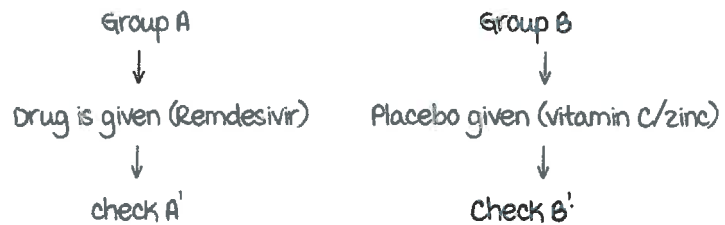
Zone of normalcy.

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P value – abnormal zone

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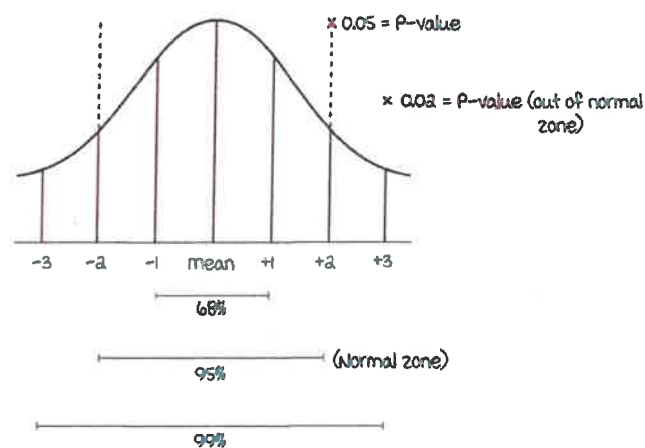
Example : Randomised clinical trial – two groups A and B



The collected data is incorporated in a machine : gives P value.

If the P value is 0.02 : Abnormal/out of the normal zone.

P value > 0.05	P value < 0.05
Normal variant	Abnormal variant
Non-significant	significant
No effect found	effect is found
Null hypothesis : Accepted	Null hypothesis : Rejected

**P value – normal zone and changes**

00:16:42

The normal zone for P value – 95% confidence interval

If the normal zone moved from 95% to 68% :

Previously non-significant becomes significant.
Chances of finding an effect increases.
The chances of reject of null hypothesis increases.
The chances of alpha error increases.

If the normal zone moves from 95% to 99% :

Previously significant becomes non-significant.
The chances of finding an effect decreases.
The chances of accepting of null hypothesis increases.
The chances of beta errors increase.

Alpha error, type I & II error

00:23:08

Definition :

It is the probability of finding an effect (just by chance)
which in reality does not exist.
It corresponds to the P value/confidence interval/limit.

Example : P value of 0.02 corresponds to α value 2%.

It means there is 2% chance of error in the study.
It also means there is 98% of confidence in the study.

68% corresponds to 32% alpha.

95% corresponds to 5% alpha.

99% corresponds to 1% alpha.

FFER : The chance of finding disease in a healthy patient.

Type I error :

Rejecting a null hypothesis, which in reality is true.

Type II error :

Accepting a null hypothesis, which is false in reality.

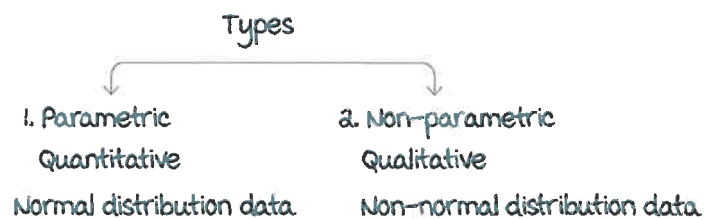
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TESTS OF SIGNIFICANCE

Statistical mathematical formula to derive a p-value.
Determines if P-value is significant or non significant.

Types of tests of significance

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Parametric test	Situation	Non-parametric test
Paired 't' test.	Single group	mc nernmar's test.
Unpaired 't' test A/M/A Independent sample 't' test.	Two groups	Chi square test (χ^2).
Analysis of variance (ANOVA)	Three or more groups	Kruskal-wallis test. Chi square for trend.

Advance tests of significance

00:08:59

- Large sample ($n > 30$) = 'z' test.
- Ordinal data : wilcoxon rank test (w/r)



- Normalcy of data : Kolmogorov smirnov test.
- Outliers : Dixon's Q test.
- Internal consistency of questionnaire : Cronbach's α score
- Compare a new test with a gold standard test : Bland altman analysis.

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- Level of agreement : KAPPA test.

Formula =
$$\frac{\text{Observed level of agreement} - \text{expected level of agreement}}{1 - \text{expected level of agreement}}$$

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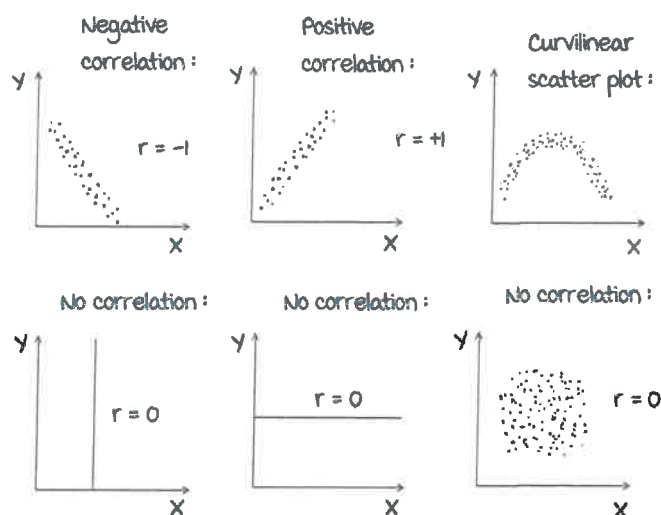
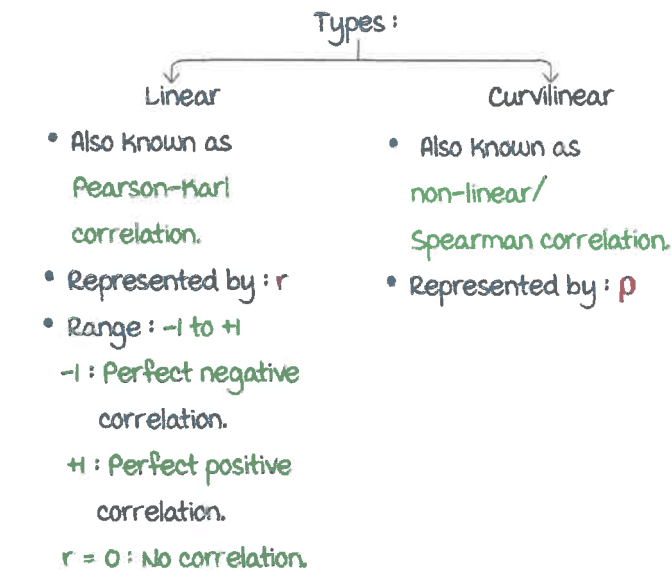
CORRELATION, REGRESSION AND SKEW

Correlation

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Relation between 2 variables.

Scatter plots are used.

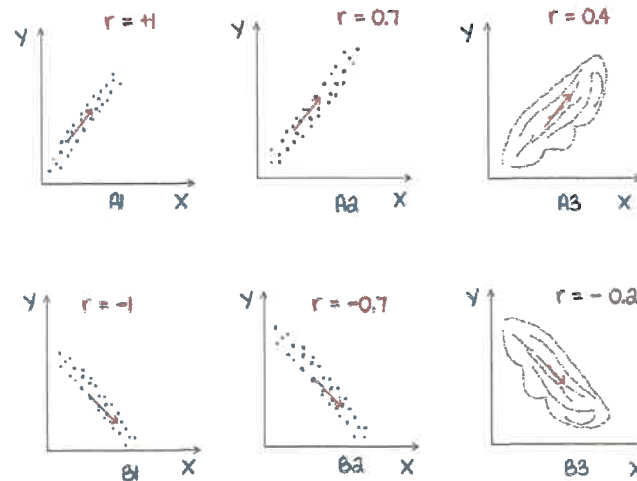


Scatter plots

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Coefficient of determination

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$+1$: Perfect positive correlation (1 unit change in X axis = 1 unit change in Y axis).

> 0.7 : Strong positive correlation.

$0.5 - 0.7$: moderately positive correlation.

< 0.5 : Weak correlation.

< 0.3 : Very weak correlation.

Coefficient of determination (CD) :

The percentage change in one variable which is accounted for by a unit change in another variable.

$$CD = r^2 \text{ in \%}$$

Regression

00:18:26

Primarily refers to prediction.

Types :

1. Linear : If variables are quantitative.

2. Logistic : If variables are qualitative.

1. univariate linear regression :

Eg : Predicting renal failure based on GFR.

2. univariate logistic regression :

Eg : Predicting MI based on obesity levels.

3. multivariate linear regression :

Eg : Predicting the renal status based on serum Na, urea, creatinine and GFR levels.