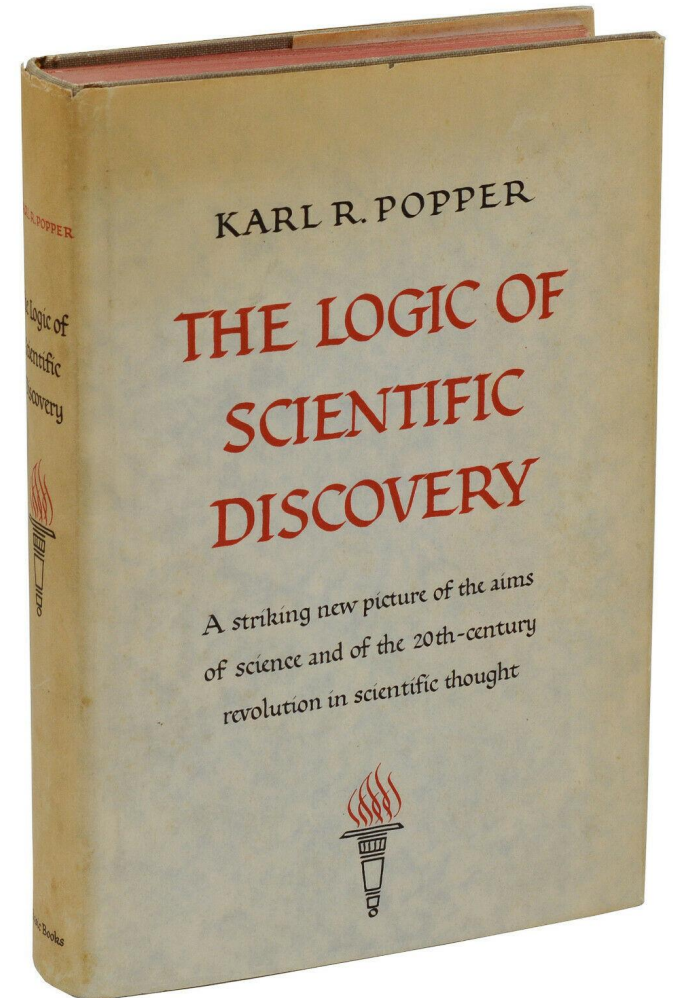


My proposal is based upon an *asymmetry* between **verifiability** and **falsifiability**; an asymmetry which results from the logical form of universal statements. For these are never derivable from singular statements, but can be contradicted by singular statements.

—Karl Popper, *The Logic of Scientific Discovery*

What is science?

a scientific theory must be
falsifiable



Reproducibility

Reproducible research means:

the ability of a researcher to duplicate the results of a prior study using the same materials as were used by the original investigator. That is, a second researcher might use the same raw data to build the same analysis files and implement the same statistical analysis in an attempt to yield the same results

<https://acmedsci.ac.uk/viewFile/56314e40aac61.pdf>

Reproducible research in practice:
all numbers in a data analysis
can be recalculated exactly
(down to stochastic variables!)
using the **code** and **raw data** provided by
the analyst.

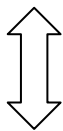
- provide raw data and code to reduce it to all stages needed to get outputs
- provide code to reproduce all figures
- provide code to reproduce all number outcomes



Probability

Frequentist interpretation

fraction of times something happens



probability of it happening

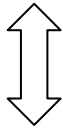
Bayesian interpretation

represents a level of certainty relating to
a potential outcome or idea:

*if I believe the coin is unfair (tricked) then
even if I get a head and a tail I will still
believe I am more likely to get heads
than tails*

Frequentist interpretation

fraction of times something happens



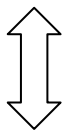
probability of it happening

$$P(E) = \text{frequency of } E$$
$$P(\text{coin} = \text{head}) = 6/11 = 0.55$$



Frequentist interpretation

fraction of times something happens



probability of it happening

$P(E) = \text{frequency of } E$

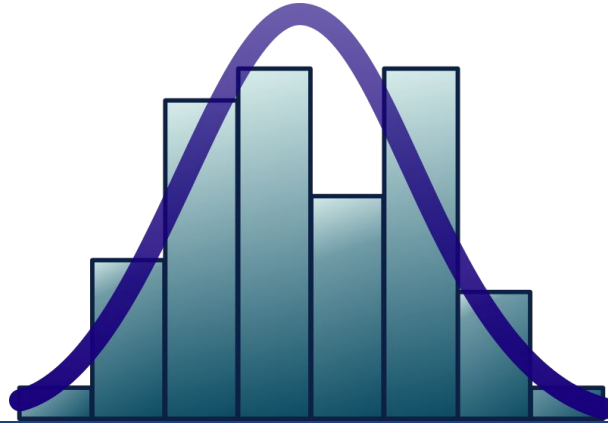
$P(\text{coin} = \text{head}) = 6/11 = 0.55$

$P(\text{coin} = \text{head}) = 49/100 = 0.49$



Summary of probabilities

Event	Probability
A	$P(A) \in [0, 1]$
not A	$P(A^c) = 1 - P(A)$
A or B	$P(A \cup B) = P(A) + P(B) - P(A \cap B)$ $P(A \cup B) = P(A) + P(B)$ if A and B are mutually exclusive
A and B	$P(A \cap B) = P(A B)P(B) = P(B A)P(A)$ $P(A \cap B) = P(A)P(B)$ if A and B are independent
A given B	$P(A B) = \frac{P(A \cap B)}{P(B)} = \frac{P(B A)P(A)}{P(B)}$



Statistics

Statistics

takes us from observing a limited
number of samples to infer on the
population

Taxonomy

Distribution: a formula (a model)

Population: all of the elements of a "family"

Sample: a finite subset of the population that you observe

Descriptive statistics - describing the central tendency

mean: (sum of all the terms)/(number of terms)

median: 50% of the distribution is to the left,
50% to the right

mode: most popular value in the distribution

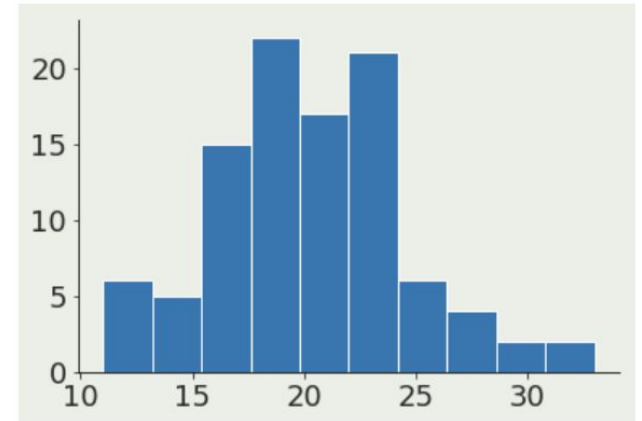
```
dist = sp.stats.poisson.rvs(size=100, mu=20)
pl.hist(dist)
print(dist.mean())
print(np.median(dist))
print(sp.stats.mode(dist))
|
```

executed in 125ms, finished 15:01:20 2019-09-09

20.06

20.0

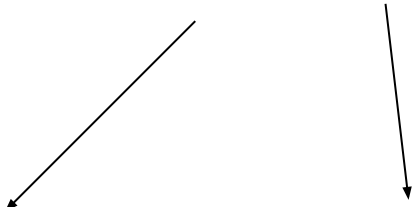
ModeResult(mode=array([18]), count=array([12]))



Taxonomy

Central tendency: mean, median, mode

Spread: variance, interquartile range

Two arrows originate from the text 'Central tendency: mean, median, mode'. One arrow points diagonally down and to the left towards the word 'variance' in the 'Spread' line. The other arrow points diagonally down and to the right towards the words 'interquartile range' in the 'Spread' line.

Descriptive statistics - measuring the spread

Variance

$$\text{Var}(X) = \text{E}[(X - \mu)^2]$$

Standard deviation

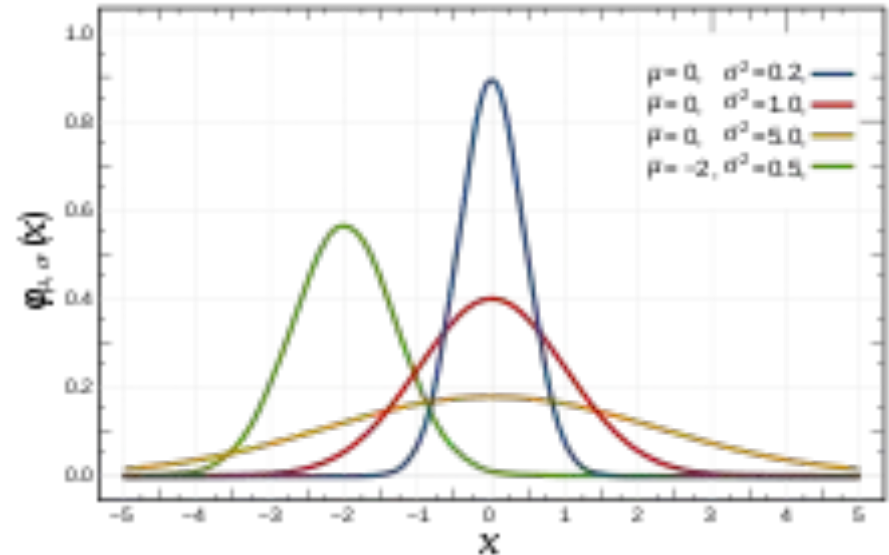
$$\sigma(X) = \sqrt{\text{E}[(X - \text{E}[X])^2]}$$

Gaussian distribution:

1σ contains 68% of the distribution

2σ contains 95% of the distribution

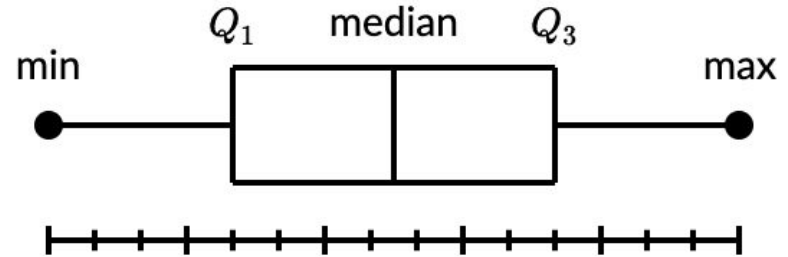
3σ contains 97.3% of the distribution



Descriptive statistics - measuring the spread

Interquartile range

Where are the limits within which X% of the distribution is contained



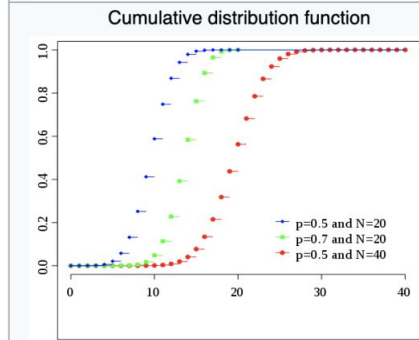
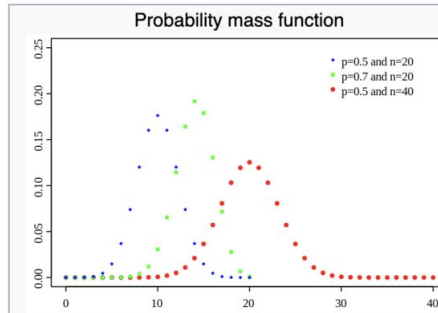
Probability distributions

Binomial

Coin toss

Fair coin = $p=0.5$ $n=1$

Binomial distribution



Notation	$B(n, p)$
Parameters	$n \in \{0, 1, 2, \dots\}$ – number of trials $p \in [0, 1]$ – success probability for each trial
Support	$k \in \{0, 1, \dots, n\}$ – number of successes
pmf	$\binom{n}{k} p^k (1-p)^{n-k}$
CDF	$I_{1-p}(n-k, 1+k)$
Mean	np
Median	$\lfloor np \rfloor$ or $\lceil np \rceil$
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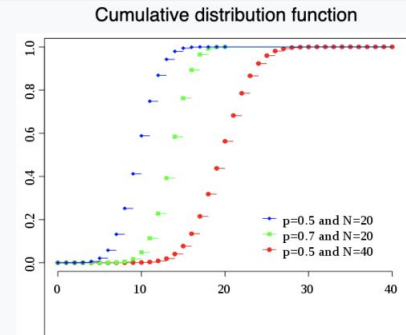
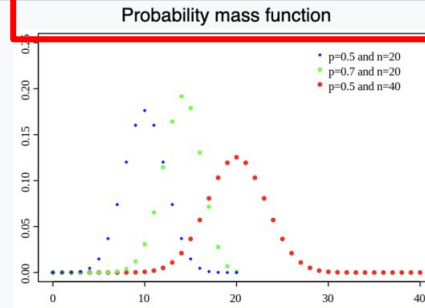
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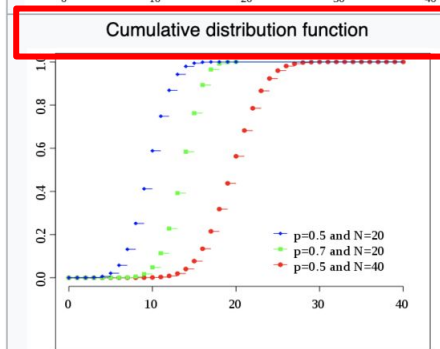
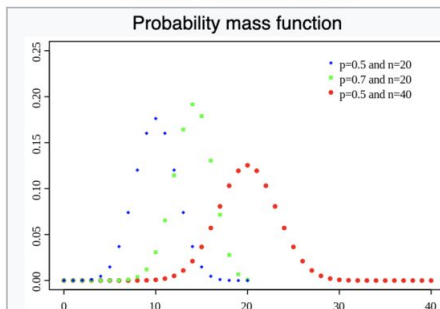
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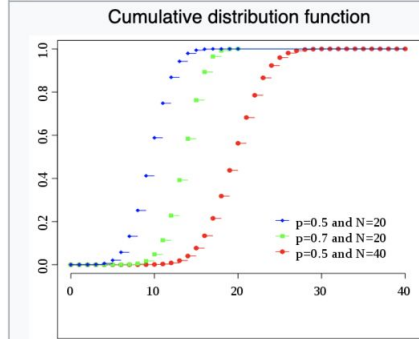
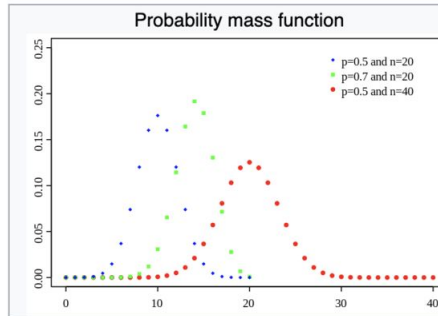
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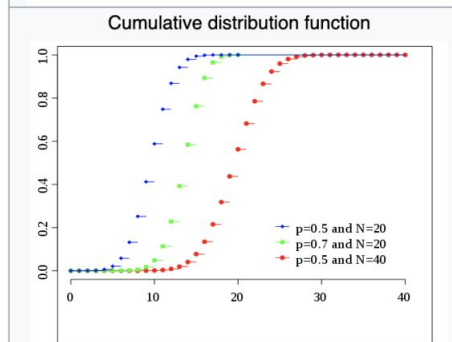
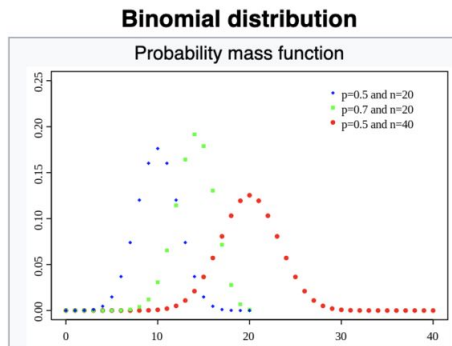
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Probability distributions

Binomial

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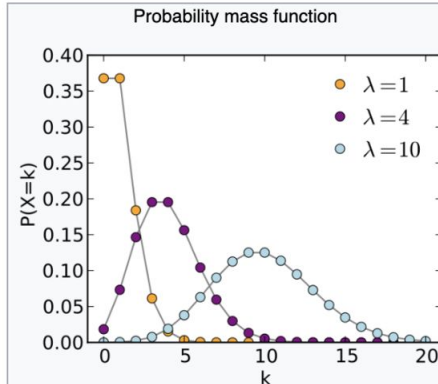
Probability distributions

Poisson

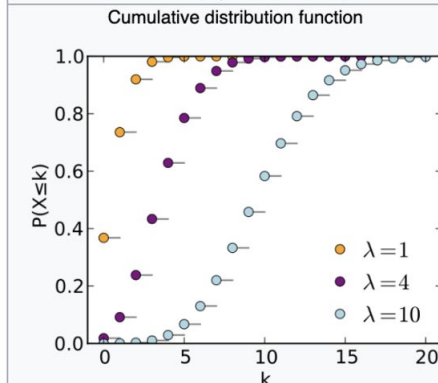
Shot noise/count noise

The innate noise in natural steady state processes (star flux, rain drops, radioactive decay...)

Poisson



The horizontal axis is the index k , the number of occurrences. λ is the expected number of occurrences, which need not be an integer. The vertical axis is the probability of k occurrences given λ . The function is defined only at integer values of k . The connecting lines are only guides for the eye.



The horizontal axis is the index k , the number of occurrences. The CDF is discontinuous at the integers of k and flat everywhere else because a variable that is Poisson distributed takes on only integer values.

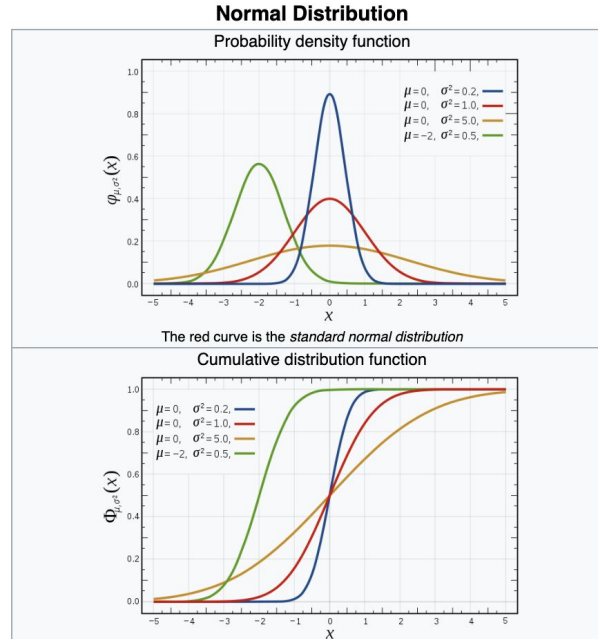
Notation	$\text{Pois}(\lambda)$
Parameters	$\lambda > 0$, (real) — rate
Support	$k \in \{0, 1, 2, \dots\}$
pmf	$\frac{\lambda^k e^{-\lambda}}{k!}$
CDF	$\frac{\Gamma([k+1], \lambda)}{[k]!}$, or $e^{-\lambda} \sum_{i=0}^{[k]} \frac{\lambda^i}{i!}$, or $Q([k+1], \lambda)$ (for $k \geq 0$, where $\Gamma(x, y)$ is the upper incomplete gamma function , $[k]$ is the floor function , and Q is the regularized gamma function)
Mean	λ
Median	$\approx \lfloor \lambda + 1/3 - 0.02/\lambda \rfloor$
Mode	$\lfloor \lambda \rfloor - 1, \lfloor \lambda \rfloor$
Variance	λ
Skewness	$\lambda^{-1/2}$
Ex. kurtosis	λ^{-1}
Entropy	$\lambda[1 - \log(\lambda)] + e^{-\lambda} \sum_{k=0}^{\infty} \frac{\lambda^k \log(k!)}{k!}$ (for large λ) $\frac{1}{2} \log(2\pi e \lambda) - \frac{1}{12\lambda} - \frac{1}{24\lambda^2} - \frac{19}{360\lambda^3} + O\left(\frac{1}{\lambda^4}\right)$
MGF	$\exp(\lambda(e^t - 1))$
CF	$\exp(\lambda(e^{it} - 1))$
PGF	$\exp(\lambda(z - 1))$
Fisher information	$\frac{1}{\lambda}$

Probability distributions

Gaussian

most common noise:

well behaved
mathematically,
symmetric, when we
can we will assume our
uncertainties are
Gaussian distributed



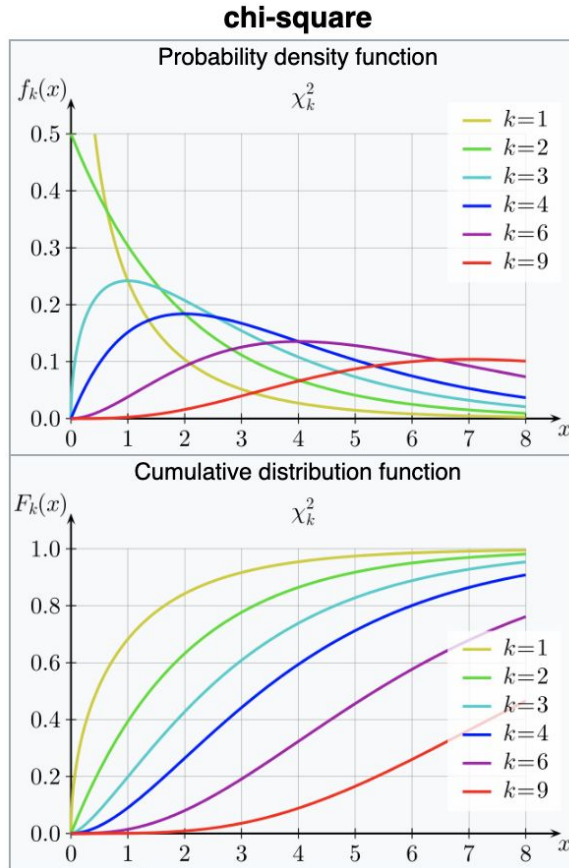
Notation	$\mathcal{N}(\mu, \sigma^2)$
Parameters	$\mu \in \mathbb{R}$ = mean (location) $\sigma^2 > 0$ = variance (squared scale)
Support	$x \in \mathbb{R}$
PDF	$\frac{1}{\sqrt{2\pi\sigma^2}} e^{-\frac{(x-\mu)^2}{2\sigma^2}}$
CDF	$\frac{1}{2} \left[1 + \operatorname{erf}\left(\frac{x-\mu}{\sigma\sqrt{2}}\right) \right]$
Quantile	$\mu + \sigma\sqrt{2} \operatorname{erf}^{-1}(2F - 1)$
Mean	μ
Median	μ
Mode	μ
Variance	σ^2
Skewness	0
Ex. kurtosis	0
Entropy	$\frac{1}{2} \log(2\pi e \sigma^2)$
MGF	$\exp(\mu t + \sigma^2 t^2 / 2)$
CF	$\exp(i\mu t - \sigma^2 t^2 / 2)$
Fisher information	$\mathcal{I}(\mu, \sigma) = \begin{pmatrix} 1/\sigma^2 & 0 \\ 0 & 2/\sigma^2 \end{pmatrix} \mathcal{I}(\mu, \sigma^2) = \begin{pmatrix} 1/\sigma^2 & 0 \\ 0 & 1/(2\sigma^4) \end{pmatrix}$
Kullback-Leibler divergence	$D_{\text{KL}}(\mathcal{N}_0 \ \mathcal{N}_1) = \frac{1}{2} \{ (\sigma_0/\sigma_1)^2 + \frac{(\mu_1 - \mu_0)^2}{\sigma_1^2} - 1 + 2 \ln \frac{\sigma_1}{\sigma_0} \}$

Probability distributions

Chi-square (X2)

turns out its extremely common

many pivotal quantities follow this distribution and thus many tests are based on this



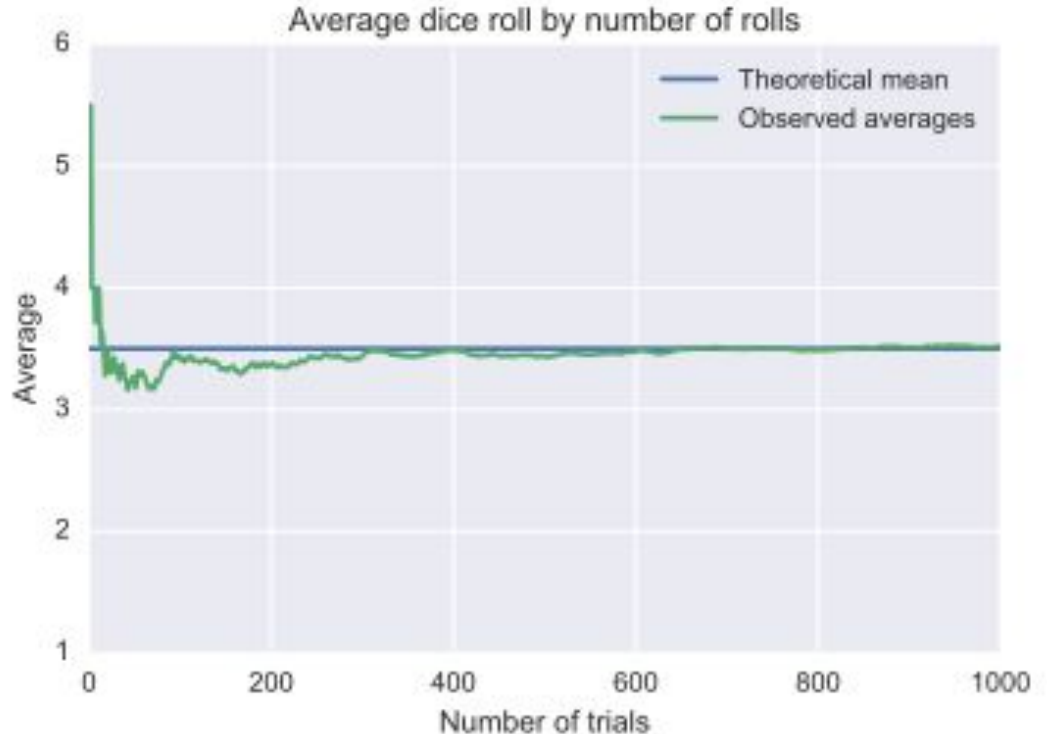
Notation	$\chi^2(k)$ or χ_k^2
Parameters	$k \in \mathbb{N}_{>0}$ (known as "degrees of freedom")
Support	$x \in (0, +\infty)$ if $k = 1$, otherwise $x \in [0, +\infty)$
PDF	$\frac{1}{2^{k/2} \Gamma(k/2)} x^{k/2-1} e^{-x/2}$
CDF	$\frac{1}{\Gamma(k/2)} \gamma\left(\frac{k}{2}, \frac{x}{2}\right)$
Mean	k
Median	$\approx k \left(1 - \frac{2}{9k}\right)^3$
Mode	$\max(k - 2, 0)$
Variance	$2k$
Skewness	$\sqrt{8/k}$
Ex. kurtosis	$\frac{12}{k}$
Entropy	$\frac{k}{2} + \ln(2\Gamma(\frac{k}{2})) + (1 - \frac{k}{2})\psi(\frac{k}{2})$
MGF	$(1 - 2t)^{-k/2}$ for $t < \frac{1}{2}$
CF	$(1 - 2it)^{-k/2}$ [1]
PGF	$(1 - 2 \ln t)^{-k/2}$ for $0 < t < \sqrt{e}$

Simulation of the different distributions in Colab

Law of large numbers

According to the law, the **average** of the results obtained from a large number of trials should be close to the **expected value** and will tend to become closer to the expected value as more trials are performed.

In the limit of $N \rightarrow \infty$ the mean of a sample of size N approaches the mean of the population μ

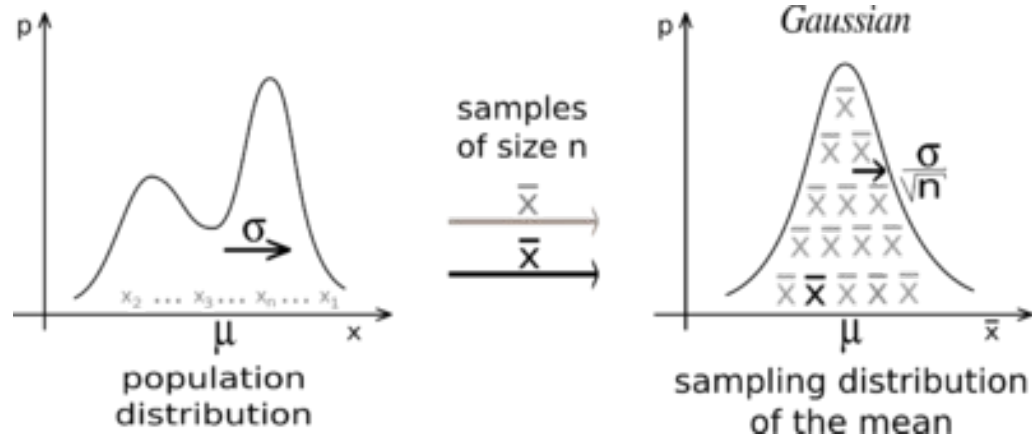


Central limit theorem

In the limit of $N \rightarrow \infty$

the sample mean \bar{x}
approaches a Normal
(Gaussian) distribution with
mean μ and standard
deviation σ

regardless of the distribution
of X



Central limit theorem in Colab

Inference

“Body weight was higher in mice fed the high-fat diet already after the first week, due to higher dietary intake in combination with lower metabolic efficiency.”

“Already during the first week after introduction of high-fat diet, body weight increased significantly more in the high-fat diet-fed mice ($+ 1.6 \pm 0.1$ g) than in the normal diet-fed mice ($+ 0.2 \pm 0.1$ g; $P < 0.001$).”

**What is $+ 1.6 \pm 0.1$ g and $+ 0.2 \pm 0.1$ g and what is $P < 0.001$?
And why are these numbers important?**

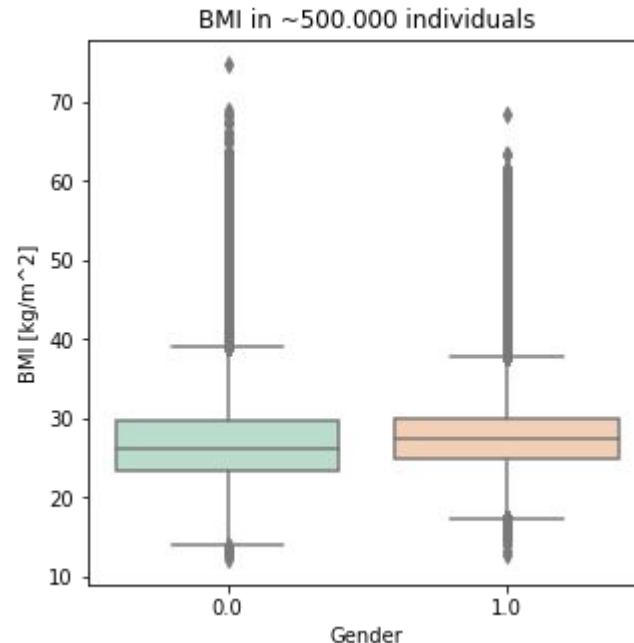
Data has variability

We need p-values and confidence intervals because the data has variability.

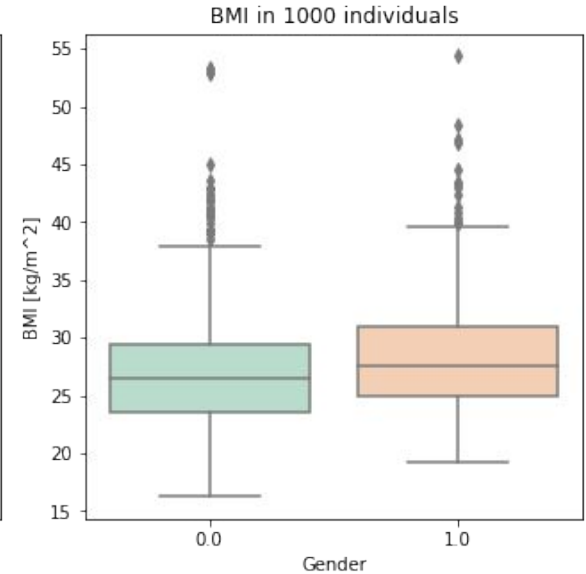
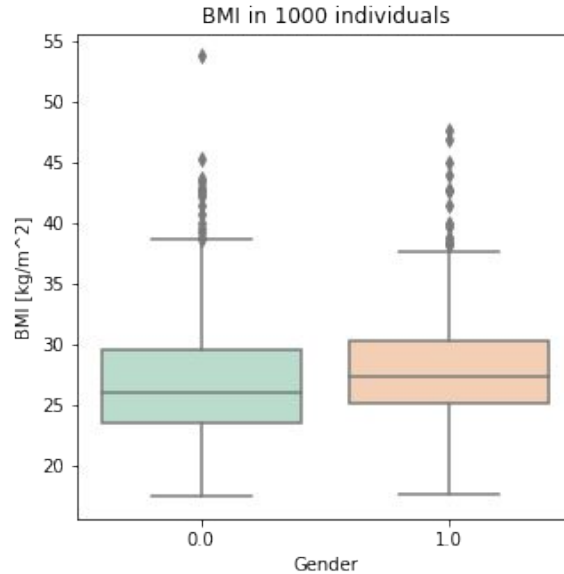
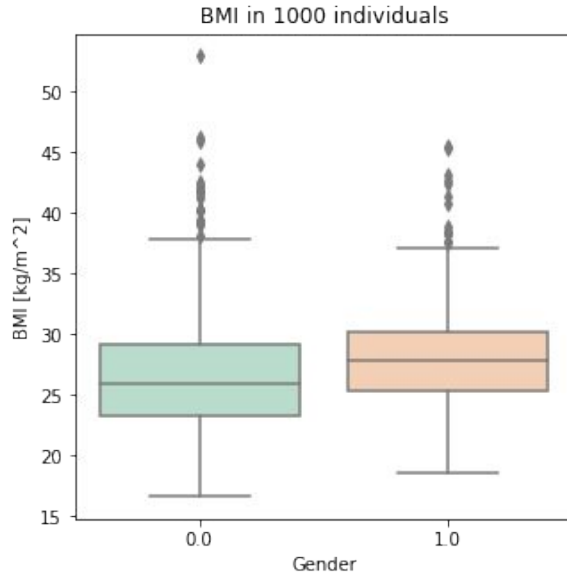
We do not have access to the **population**, we are just sampling it and inferring the characteristics of the population based on the values of its sample.

Every time we resample the population, we get different descriptive statistics (mean, standard deviation, etc).

In particular, the means is a random variable.



Random variables when sampling a population



mean males: 28.065288965238878
mean females: 26.76744359319031

mean males: 27.9795995248824
mean females: 26.92432748169731

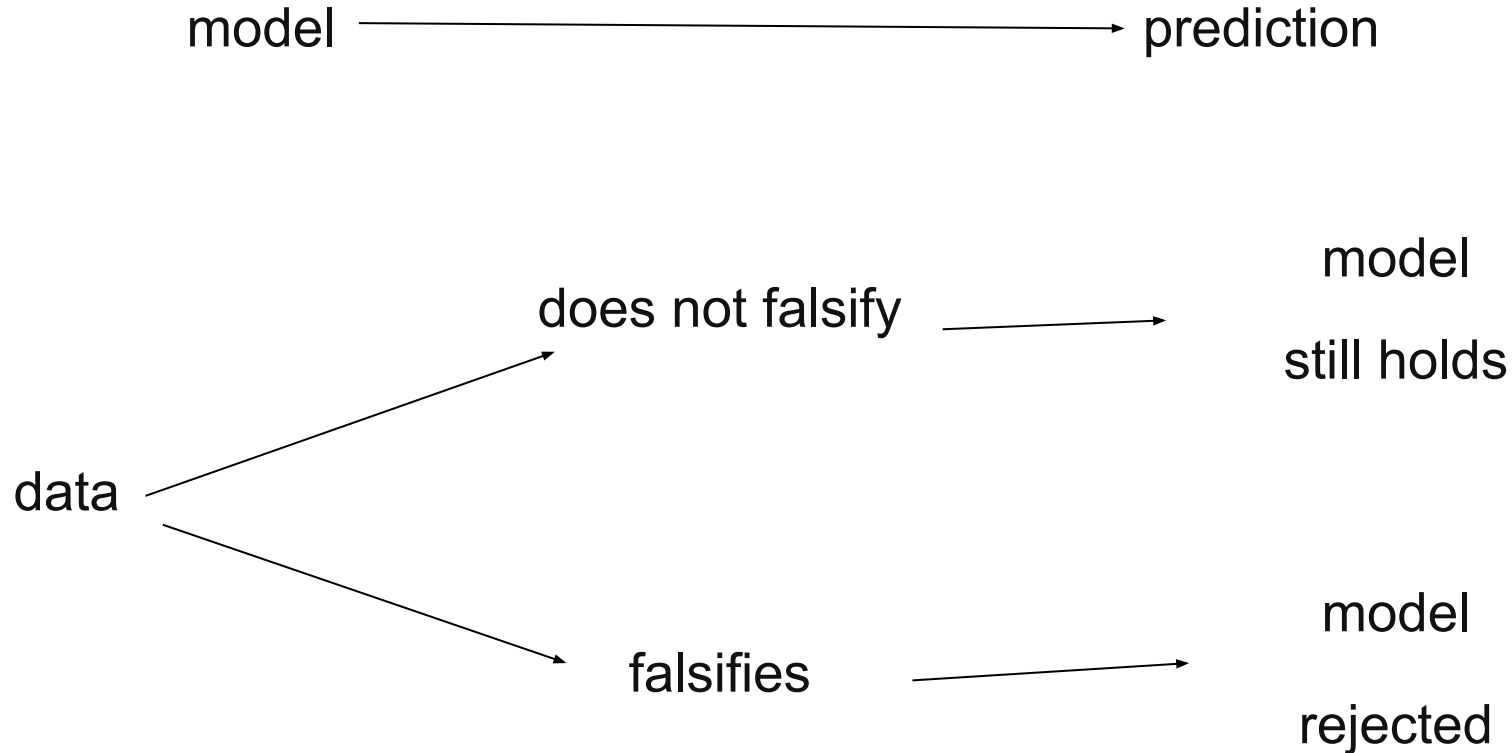
mean males: 28.379904688459526
mean females: 27.187517814945377

The Null Hypothesis

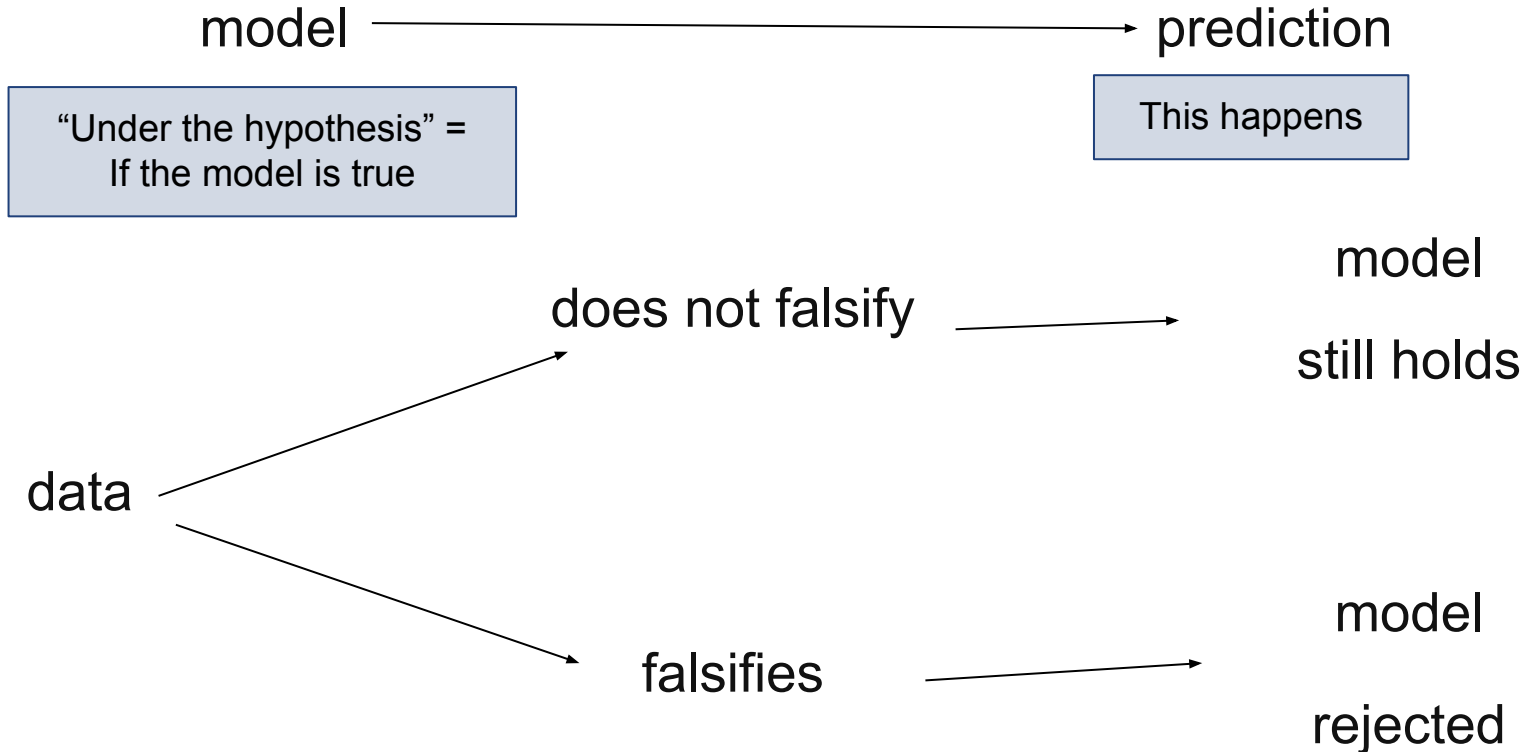
How do we know that males and females have really different BMI, and that is not due to a random effect of the sampling?

Null Hypothesis Rejection Testing

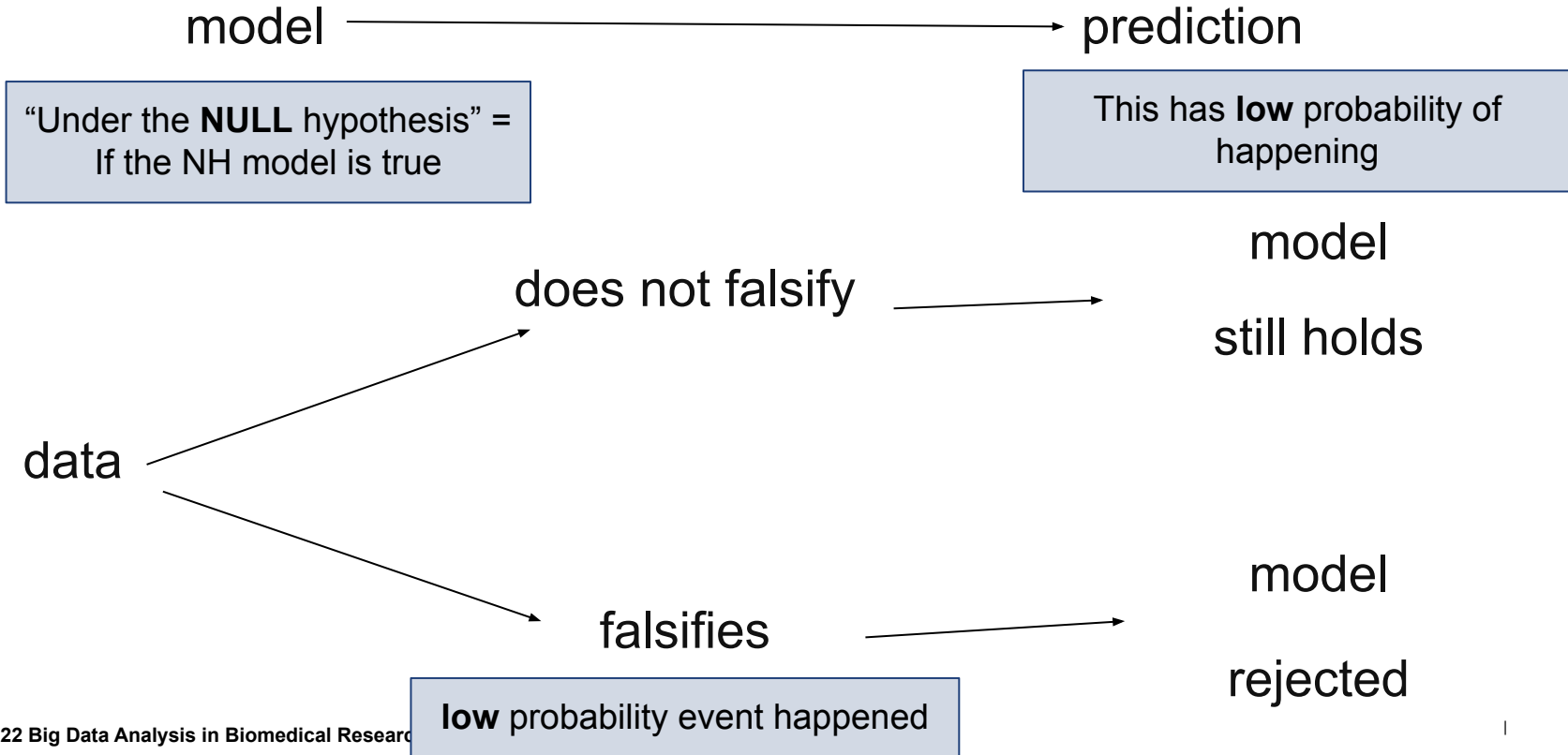
Null Hypothesis Rejection Testing



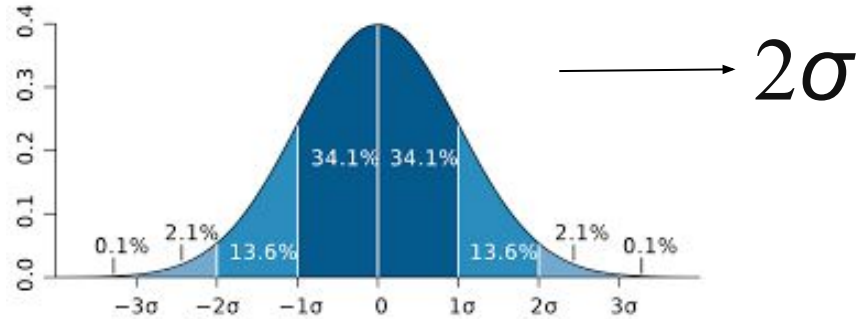
Null Hypothesis Rejection Testing



Null Hypothesis Rejection Testing



Null Hypothesis Rejection Testing



rejected at 95%

0.05 p-value

5% confidence

does not falsify

model
still holds

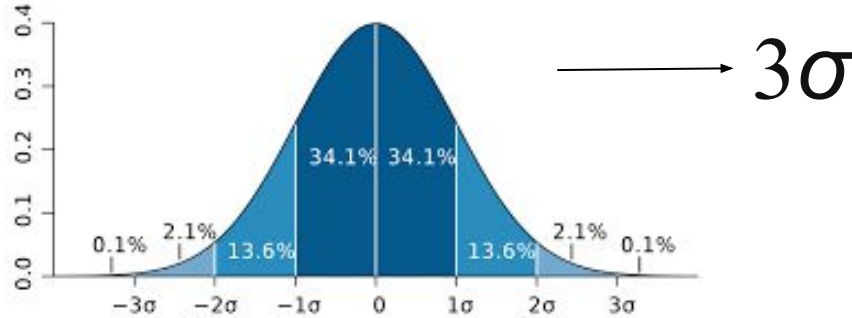
data

falsifies

model
rejected

low probability event happened

Null Hypothesis Rejection Testing



rejected at 99.7%

0.003 p-value

0.3% confidence

does not falsify

model
still holds

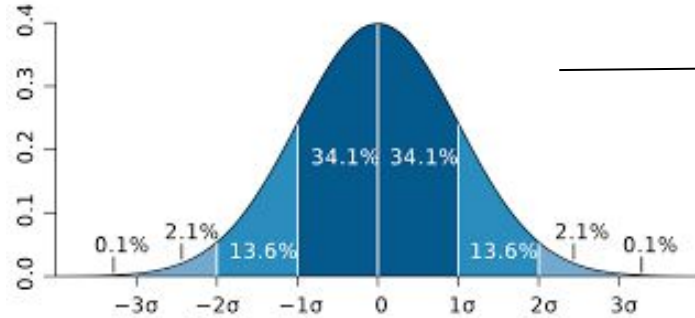
data

falsifies

model
rejected

low probability event happened

Null Hypothesis Rejection Testing



rejected at 99.99..%

3-e7 p-value

3-e5% confidence

5σ

does not falsify

model
still holds

data

falsifies

model
rejected

low probability event happened

Null Hypothesis Rejection Testing

formulate the Null as the comprehensive opposite of your theory

model → prediction

“Under the **NULL** hypothesis” =
If the proposed model is
FALSE

This has **low** probability of
happening

does not falsify
alternative

model
rejected

data

Falsifies
alternative

model
still holds

low probability event happened

Example

Earth is flat is Null

not flat

Earth is ~~round~~ is Alternative

We reject the Null hypothesis that the Earth is flat
($p=0.05$)

Null Hypothesis Rejection Testing

1. Formulate your prediction (Null Hypothesis)
2. Identify all alternative outcomes (Alternative hypothesis):
 - a. If all alternatives to our model are ruled out, then our model must hold
3. Set confidence threshold (2sigma, 0.05 p-value, 95% alpha threshold)
4. Find a measurable quantity which under the Null has a known distribution:
 - a. If a quantity follows a known distribution, once I measure its value I can know what the probability of getting that value actually is.
 - b. Different statistics:
 - i. X2 statistics: difference between expectation and reality squared
 - ii. Z statistics: difference between means
 - iii. K-S statistics: maximum distance of cumulative distributions
5. Calculate the pivotal quantities
6. Calculate the probability of value obtained for the pivotal quantity under the Null

If probability < p-value: reject Null

Common tests and pivotal quantities



Z-test

Is the mean of a sample with known variance the same as that of a known population?

Pivotal quantity

$$Z = \frac{(\bar{X} - \mu_0)}{s}$$

Sample mean Population mean Sample variance

$$Z \sim N(\mu=0, \sigma=1)$$

Z-test

Is the mean of a sample with known variance the same as that of a known population?

Pivotal quantity

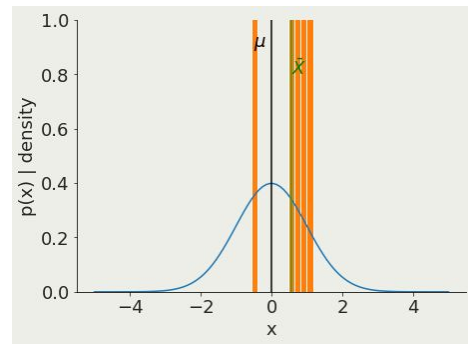
$$Z = \frac{(\bar{X} - \mu_0)}{s}$$

Sample variance

Sample mean

Population mean

$$Z \sim N(\mu=0, \sigma=1)$$



why do we need a test? why not just measuring the means and seeing if they are the same?

Z-test

Is the mean of a sample with known variance the same as that of a known population?

Pivotal quantity

$$Z = \frac{(\bar{X} - \mu_0)}{s}$$

Sample variance

Sample mean

Population mean

$$Z \sim N(\mu=0, \sigma=1)$$

The Z test provides a trivial interpretation of the measured quantity: the Z value is exactly the distance for the mean of the standard distribution of possible outcomes in units of standard deviation

so a result of 0.13 means we are 0.13 standard deviations to the mean ($p > 0.05$)

t-test

Are the means of 2 samples significantly different?

Pivotal quantity

$$t = \frac{\bar{X}_1 - \bar{X}_2}{s_{\bar{\Delta}}}$$

unbiased
variance
estimator

$$s_{\bar{\Delta}} = \sqrt{\frac{s_1^2}{n_1} + \frac{s_2^2}{n_2}}$$

size of
sample

$$t \sim \text{Student's } t \left(\text{d. f.} = \frac{\left(\frac{s_1^2}{n_1} + \frac{s_2^2}{n_2} \right)^2}{\frac{(s_1^2/n_1)^2}{n_1-1} + \frac{(s_2^2/n_2)^2}{n_2-1}} \right)$$

t-test

Are the means of 2 samples significantly different?

Pivotal quantity

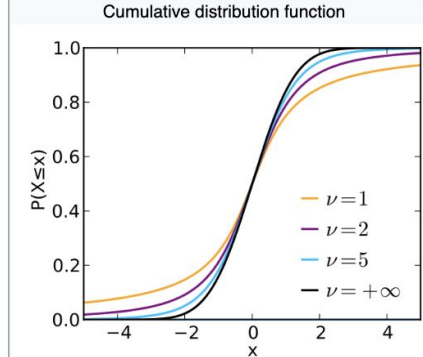
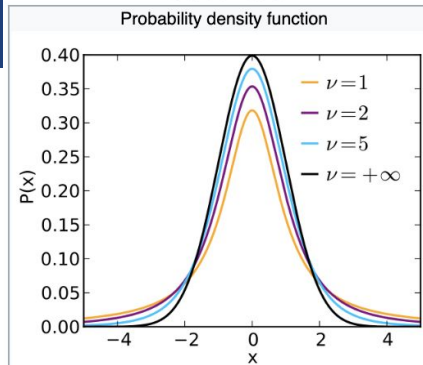
$$t = \frac{\bar{X}_1 - \bar{X}_2}{s_{\bar{\Delta}}}$$

unbias
variance
estimator

$$s_{\bar{\Delta}} = \sqrt{\frac{s_1^2}{n_1} + \frac{s_2^2}{n_2}}$$

size of
sample

Student's t



Parameters	$\nu > 0$ degrees of freedom (real)
Support	$x \in (-\infty, \infty)$
PDF	$\frac{\Gamma(\frac{\nu+1}{2})}{\sqrt{\nu\pi} \Gamma(\frac{\nu}{2})} \left(1 + \frac{x^2}{\nu}\right)^{-\frac{\nu+1}{2}}$
CDF	$\frac{1}{2} + x\Gamma\left(\frac{\nu+1}{2}\right) \times \frac{{}_2F_1\left(\frac{1}{2}, \frac{\nu+1}{2}; \frac{3}{2}; -\frac{x^2}{\nu}\right)}{\sqrt{\pi\nu} \Gamma(\frac{\nu}{2})}$ where ${}_2F_1$ is the hypergeometric function
Mean	0 for $\nu > 1$, otherwise undefined

Parameters	$\nu > 0$ degrees of freedom (real)
Support	$x \in (-\infty, \infty)$
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Mean	0 for $\nu > 1$, otherwise undefined
Median	0
Mode	0
Variance	$\frac{\nu}{\nu-2}$ for $\nu > 2$, ∞ for $1 < \nu \leq 2$, otherwise undefined

t-test

Are the means of 2 samples significantly different?

Pivotal quantity

$$t = \frac{\bar{X}_1 - \bar{X}_2}{s_{\bar{\Delta}}}$$

unbias
variance
estimator

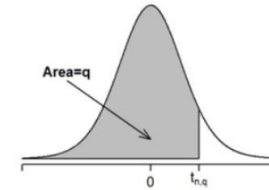
$$s_{\bar{\Delta}} = \sqrt{\frac{s_1^2}{n_1} + \frac{s_2^2}{n_2}}$$

size
sample

To interpret the outcome of a t-test I have to figure out the probability of a given p

Quartiles of the t Distribution

The table gives the value if $t_{n,q}$ - the q th quantile of the t distribution for n degrees of freedom



	$q = 0.6$	0.75	0.9	0.95	0.975	0.99	0.995	0.9975	0.999	0.9995
$n = 1$	0.3249	1.0000	3.078	6.314	12.706	31.821	63.657	127.321	318.309	636.619
2	0.2887	0.8165	1.886	2.920	4.303	6.965	9.925	14.089	22.327	31.599
3	0.2767	0.7649	1.638	2.353	3.182	4.541	5.841	7.453	10.215	12.924
4	0.2707	0.7407	1.533	2.132	2.776	3.747	4.604	5.598	7.173	8.610
5	0.2672	0.7267	1.476	2.015	2.571	3.365	4.032	4.773	5.893	6.869
6	0.2648	0.7176	1.440	1.943	2.447	3.143	3.707	4.317	5.208	5.959
7	0.2632	0.7111	1.415	1.895	2.365	2.998	3.499	4.029	4.785	5.408
8	0.2619	0.7064	1.397	1.860	2.306	2.896	3.355	3.833	4.501	5.041
9	0.2610	0.7027	1.383	1.833	2.262	2.821	3.250	3.690	4.297	4.781
10	0.2602	0.6998	1.372	1.812	2.228	2.764	3.169	3.581	4.144	4.587
11	0.2596	0.6974	1.363	1.796	2.201	2.718	3.106	3.497	4.025	4.437
12	0.2590	0.6955	1.356	1.782	2.179	2.681	3.055	3.428	3.930	4.318
13	0.2586	0.6938	1.350	1.771	2.160	2.650	3.012	3.372	3.852	4.221
14	0.2582	0.6924	1.345	1.761	2.145	2.624	2.977	3.326	3.787	4.140

χ^2 test

are the data what is expected from the model - there are a few χ^2 tests. The one here is the "Pearson's χ^2 tests"

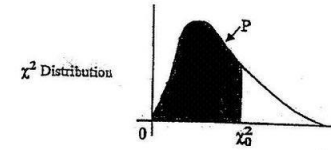
Pivotal quantity

$$\chi^2 \equiv \sum_i \frac{(f(x_i) - y_i)^2}{\sigma_i^2}$$

model \swarrow σ_i^2 \searrow observation
 uncertainty

$$\chi^2 \sim \chi^2(df = n - 1)$$

N of observations \nearrow
 N parameters model \searrow



The table below gives the value χ_0^2 for which $P[\chi^2 < \chi_0^2] = P$ for a given number of degrees of freedom and a given value of P.

Degrees of Freedom	Values of P									
	0.005	0.010	0.025	0.050	0.100	0.900	0.950	0.975	0.990	0.995
1	---	---	0.001	0.004	0.016	2.706	3.841	5.024	6.635	7.879
2	0.01	0.020	0.051	0.103	0.211	4.605	5.991	7.378	9.210	10.597
3	0.072	0.115	0.216	0.352	0.584	6.251	7.815	9.348	11.345	12.838
4	0.207	0.297	0.484	0.711	1.064	7.779	9.488	11.143	13.277	14.860
5	0.412	0.554	0.831	1.145	1.610	9.236	11.070	12.833	15.086	16.750
6	0.676	0.872	1.237	1.635	2.204	10.645	12.592	14.449	16.812	18.548
7	0.989	1.239	1.690	2.167	2.833	12.017	14.067	16.013	18.475	20.278
8	1.344	1.646	2.180	2.733	3.490	13.362	15.507	17.535	20.090	21.955
9	1.735	2.088	2.700	3.325	4.168	14.684	16.919	19.023	21.666	23.589
10	2.156	2.558	3.247	3.940	4.865	15.987	18.307	20.483	23.209	25.188
11	2.603	3.053	3.816	4.575	5.578	17.275	19.675	21.920	24.725	26.757
12	3.074	3.571	4.404	5.226	6.304	18.549	21.026	23.337	26.217	28.300
13	3.565	4.107	5.009	5.892	7.042	19.812	22.362	24.736	27.688	29.819
14	4.075	4.660	5.629	6.571	7.790	21.064	23.685	26.119	29.141	31.319
15	4.601	5.229	6.262	7.261	8.547	22.307	24.996	27.488	30.578	32.801
16	5.142	5.812	6.908	7.962	9.312	23.542	26.296	28.845	32.000	34.267
17	5.697	6.408	7.564	8.672	10.085	24.769	27.587	30.191	33.409	35.718
18	6.265	7.015	8.231	9.390	10.865	25.989	28.869	31.526	34.805	37.156
19	6.844	7.633	8.907	10.117	11.651	27.204	30.144	32.852	36.191	38.582
20	7.434	8.260	9.591	10.851	12.443	28.412	31.410	34.170	37.566	39.997

Kolmogorov–Smirnov test

Kolmogorov-Smirnov test:

do two samples come from the same parent distribution?

Pivotal quantity

$$D_n = \sup_x |F_n(x) - F(x)|$$

Cumulative
distribution 1

Cumulative
distribution 2

