

CAUSATION, 18(1): 5-99

DOI:10.5281/zenodo.6816088

Received: July 10, 2022 Accepted: July 10, 2022 Published: July 10, 2022

Deutsche Nationalbibliothek Frankfurt

Conditions and study design

Research article

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Abstract

Background:

It seems like that the more is published on a single medical issue, the more it is difficult to obtain a clear, logically consistent and comprehensive information on the given medical topic. This is one of the reasons, why it is necessary to apply stringent standards and rigorous and strict study design criteria.

Methods:

In spite of all the differences, the various types of studies have at least one thing in common: find out, what is truth?

Results:

The impact of study design on necessary conditions and on sufficient conditions has been reinvestigated. The design of a study investigating necessary or sufficient condition should ensure as much as possible an index of unfairness as near as possible to 0.

Conclusion:

Investigations of necessary or sufficient conditions require an index of unfairness equal to 0.

Keywords: Study design; Index of unfairness; Conditions; Cause; Effect; Causal relationship k; Causality; Causation

1. Introduction

Often, medical research studies are used to look at factors or exposures which might potentially be related with diseases or outcomes. In point of fact, several types of studies are known but commonly divided into experimental studies (randomized clinical trials or randomized control trials (RCT)) and observational studies (i.e. cross-sectional; case-control and cohort studies). Nonetheless, each study design has both advantages and disadvantages. In an randomized clinical trials ¹ or randomized control trials, an experimental group gets the treatment/exposure (verum group or intervention group). In contrast to the intervention group, the control group of a randomized clinical trial receives no treatment or a placebo treatment (placebo group) or another standard of care treatment. The effect of an intervention

¹Munnangi S, Boktor SW. Epidemiology Of Study Design. 2022 Apr 28. In: StatPearls [Internet]. Treasure Island (FL): StatPearls Publishing; 2022 Jan–. PMID: 29262004.

is assessed. Observational or retrospective or case-control studies ² are used to establish a relationship between exposures and outcomes, but are of minor importance in order to establish causation. Therefore, when investigating a certain research issue, a researcher should put a lot of effort in the choice of an appropriate study type and on several aspect of the design of a medical study. Theoretically, let us investigate a certain research question for the sake of the argument only by an observational study and equally by an experimental study under optimal and perfectly ideal conditions. After the investigations are finished, we compare the results of the experimental study with the results of the observational study. Theoretically, it appears to be highly doubtful whether research results obtained by these two studies are allowed to differ or to deviate from one another. In addition, a relationship between an event A and another event B, assumed that the same is given in reality, is existing independently of any study design. Therefore, even if the impact of (an inappropriate) study design on the results of medical research cannot be ignored, it is not the study design itself which determines a relationship between certain events. It is therefore more than reasonable to demand that the design of a study must ensure that different studies on a single topic can be compared with each other. Especially, independent of any type of a study, different studies should be able to achieve the same research results on the same single topic. The question immediately arises and must be asked, under what conditions can the design of a study ensure comparability of studies independently of the type of a study, including repeatability, reproducibility, safety et cetera?

2. Material and methods

Scientific knowledge and objective reality are more than only interrelated. It cannot be repeated often enough that objective reality or processes of objective reality is the foundation of any scientific knowledge. Our human experience teaches us however that seen by light, grey is never merely simply grey, and looked at from different angles, many paths may lead to climb up a certain mountain. In general, it is appropriate to ensure as much as possible a broader consideration of a research question and to take into account the different facets and viewpoints of an issue investigated in order to reach a goal.

²Tenny S, Kerndt CC, Hoffman MR. Case Control Studies. 2022 Mar 28. In: StatPearls [Internet]. Treasure Island (FL): StatPearls Publishing; 2022 Jan–. PMID: 28846237.

2.1. Methods

Definitions should help us to provide and assure a systematic approach to a scientific issue. It also goes without the need of further saying that a definition as such need to be logically consistent and correct.

2.1.1. Random variables

As highlighted especially by Albert Einstein (1879-1955) and his coworkers Boris Yakovlevich Podolsky (1896-1966) and Nathan Rosen (1909-1995), "... objective reality ... is independent of any theory ... "3 (see Einstein et al., 1935, p. 777), objective reality is independent of any observer and of any perceiving subject, objective reality is independent of any measurements. Let us carry this point to epistemological extremes, objective reality is existing independently and outside of human mind and consciousness. However, in its own self-sameness objective reality is different from a random variable too and self-contradictory. Nonetheless, in its difference, in its own contradiction, a random variable itself is self-identical and is in its own self a transition of itself into the other of itself and vice versa. Lastly, a random variable as such is in its own self the opposite of itself. More or less, a random variable is in its own self the unity of identity and difference and finds its own completion in the determinate relationship of self-identity and difference. A random variable as such is in its own self self-identical and different. This has at least a twofold aspect, identity and difference constitute the determinations of a random variable itself. These two moments of a random variable which are merely different in one and the same identity are constituting as moments of difference the determinations of an opposition too. A self-identical and a different constitute equally the interior nature of itself in relation to one another. The self-identical, determined with reference to an otherness, has within itself the reference-to-other which is the determinateness of the self-identical itself. The difference contains within itself the reference to its non-being, to identity, and vice versa. Identity contains within itself the reference to its non-being, to difference. However, a random variable as such is itself and its other and the identity of difference with itself is at the end a self-reference too. Consequently, a random variable as such has its own determinateness not in an other, but in its own self, it is self-referred, while the reference to its other manifests itself as a self-reference. The other of itself which a random variable as such contains is also the non-being of that in which it is supposed to be contained only as a moment. A random variable as such therefore is, only in so far as its non-being is, and is in an identical relationship with it. The moments of a random variable are different in one and the same identity and as moments of difference are constituting the determinations of an opposition. Closer consideration shows that a random variable as such is only in so far as the same contains a reference to its non-being, to its own other moment (i.e. local hidden variable). A self-identical which is equally a different too is thus far determining an opposition as such. While the one is not as yet self-identical, the other is not as yet different. However, both are different to one another. Nonetheless, the indifference of a random variable as such towards another random variable distinguished from the same has no influence on the fact that a random variable as such is in its own self the unity of identity and difference. At the end, a

³Einstein, A; B Podolsky; N Rosen (1935-05-15). "Can Quantum-Mechanical Description of Physical Reality be Considered Complete?" (PDF). Physical Review. 47 (10): 777–780. DOI:https://doi.org/10.1103/PhysRev.47.777

random variable as such is, only in so far as the other is; it is what it is, through the other, through its own non-being. A random variable is, in so far as the other is not; it is what it is, through the non-being of its own other.

The notion something is widely taken for granted as a foundation of axioms, theorems and theories. But, very broadly put, there are many different kinds of very concrete, single entities with real world implications. Thus far, what is something, what is its own other? In the most general way, there are circumstances where something and its own other existing independently and outside of human mind and consciousness is described mathematically by the notion random variable. Let a **random variable** (Gosset, 1914) X denote something like a function defined on a probability space, which itself maps from the sample space (Neyman and Pearson, 1933) to the real numbers.

2.1.1.1. The Expectation of a Random Variable

Definition 2.1 (**The First Moment Expectation of a Random Variable**). Summaries of an entire distribution of a random variable (see Kolmogorov, Andreĭ Nikolaevich, 1950, p. 22) X, such as the expected value, or average value, are useful in order to identify where X is expected to be without describing the entire distribution. For practical and other reasons, we shall limit ourselves here to discrete random variables, while the basic properties of the expectation value of a random variable X will not be investigated. Thus far, let X be a discrete random variable with the probability p(X). The relationship between the first moment expectation value (see Huygens and van Schooten, 1657, Kolmogorov, Andreĭ Nikolaevich, 1950, LaPlace, 1812, Whitworth, 1901) of X, denoted by E(X), and the probability p(X), is given by the equation:

$$E(X) \equiv X \times p(X)$$

$$\equiv \Psi(X) \times X \times \Psi^{*}(X)$$
(1)

where $\Psi(X)$ is the wave-function (see Born, 1926, Schrödinger, Erwin Rudolf Josef Alexander, 1926) of X, $\Psi^*(X)$ is the complex conjugate wave-function of X. Under conditions where $p(X) \equiv +1$ equation 1 (see p. 9) becomes

$$E\left(X\right) \equiv X\tag{2}$$

but not general. The first moment expectation value squared of a random variable X follows as

$$E(X)^{2} \equiv p(X) \times X \times p(X) \times X$$

$$\equiv p(X) \times p(X) \times X \times X$$

$$\equiv (p(X) \times X)^{2}$$

$$\equiv E(X) \times E(X)$$
(3)

The ongoing progress with artificial intelligence has the potential to transform human society far beyond any imaginable border of human recognition and can help even to solve problems that otherwise would not be tractable. No wonder, scientist and systems are confronted with large volumes of data (big data) of various natures and from different sources. The use of tensor technology can simplify and

accelerate Big data analysis. In other words, let $X_{kl\mu\nu}$... denote an n-th index co-variant tensor with the probability $p(X_{kl\mu\nu}...)$. The first moment expectation value (see Huygens and van Schooten, 1657, Kolmogorov, Andreĭ Nikolaevich, 1950, LaPlace, 1812, Whitworth, 1901) of $X_{kl\mu\nu}...$, denoted by $E(X_{kl\mu\nu}...)$, is a number defined as follows:

$$E\left(X_{\mathrm{kl}\mu\nu\dots}\right) \equiv p\left(X_{\mathrm{kl}\mu\nu\dots}\right) \times X_{\mathrm{kl}\mu\nu\dots} \equiv p\left(X_{\mathrm{kl}\mu\nu\dots}\right) \cap X_{\mathrm{kl}\mu\nu\dots} \tag{4}$$

while \times or \cap might denote the commutative multiplications of tensors. The first moment expectation value squared of a random variable X follows as

$${}^{2}E\left(X_{\mathrm{kl}\mu\nu\dots}\right) \equiv p\left(X_{\mathrm{kl}\mu\nu\dots}\right) \times X_{\mathrm{kl}\mu\nu\dots} \times p\left(X_{\mathrm{kl}\mu\nu\dots}\right) \times X_{\mathrm{kl}\mu\nu\dots}$$

$$\equiv p\left(X_{\mathrm{kl}\mu\nu\dots}\right) \times p\left(X_{\mathrm{kl}\mu\nu\dots}\right) \times X_{\mathrm{kl}\mu\nu\dots} \times X_{\mathrm{kl}\mu\nu\dots}$$

$$\equiv {}^{2}\left(p\left(X_{\mathrm{kl}\mu\nu\dots}\right) \times X_{\mathrm{kl}\mu\nu\dots}\right)$$

$$\equiv E\left(X_{\mathrm{kl}\mu\nu\dots}\right) \times E\left(X_{\mathrm{kl}\mu\nu\dots}\right)$$
(5)

Definition 2.2 (The Second Moment Expectation of a Random Variable). The second (see Kolmogorov, Andreĭ Nikolaevich, 1950, p. 42) moment expectation value (or more or less arithmetic mean) of a (large) number of independent realizations of a random variable X follows as:

$$E(X^{2}) \equiv p(X) \times X^{2}$$

$$\equiv (p(X) \times X) \times X$$

$$\equiv E(X) \times X$$

$$\equiv X \times E(X)$$
(6)

From the point of view of tensor algebra it is

$$E\left(^{2}X_{kl\mu\nu\dots}\right) \equiv p\left(X_{kl\mu\nu\dots}\right) \times ^{2}X_{kl\mu\nu\dots}$$

$$\equiv \left(p\left(X_{kl\mu\nu\dots}\right) \times X_{kl\mu\nu\dots}\right) \times X_{kl\mu\nu\dots}$$

$$\equiv E\left(X_{kl\mu\nu\dots}\right) \times X_{kl\mu\nu\dots}$$

$$\equiv X_{kl\mu\nu\dots} \times E\left(X_{kl\mu\nu\dots}\right)$$
(7)

Definition 2.3 (The n-th Moment Expectation of a Random Variable). The n-th (see Barukčić, 2020a, 2021c) moment expectation value of a (large) number of independent realizations of a random variable X follows as:

$$E(X^{n}) \equiv p(X) \times X^{n}$$

$$\equiv (p(X) \times X) \times X^{n-1}$$

$$\equiv E(X) \times X^{n-1}$$
(8)

2.1.1.2. Probability of a Random Variable What is the nature of the probability of an event, or what is the relationship between probability and geometry or between the probability of an event and notions like false or true. At a first pass, various authors answer this question, one way or another. For authors like De Morgan, probability is only a degree of confidence, or credences or of belief. "By degree of probability, we really mean, or ought to mean, degree of belief" (see De Morgan, 1847, p. 172). Such a purely subjective (or personalist or Bayesian (see Bayes, 1763)) interpretation of probabilities as degrees of confidence, or credences finds its own scientific opposition, moreover, in Kolmogorov's axiomatization of probability theory. However, perhaps we can do better, then, to think that Kolmogorov's axiomatization of probability theory is the last word spoken on probability theory. Nobody seriously considers that Kolmogorov's conceptual apparatus of probability theory has solved the basic problem of any probability theory, the relationship between classical logic or geometry and probability theory. One very massive disadvantage of Kolmogorov's axiomatization of probability theory is that it is very silent especially on this issue. Any unification of geometry and probability theory into one unique mathematical framework might prove very difficult as long as we rely purely on Kolmogorov's understanding of probability theory. It's not surprising that the probability of an event bear at least directly, and sometimes indirectly, upon central philosophical and scientific concerns. A correct understanding of probability is one of the most important foundational scientific problems. Now let us strengthen our position with respect to the probability of an event. In our understanding, the probability of an event is something objectively and real. The probability of an event is the truth value of something or the degree to which something, i.e. a random variable X, is determined by its own expectation value. The probability p(X) of a random variable X follows as (see equation 1)

$$p(X) \equiv \frac{X \times p(X)}{X} \equiv \frac{E(X)}{X} \equiv p(X)$$

$$\equiv \frac{X \times X \times p(X)}{X \times X} \equiv \frac{X \times E(X)}{X \times X} \equiv \frac{E(X^{2})}{X^{2}}$$

$$\equiv \frac{E(X)}{X} \equiv \frac{E(X) \times E(X)}{X \times E(X)} \equiv \frac{E(X)^{2}}{E(X^{2})}$$

$$\equiv \frac{E(X)}{X} \equiv \frac{E(X) \times E(\underline{X})}{X \times E(\underline{X})} \equiv \frac{\sigma(X)^{2}}{X \times X \times (1 - p(X))} \equiv \frac{\sigma(X)^{2}}{E(\underline{X}^{2})}$$

$$\equiv \Psi(X) \times \Psi^{*}(X)$$
(9)

where $\Psi(X)$ is the wave-function of X, $\Psi^*(X)$ is the complex conjugate wave-function of X. As soon as the probability p(X) of an event X is determined, the probability of its own other, 1 - p(X), the

complementary of X, the opposite of X, anti X, is determined too. We obtain

$$1 - p(X) \equiv 1 - \frac{X \times p(X)}{X} \equiv 1 - \frac{E(X)}{X} \equiv \frac{X}{X} - \frac{E(X)}{X} \equiv \frac{X - E(X)}{X} \equiv \frac{E(\underline{X})}{X} \equiv p(\underline{X})$$

$$\equiv 1 - \frac{X \times X \times p(X)}{X \times X} \equiv 1 - \frac{X \times E(X)}{X \times X} \equiv 1 - \frac{E(X^2)}{X^2} \equiv \frac{X^2}{X^2} - \frac{E(X^2)}{X^2} \equiv \frac{X^2 - E(X^2)}{X^2}$$

$$\equiv 1 - \frac{E(X)}{X} \equiv 1 - \frac{E(X) \times E(X)}{X \times E(X)} \equiv 1 - \frac{E(X)^2}{E(X^2)}$$

$$\equiv 1 - \frac{E(X)}{X} \equiv 1 - \frac{E(X) \times E(\underline{X})}{X \times E(\underline{X})} \equiv 1 - \frac{\sigma(X)^2}{X \times X \times (1 - p(X))} \equiv 1 - \frac{\sigma(X)^2}{E(\underline{X}^2)}$$

$$\equiv 1 - \Psi(X) \times \Psi^*(X)$$
(10)

2.1.1.3. Variance of a Random Variable

Definition 2.4 (**The Variance of a Random Variable**). Johann Carl Friedrich Gauß (1777-1855) introduced the normal distribution and the error of mean squared in his 1809 monograph (see Gauß, Carl Friedrich, 1809). In the following, Karl Pearson (1857-1936) coined the term "standard deviation" in 1893. Pearson is writing: "Then σ will be termed its standard-deviation (error of mean square)." (see Pearson, 1894, p. 80). Finally, the term variance was introduced by Sir Ronald Aylmer Fisher (1890-1962) in the year 1918.

"The ... deviations of a ... measurement from its mean ... may be ... measured by the standard deviation corresponding to the square root of the mean square error ... It is ... desirable in analysing the causes ... to deal with the square of the standard deviation as the measure of variability. We shall term this quantity the Variance..."

(see Fisher, Ronald Aylmer, 1919, p. 399)

The deviation of a random variable X from its population mean or sample mean E(X) has a central role in statistics and is one important measure of dispersion. The variance $\sigma(X)^2$ (see Kolmogorov, Andreĭ Nikolaevich, 1950, p. 42), the second central moment of a distribution, is the expectation value of the squared deviation of a random variable X from its own expectation value E(X) and is determined in general as (see equation 6):

$$\sigma(X)^{2} \equiv E(X^{2}) - E(X)^{2}$$

$$\equiv (X \times E(X)) - E(X)^{2}$$

$$\equiv E(X) \times (X - E(X))$$

$$\equiv E(X) \times E(X)$$
(11)

while $E(\underline{X}) \equiv X - E(X)$. From the point of view of tensor algebra, it is

$${}^{2}\sigma\left(X_{\mathrm{kl}\mu\nu\ldots}\right) \equiv E\left({}^{2}X_{\mathrm{kl}\mu\nu\ldots}\right) - {}^{2}E\left(X_{\mathrm{kl}\mu\nu\ldots}\right)$$

$$\equiv \left(X_{\mathrm{kl}\mu\nu\ldots} \times E\left(X_{\mathrm{kl}\mu\nu\ldots}\right)\right) - {}^{2}E\left(X_{\mathrm{kl}\mu\nu\ldots}\right)$$

$$\equiv E\left(X_{\mathrm{kl}\mu\nu\ldots}\right) \times \left(X_{\mathrm{kl}\mu\nu\ldots} - E\left(X_{\mathrm{kl}\mu\nu\ldots}\right)\right)$$

$$\equiv E\left(X_{\mathrm{kl}\mu\nu\ldots}\right) \times E\left(X_{\mathrm{kl}\mu\nu\ldots}\right)$$

$$\equiv E\left(X_{\mathrm{kl}\mu\nu\ldots}\right) \times E\left(X_{\mathrm{kl}\mu\nu\ldots}\right)$$
(12)

while $E\left(\underline{X}_{\mathrm{kl}\mu\nu}...\right) \equiv X_{\mathrm{kl}\mu\nu}... - E\left(X_{\mathrm{kl}\mu\nu}...\right)$. As demonstrated by equation 12, variance depends not just on the expectation value of what has actually been observed $E\left(\left(X_{\mathrm{kl}\mu\nu}...\right)\right)$, but also on the expectation value that could have been observed but were not $\left(E\left(\underline{X}_{\mathrm{kl}\mu\nu}...\right)\right)$. There are circumstances in quantum mechanics where this fact is called the local hidden variable. Even if his might strike us

as peculiar, variance ⁴ is primarily a mathematical method which is of use in order to evaluate specific hypotheses in the light of some empirical facts. However, as a mathematical tool or method, variance is also a scientific description of a certain part of objective reality too. In this context, as a general mathematical principle, one fundamental meaning of variance is to provide a logically consistent link between something and its own other, between X and anti X.

"The variance in this sense is a measure of the inner contradictions of a random variable, of changes, of struggle within this random variable itself, or the greater $\sigma(X)^2$ of a random variable, the greater the inner contradictions of this random variable."

(see Barukčić, 2006a, p. 57)

All things considered, we can safely say that, on the whole, **the variance is a mathematical description of the philosophical notion of the inner contradiction of a random variable X** (see Hegel, Georg Wilhelm Friedrich, 1812a, 1813, 1816). Based on equation 11, it is

$$E\left(X^{2}\right) \equiv E\left(X\right)^{2} + \sigma\left(X\right)^{2} \tag{13}$$

or

$$\frac{E(X)^{2}}{E(X^{2})} + \frac{\sigma(X)^{2}}{E(X^{2})} \equiv p(X) + \frac{\sigma(X)^{2}}{E(X^{2})} \equiv +1$$
 (14)

In other words, the variance (see Barukčić, 2006b) of a random variable is a determining part of the probability of a random variable. The wave function Ψ follows in general, as

$$\Psi(X) \equiv \frac{1}{\Psi^{*}(X)} - \frac{\sigma(X)^{2}}{(\Psi^{*}(X) \times E(X^{2}))}$$

$$\equiv \frac{(E(X^{2}) - \sigma(X)^{2})}{(\Psi^{*}(X) \times E(X^{2}))}$$

$$\equiv \frac{1}{(\Psi^{*}(X) \times E(X^{2}))} \times (E(X^{2}) - \sigma(X)^{2})$$

$$\equiv \frac{1}{(\Psi^{*}(X) \times E(X^{2}))} \times E(X)^{2}$$

$$\equiv \frac{1}{(\Psi^{*}(X) \times E(X^{2}))} \times E(X)^{2}$$

$$\equiv \frac{1}{\Psi^{*}(X)} \times \frac{E(X)^{2}}{E(X^{2})}$$

$$\equiv \frac{1}{\Psi^{*}(X) \times X} \times E(X)$$
(15)

The wave function (see Born, 1926) of a quantum-mechanical system is a central determining part of the Schrödinger wave equation (see Schrödinger, Erwin Rudolf Josef Alexander, 1926, 1929, 1952).

⁴Romeijn, Jan-Willem, "Philosophy of Statistics", The Stanford Encyclopedia of Philosophy (Spring 2022 Edition), Edward N. Zalta (ed.), forthcoming URL = https://plato.stanford.edu/archives/spr2022/entries/statistics/.

Definition 2.5 (The First Moment Expectation of a Random Variable of \underline{X} (anti X)). In general, let E(X) be defined as

$$E(\underline{X}) \equiv X - E(X) \equiv X - (X \times p(X)) \equiv X \times (+1 - p(X)) \tag{16}$$

and denote an expectation value of a (discrete) random variable anti X with the probability

$$p(\underline{X}) \equiv 1 - p(X) \tag{17}$$

The first moment expectation value (see Huygens and van Schooten, 1657, Kolmogorov, Andreĭ Nikolaevich, 1950, LaPlace, 1812, Whitworth, 1901) of anti X, denoted as $E(\underline{X})$, is a number defined as follows:

$$E(X) \equiv X - (X \times p(X)) \equiv X \times (1 - p(X)) \equiv X \times p(X) \tag{18}$$

The first moment expectation value squared of a random variable anti X follows as

$$E(\underline{X})^{2} \equiv p(\underline{X}) \times X \times p(\underline{X}) \times X$$

$$\equiv p(\underline{X}) \times p(\underline{X}) \times X \times X$$

$$\equiv (p(\underline{X}) \times X)^{2}$$

$$\equiv E(\underline{X}) \times E(\underline{X})$$
(19)

Definition 2.6 (The Second Moment Expectation of a Random Variable of \underline{X} (anti X)). The second (see Kolmogorov, Andreĭ Nikolaevich, 1950, p. 42) moment expectation value (or more or less arithmetic mean) of a (large) number of independent realizations of a random variable anti X follows as:

$$E(\underline{X}^{2}) \equiv p(\underline{X}) \times X^{2}$$

$$\equiv (p(\underline{X}) \times X) \times X$$

$$\equiv E(\underline{X}) \times X$$

$$\equiv X \times E(\underline{X})$$
(20)

Definition 2.7 (The n-th Moment Expectation of a Random Variable of \underline{X} (anti X)). The n-th (see Barukčić, 2020a, 2021c) moment expectation value of a (large) number of independent realizations of a random variable anti X follows as:

$$E(\underline{X}^{n}) \equiv p(\underline{X}) \times X^{n}$$

$$\equiv (p(\underline{X}) \times X) \times X^{n-1}$$

$$\equiv E(\underline{X}) \times X^{n-1}$$
(21)

Definition 2.8 (The Co-Variance of a Random Variable). Sir Ronald Aylmer Fisher (1890 -1962) introduced the term covariance (see Bailey, 1931) in the year 1930 in his book as follows:

"It is obvious too that where a considerable fraction of the variance is contributed by chance causes, the variance of any group of individuals will be inflated in comparison with the covariances between related groups ..."

(see Fisher, Ronald Aylmer, 1930, p. 195)

In general, the co-variance is defined as given by equation 22.

$$\sigma(X,Y) \equiv E(X,Y) - (E(X) \times E(Y)) \tag{22}$$

From the point of view of tensor algebra, it is

$$\sigma\left(X_{kl\mu\nu\dots},Y_{kl\mu\nu\dots}\right) \equiv E\left(X_{kl\mu\nu\dots},Y_{kl\mu\nu\dots}\right) - \left(E\left(X_{kl\mu\nu\dots}\right) \times E\left(Y_{kl\mu\nu\dots}\right)\right) \tag{23}$$

2.1.2. Bernoulli distribution

A single event distribution is more or less a discrete probability distribution of any random variable X which takes a certain (observer independent) single value X_t at a **Bernoulli trial** (Uspensky, 1937, p. 45) (period of time) t with the probability $p(X_t)$. The same random variable X takes a certain single anti value X_t at a Bernoulli trial (period of time) t with the probability 1-p(X_t). There are conditions in nature where a random variable X can take only the values either +0 or +1 (see Birnbaum, 1961). Under these conditions, the random variable X takes the value 1 with probability $p(X_t = +1)$ and the value 0 with probability $q(X_t = +0) = 1 - p(X_t = +1)$ while the single event distribution passes over into the **Bernoulli distribution**, named after Swiss mathematician Jacob Bernoulli (Bernoulli, 1713). Less formally, many times, the Bernoulli distribution is represented by a (possibly not biased) coin toss where 1 and 0 would represent 'heads' and 'tails' (or vice versa), respectively. However, the relationship between random variables (Gosset, 1914) can be investigated by many (Gosset, 1908) methods, including the tools of probability theory, too.

Definition 2.9 (Two by two table of single event random variables).

The two by two or contingency table which has been introduced by Karl Pearson (Pearson, 1904b) in 1904 harbours still a large variety of topics and debates. Central to this is the problem to apply the laws of classical logic on data sets, which concerns the justification of inferences which extrapolate from sample data to general facts. Nevertheless, a contingency table is still an appropriate theoretical model too for studying the relationships between random variables, including *Bernoulli* (*Bernoulli*, 1713) (i.e. +0/+1) distributed random variables existing or occurring at the same *Bernoulli trial* (Uspensky, 1937) (period of time) t.

In this context, let a random variable A at the *Bernoulli trial* (Uspensky, 1937) (period of time) t, denoted by A_t, indicate a risk factor, a condition, a cause et cetera and occur or exist with the probability

 $p(A_t)$ at the *Bernoulli trial* (Uspensky, 1937) (period of time) t. Let $E(A_t)$ denote the expectation value of A_t . In general it is

$$p(A_{t}) \equiv p(a_{t}) + p(b_{t}) \tag{24}$$

The expectation value $E(A_t)$ follows as

$$E(A_{t}) \equiv A_{t} \times p(A_{t})$$

$$\equiv A_{t} \times (p(a_{t}) + p(b_{t}))$$

$$\equiv (A_{t} \times p(a_{t})) + (A_{t} \times p(b_{t}))$$

$$\equiv E(a_{t}) + E(b_{t})$$
(25)

Under conditions of +0/+1 distributed Bernoulli random variables it is

$$E(A_{t}) \equiv A_{t} \times p(A_{t})$$

$$\equiv (+0+1) \times p(A_{t})$$

$$\equiv p(A_{t})$$

$$\equiv p(a_{t}) + p(b_{t})$$
(26)

Furthermore, it is

$$p(\underline{A}_{t}) \equiv p(c_{t}) + p(d_{t}) \equiv (1 - p(A_{t})) \tag{27}$$

The expectation value $E(\underline{A}_t)$ is given as

$$E(\underline{A}_{t}) \equiv A_{t} \times (1 - p(A_{t}))$$

$$\equiv A_{t} \times (p(c_{t}) + p(d_{t}))$$

$$\equiv (A_{t} \times p(c_{t})) + (A_{t} \times p(d_{t}))$$

$$\equiv E(c_{t}) + E(d_{t})$$
(28)

Under conditions of +0/+1 distributed Bernoulli random variables we obtain

$$E(\underline{A}_{t}) \equiv A_{t} \times (1 - p(A_{t}))$$

$$\equiv (+0 + 1) \times (1 - p(A_{t}))$$

$$\equiv (1 - p(A_{t}))$$

$$\equiv p(c_{t}) + p(d_{t})$$
(29)

Let a random variable B at the *Bernoulli trial* (Uspensky, 1937) (period of time) t, denoted by B_t , indicate an outcome, a conditioned, an effect et cetera and occur or exist with the probability $p(B_t)$ at the *Bernoulli trial* (Uspensky, 1937) (period of time) t. Let $E(B_t)$ denote the expectation value of B_t . In general it is

$$p(B_t) \equiv p(a_t) + p(c_t) \tag{30}$$

The expectation value $E(B_t)$ is given by the equation

$$E(B_{t}) \equiv B_{t} \times p(B_{t})$$

$$\equiv B_{t} \times (p(a_{t}) + p(c_{t}))$$

$$\equiv (B_{t} \times p(a_{t})) + (B_{t} \times p(c_{t}))$$

$$\equiv E(a_{t}) + E(c_{t})$$
(31)

Under conditions of +0/+1 distributed Bernoulli random variables it is

$$E(B_{t}) \equiv B_{t} \times p(B_{t})$$

$$\equiv (+0+1) \times p(B_{t})$$

$$\equiv p(B_{t})$$

$$\equiv p(a_{t}) + p(c_{t})$$
(32)

Furthermore, it is

$$p(\underline{B}_{t}) \equiv p(b_{t}) + p(d_{t}) \equiv (1 - p(B_{t})) \tag{33}$$

The expectation value $E(B_t)$ is given by the equation

$$E(\underline{B}_{t}) \equiv B_{t} \times (1 - p(B_{t}))$$

$$\equiv B_{t} \times (p(b_{t}) + p(d_{t}))$$

$$\equiv (B_{t} \times p(b_{t})) + (B_{t} \times p(d_{t}))$$

$$\equiv E(b_{t}) + E(d_{t})$$
(34)

Under conditions of +0/+1 distributed Bernoulli random variables it is

$$E(\underline{B}_{t}) \equiv B_{t} \times (1 - p(B_{t}))$$

$$\equiv (+0 + 1) \times (1 - p(B_{t}))$$

$$\equiv (1 - p(B_{t}))$$

$$\equiv p(b_{t}) + p(d_{t})$$
(35)

Let $p(a_t) = p(A_t \land B_t)$ denote the joint probability distribution of A_t and B_t at the same Bernoulli trial (period of time) t. In general, it is

$$E(a_{t}) \equiv E(A_{t} \wedge B_{t})$$

$$\equiv (A_{t} \times B_{t}) \times p(A_{t} \wedge B_{t})$$

$$\equiv (A_{t} \times B_{t}) \times p(a_{t})$$
(36)

Under conditions of +0/+1 distributed Bernoulli random variables, it is

$$E(a_{t}) \equiv E(A_{t} \wedge B_{t})$$

$$\equiv (A_{t} \times B_{t}) \times p(A_{t} \wedge B_{t})$$

$$\equiv ((+0+1) \times (+0+1)) \times p(A_{t} \wedge B_{t})$$

$$\equiv p(A_{t} \wedge B_{t})$$

$$\equiv p(a_{t})$$
(37)

Let $p(b_t) = p(A_t \land \neg B_t)$ denote the joint probability distribution of A_t and not B_t at the same Bernoulli trial (period of time) t. In general, it is

$$E(b_{t}) \equiv E(A_{t} \wedge \neg B_{t})$$

$$\equiv (A_{t} \times \neg B_{t}) \times p(A_{t} \wedge \neg B_{t})$$

$$\equiv (A_{t} \times \neg B_{t}) \times p(b_{t})$$
(38)

Under conditions of +0/+1 distributed Bernoulli random variables, it is

$$E(b_{t}) \equiv E(A_{t} \wedge \neg B_{t})$$

$$\equiv (A_{t} \times \neg B_{t}) \times p(A_{t} \wedge \neg B_{t})$$

$$\equiv ((+0+1) \times (+0+1)) \times p(A_{t} \wedge \neg B_{t})$$

$$\equiv p(A_{t} \wedge \neg B_{t})$$

$$\equiv p(b_{t})$$
(39)

Let $p(c_t) = p(\neg A_t \land B_t)$ denote the joint probability distribution of not A_t and B_t at the same Bernoulli trial (period of time) t. In general, it is

$$E(c_{t}) \equiv E(\neg A_{t} \wedge B_{t})$$

$$\equiv (\neg A_{t} \wedge B_{t}) \times p(\neg A_{t} \wedge B_{t})$$

$$\equiv (\neg A_{t} \wedge B_{t}) \times p(c_{t})$$

$$(40)$$

Under conditions of +0/+1 distributed Bernoulli random variables, it is

$$E(c_{t}) \equiv E(\neg A_{t} \wedge B_{t})$$

$$\equiv (\neg A_{t} \times B_{t}) \times p(\neg A_{t} \wedge B_{t})$$

$$\equiv ((+0+1) \times (+0+1)) \times p(\neg A_{t} \wedge B_{t})$$

$$\equiv p(\neg A_{t} \wedge B_{t})$$

$$\equiv p(c_{t})$$

$$(41)$$

Let $p(d_t) = p(\neg A_t \land \neg B_t)$ denote the joint probability distribution of not A_t and not B_t at the same Bernoulli trial (period of time) t. In general, it is

$$E(d_{t}) \equiv E(\neg A_{t} \times \neg B_{t})$$

$$\equiv (\neg A_{t} \times \neg B_{t}) \times p(\neg A_{t} \wedge \neg B_{t})$$

$$\equiv (\neg A_{t} \times \neg B_{t}) \times p(d_{t})$$

$$(42)$$

Under conditions of +0/+1 distributed Bernoulli random variables, it is

$$E(d_{t}) \equiv E(\neg A_{t} \wedge \neg B_{t})$$

$$\equiv (\neg A_{t} \times \neg B_{t}) \times p(\neg A_{t} \wedge \neg B_{t})$$

$$\equiv ((+0+1) \times (+0+1)) \times p(\neg A_{t} \wedge \neg B_{t})$$

$$\equiv p(\neg A_{t} \wedge \neg B_{t})$$

$$\equiv p(d_{t})$$

$$(43)$$

In general, it is

$$p(a_t) + p(b_t) + p(c_t) + p(d_t) \equiv +1$$
 (44)

Table 1 provide us with an overview of the definitions above.

In our understanding, it is

$$p(B_t) + p(\Lambda_t) \equiv p(a_t) + p(c_t) + p(\Lambda_t) \equiv p(a_t) + p(b_t) \equiv p(A_t)$$

$$(45)$$

 $\begin{array}{c|cccc} & & & & & & & & \\ & & & & & & \\ \hline Condition & & & & & \\ \hline Condition & TRUE & p(a_t) & p(b_t) & p(A_t) \\ A_t & FALSE & p(c_t) & p(d_t) & p(\underline{A}_t) \\ & & & & & \\ \hline p(B_t) & p(B_t) & +1 \\ \hline \end{array}$

Table 1. The two by two table of Bernoulli random variables

or

$$p(c_{t}) + p(\Lambda_{t}) \equiv p(b_{t}) \tag{46}$$

Under conditions of Einstein's general theory of relativity, Λ denotes the Einstein cosmological (Einstein, 1917) 'constant'.

2.1.3. Binomial and Anti-binomial distribution

The binomial (see Pearson, 1895, p. 351) distribution (see Cramér, 1937) with parameters n and p has been developed by the Swiss mathematician Jakob Bernoulli (1655-1705) in a proof published in his 1713 book Ars Conjectandi (see Bernoulli, 1713) Part 1. In probability theory and statistics, the probability of getting exactly k successes in n independent Bernoulli trials is given by the probability mass function as

$$p(X_{t} = k) \equiv \binom{n}{k} \times p^{k} \times q^{n-k}$$
(47)

while is $\binom{n}{k} = \frac{n!}{k!(n-k)!}$ the binomial coefficient. The Anti-binomial distribution, the other or the complementary of a Binomial distribution, denoted as $p(X_t = \underline{k})$ is given as:

$$p(X_{t} = \underline{k}) = 1 - p(X_{t} = k) \equiv 1 - \binom{n}{k} \times p^{k} \times q^{n-k}$$
(48)

The variance of a Binomial distributed event is given as

$$\sigma(k)^{2} \equiv k \times k \times p(k) \times p(\underline{k})$$

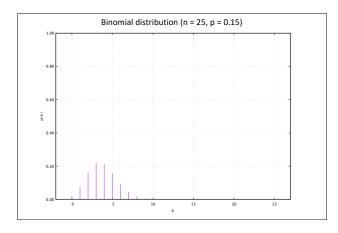
$$\equiv (k) \times (k) \times \left(\binom{n}{k} \times p^{k} \times q^{n-k} \right) \times \left(1 - \left(\binom{n}{k} \times p^{k} \times q^{n-k} \right) \right)$$
(49)

The relationship between Binomial distribution and Anti-binomial distribution is illustrated by fig. 1 and fig. 2 in more detail.

As known, the cumulative distribution function is given as

$$p(X_{t} \le k) \equiv 1 - p(X_{t} > k) \equiv \sum_{t=0}^{k} {n \choose t} \cdot p^{t} \cdot q^{n-t}$$

$$(50)$$



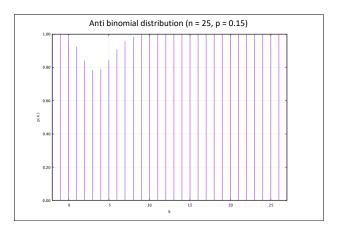


Figure 1. Binomial distribution.

Figure 2. Anti-binomial dis.

or as

$$p(X_t > k) \equiv 1 - p(X_t \le k) \equiv 1 - \sum_{t=0}^{k} {n \choose t} \cdot p^t \cdot q^{n-t}$$
 (51)

Furthermore, it is

$$p(X_{t} < k) \equiv 1 - p(X_{t} \ge k) \equiv \sum_{t=0}^{k-1} \binom{n}{t} \cdot p^{t} \cdot q^{n-t}$$

$$(52)$$

or

$$p(X_{t} \ge k) \equiv 1 - p(X_{t} < k) \equiv 1 - \sum_{t=0}^{k-1} {n \choose t} \cdot p^{t} \cdot q^{n-t}$$
(53)

The binomial distribution is the mathematical foundation of a binomial test. The random variable X_t is counting for different things. The discrete geometric (see Feller, 1950, p. 61) distribution describes under certain circumstances the number of Bernoulli trials needed to get one success. The probability that the first occurrence of success requires k independent trials, each with success probability p, is given by the equation

$$p(X_t = k) \equiv p \cdot q^{k-1} \tag{54}$$

The negative (see Fisher, 1941, Haldane, 1941) binomial probability is a discrete probability distribution which defines the number of successes (k) in a sequence of independent and identically distributed Bernoulli trials (n) before a specified (non-random) number of failures (denoted r) occurs. The probability mass function of the negative binomial distribution is

$$p(X_{t} = r) \equiv {k+r-1 \choose k-1} p^{k} \cdot q^{r}$$
(55)

where k is the number of successes, r is the number of failures, and p is the probability of success.

Definition 2.10 (Expectation value and variance of a binomial random variable).

The variance(see Pearson, 1904a, p. 66) of a binomial distributed random variable with parameters n, the number of independent experiments each asking a yes–no question and p, the probability of a

single event, is defined in contrast to Pearson (see Barukčić, 2022c) as

$$\sigma(X_{t})^{2} \equiv N \times N \times p(X_{t}) \times (1 - p(X_{t}))$$
(56)

Definition 2.11 (Two by two table of Binomial random variables).

Let a, b, c, d, A, \underline{A} , B, and \underline{B} denote expectation values. Under conditions where the probability of an event, an outcome, a success et cetera is **constant** from Bernoulli trial to Bernoulli trial t, it is

$$A = N \times E(A_{t})$$

$$\equiv N \times (A_{t} \times p(A_{t}))$$

$$\equiv N \times (p(A_{t}) + p(B_{t}))$$

$$\equiv N \times p(A_{t})$$
(57)

and

$$B = N \times E(B_{t})$$

$$\equiv N \times (B_{t} \times p(B_{t}))$$

$$\equiv N \times (p(A_{t}) + p(c_{t}))$$

$$\equiv N \times p(B_{t})$$
(58)

where N might denote the population or even the sample size. Furthermore, it is

$$a \equiv N \times (E(A_{t})) \equiv N \times (p(A_{t})) \tag{59}$$

and

$$b \equiv N \times (E(B_{t})) \equiv N \times (p(B_{t})) \tag{60}$$

and

$$c \equiv N \times (E(c_{t})) \equiv N \times (p(c_{t})) \tag{61}$$

and

$$d \equiv N \times (E(d_{t})) \equiv N \times (p(d_{t})) \tag{62}$$

and

$$a+b+c+d \equiv A+\underline{A} \equiv B+\underline{B} \equiv N \tag{63}$$

Table 2 provide us again an overview of a two by two contingency (see also Pearson, 1904b, p. 33) table of Binomial random variables.

Table 2. The two by two table of Binomial random variables

	Conditioned B _t			
		TRUE	FALSE	
Condition	TRUE	a	b	Α
A_t	FALSE	c	d	<u>A</u>
		В	В	N

"Such a table is termed a contingency table, and the ultimate scientific statement of description of the relation between two things can always be thrown back upon such a contingency table ... Once the reader realizes the nature of such a table, he will have grasped the essence of the conception of association between cause and effect, and the nature of its ideal limit in causation."

(see also Pearson, 1911, p. 159)

2.1.4. Poisson and Anti-Poisson distribution

The Poisson distribution (see also Poisson, 1829, pp. 261-262) is a discrete distribution (with k = 0, 1, 2, 3, ...) which depends on λ , the mean number of occurrences expected (see also Poisson and Poisson, 1837, pp. 205) while there is no specified number n of possible tries. The probability of a given number of events occurring in a fixed interval of time or space under the condition that these events occur with a known constant mean rate and independently of the time since the last event, is calculated as

$$p(k) = \left(\frac{\lambda^k}{k!}\right) \times e^{-\lambda} \tag{64}$$

Cavalry men being killed by a kick of a horse (see also von Bortkiewitsch, 1898) is a famous example of Poisson distribution. The Anti-Poisson distribution, the other of the Poisson distribution or the complementary of the Poisson distribution, denoted as p(k) is given as

$$p(\underline{k}) = 1 - p(k) = 1 - \left(\frac{\lambda^k}{k!}\right) \times e^{-\lambda}$$
(65)

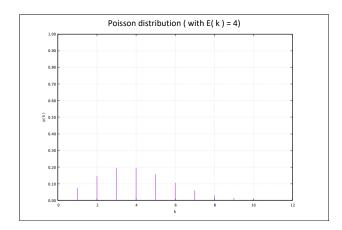
The variance of a Poisson distributed event is given as

$$\sigma(k)^{2} \equiv k \times k \times p(k) \times p(\underline{k})$$

$$\equiv (k) \times (k) \times \left(\left(\frac{\lambda^{k}}{k!} \right) \times e^{-\lambda} \right) \times \left(1 - \left(\left(\frac{\lambda^{k}}{k!} \right) \times e^{-\lambda} \right) \right)$$
(66)

The relationship between the Poisson distribution and the Anti Poisson distribution is illustrated by fig. 3 and fig. 4 in more detail.

Bombing of London during World War II by Germans



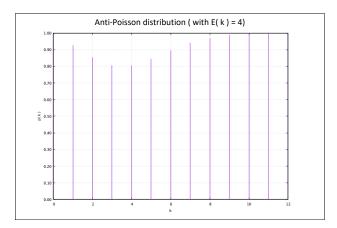


Figure 3. Poisson distribution.

Figure 4. Anti-poisson dis.

During World War II, London was bombed 5 , 6 by Germans. In order to determine (Clarke, 1946), whether the Germans were bombing London randomly or could target specific areas, London was divided into a grid consisting of 576 equal squares, each square of area 0.25 square kilometres. The number of squares with k = 0, 1, ... bombs that landed in each grid square was counted. Over the period considered, the total number of bombs within the area of London involved was 537. The data are illustrated by fig. 3.

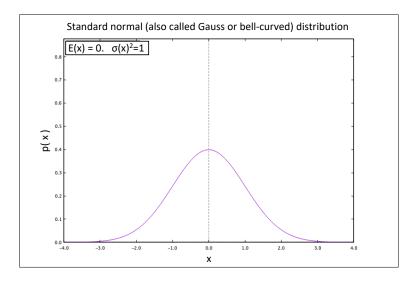
Table 3. Bombing of London during World War II by Germans

Number of flying bombs	Observed number	Expected number	
per square (k)	of squares	of squares (Poisson)	
0	229	226.74	
1	211	211.39	
2	93	98.54	
3	35	30.62	
4	7	7.14	
5 and over	1	1.57	
	576	576	

The closeness of fit with the Poisson distribution is obvious and has been tested by the $\tilde{\chi}^2$ goodness of fit test.

⁵Clarke, R. D. (1946). An application of the Poisson distribution. Journal of the Institute of Actuaries, 72(3), 481-481.

⁶Clarke, R. D. (1946). An application of the Poisson distribution. Journal of the Institute of Actuaries, 72(3), 481-481.



Normal distribution with E(x) = 0 and $\sigma(x)^2 = 1$

Figure 5. Normal distribution

2.1.5. Normal and Anti-normal distribution

The origins of the normal distribution, also known as the Gaussian distribution, the second law of Laplace, the law of error et cetera, has been studied at least since the 18th century and can be traced back even to a French mathematician Abraham de Moivre. Johann Carl Friedrich Gauß's (1777-1855) presented 1809 the normal distribution (see Gauß, Carl Friedrich, 1809, p. 244) while illustrating the method of least squares. In the following, Karl Pearson (1857-1936) popularised a new name for Gauß distribution. Pearson wrote: "A frequency-curve, which for practical purposes, can be represented by the error curve, will for the remainder of this paper be termed a normal curve." (see Pearson, 1894, p. 72).

$$p(\mathbf{R}X_{t}) = \left(\frac{1}{\sqrt{2\pi \times \sigma(\mathbf{R}X_{t})^{2}}}\right) e^{-\frac{(\mathbf{R}X_{t} - E(\mathbf{R}X_{t}))^{2}}{2 \times \sigma(\mathbf{R}X_{t})^{2}}}$$

$$(67)$$

The standard normal distribution is illustrated by figure 5.

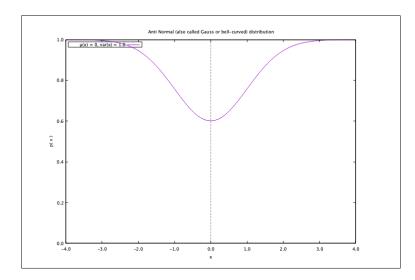
Sir Ronald Aylmer Fisher (1890-1962) ⁷, a very influential statistician of the first half of the 20th century, presented the case of a normal (see Fisher, Ronald Aylmer, 1912, p. 157) distribution with non-zero mean (see Fisher, Ronald Aylmer, 1920, p. 758) as a typical case. The probability density function (pdf) of an anti-normal distribution is given as

$$p(\underline{X}_{t}) = 1 - \left(\frac{1}{\sqrt{2\pi \times \sigma(\underline{X}_{t})^{2}}}\right) e^{-\frac{(\underline{X}_{t} - E(\underline{X}_{t}))^{2}}{2 \times \sigma(\underline{X}_{t})^{2}}}$$
(68)

as illustrated by figure 6. In general, it is

$$p(_{R}X_{t}) + p(_{R}X_{t}) = 1 (69)$$

⁷R. A. Fisher Digital Archive, The University of Adelaide. 5005 AUSTRALIA. copyright@adelaide.edu.au



Anti-normal distribution with E(x) = 0 and $\sigma(x)^2 = 1$

Figure 6. Anti-normal distribution

The variance of a Gaussian distributed random variable is given as

$$\sigma(_{R}X_{t})^{2} \equiv {_{R}X_{t}} \times {_{R}X_{t}} \times p(_{R}X_{t}) \times p(_{R}X_{t})$$

$$\equiv {_{R}X_{t}} \times {_{R}X_{t}} \times \left(\left(\frac{1}{\sqrt{2\pi \times \sigma(_{R}X_{t})^{2}}}\right) e^{-\frac{\left(_{R}X_{t} - E(_{R}X_{t})\right)^{2}}{2 \times \sigma(_{R}X_{t})^{2}}}\right) \times \left(1 - \left(\left(\frac{1}{\sqrt{2\pi \times \sigma(_{R}X_{t})^{2}}}\right) e^{-\frac{\left(_{R}X_{t} - E(_{R}X_{t})\right)^{2}}{2 \times \sigma(_{R}X_{t})^{2}}}\right)\right)$$
(70)

Under conditions where $E(_RX_t) = 0$ and $\sigma(_RX_t)^2 = 1$, equation 70 becomes

$$\sigma\left({}_{R}X_{t}\right)^{2} \equiv {}_{R}X_{t} \times {}_{R}X_{t} \times p\left({}_{R}X_{t}\right) \times p\left({}_{R}X_{t}\right)$$

$$\equiv {}_{R}X_{t} \times {}_{R}X_{t} \times \left(\left(\frac{1}{\sqrt{2\pi \times \sigma({}_{R}X_{t})^{2}}}\right) e^{-\frac{\left({}_{R}X_{t} - E\left({}_{R}X_{t}\right)\right)^{2}}{2 \times \sigma({}_{R}X_{t})^{2}}}\right) \times \left(1 - \left(\left(\frac{1}{\sqrt{2\pi \times \sigma({}_{R}X_{t})^{2}}}\right) e^{-\frac{\left({}_{R}X_{t} - E\left({}_{R}X_{t}\right)\right)^{2}}{2 \times \sigma({}_{R}X_{t})^{2}}}\right)\right)$$

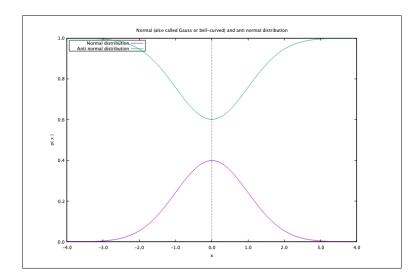
$$\equiv {}_{R}X_{t} \times {}_{R}X_{t} \times \left(\left(\frac{1}{\sqrt{2\pi \times 1}}\right) e^{-\frac{\left({}_{R}X_{t} - 0\right)^{2}}{2 \times 1}}\right) \times \left(1 - \left(\left(\frac{1}{\sqrt{2\pi \times 1}}\right) e^{-\frac{\left({}_{R}X_{t} - 0\right)^{2}}{2 \times 1}}\right)\right)$$

$$\equiv {}_{R}X_{t} \times {}_{R}X_{t} \times \left(\left(\frac{1}{\sqrt{2\pi }}\right) e^{-\frac{\left({}_{R}X_{t}\right)^{2}}{2}}\right) \times \left(1 - \left(\left(\frac{1}{\sqrt{2\pi }}\right) e^{-\frac{\left({}_{R}X_{t} - 0\right)^{2}}{2}}\right)\right)$$

Standard normal distribution

In general, a normal distribution with mean 0 and variance 1 is called the standard normal distribution. Modern publications often write the density function for the standard normal distribution, 'bell-shaped curve', as

$$p(z) = \left(\frac{1}{\sqrt{2\pi}}\right)e^{-\frac{z^2}{2}}\tag{72}$$



Normal and anti-normal distribution with E(x) = 0 and $\sigma(x)^2 = 1$

Figure 7. Normal and anti-normal distribution

The density function for the anti-standard normal distribution is given as

$$p(\underline{z}) = 1 - p(z) = 1 - \left(\frac{1}{\sqrt{2\pi}}\right)e^{-\frac{z^2}{2}}$$
 (73)

It is

$$p(z) + p(z) = 1 \tag{74}$$

and is illustrated by figure 7.

Truman Lee Kelley (1884–1961) introduced statistical methods into psychological studies 8 and defined the z-score (see Kelley, 1924, p. 115). In mathematical statistics, a random variable $_RX_t$ is standardised by subtracting its expected value $E(_RX_t)$ and dividing the difference by its standard deviation $\sigma(_RX_t)$. The z-score or standard score, denoted as $z(_RX_t)$, is defined as

$$z({}_{R}X_{t}) = \frac{({}_{R}X_{t} - E({}_{R}X_{t}))}{\sigma({}_{R}X_{t})}$$

$$(75)$$

Simply put, a z-score (also called a standard score) describes how many standard deviations a given quantum mechanical observable or a random variable lies above or below a specific value. Equation 75 changes to

$$z({_{R}X_{t}})^{2} = \frac{({_{R}X_{t}} - E({_{R}X_{t}}))^{2}}{\sigma({_{R}X_{t}})^{2}} = \frac{E({_{R}X_{t}})^{2}}{E({_{R}X_{t}}) \times E({_{R}X_{t}})} = \frac{E({_{R}X_{t}})}{E({_{R}X_{t}})} = \frac{{_{R}X_{t}} \times (1 - p({_{R}X_{t}}))}{{_{R}X_{t}} \times p({_{R}X_{t}})} = \frac{(1 - p({_{R}X_{t}}))}{p({_{R}X_{t}})} = \frac{(1 - p({_{R}X_{t}}))}{p({_{R}X_{t}$$

Equation 76 simplifies as

$$E(_{\mathbf{R}}\underline{X}_{\mathbf{t}}) = z(_{\mathbf{R}}X_{\mathbf{t}})^{2} \times E(_{\mathbf{R}}X_{\mathbf{t}})$$
(77)

⁸McClure WE. Speed and Accuracy of the Feebleminded on Performance Tests. Psychol Clin. 1931 Feb;19(9):265-274. PMID: 28909304; PMCID: PMC5138284.

We can imagine drawing figure 7 in n dimensions. Under these circumstances we would obtain something similar to an **Einstein–Rosen bridge** or Einstein–Rosen wormhole ⁹ formulated in terms of the framework of probability theory. Attention should be drawn to circumstances especially of quantum mechanics, where $E(_{R}X_{t})$ indicates something like the expectation value of a 'local hidden variable '. Equation 77 changes slightly. It is

$$_{\mathbf{R}}X_{\mathbf{t}} \times (1 - p(_{\mathbf{R}}X_{\mathbf{t}})) = z(_{\mathbf{R}}X_{\mathbf{t}})^{2} \times _{\mathbf{R}}X_{\mathbf{t}} \times p(_{\mathbf{R}}X_{\mathbf{t}})$$

$$\tag{78}$$

and

$$(1 - p(_{R}X_{t})) = z(_{R}X_{t})^{2} \times p(_{R}X_{t})$$
(79)

Equation 79 is rearranged as

$$1 = z(_{R}X_{t})^{2} \times p(_{R}X_{t}) + p(_{R}X_{t})$$
(80)

or

$$1 = (z(_{R}X_{t})^{2} + 1) \times p(_{R}X_{t})$$
(81)

At the end, it follows that

$$p(_{R}X_{t}) = \frac{1}{z(_{R}X_{t})^{2} + 1}$$
(82)

From equation 76 follows that

$$z({_{R}X_{t}})^{2} = \frac{({_{R}X_{t}} - E({_{R}X_{t}}))^{2}}{\sigma({_{R}X_{t}})^{2}} = \frac{E({_{R}\underline{X}_{t}})^{2}}{E({_{R}X_{t}}) \times E({_{R}\underline{X}_{t}})} = \frac{E({_{R}\underline{X}_{t}})}{E({_{R}X_{t}})} = \frac{E({_{R}\underline{X}_{t}}) \times E({_{R}X_{t}})}{E({_{R}X_{t}}) \times E({_{R}X_{t}})} = \frac{\sigma({_{R}X_{t}})^{2}}{E({_{R}X_{t}})^{2}}$$
(83)

Thus far, it is equally

$$\sigma(_{R}X_{t})^{2} = z(_{R}X_{t})^{2} \times E(_{R}X_{t})^{2}$$
(84)

or

$$\sigma(_{R}X_{t}) = z(_{R}X_{t}) \times E(_{R}X_{t})$$
(85)

Per definition, it is

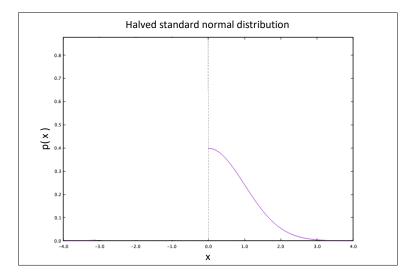
$$E(_{R}X_{t}) = \frac{\sigma(_{R}X_{t})}{z(_{R}X_{t})}$$
(86)

The probability density of a halved normal distribution for positive x is given as

$$p({}_{\mathbf{R}}X_{\mathbf{t}}) = \left(\frac{2}{\sqrt{2\pi \times \sigma({}_{\mathbf{R}}X_{\mathbf{t}})^2}}\right) e^{-\frac{\left({}_{\mathbf{R}}X_{\mathbf{t}} - E({}_{\mathbf{R}}X_{\mathbf{t}})\right)^2}{2 \times \sigma({}_{\mathbf{R}}X_{\mathbf{t}})^2}}$$
(87)

and illustrated by figure 8.

⁹Cramer JG, Forward RL, Morris MS, Visser M, Benford G, Landis GA. Natural wormholes as gravitational lenses. Phys Rev D Part Fields. 1995 Mar 15;51(6):3117-3120. doi: 10.1103/physrevd.51.3117. PMID: 10018782.



Halved normal distribution with E(x) = 0 and $\sigma(x)^2 = 1$

Figure 8. Halved normal distribution

2.1.6. Independence

Definition 2.12 (Independence).

The philosophical, mathematical (Kolmogoroff, Andreĭ Nikolaevich, 1933) and physical (Einstein, 1948) et cetera concept of independence is of fundamental(Kolmogoroff, Andreĭ Nikolaevich, 1933) importance in (natural) sciences as such. Therefore, it is appropriate to investigate the concept of independence as completely as possible. In fact, de Moivre sums it up in his book The Doctrine of Chances (see also Moivre, 1718). "Two Events are independent, when they have no connexion one with the other, and that the happening of one neither forwards nor obstructs the happening of the other. Two events are **dependent**, when they are so connected together as that the Probability of either's happening is alter'd by the happening of the other. "(see also Moivre, 1756, p. 6) We should consider Kolmogorov's position on independence before the mind's eve too. "The concept of mutual independence of two or more experiments holds, in a certain sense, a central position in the theory of probability." (see also Kolmogorov, Andreĭ Nikolaevich, 1950, p. 8) Furthermore, it is insightful to recall even Einstein's theoretical approach to the concept of independence. "Ohne die Annahme einer · · · Unabhängigkeit der · · · Dinge voneinander · · · wäre physikalisches Denken · · · nicht möglich." (Einstein, 1948). In general, an event A_t at the Bernoulli trial t need not, but can be independent of the existence or of the occurrence, of another event B_t at the same Bernoulli trial t. De Moivre brings it to the point. "From what has been said, it follows, that if a Fraction expresses the Probability of an Event, and another Fraction the Probability of another Event, and those two Events are independent; the Probability that both those Events will Happen, will be the Product of those two Fractions." (see also Moivre, 1718, p. 4). Mathematically, in terms of probability theory, independence (Kolmogoroff, Andreĭ Nikolaevich, 1933) of events at the same (period of) time (i.e. Bernoulli trial) t is defined as

$$p(A_{t} \wedge B_{t}) \equiv p(A_{t}) \times p(B_{t}) \equiv p(a_{t})$$

$$\equiv \frac{\sum_{t=1}^{N} (A_{t} \wedge B_{t})}{N} \equiv \frac{N \times (p(a_{t}))}{N} \equiv 1 - p(A_{t} \mid B_{t}) \equiv 1 - p(A_{t} \uparrow B_{t})$$
(88)

while $p(A_t \cap B_t)$ is the joint probability of the events A_t and B_t at a same Bernoulli trial t, $p(A_t)$ is the probability of an event A_t at a same Bernoulli trial t, and $p(B_t)$ is the probability of an event B_t at a same Bernoulli trial t. With respect to a two-by-two table, under conditions of independence, it is

$$p(b_t) \equiv p(A_t) \times p(B_t) \tag{89}$$

or

$$p(c_{t}) \equiv p(A_{t}) \times p(B_{t}) \tag{90}$$

and

$$p(d_{t}) \equiv p(\underline{A}_{t}) \times p(\underline{B}_{t}) \tag{91}$$

Example. In a narrower sense, the conditio sine qua non relationship concerns itself at the end only with the case whether the presence of an event A_t (condition) enables or guarantees the presence of another event B_t (conditioned). Thus far, as a result of the thoughts before, another question worth asking concerns the relationship between the independence of an event A_t (a condition) and another event B_t (conditioned) and the necessary condition relationship. To be confronted with the danger of bias and equally with the burden of inappropriate conclusions drawn, another fundamental question at this stage is whether is it possible that an event A_t (a condition) is a necessary condition of event B_t (conditioned) even under circumstances where the event A_t (a condition) (a necessary condition) is independent of an event B_t (conditioned)? Meanwhile, this question is more or less already answered to the negative (Barukčić, 2018b). An event A_t which is a necessary condition of another event B_t is equally an event without which another event (B_t) could not be, could not occur, and implies as such already a kind of dependence. However, it is not mandatory that such a kind of dependence is a causal one. It is remarkable that data which provide evidence of a significant conditio sine qua non relationship between two events like A_t and B_t and equally support the hypothesis that A_t and B_t are independent of each other are more or less self-contradictory and of very restricted or of none value for further analysis. In fact, if the opposite view would be taken as plausible, contradictions are more or less inescapable.

2.1.7. Dependence

Definition 2.13 (Dependence).

Whilst it may be true that the occurrence of an event A_t does not affect the occurrence of an other event B_t the contrary is of no minor importance. Under these other conditions, events, trials and

random variables et cetera are dependent on each other too. The dependence of events (Barukčić, 1989, p. 57-61) is defined as

$$p\left(\underbrace{A_{t} \wedge B_{t} \wedge C_{t} \wedge \dots}_{n \ random \ variables}\right) \equiv \sqrt[n]{p(A_{t}) \times p(B_{t}) \times p(C_{t}) \times \dots}_{n \ random \ variables}$$
(92)

2.1.8. Sensitivity and specificity

Definition 2.14 (Sensitivity and specificity).

A (medical) test should measure what is supposed to measure. However, the extent to which a test measures what it is supposed to measure varies and is seldom equal to 100 %. In other words, it is necessary to check once and again the accuracy or the validity of a test, we have to fight it out in detail. In clinical practice, the concept of sensitivity and specificity is commonly used to quantify the diagnostic ability of a (medical) test. Sensitivity and specificity were introduced by the American ¹⁰, ¹¹, ¹², ¹³ biostatistician Jacob Yerushalmy (see also Yerushalmy, 1947) in the year 1947. The interior logic of sensitivity and specificity is best illustrated using a conventional two- by-two (2 x 2) table (see table 4).

Table 4. Sensitivity and specificity

		Disease B _t		
		present	absent	
Test	positive	a (true positive)	b (false positive)	A
A_{t}	negative	c (false negative)	d (true negative)	<u>A</u>
		В	<u>B</u>	N

The ability of a positive test (A_t) to correctly classify an individual as diseased (B_t) is defined as the proportion of true positives that are correctly identified by the test (a) divided by the individuals being truly diseased (B_t) . In general, sensitivity follows as

Sensitivity
$$(A \mid B) \equiv p(a \mid B) \equiv \frac{a}{B}$$
 (93)

The specificity of a test is the ability of a negative test (\underline{A}_t) to correctly classify an individual as not diseased (\underline{B}_t and is defined as the proportion of true negative that are correctly identified by the test (d) divided by the individuals being truly not diseased (\underline{B}_t). In general, specificity is given by the equation

Specificity
$$(\underline{A}, \underline{B}) \equiv p(d \mid \underline{B}) \equiv \frac{d}{\underline{B}}$$
 (94)

The positive predictive value (PPV) is defined as

$$PPV(A,B) \equiv \frac{a}{a+b} \tag{95}$$

¹⁰Yerushalmy Jacob. Statistical problems in assessing methods of medical diagnosis, with special reference to X-ray techniques. Public Health Rep. 1947 Oct 3;62(40):1432-49. PMID: 20340527.

¹¹Galen RS, Gambino SR. Beyond normality-the predictive value and efficiency of medical diagnosis. New York: NY:Wiley; 1975.

¹²Altman DG, Bland JM. Diagnostic tests. 1: Sensitivity and specificity. BMJ. 1994 Jun 11;308(6943):1552. doi: 10.1136/bmj.308.6943.1552. PMID: 8019315; PMCID: PMC2540489.

¹³Parikh R, Mathai A, Parikh S, Chandra Sekhar G, Thomas R. Understanding and using sensitivity, specificity and predictive values. Indian J Ophthalmol. 2008 Jan-Feb;56(1):45-50. doi: 10.4103/0301-4738.37595. PMID: 18158403; PMCID: PMC2636062.

The negative predictive value (NPV) is defined as

$$NPV(A,B) \equiv \frac{d}{c+d} \tag{96}$$

Example.

The importance of sensitivity and specificity in any research should certainly not be underestimated. However, it is essential not to lose sight of the major advantages and limitations ¹⁴ of these measures. In the following, in order to avoid misconceptions about sensitivity, specificity et cetera, let us consider a test with a sensitivity of 95 % and a specificity of 95 %. A two-by-two table is used as an illustration (see table 5).

Table 5. Sensitivity and specificity

		Disease B _t		
		present	absent	
Test	positive	95	5	100
A_{t}	negative	5	95	100
		100	100	200

Sensitivity is calculated as

Sensitivity
$$(A \mid B) \equiv p(a \mid B) \equiv 100 \times \frac{a}{B} \equiv \frac{95}{100} \equiv 95\%$$
 (97)

There are at least two kinds of medical tests, diagnostic tests and screening tests. Depending on the type of medical test, there are other logical implications. A screening test should correctly identify all people who suffer from a certain disease or all people with a certain outcome. Therefore, the sensitivity of a screening test should be at best 100 %. Under these conditions, we obtain **without** positive test **no** disease/outcome present. However, confusion should be avoided with regard to the adequacy and usefulness of the sensitivity of a screening test. The sensitivity of a test does not take into account events which are false positive (b) or which are true negative (d), the meaning of these events is ignored completely by sensitivity. Therefore, sensitivity is blind on one eye since its inception and underestimates the extent to which a screening test is able to identify the likely presence of a condition of interest. We calculated a 95 % sensitivity while the true possibility of the test to detect a disease is (see table 5)

SINE
$$(A,B) \equiv 100 \times \frac{a+b+d}{N} \equiv \frac{95+5+95}{200} \equiv 97.5\%$$
 (98)

In a way similar to sensitivity, specificity is not much better. Diagnostic tests are able to identify people who do not have a certain condition. Specificity is calculated as

Specificity
$$(\underline{A} \mid \underline{B}) \equiv p(d \mid \underline{B}) \equiv 100 \times \frac{d}{B} \equiv \frac{95}{100} \equiv 95\%$$
 (99)

¹⁴Trevethan R. Sensitivity, Specificity, and Predictive Values: Foundations, Pliabilities, and Pitfalls in Research and Practice. Front Public Health. 2017 Nov 20;5:307. doi: 10.3389/fpubh.2017.00307. PMID: 29209603; PMCID: PMC5701930.

However, specificity does not take into account any individuals who suffer from a disease, who do have the condition and is well-known for being imperfect because of this fact too. Specificity underestimates the possibility of a diagnostic test to detect a disease. Above, the specificity has been calculated as being 95 %. In point of fact, the ability of the test to detect a disease or the relationship **if** test positive **then** disease present is much better and has to be calculated as (see table 5)

$$IMP(A,B) \equiv \frac{a+c+d}{N} \equiv \frac{95+5+95}{200} \equiv 97.5\%$$
 (100)

As can be seen, the test detected the disease in 97.5 % while specificity allows only 95 %. How valuable is such a measure epistemologically? Measures like sensitivity and specificity are blurring of the issue, do risk leading us astray and disorient us systematically again and again. These measures should be abandoned.

2.1.9. Odds ratio (OR)

Definition 2.15 (Odds ratio (OR)).

Odds ratios as an appropriate measure for estimating the relative risk have become widely used in medical reports of case-control studies. The odds ratio(Fisher, 1935, p. 50) is defined(Cox, 1958) as the ratio of the odds of an event occurring in one group with respect to the odds of its occurring in another group. Odds(Yule and Pearson, 1900, p. 273) ratio (OR) is a measure of association which quantifies the relationship between two binomial distributed random variables (exposure vs. outcome) and is related to Yule's (Yule and Pearson, 1900, p. 272) Q(Yule, 1912, p. 585/586). Two events A_t and B_t are regarded as independent if $(A_t, B_t) = 1$. Let

 a_t = number of persons exposed to A_t and with disease B_t

 b_t = number of persons exposed to A_t but without disease \underline{B}_t

 c_t = number of persons unexposed \underline{A}_t but with disease B_t

 d_t = number of persons unexposed \underline{A}_t : and without disease \underline{B}_t

 a_t+c_t = total number of persons with disease B_t (case-patients)

 b_t+d_t = total number of persons without disease \underline{B}_t (controls).

Hereafter, consider the table 6. The odds' ratio (OR) is defined as

Table 6. The two by two table of random variables

		Conditioned/Outcome B _t		
		TRUE	FALSE	
Condition/Exposure	TRUE	a _t	b _t	$\overline{A_t}$
A_{t}	FALSE	c_{t}	d_t	$\underline{\mathbf{A}}_{t}$
		B _t	$\underline{\mathbf{B}}_{t}$	N_{t}

$$OR(A_{t}, B_{t}) \equiv \left(\frac{a_{t}}{b_{t}}\right) / \left(\frac{c_{t}}{d_{t}}\right)$$

$$\equiv \left(\frac{a_{t} \times d_{t}}{b_{t} \times c_{t}}\right)$$
(101)

Remark 2.1. Odds ratios can support logical fallacies and cause difficulties in drawing logically consistent conclusions. The chorus of voices is growing, which demand the immediate ending(Knol, 2012, Sackett, DL and Deeks, JJ and Altman, DG, 1996) of any use of Odds ratio.

Under conditions where (b = 0), the measure of association odds ratio will collapse, because we need to divide by zero, as can be seen at eq. 101. However, according to today's rules of mathematics,

a division by zero is neither allowed nor generally accepted as possible. It does no harm to remind ourselves that in the case b = 0 the event A_t is a sufficient condition of B_t . In other words, odds ratio is not able to recognize elementary relationships of objective reality. In fact, it would be a failure not to recognize how dangerous and less valuable odds ratio is.

Under conditions where (c = 0) odds ratio collapses too, because we need again to divide by zero, as can be seen at eq. 101. However, and again, today's rules of mathematics don't allow us a division by zero. In point of fact, in the case c = 0 it is more than necessary to point out that A_t is a necessary condition of B_t . In other words, odds ratio or the cross-product ratio is not able to recognize elementary relationships of nature like necessary conditions. We can and need to overcome all the epistemological obstacles as backed by odds ratio entirety. Sooner rather than later, we should give up this measure of relationship completely.

2.1.10. Relative risk (RR)

2.1.10.1. Relative risk (RR_{nc})

Definition 2.16 (Relative risk (RR_{nc})).

The degree of association between the two binomial variables can be assessed by a number of very different coefficients, the relative (Cornfield, 1951, Sadowsky et al., 1953) risk is one(Barukčić, 2021d) of them. In general, relative risk RR_{nc}, which provides some evidence of a necessary condition, is defined as

$$RR(A_{t}, B_{t})_{nc} \equiv \frac{\frac{p(a_{t})}{p(A_{t})}}{\frac{p(c_{t})}{p(NotA_{t})}}$$

$$\equiv \frac{p(a_{t}) \times p(NotA_{t})}{p(c_{t}) \times p(A_{t})}$$

$$\equiv \frac{N \times p(a_{t}) \times N \times p(NotA_{t})}{N \times p(c_{t}) \times N \times p(A_{t})}$$

$$\equiv \frac{a_{t} \times (NotA_{t})}{c_{t} \times A_{t}}$$

$$\equiv \frac{EER(A_{t}, B_{t})}{CER(A_{t}, B_{t})}$$
(102)

That what scientist generally understand by relative risk is the ratio of a probability of an event occurring with an exposure versus the probability of an event occurring without an exposure. In other words,

relative risk = (probability(event in exposed group)) / (probability(the same event in not exposed group)).

A $RR(A_t, B_t)$ = +1 means that exposure does not affect the outcome or both are independent of each other while $RR(A_t, B_t)$ less than +1 means that the risk of the outcome is decreased by the exposure. In this context, an $RR(A_t, B_t)$ greater than +1 denotes that the risk of the outcome is increased by the exposure. Widely known problems with odds ratio and relative risk are already documented in literature.

2.1.10.2. Relative risk (RR (sc))

Definition 2.17 (Relative risk (RR (sc))).

The relative risk (sc), which provides some evidence of a sufficient condition, is calculated from the point of view of an outcome and is defined as

$$RR(A_{t}, B_{t})_{sc} \equiv \frac{\frac{p(a_{t})}{p(B_{t})}}{\frac{p(b_{t})}{p(NotB_{t})}}$$

$$\equiv \frac{p(a_{t}) \times p(NotB_{t})}{p(b_{t}) \times p(B_{t})}$$

$$\equiv \frac{N \times p(a_{t}) \times N \times p(NotB_{t})}{N \times p(b_{t}) \times N \times p(B_{t})}$$

$$\equiv \frac{a_{t} \times (NotB_{t})}{b_{t} \times B_{t}}$$

$$\equiv \frac{OPR(A_{t}, B_{t})}{CPR(A_{t}, B_{t})}$$
(103)

2.1.10.3. Relative risk reduction (RRR)

Definition 2.18 (Relative risk reduction (RRR)).

$$RRR(A_{t}, B_{t}) \equiv \frac{CER(A_{t}, B_{t}) - EER(A_{t}, B_{t})}{CER(A_{t}, B_{t})}$$

$$= 1 - RR(A_{t}, B_{t})$$
(104)

2.1.10.4. Vaccine efficacy (VE)

Definition 2.19 (Vaccine efficacy (VE)).

Vaccine efficacy is defined as the percentage reduction of a disease in a vaccinated group of people as compared to an unvaccinated group of people.

$$VE(A_{t}, B_{t}) \equiv 100 \times (1 - RR(A_{t}, B_{t}))$$

$$\equiv 100 \times \left(\frac{CER(A_{t}, B_{t}) - EER(A_{t}, B_{t})}{CER(A_{t}, B_{t})}\right)$$
(105)

Historically, vaccine efficacy has been designed to evaluate the efficacy of a certain vaccine by Greenwood and Yule in 1915 for the cholera and typhoid vaccines(Greenwood and Yule, 1915) and best measured using double-blind, randomized, clinical controlled trials. However, the calculated vaccine efficacy is depending too much on the study design, can lead to erroneous conclusions and is only of very limited value.

2.1.10.5. Experimental event rate (EER)

Definition 2.20 (Experimental event rate (EER)).

$$EER(A_t, B_t) \equiv \frac{p(a_t)}{p(A_t)} = \frac{a_t}{a_t + b_t}$$
(106)

Definition 2.21 (Control event rate (CER)).

$$CER(A_{t}, B_{t}) \equiv \frac{p(c_{t})}{p(\underline{A}_{t})} = \frac{c_{t}}{c_{t} + d_{t}}$$
(107)

2.1.10.6. Absolute risk reduction (ARR)

Definition 2.22 (Absolute risk reducation (ARR)).

$$ARR(A_{t}, B_{t}) \equiv \frac{p(c_{t})}{p(\underline{A}_{t})} - \frac{p(a_{t})}{p(A_{t})}$$

$$= \frac{c_{t}}{c_{t} + d_{t}} - \frac{a_{t}}{a_{t} + b_{t}}$$

$$= CER(A_{t}, B_{t}) - EER(A_{t}, B_{t})$$
(108)

2.1.10.7. Absolute risk increase (ARI)

Definition 2.23 (Absolute risk increase (ARI)).

$$ARI(A_{t}, B_{t}) \equiv \frac{p(a_{t})}{p(A_{t})} - \frac{p(c_{t})}{p(\underline{A}_{t})}$$

$$= EER(A_{t}, B_{t}) - CER(A_{t}, B_{t})$$
(109)

2.1.10.8. Number needed to treat (NNT)

Definition 2.24 (Number needed to treat (NNT)).

$$NNT(A_{t}, B_{t}) \equiv \frac{1}{CER(A_{t}, B_{t}) - EER(A_{t}, B_{t})}$$
(110)

An ideal number needed to treat(Cook and Sackett, 1995, Laupacis et al., 1988), mathematically the reciprocal of the absolute risk reduction, is NNT = 1. Under these circumstances, everyone improves with a treatment, while no one improves with control. A higher number needed to treat indicates more or less a treatment which is less effective.

2.1.10.9. Number needed to harm (NNH)

Definition 2.25 (Number needed to harm (NNH)).

$$NNH\left(A_{t}, B_{t}\right) \equiv \frac{1}{EER\left(A_{t}, B_{t}\right) - CER\left(A_{t}, B_{t}\right)} \tag{111}$$

The number needed to harm (Massel and Cruickshank, 2002), mathematically the inverse of the absolute risk increase, indicates at the end how many patients need to be exposed to a certain factor, in order to observe a harm in one patient that would not otherwise have been harmed.

2.1.10.10. Outcome prevalence rate (OPR)

Definition 2.26 (Outcome prevalence rate (OPR)).

$$OPR(A_t, B_t) \equiv \frac{p(a_t)}{p(B_t)} = \frac{a_t}{a_t + c_t}$$
(112)

2.1.10.11. Control prevalence rate (CPR)

Definition 2.27 (Control prevalence rate (CPR)).

$$CPR(A_{t}, B_{t}) \equiv \frac{p(b_{t})}{p(B_{t})} = \frac{b_{t}}{b_{t} + d_{t}}$$

$$(113)$$

Bias and confounding is present to some degree in all research. In order to assess the relationship of exposure with a disease or an outcome, a fictive control group (i.e. of newborn or of young children et cetera) can be of use too. Under certain circumstances, even a CPR = 0 is imaginable.

2.1.10.12. Absolute prevalence reduction (APR)

Definition 2.28 (Absolute prevalence reduction (APR)).

$$APR(A_t, B_t) \equiv CPR(A_t, B_t) - OPR(A_t, B_t)$$
(114)

2.1.10.13. Absolute prevalence increase (API)

Definition 2.29 (Absolute prevalence increase (API)).

$$API(A_{t}, B_{t}) \equiv OPR(A_{t}, B_{t}) - CPR(A_{t}, B_{t})$$
(115)

2.1.10.14. Relative prevalence reduction (RPR)

Definition 2.30 (Relative prevalence reduction (RPR)).

$$RPR(A_{t}, B_{t}) \equiv \frac{CPR(A_{t}, B_{t}) - OPR(A_{t}, B_{t})}{CPR(A_{t}, B_{t})}$$

$$= 1 - RR(A_{t}, B_{t})_{sc}$$
(116)

2.1.10.15. The index NNS

Definition 2.31 (The index NNS).

$$NNS(A_{t}, B_{t}) \equiv \frac{1}{CPR(A_{t}, B_{t}) - OPR(A_{t}, B_{t})}$$

$$(117)$$

Mathematically, the index NNS is the reciprocal of the absolute prevalence reduction.

2.1.10.16. The index NNI

Definition 2.32 (The index NNI).

$$NNI(A_{t}, B_{t}) \equiv \frac{1}{OPR(A_{t}, B_{t}) - CPR(A_{t}, B_{t})}$$

$$(118)$$

Mathematically, the index NNI is the reciprocal of the absolute prevalence increase.

2.1.11. Index of relationship (IOR)

Definition 2.33 (Index of relationship (IOR)).

Due to several reasons, it is not always easy to identify the unique characteristics between two events like A_t and B_t . And more than that, it is difficult to decide what to do, and much more difficult to know in which direction one should think and which decision is right. Sometimes it is helpful to know at least something about the direction of the relationship between two events like A_t and B_t . Under conditions where $p(a_t) = p(A_t \wedge B_t)$, the index of relationship(Barukčić, 2021b), abbreviated as IOR, is defined as

$$IOR(A_{t}, B_{t}) \equiv \left(\frac{p(A_{t} \wedge B_{t})}{p(B_{t}) \times p(A_{t})}\right) - 1$$

$$\equiv \left(\frac{p(a_{t})}{p(B_{t}) \times p(A_{t})}\right) - 1$$

$$\equiv \left(\left(\frac{N \times N \times p(a_{t})}{N \times p(B_{t}) \times N \times p(A_{t})}\right) - 1\right)$$

$$\equiv \left(\left(\frac{N \times a}{A \times B}\right) - 1\right)$$
(119)

where $p(A_t)$ denotes the probability of an event A_t at the Bernoulli trial t and $p(B_t)$ denotes the probability of another event B_t at the same Bernoulli trial t while $p(a_t)$ denotes the joint probability of $p(A_t \text{ AND } B_t)$ at the same Bernoulli trial t and a, A and B may denote the expectation values.

Definition 2.34 (Multi dimensional index of relationship (NIOR)).

The multi dimensional index of relationship (NIOR) is defined as

$$NIOR(A_{t}, B_{t}) \equiv \left(\frac{N^{k} \times p({}_{1}A_{t} \wedge {}_{2}A_{t} \cdots {}_{k}A_{t})}{N \times (p({}_{1}A_{t})) N \times (p({}_{2}A_{t})) \cdots N \times (p({}_{k}A_{t}))}\right) - 1$$

$$\equiv \left(\frac{N^{k-1} \times E({}_{1}A_{t} \wedge {}_{2}A_{t} \cdots {}_{k}A_{t})}{E({}_{1}A_{t}) \times E({}_{2}A_{t}) \cdots \times E({}_{k}A_{t})}\right) - 1$$

$$(120)$$

where N is the sample size and $p({}_{1}A_{t} \wedge {}_{2}A_{t} \cdots {}_{k}A_{t})$ is the joint distribution function.

However, there might exist circumstances where a multi dimensional index of relationship might take the form of the following equation.

$$NIOR(A_{t}, B_{t}) \equiv \left(\frac{{}_{1}N \times {}_{2}N \times \cdots {}_{k}N \times p({}_{1}A_{t} \wedge {}_{2}A_{t} \cdots {}_{k}A_{t})}{({}_{1}N \times p({}_{1}A_{t})) \times ({}_{2}N \times p({}_{2}A_{t})) \cdots \times ({}_{k}N \times p({}_{k}A_{t}))}\right) - 1$$

$$\equiv \left(\frac{{}_{1}N \times {}_{2}N \times \cdots {}_{k}N \times p({}_{1}A_{t} \wedge {}_{2}A_{t} \cdots {}_{k}A_{t})}{E({}_{1}A_{t}) \times E({}_{2}A_{t}) \cdots \times E({}_{k}A_{t})}\right) - 1$$

$$(121)$$

2.2. Conditions

Even if a condition and a cause are deeply related, there are circumstances where a sharp distinction between a cause and a condition is necessary. However, exactly this has been denied by John Stuart Mill's (1806-1873) regularity view of causality (see Mill, 1843b). What might seem to be a theoretical difficulty for many authors is none for Mill. Mill simply reduced a cause to a condition and claimed that "... the real cause of the phenomenon is the assemblage of all its conditions." (see Mill, 1843a, p. 403)

2.2.1. Exclusion relationship

Definition 2.35 (Exclusion relationship [*EXCL*]).

Mathematically, the exclusion(see also Barukčić, 2021a) relationship 15 (EXCL), denoted by p(A_t | B_t) in terms of statistics and probability theory, is defined(see also Barukčić, 1989, p. 68-70) as

$$p(A_{t} | B_{t}) \equiv p(A_{t} \uparrow B_{t})$$

$$\equiv p(b_{t}) + p(c_{t}) + p(d_{t})$$

$$\equiv \frac{N \times (p(b_{t}) + p(c_{t}) + p(d_{t}))}{N}$$

$$\equiv \frac{\sum_{t=1}^{N} (\underline{A}_{t} \vee \underline{B}_{t})}{N} \equiv \frac{b + c + d}{N}$$

$$\equiv \frac{b + \underline{A}}{N}$$

$$\equiv \frac{c + \underline{B}}{N}$$

$$\equiv +1$$

$$(122)$$

Based on the 1913 Henry Maurice Sheffer (1882-1964) relationship, the Sheffer stroke(Nicod, 1917, Sheffer, 1913) usually denoted by \uparrow , it is $p(A_t \land B_t) \equiv 1 - p(A_t \mid B_t)$ (see table 7).

Table 7. A_t excludes B_t and vice versa.

	Conditioned (COVID-19) B _t			
		TRUE	FALSE	
Condition (Vaccine)	TRUE	+0	p(b _t)	$p(A_t)$
A_{t}	FALSE	$p(c_t)$	$p(d_t)$	$p(\underline{A}_t)$
		$p(B_t)$	$p(\underline{B}_t)$	+1

¹⁵Barukčić, Ilija. (2021). Mutually exclusive events. Causation, 16(11), 5–57. https://doi.org/10.5281/zenodo.5746415

Example 2.1. Pfizer Inc. and BioNTech SE announced on Monday, November 09, 2020 - 06:45am results from a Phase 3 COVID-19 vaccine trial with 43.538 participants which provides evidence that their vaccine (BNT162b2) is preventing COVID-19 in participants without evidence of prior SARS-CoV-2 infection. In toto, 170 confirmed cases of COVID-19 were evaluated, with 8 in the vaccine group versus 162 in the placebo group. The exclusion relationship can be calculated as follows.

$$p(Vaccine : BNT 162b2 \mid COVID - 19(infection)) \equiv p(b_t) + p(c_t) + p(d_t)$$

$$\equiv 1 - p(a_t)$$

$$\equiv 1 - \left(\frac{8}{43538}\right)$$

$$\equiv +0.99981625$$
(123)

with a P Value = 0,000184.

Following Kolmogorov's definition of an n-dimensional probability density (see also Kolmogorov, Andreĭ Nikolaevich, 1950, p. 26) of random variables A_t, B_t et cetera at the point t, we obtain

$$p(A_{t} | B_{t}) \equiv p(\underline{A}_{t} \cup \underline{B}_{t})$$

$$\equiv 1 - p(A_{t} \cap B_{t})$$

$$\equiv 1 - \int_{-\infty}^{A_{t}} \int_{-\infty}^{B_{t}} f(A_{t}, B_{t}) dA_{t} dB_{t}$$

$$\equiv +1$$
(124)

while $p(A_t | B_t)$ would denote the cumulative distribution function of random variables and $f(A_t, B_t)$ is the joint density function.

2.2.2. Observational study and exclusion relationship

Under conditions of an observational study, the exclusion relationship follows approximately(see Barukčić, 2021a) as

$$p(A_t \mid B_t) \equiv p(A_t \uparrow B_t) \ge 1 - \frac{p(a_t)}{p(B_t)}$$
(125)

2.2.3. Experimental study and exclusion relationship

Under conditions of an experimental study, the exclusion relationship follows approximately(see Barukčić, 2021a) as

$$p(A_t \mid B_t) \equiv p(A_t \uparrow B_t) \ge 1 - \frac{p(a_t)}{p(A_t)}$$
(126)

2.2.4. The goodness of fit test of an exclusion relationship

Definition 2.36 (The $\tilde{\chi}^2$ goodness of fit test of an exclusion relationship).

Under some well known circumstances, testing hypothesis about an exclusion relationship $p(A_t \mid B_t)$ is possible by the chi-square distribution (also chi-squared or $\tilde{\chi}^2$ -distribution) too. The $\tilde{\chi}^2$ goodness of fit test of an exclusion relationship with degree of freedom (d. f.) of d. f. = 1 is calculated as

$$\tilde{\chi}^{2}_{\text{Calculated}}((A_{t} \mid B_{t}) \mid A) \equiv \frac{(b - (a + b))^{2}}{A} + \frac{((c + d) - \underline{A})^{2}}{\underline{A}}$$

$$\equiv \frac{a^{2}}{A} + 0$$

$$\equiv \frac{a^{2}}{A}$$

$$(127)$$

or equally as

$$\tilde{\chi}^{2}_{\text{Calculated}}\left(\left(A_{t} \mid B_{t}\right) \mid B\right) \equiv \frac{\left(c - (a + c)\right)^{2}}{B} + \frac{\left(\left(b + d\right) - \underline{B}\right)^{2}}{\underline{B}}$$

$$\equiv \frac{a^{2}}{B} + 0$$

$$\equiv \frac{a^{2}}{B}$$
(128)

and can be compared with a theoretical chi-square value at a certain level of significance α . The $\tilde{\chi}^2$ -distribution equals zero when the observed values are equal to the expected/theoretical values of an exclusion relationship/distribution $p(A_t \mid B_t)$, in which case the null hypothesis has to be accepted. Yate's (Yates, 1934) continuity correction was not used under these circumstances.

2.2.5. The left-tailed p Value of an exclusion relationship

Definition 2.37 (The left-tailed p Value of an exclusion relationship).

It is known that as a sample size, N, increases, a sampling distribution of a special test statistic approaches the normal distribution (central limit theorem). Under these circumstances, the left-tailed

(lt) p Value (Barukčić, 2019d) of an exclusion relationship can be calculated as follows.

$$pValue_{lt}(A_t \mid B_t) \equiv 1 - e^{-(1 - p(A_t \mid B_t))}$$

$$\equiv 1 - e^{-(a/N)}$$
(129)

A low p-value may provide some evidence of statistical significance.

2.2.6. Neither nor conditions

Definition 2.38 (Neither A_t nor B_t conditions [NOR]).

Mathematically, a neither A_t nor B_t condition (or rejection according to the French philosopher and logician Jean George Pierre Nicod (1893-1924), i.e. Jean Nicod's statement (Nicod, 1924)) relationship (NOR), denoted by $p(A_t \downarrow B_t)$ in terms of statistics and probability theory, is defined (Barukčić, 1989, p. 68-70) as

$$p(A_{t} \downarrow B_{t}) \equiv p(d_{t})$$

$$\equiv \frac{N - \sum_{t=1}^{N} (A_{t} \lor B_{t})}{N} \equiv \frac{\sum_{t=1}^{N} (\underline{A}_{t} \land \underline{B}_{t})}{N} \equiv \frac{N \times (p(d_{t}))}{N}$$

$$\equiv \frac{d}{N}$$

$$\equiv +1$$
(130)

2.2.7. The Chi square goodness of fit test of a neither nor condition relationship

Definition 2.39 (The $\tilde{\chi}^2$ goodness of fit test of a neither A_t nor B_t condition relationship).

A neither A_t nor B_t condition relationship $p(A_t \downarrow B_t)$ can be tested by the chi-square distribution (also chi-squared or $\tilde{\chi}^2$ -distribution). The $\tilde{\chi}^2$ goodness of fit test of a neither A_t nor B_t condition relationship with degree of freedom (d. f.) of d. f. = 1 may be calculated as

$$\tilde{\chi}^{2}_{\text{Calculated}}((A_{t} \downarrow B_{t}) \mid A) \equiv \frac{(d - (c + d))^{2}}{\frac{\underline{A}}{A}} + \frac{((a + b) - A)^{2}}{A}$$

$$\equiv \frac{c^{2}}{\underline{A}} + 0$$
(131)

or equally as

$$\tilde{\chi}^{2}_{\text{Calculated}}((A_{t} \downarrow B_{t}) \mid B) \equiv \frac{(d - (b + d))^{2}}{\underline{B}} + \frac{((a + c) - B)^{2}}{B}$$

$$\equiv \frac{b^{2}}{\underline{B}} + 0$$
(132)

Yate's (Yates, 1934) continuity correction has not been used in this context.

2.2.8. The left-tailed p Value of a neither nor B condition relationship

Definition 2.40 (The left-tailed p Value of a neither A_t nor B_t condition relationship).

The left-tailed (lt) p Value (Barukčić, 2019d) of a neither A_t nor B_t condition relationship can be calculated as follows.

$$pValue_{lt}(A_{t} \downarrow B_{t}) \equiv 1 - e^{-(1 - p(A_{t} \downarrow B_{t}))}$$

$$\equiv 1 - e^{-p(A_{t} \lor B_{t})}$$

$$\equiv 1 - e^{-((a+b+c)/N)}$$
(133)

where \vee may denote disjunction or logical inclusive or. In this context, a low p-value indicates again a statistical significance. In general, it is $p(A_t \vee B_t) \equiv 1 - p(A_t \downarrow B_t)$ (see table 8).

Table 8. Neither A_t nor B_t relationship.

	Conditioned B _t			
		YES	NO	
Condition A _t	YES	0	0	0
	NO	0	1	1
		0	1	1

2.2.9. Necessary condition

Definition 2.41 (Necessary condition [Conditio sine qua non]).

Despite the most extended efforts, the current state of research on conditions and conditioned is still incomplete and very contradictory. However, even thousands of years ago and independently of any human mind and consciousness, water has been and is still a necessary (see Barukčić, 2022b) condition for (human) life. **Without** water, there has been and there is **no** (human) life¹⁶. It comes therefore as no surprise that one of the first documented attempts to present a rigorous theory of conditions and causation (see also Aristotle, of Stageira (384-322 B.C.E), 1908, Metaphysica III 2 997a 10 and 13/14) came from the Greek philosopher and scientist Aristotle (384-322 BCE). Thus far, it is amazing that Aristotle himself made already a strict distinction between conditions and causes. Taking Aristotle very seriously, it is necessary to consider that

"... everything which has a potency in question has the potency ... of acting ... not in all circumstances but on certain conditions ... " (see also Aristotle, of Stageira (384-322 B.C.E), 1908, Metaphysica IX 5 1048a 14-19)

Before going into details, Aristotle went on to define the necessary condition as follows.

```
"... necessary ... means ...
without ... a condition, a thing cannot live ... "
```

(see also Aristotle, of Stageira (384-322 B.C.E), 1908, Metaphysica V 2 1015a 20-22)

In point of fact, Aristotle developed a theory of conditions and causality commonly referred to as the doctrine of four causes. Many aspects and general features of Aristotle's logical concept of causality are meanwhile extensively and critically debated in secondary literature. However, even if the Greek philosophers Heraclitus, Plato, Aristotle et cetera numbers among the greatest philosophers of all time, the philosophy has evolved. Scientific knowledge and objective reality are deeply interrelated and cannot be reduced only to Greek philosophers like Aristotle. Among many other issues, the specification of necessary conditions has traditionally been part of the philosopher's investigations of different phenomena. However, behind the need of a detailed evidence, it is justified to consider that philosophy or philosophers as such certainly do not possess **a monopoly on the truth** and other areas such as medicine as well as other sciences and technology may transmit truths as well and may be of help to move beyond one's self enclosed unit. Seemingly, **the law's concept of causation** justifies to say few words on this subject, to put some light on some questions. Are there any criteria in law for deciding whether one action or an event A_t has caused another (generally harmful) event B_t? What are these criteria? May causation in legal contexts differ from causation outside the law, for example, in science

¹⁶Barukčić, Ilija. (2022). Conditio sine qua non (Version 1). Zenodo. https://doi.org/10.5281/zenodo.5854744

or in our everyday life and to what extent? Under which circumstances is it justified to tolerate such differences as may be found to exist? To understand just what is the law's concept of causation, it is useful to re-consider how the highest court of states is dealing with causation. In the case Hayes v. Michigan Central R. Co., 111 U.S. 228, the U.S. Supreme Court defined 1884 conditio sine qua non as follows: "... causa sine qua non - a cause which, if it had not existed, the injury would not have taken place". (Justice Matthews, Mr., 1884) The German Bundesgerichtshof für Strafsachen stressed once again the importance of conditio sine qua non relationship in his decision by defining the following: "Ursache eines strafrechtlich bedeutsamen Erfolges jede Bedingung, die nicht hinweggedacht werden kann, ohne daß der Erfolg entfiele" (Bundesgerichtshof für Strafsachen, 1951) Another lawyer elaborated on the basic issue of identity and difference between cause and condition. Von Bar was writing: "Die erste Voraussetzung, welche erforderlich ist, damit eine Erscheinung als die Ursache einer anderen bezeichnet werden könne, ist, daß jene eine der Bedingungen dieser sein. Würde die zweite Erscheinung auch dann eingetreten sein, wenn die erste nicht vorhanden war, so ist sie in keinem Falle Bedingung und noch weniger Ursache. Wo immer ein Kausalzusammenhang behauptet wird, da muß er wenigstens diese Probe aushalten ... Jede Ursache ist nothwendig auch eine Bedingung eines Ereignisses; aber nicht jede Bedingung ist Ursache zu nennen."(Bar, Carl Ludwig von, 1871) Von Bar's position translated into English: The first requirement, which is required, thus that something could be called as the cause of another, is that the one has to be one of the conditions of the other. If the second something had occurred even if the first one did not exist, so it is by no means a condition and still less a cause. Wherever a causal relationship is claimed, the same must at least withstand this test... Every cause is necessarily also a condition of an event too; but not every condition is cause too. Thus far, let us consider among other the following in order to specify necessary conditions from another, probabilistic point of view. An event (i.e. A_t) which is a necessary condition of another event or outcome (i.e. B_t) must be given, must be present for a conditioned, for an event or for an outcome B_t to occur. A necessary condition (i.e. A_t) is a requirement which need to be fulfilled at every single Bernoulli trial t, in order for a conditioned or an outcome (i.e. B_t) to occur, but it alone does not determine the occurrence of such an event. In other words, if a necessary condition (i.e. A_t) is given, an outcome (i.e. B_t) need not to occur. In contrast to a necessary condition, a 'sufficient' condition is the one condition which 'guarantees' that an outcome will take place or will occur for sure. Under which conditions we may infer about the unobserved and whether observations made are able at all to justify predictions about potential observations which have not yet been made or even general claims which my go even beyond the observed (the 'problem of induction') is not the issue of the discussion at this point. Besides of the principal necessity of meeting such a challenge, a necessary condition of an event can but need not be at the same Bernoulli trial t a sufficient condition for an event to occur. However, theoretically, it is possible that an event or an outcome is determined by many necessary conditions. Let us focus to some extent on what this means, or in other words how much importance can we attribute to such a special case. Example. A human being cannot live without oxygen. A human being cannot live without water. A human being cannot live without a brain. A human being cannot live without kidneys. A human being cannot live without ... et cetera. Thus far, even if oxygen is given, if a brain is given ... et cetera, without water a human being will not survive on the long run. This example is of use to reach the following conclusion. Although it might seem somewhat paradoxical at first sight, even under circumstances where a condition or an outcome depends on several different necessary conditions it is particularly important that every single of these necessary conditions for itself must be given otherwise the conditioned (i.e. the outcome) will not occur. Mathematically, the necessary condition (SINE) relationship, denoted by $p(A_t \leftarrow B_t)$ in terms of statistics and probability theory, is defined (Barukčić, 1989, p. 15-28) as

$$p(A_{t} \leftarrow B_{t}) \equiv p(A_{t} \vee \underline{B}_{t}) \equiv \frac{\sum_{t=1}^{N} (A_{t} \vee \underline{B}_{t})}{N} \equiv \frac{(A_{t} \vee \underline{B}_{t}) \times p(A_{t} \vee \underline{B}_{t})}{(A_{t} \vee \underline{B}_{t})}$$

$$\equiv p(a_{t}) + p(b_{t}) + p(d_{t})$$

$$\equiv \frac{N \times (p(a_{t}) + p(b_{t}) + p(d_{t}))}{N} \equiv \frac{E(A_{t} \leftarrow B_{t})}{N}$$

$$\equiv \frac{a + b + d}{N} \equiv \frac{E(A_{t} \vee \underline{B}_{t})}{N}$$

$$\equiv \frac{A + d}{N} \equiv \frac{E(A_{t} \leftarrow B_{t})}{N}$$

$$\equiv \frac{a + \underline{B}}{N} \equiv \frac{E(A_{t} \vee \underline{B}_{t})}{N}$$

$$\equiv +1$$

$$(134)$$

where $E(A_t \leftarrow B_t) \equiv E(A_t \lor \underline{B}_t)$ indicates the expectation value of the necessary condition. In general, it is $p(A_t \multimap B_t) \equiv 1 - p(A_t \leftarrow B_t)$ (see Table 9).

Table 9. Necessary condition.

	Conditioned B _t			
		TRUE	FALSE	
Condition	TRUE	p(a _t)	p(b _t)	p(A _t)
A_t	FALSE	+0	$p(d_t)$	$p(\underline{A}_t)$
		p(B _t)	$p(\underline{B}_t)$	+1

A necessary condition A_t is characterised itself by the property that another event B_t will not occur if A_t is not given, if A_t did not occur (Barukčić, 1989, 1997, 2005, 2016b, 2017b,c, 2020a,b,c,d, Barukčić and Ufuoma, 2020a). Taking into account Kolmogorov's definition of an n-dimensional probability density (see also Kolmogorov, Andreĭ Nikolaevich, 1950, p. 26) of random variables A_t , B_t et cetera at the (period of) time t, we obtain

$$p(A_{t} \leftarrow B_{t}) \equiv +1$$

$$\equiv +1 - p(c_{t})$$

$$\equiv +1 - p(\underline{A}_{t} \cap B_{t})$$

$$\equiv \left(\int_{-\infty}^{A_{t}} \int_{-\infty}^{B_{t}} f(A_{t}, B_{t}) dA_{t} dB_{t}\right) + \left(1 - \int_{-\infty}^{B_{t}} f(B_{t}) dB_{t}\right)$$
(135)

while $p(A_t \leftarrow B_t)$ would denote the cumulative distribution function of random variables of a necessary condition. Another adequate formulation of a necessary condition is possible too. If certain conditions

are met, then necessary conditions and sufficient conditions are one way or another converses of each other, too. It is

$$p(A_{t} \leftarrow B_{t}) \equiv \underbrace{(A_{t} \vee \underline{B_{t}})}_{\text{(Necessary condition)}} \equiv \underbrace{(\underline{B_{t}} \vee A_{t})}_{\text{(Sufficient condition)}} \equiv p(B_{t} \rightarrow A_{t})$$
(136)

These relationships are illustrated by the following tables.

Table 10. Without A_t no B_t

 $\begin{array}{c|cccc} & & & & B_t \\ & & TRUE & FALSE \\ \hline A_t & TRUE & a_t & b_t & A_t \\ & FALSE & c_t = 0 & d_t & \underline{A}_t \\ \hline & B_t & \underline{B}_t & +1 \\ \hline \end{array}$

Table 11. If B_t then A_t

		1		
		TRUE	FALSE	
$\overline{\mathbf{B}_{t}}$	TRUE	a _t	$c_t = 0$	B _t
	FALSE	b_t	d_t	$\underline{\mathbf{B}}_{t}$
		A_t	$\underline{\mathbf{A}}_{t}$	+1

There are circumstances under which

$$p(A_{t} \leftarrow B_{t}) \equiv \underbrace{(A_{t} \vee \underline{B}_{t})}_{\text{(Nessessary condition)}} \equiv \underbrace{(\underline{A}_{t} \vee B_{t})}_{\text{(Sufficient condition)}} \equiv p(A_{t} \rightarrow B_{t})$$

$$(137)$$

However, equation 136 does not imply the relationship of equation 137 under any circumstances.

Example I.

A wax candle is characterised by various properties, but is also subject to certain conditions. **Without** sufficient amounts of gaseous oxygen **no** burning wax candle, gaseous oxygen is a necessary condition of a burning candle. However, the converse relationship **if** burning wax candle, **then** sufficient amounts of gaseous oxygen are given is is at the same (period of) time t / Bernoulli trial t true. The following tables are illustrating these relationships.

Table 12. Without gaseous oxygen no burning candle

	Burning candle			
		TRUE	FALSE	
Gaseous	TRUE	a _t	b _t	A _t
oxygen	FALSE	$c_t = 0$	d_t	$\underline{\mathbf{A}}_{t}$
		B_t	$\underline{\mathbf{B}}_{t}$	+1

Table 13. If burning candle then gaseous oxygen

		Gaseou	s oxygen	
		TRUE	FALSE	
Burning	TRUE	a _t	$c_t = 0$	B _t
candle	FALSE	b_t	d_t	$\underline{\mathbf{B}}_{t}$
		At	At	+1

Example II.

Once again, a human being cannot live without water. A human being cannot live without gaseous oxygen, et cetera. Water itself is a necessary condition for human life. However, gaseous oxygen is a necessary condition for human life too. Thus far, even if water is given and even if water is a necessary condition for human life, without gaseous oxygen there will be no human life. In general, if a conditioned or an outcome B_t depends on the necessary condition A_t and equally on numerous other

necessary conditions, an event B_t will not occur if A_t itself is not given independently of the occurrence of other necessary conditions.

Example III.

Another different aspect of a necessary condition relationship is appropriate to be focused upon here. As a direct consequence of a necessary condition **without** sufficient amounts of gaseous oxygen **no** burning wax candle is a special case of an exclusion relationship. The absence of sufficient amounts of gaseous oxygen A_t excludes (see Barukčić, 2021a) a burning wax candle B_t . Thus far, if we want to stop the burning of a wax candle, we would have to significantly reduce the amounts of gaseous oxygen A_t . Under these conditions, a wax candle will stop burning. The following tables (table 14 and table 15) may illustrate this aspect of a necessary condition in more detail.

Table 14. Without gaseous oxygen no burning candle

	Burning candle			
		TRUE	FALSE	
Gaseous	TRUE	a _t	b _t	At
oxygen	FALSE	$c_t = 0$	d_t	$\underline{\mathbf{A}}_{t}$
		B _t	$\underline{\mathbf{B}}_{t}$	+1

Table 15. Absent gaseous oxygen excludes burning wax candle

	Burning candle			
		TRUE	FALSE	
Gaseous	FALSE	$c_t = 0$	d _t	B _t
oxygen	TRUE	a_{t}	b_t	$\underline{\mathbf{B}}_{t}$
		At	At	+1

The necessary condition relationship follows approximately (see Barukčić, 2022b) as

$$p(A_{\mathsf{t}} \leftarrow B_{\mathsf{t}}) \ge 1 - \frac{p(c_{\mathsf{t}})}{p(B_{\mathsf{t}})} \tag{138}$$

and as

$$p(A_{\mathsf{t}} \leftarrow B_{\mathsf{t}}) \ge 1 - \frac{p(c_{\mathsf{t}})}{p(A_{\mathsf{t}})} \tag{139}$$

2.2.10. The Chi-square goodness of fit test of a necessary condition relationship

Definition 2.42 (The $\tilde{\chi}^2$ goodness of fit test of a necessary condition relationship).

Under some well known circumstances, hypothesis about the conditio sine qua non relationship $p(A_t \leftarrow B_t)$ can be tested by the chi-square distribution (also chi-squared or χ^2 -distribution), first described by the German statistician Friedrich Robert Helmert (Helmert, 1876) and later rediscovered by Karl Pearson (Pearson, 1900) in the context of a goodness of fit test. The $\tilde{\chi}^2$ goodness of fit test of a conditio sine qua non relationship with degree of freedom (d. f.) of d. f. = 1 is calculated as

$$\tilde{\chi}^{2}_{\text{Calculated}}(A_{t} \leftarrow B_{t} \mid B) \equiv \frac{(a - (a + c))^{2}}{B} + \frac{((b + d) - \underline{B})^{2}}{\underline{B}}$$

$$\equiv \frac{c^{2}}{B} + 0$$

$$\equiv \frac{c^{2}}{B}$$

$$\equiv \frac{c^{2}}{B}$$
(140)

or equally as

$$\tilde{\chi}^{2}_{\text{Calculated}}(A_{t} \leftarrow B_{t} \mid \underline{A}) \equiv \frac{(d - (c + d))^{2}}{\underline{A}} + \frac{((a + b) - A)^{2}}{A}$$

$$\equiv \frac{c^{2}}{\underline{A}} + 0$$

$$\equiv \frac{c^{2}}{A}$$
(141)

and can be compared with a theoretical chi-square value at a certain level of significance α . It has not yet been finally clarified whether the use of Yate's (Yates, 1934) continuity correction is necessary at all.

2.2.11. The left-tailed p Value of the conditio sine qua non relationship

Definition 2.43 (The left-tailed p Value of the conditio sine qua non relationship).

The left-tailed (lt) p Value (Barukčić, 2019d) of the conditio sine qua non relationship can be calculated as follows.

$$pValue_{lt}(A_t \leftarrow B_t) \equiv 1 - e^{-(1 - p(A_t \leftarrow B_t))}$$

$$\equiv 1 - e^{-(c/N)}$$
(142)

2.2.12. Sufficient condition

Definition 2.44 (Sufficient condition [Conditio per quam]).

Mathematically, the sufficient (Barukčić, 2021c, p. 68-70) condition (see Barukčić, 2022a) (IMP) relationship, denoted by $p(A_t \to B_t)$ in terms of statistics and probability theory, is defined (Barukčić, 1989, p. 68-70) as

$$p(A_{t} \to B_{t}) \equiv p(\underline{A}_{t} \lor B_{t}) \equiv \frac{\sum_{t=1}^{N} (\underline{A}_{t} \lor B_{t})}{N} \equiv \frac{(\underline{A}_{t} \lor B_{t}) \times p(\underline{A}_{t} \lor B_{t})}{(\underline{A}_{t} \lor B_{t})}$$

$$\equiv p(a_{t}) + p(c_{t}) + p(d_{t})$$

$$\underbrace{\frac{N \times (p(a_{t}) + p(c_{t}) + p(d_{t}))}{N}}_{N}$$

$$\equiv \frac{a + c + d}{N} \equiv \frac{E(\underline{A}_{t} \lor B_{t})}{N}$$

$$\equiv \frac{B + d}{N} \equiv \frac{E(A_{t} \to B_{t})}{N}$$

$$\equiv \frac{a + \underline{A}}{N}$$

$$\equiv +1$$

$$(143)$$

In general, it is $p(A_t > -B_t) \equiv 1 - p(A_t \rightarrow B_t)$ (see Table 16).

2.2.12.1. Mackie's INUS Condition John Leslie Mackie (1917-1981) critically examined the theories of causation of various (see Ducasse, 1926) philosophers such as Hume (Book I, Part III, of the Treatise) (see Mackie, 1974, pp. 3-28), Kant (as well as Kantian approaches offered by Strawson and Bennett), Mill and other. Mackie rightly claims that Hume's regularity theory of causation offer only an incomplete picture of the nature of causation. Mackie writes: "It seems appropriate to begin by examining and criticizing it, so that we can take over from it whatever seems to be defensible but develop an improved account by correcting its errors and deficiencies." (see Mackie, 1974, p. 3). Nonetheless, in his trial to develop an improved account of Hume's theory of causation, Mackie's own account of the nature of causation follows Hume's principles of causation very closely (see Mackie, 1974, pp. 3-28). Mackie himself proposed already in 1965 that "the so-called cause is ... an insufficient but necessary part of a condition which is itself unnecessary but sufficient for the result ... let us call such a condition ... an INUS condition." (see Mackie, 1965, p. 245). However Mackie's account needs modification, and can be modified and when it is modified we can explain much more satisfactorily what Mackie ordinarily take to be a cause. Mackie is of the opinion that "... cause is ... part of a condition ... " (see Mackie, 1965, p. 245) and that "... a condition ... is ... unnecessary but sufficient for the result [i. e. effect, author]. " (see Mackie, 1965, p. 245). To put it very simply one could say that Mackie reduces a cause to a sufficient condition, "... cause is ... a condition which is itself ... sufficient ... " (see Mackie, 1965, p. 245). Indeed, there are circumstances, where several

different events ¹⁷ might be necessary or sufficient et cetera at the same time in order to determine a compound/complex sufficient condition relationship. Thus far, it seems appropriate to take over from Mackie's INUS condition whatever seems to be acceptable but to develop an improved account by correcting its deficiencies and errors in order to do justice to the complexity of affairs. Equation 144 illustrates one real-world example of a compound/complex sufficient condition relationship in more detail.

$$p(((_{1}X_{t} \wedge_{2}X_{t} \wedge_{3}X_{t} \wedge \cdots) \wedge A_{t}) \rightarrow B_{t}) \equiv p\left(\underbrace{((_{1}X_{t} \wedge_{2}X_{t} \wedge_{3}X_{t} \wedge \cdots) \wedge A_{t})}_{N} \vee B_{t}\right)$$

$$\equiv \sum_{t=1}^{N} \underbrace{\left(\underbrace{((_{1}X_{t} \wedge_{2}X_{t} \wedge_{3}X_{t} \wedge \cdots) \wedge A_{t})}_{N} \vee B_{t}\right)}_{N}$$

$$\equiv +1$$

$$(144)$$

Again, taking into account Kolmogorov's definition of an n-dimensional probability density (see also Kolmogorov, Andreĭ Nikolaevich, 1950, p. 26) of random variables A_t , B_t et cetera at the (period of) time t, we obtain

$$p(A_{t} \to B_{t}) \equiv +1$$

$$\equiv +1 - p(b_{t})$$

$$\equiv +1 - p(A_{t} \cap \underline{B}_{t})$$

$$\equiv \left(\int_{-\infty}^{A_{t}} \int_{-\infty}^{B_{t}} f(A_{t}, B_{t}) dA_{t} dB_{t}\right) + \left(1 - \int_{-\infty}^{A_{t}} f(A_{t}) dA_{t}\right)$$

$$(145)$$

while $p(A_t \to B_t)$ would denote the cumulative distribution function of random variables of a sufficient condition. Another adequate formulation of a sufficient condition is possible too.

Table 16. Sufficient condition.

	Conditioned B _t			
		TRUE	FALSE	
Condition	TRUE	p(a _t)	+0	p(A _t)
A_{t}	FALSE	$p(c_t)$	$p(d_t)$	$p(\underline{A}_t)$
		p(B _t)	$p(\underline{B}_t)$	+1

Remark 2.2. A sufficient condition A_t is characterized by the property that another event B_t will occur if A_t is given, if A_t itself occured (Barukčić, 1989, 1997, 2005, 2016b, 2017b,c, 2020a,b,c,d, Barukčić and Ufuoma, 2020a). **Example**. The ground, the streets, the trees, human beings and many other objects too will become wet during heavy rain. Especially, **if** it is raining (event A_t), **then** human beings will become wet (event B_t). However, even if this is a common human wisdom, a human being equipped with an appropriate umbrella (denoted by R_t) need not become wet even during heavy rain. An appropriate umbrella (R_t) is similar to an event with the potential to counteract the occurrence of

¹⁷Barukčić, Ilija. (2022). Conditio per quam. Causation, 17(3), 5–86. https://doi.org/10.5281/zenodo.6369831

another event (B_t) and can be understood something as an **anti-dot** of another event. In other words, an appropriate umbrella is an antidote of the effect of rain on human body, an appropriate umbrella has the potential to protect humans from the effect of rain on their body. It is a good rule of thumb that the following relationship

$$p(A_t \to B_t) + p(R_t \land B_t) \equiv +1 \tag{146}$$

indicates that R_t is an antidote of A_t . However, taking a shower, swimming in a lake et cetera may make human hair wet too. More than anything else, however, these events does not affect the final outcome, the effect of raining on human body.

The approximate (see Barukčić, 2022a) value of the material implication is given as

$$p(A_{\mathsf{t}} \to B_{\mathsf{t}}) \ge 1 - \frac{p(b_{\mathsf{t}})}{p(A_{\mathsf{t}})} \tag{147}$$

and alternatively as

$$p(A_{t} \to B_{t}) \ge 1 - \frac{p(b_{t})}{p(\underline{B}_{t})} \tag{148}$$

2.2.13. The Chi square goodness of fit test of a sufficient condition relationship

Definition 2.45 (The $\tilde{\chi}^2$ goodness of fit test of a sufficient condition relationship).

Under some well known circumstances, testing hypothesis about the conditio per quam relationship $p(A_t \to B_t)$ is possible by the chi-square distribution (also chi-squared or $\tilde{\chi}^2$ -distribution) too. The $\tilde{\chi}^2$ goodness of fit test of a conditio per quam relationship with degree of freedom (d. f.) of d. f. = 1 is calculated as

$$\tilde{\chi}^{2}_{\text{Calculated}}(A_{t} \to B_{t} \mid A) \equiv \frac{(a - (a + b))^{2}}{A} + \frac{((c + d) - \underline{A})^{2}}{\underline{A}}$$

$$\equiv \frac{b^{2}}{A} + 0$$

$$\equiv \frac{b^{2}}{A}$$

$$(149)$$

or equally as

$$\tilde{\chi}^{2}_{\text{Calculated}}(A_{t} \to B_{t} \mid \underline{B}) \equiv \frac{(d - (b + d))^{2}}{\underline{B}} + \frac{((a + c) - B)^{2}}{B}$$

$$\equiv \frac{b^{2}}{\underline{B}} + 0$$

$$\equiv \frac{b^{2}}{\underline{B}}$$

$$(150)$$

and can be compared with a theoretical chi-square value at a certain level of significance α . The $\tilde{\chi}^2$ -distribution equals zero when the observed values are equal to the expected/theoretical values of the conditio per quam relationship/distribution $p(A_t \to B_t)$, in which case the null hypothesis is accepted. Yate's (Yates, 1934) continuity correction has not been used in this context.

2.2.14. The left-tailed p Value of the conditio per quam relationship

Definition 2.46 (The left-tailed p Value of the conditio per quam relationship).

The left-tailed (lt) p Value (Barukčić, 2019d) of the conditio per quam relationship can be calculated as follows.

$$pValue_{lt}(A_t \to B_t) \equiv 1 - e^{-(1 - p(A_t \to B_t))}$$

$$\equiv 1 - e^{-(b/N)}$$
(151)

Again, a low p-value indicates a statistical significance.

2.2.15. Necessary and sufficient conditions

Definition 2.47 (Necessary and sufficient conditions [EQV]).

The necessary and sufficient condition (EQV) relationship, denoted by $p(A_t \leftrightarrow B_t)$ in terms of

statistics and probability theory, is defined(Barukčić, 1989, p. 68-70) as

$$p(A_{t} \leftrightarrow B_{t}) \equiv \frac{\sum_{t=1}^{N} ((A_{t} \vee \underline{B}_{t}) \wedge (\underline{A}_{t} \vee B_{t}))}{N}$$

$$\equiv p(a_{t}) + p(d_{t})$$

$$\equiv \frac{N \times (p(a_{t}) + p(d_{t}))}{N}$$

$$\equiv \frac{a+d}{N}$$

$$\equiv +1$$

$$(152)$$

2.2.16. The Chi square goodness of fit test of a necessary and sufficient condition relationship

Definition 2.48 (The $\tilde{\chi}^2$ goodness of fit test of a necessary and sufficient condition relationship).

Even the necessary and sufficient condition relationship $p(A_t \leftrightarrow B_t)$ can be tested by the chi-square distribution (also chi-squared or $\tilde{\chi}^2$ -distribution) too. The $\tilde{\chi}^2$ goodness of fit test of a necessary and sufficient condition relationship with degree of freedom (d. f.) of d. f. = 1 is calculated as

$$\tilde{\chi}^{2}_{\text{Calculated}}(A_{t} \leftrightarrow B_{t} \mid A) \equiv \frac{(a - (a + b))^{2}}{A} + \frac{d - ((c + d))^{2}}{\frac{A}{A}}$$

$$\equiv \frac{b^{2}}{A} + \frac{c^{2}}{A}$$
(153)

or equally as

$$\tilde{\chi}^{2}_{\text{Calculated}}(A_{t} \leftrightarrow B_{t} \mid B) \equiv \frac{(a - (a + c))^{2}}{B} + \frac{d - ((b + d))^{2}}{\underline{B}}$$

$$\equiv \frac{c^{2}}{B} + \frac{b^{2}}{B}$$
(154)

The calculated $\tilde{\chi}^2$ goodness of fit test of a necessary and sufficient condition relationship can be compared with a theoretical chi-square value at a certain level of significance α . Under conditions where the observed values are equal to the expected/theoretical values of a necessary and sufficient condition relationship/distribution $p(A_t \leftrightarrow B_t)$, the $\tilde{\chi}^2$ -distribution equals zero. It is to be cleared whether Yate's (Yates, 1934) continuity correction should be used at all.

2.2.17. The left-tailed p Value of a necessary and sufficient condition relationship

Definition 2.49 (The left-tailed p Value of a necessary and sufficient condition relationship).

The left-tailed (lt) p Value (Barukčić, 2019d) of a necessary and sufficient condition relationship can be calculated as follows.

$$pValue_{lt}(A_t \leftrightarrow B_t) \equiv 1 - e^{-(1 - p(A_t \leftrightarrow B_t))}$$

$$\equiv 1 - e^{-((b+c)/N)}$$
(155)

In this context, a low p-value indicates again a statistical significance. Table 17 may provide an overview of the theoretical distribution of a necessary and sufficient condition.

Table 17. Necessary and sufficient condition.

2.2.18. Either or conditions

Definition 2.50 (Either A_t or B_t conditions [NEQV]).

Mathematically, an either A_t or B_t condition relationship (NEQV), denoted by $p(A_t > < B_t)$ in terms of statistics and probability theory, is defined(Barukčić, 1989, p. 68-70) as

$$p(A_{t} > < B_{t}) \equiv \frac{\sum_{t=1}^{N} ((A_{t} \wedge \underline{B}_{t}) \vee (\underline{A}_{t} \wedge B_{t}))}{N}$$

$$\equiv p(b_{t}) + p(c_{t})$$

$$\equiv \frac{N \times (p(b_{t}) + p(c_{t}))}{N}$$

$$\equiv \frac{b+c}{N}$$

$$\equiv +1$$

$$(156)$$

It is $p(A_t > < B_t) \equiv 1 - p(A_t < > B_t)$ (see Table 18).

Table 18. Either A_t or B_t relationship.

	Conditioned B _t			
		YES	NO	
Condition A _t	YES	0	1	1
	NO	1	0	1
		1	1	2

2.2.19. The Chi-square goodness of fit test of an either or condition relationship

Definition 2.51 (The $\tilde{\chi}^2$ goodness of fit test of an either or condition relationship).

An either or condition relationship $p(A_t > < B_t)$ can be tested by the chi-square distribution (also chi-squared or $\tilde{\chi}^2$ -distribution) too. The $\tilde{\chi}^2$ goodness of fit test of an either or condition relationship with degree of freedom (d. f.) of d. f. = 1 is calculated as

$$\tilde{\chi}^{2}_{\text{Calculated}}((A_{t} > < B_{t}) | A) \equiv \frac{(b - (a+b))^{2}}{A} + \frac{c - ((c+d))^{2}}{\frac{A}{A}}$$

$$\equiv \frac{a^{2}}{A} + \frac{d^{2}}{A}$$
(157)

or equally as

$$\tilde{\chi}^{2}_{\text{Calculated}}\left(\left(A_{t} > < B_{t}\right) \mid B\right) \equiv \frac{\left(c - (a + c)\right)^{2}}{B} + \frac{b - \left(\left(b + d\right)\right)^{2}}{\frac{B}{B}}$$

$$\equiv \frac{a^{2}}{B} + \frac{d^{2}}{B}$$
(158)

Yate's (Yates, 1934) continuity correction has not been used in this context.

2.2.20. The left-tailed p Value of an either or condition relationship

Definition 2.52 (The left-tailed p Value of an either or condition relationship).

The left-tailed (lt) p Value (Barukčić, 2019d) of an either or condition relationship can be calculated as follows.

$$pValue_{lt}(A_{t} > < B_{t}) \equiv 1 - e^{-(1 - p(A_{t} > - < B_{t}))}$$

$$\equiv 1 - e^{-((a+d)/N)}$$
(159)

In this context, a low p-value indicates again a statistical significance.

2.3. Causation

2.3.1. Causation in general

The history of the denialism of causality in Philosophy, Mathematics, Statistics, Physics et cetera is very long. We only recall David Hume's (1711-1776) account of causation and his inappropriate reduction of the cause-effect relationship to a simple habitual connection in human thinking or Immanuel Kant's (1724-1804) initiated trial to consider causality as nothing more but a 'a priori' given category (Langsam, 1994) in human reasoning and other similar attempts too.

It is worth noting in this context that especially Karl Pearson (1857 - 1936) himself has been engaged in a long lasting and never-ending crusade against causation too. "Pearson categorically denies the need for an independent concept of causal relation beyond correlation ... he *exterminated* causation from statistics before it had a chance to take root "(see Pearl, 2000, p. 340).

At the beginning of the 20th century notable proponents of **conditionalism** like the German anatomist and pathologist David Paul von Hansemann (Hansemann, David Paul von, 1912) (1858 - 1920) and the biologist and physiologist Max Richard Constantin Verworn(Verworn, 1912) (1863 - 1921) started a new attack(Kröber, 1961) on the principle of causality. In his essay "Kausale und konditionale Weltanschauung" Verworn(Verworn, 1912) presented "an exposition of 'conditionism' as contrasted with 'causalism,' (Unknown, 1913) while ignoring cause and effect relationships completely. "**Das Ding ist also identisch mit der Gesamtheit seiner Bedingungen**." (Verworn, 1912) However, Verworn's goal to exterminate causality completely out of science was hindered by the further development of research.

The history of futile attempts to refute **the principle of causality** culminated in a publication by the German born physicist Werner Karl Heisenberg (1901 - 1976). Heisenberg put forward an illogical, inconsistent and confusing uncertainty principle which opened the door to wishful thinking and logical fallacies in physics and in science as such. Heisenberg's unjustified reasoning ended in an act of a manifestly unfounded conclusion: "Weil alle Experimente den Gesetzen der Quantenmechanik und damit der Gleichung (1) unterworfen sind, so wird durch die Quantenmechanik die Ungültigkeit des Kausalgesetzes definitiv festgestellt." (Heisenberg, Werner Karl, 1927) while 'Gleichung (1)'denotes Heisenberg's uncertainty principle. Einstein's himself, a major contributor to quantum theory and in the same respect a major critic of quantum theory, disliked Heisenberg's uncertainty principle fundamentally while Einstein's opponents used Heisenberg's Uncertainty Principle against Einstein. After the End of the German Nazi initiated Second World War with unimaginable brutality and high human losses and a death toll due to an industrially organised mass killing of people by the German Nazis which did not exist in this way before, Werner Heisenberg visited Einstein in Princeton (New Jersey, USA) in October 1954 (Neffe, 2006). Einstein agreed to meet Heisenberg only for a very short period of time but their encounter lasted longer. However, there where not only a number of differences between Einstein and Heisenberg, these two physicists did not really loved each other. "Einstein remarked that the inventor of the uncertainty principle was a 'big Nazi'... "(Neffe, 2006) Albert Einstein (1879 - 1955) took again the opportunity to refuse to endorse **Heisenberg's uncertainty principle**

as a fundamental law of nature and rightly too. Meanwhile, Heisenberg's uncertainty principle is refuted (see Barukčić, 2011a, 2014, 2016a) for several times but still not exterminated completely out of physics and out of science as such.

In contrast to such extreme anti-causal positions as advocated by Heisenberg and the **Copenhagen interpretation of quantum mechancis**, the search for a (mathematical) solution of *the issue of causal inferences* is as old as human mankind itself ("*i. e. Aristotle's Doctrine of the Four Causes*") (Hennig, 2009) even if there is still little to go on.

It is appropriate to specify especially the position of D'Holbach (Holbach, Paul Henri Thiry Baron de, 1770). D'Holbach (1723-1789) himself linked cause and effect or causality as such to changes. "Une cause, est un être qui e met un autre en mouvement, ou qui produit quelque changement en lui. L'effet est le changement qu'un corps produit dans un autre ..." (Holbach, Paul Henri Thiry Baron de, 1770). D'Holbach infers in the following: "De l'action et de la réaction continuelle de tous les êtres que la nature renferme, il résulte une suite de causes et d'effets ..." (Holbach, Paul Henri Thiry Baron de, 1770).

With more or less meaningless or none progress on the matter in hand even in the best possible conditions, it is not surprising that authors are suggesting more and more different approaches and models for causal inference. Indeed, the hope is justified that logically consistent *statistical methods* of causal inference can help scientist to achieve so much with so little.

One of the methods of causal inference in Bio-sciences are based on the known Henle(Henle, 1840) (1809–1885) - Koch(Koch, 1878) (1843–1910) postulates (Carter, 1985) which are applied especially for the identification of a causative agent of an (infectious) disease. However, the pathogenesis of most chronic diseases is more or less very complex and potentially involves the interaction of several factors. In practice, from the 'pure culture' requirement of the Henle-Koch postulates insurmountable difficulties may emerge. In light of subsequent developments (PCR methodology, immune antibodies et cetera) it is appropriate to review the full validity of the Henle-Koch postulates in our days.

In 1965, Sir Austin Bradford Hill (Hill, 1965) published nine criteria (the 'Bradford Hill Criteria') in order to determine whether observed epidemiological associations are causal. Somewhat worrying, is at least the fact that, Hill's "... fourth characteristic is the temporal relationship of the association" and so-to-speak just a reformulation of the 'post hoc ergo propter hoc'(Barukčić, 1989, Woods and Walton, 1977) logical fallacy through the back-door and much more then this. It is questionable whether association as such can be treated as being identical with causation. Unfortunately, due to several reasons, it seems therefore rather problematic to rely on Bradford Hill Criteria carelessly.

Meanwhile, several other and competing mathematical or statistical approaches for causal inference have been discussed by various modern authors (Barukčić, 1989, 1997, 2005, 2016b, 2017a,c, Bohr, 1937, Chisholm, 1946, Dempster, 1990, Espejo, 2007, Goodman, 1947, Granger, 1969, Hessen, Johannes, 1928, Hesslow, 1976, 1981, Korch, Helmut, 1965, Lewis, David Kellogg, 1973, 1974, Pearl, 2000, Schlick, Friedrich Albert Moritz, 1931, Spohn, 1983, Suppes, 1970, Todd, 1968, Zesar, 2013) or even established (Barukčić, 1989, 1997, 2005, 2016b, 2017a,c). Nevertheless, the question is still

not answered, is it at all possible to establish a cause effect relationship between two factors while applying only certain statistical (Sober, 2001) methods?

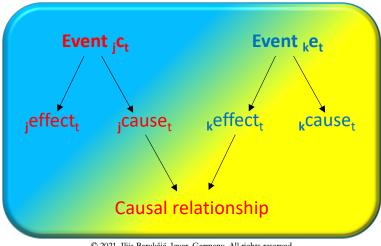
2.3.2. Cause and effect

Besides all, there are several further aspects of causation for which our attention so far has not been adequately fixed in this context. In the causal relationship, cause and effect are united, a cause is an effect and an effect is a cause.

"Thus, in the causal relation, cause and effect are inseparable; a cause which had no effect would not be a cause, just as an effect which had no cause would no longer be an effect."

(see Hegel, Georg Wilhelm Friedrich, 1991, p. 151)

The unity of cause and effect is a unity of two which are not the same. Cause and effect as inseparable in the causal relation are at the same time mutually related as sheer others; each of both as united in its own self to the other of itself is able to passes over into its own other and vice versa. Yet, to approach from a different point of view, a cause and an effect are separated in the same relation too, a cause is not an effect and an effect is not a cause, both are different in the same relation.



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"Therefore, though the cause has an effect and is at the same time itself effect,

and the effect not only has a cause but is also itself cause.

yet the effect which the cause has, and the effect which it is, are different.

as are also the cause which the effect has, and the cause which it is." (see Hegel, Georg Wilhelm Friedrich, 1991, p. 565/566)

2.3.2.1. What is a cause, what is an effect? An important fact to which we must pay attention here is that in a causal relation, under certain circumstances, an individual cause and an individual effect are related to each other in their own particular way. An effect which vanishes in its own cause in the same respect equally becomes again in it and vice versa. A cause which is merely extinguished in its own effect becomes again in the same. In fact, each of these determinations presupposes in its own other its own self and constitutes the intimate tie between an individual cause and its own individual

effect. Thus far, under conditions of a **positive** causal relationship k, an event U_t which is for sure a cause of another event W_t is at the same time t a necessary and sufficient condition of an event W_t . Table 19 may illustrate this relationship. A matter of great theoretical importance is the fundamental

Table 19. What is the cause, what is the effect?

	Effect W _t			
		TRUE	FALSE	
Cause	TRUE	+1	+0	p(U _t)
U_t	FALSE	+0	+1	$p(\underline{\boldsymbol{U}}_t)$
		$p(W_t)$	$p(\underline{W}_t)$	+1

relationship between a cause and a condition. Are both, cause and condition, at the end identical? As of now, following Mill (see Mill, 1843a, p. 403), Verworn (see Verworn, 1912), Mackie and others, we can give a clear 'Yes'in reply to this question: "... cause is ... a condition which is itself ... *sufficient* ... " (see Mackie, 1965, p. 245). However, this issue is not as simple as it sounds, according to Mackie. Thus far, it is essential to eliminate some errors. Indeed, there are circumstances where a cause and a condition are identical, a cause and a condition are equivalent. However, as outlined in this publication, both, a cause and a condition, are different too and a cause and a condition are not identical either.

"Jede Ursache ist nothwendig auch eine Bedingung eines Ereignisses; aber nicht jede Bedingung ist Ursache zu nennen."

(see Bar, Carl Ludwig von, 1871, p. 4)

The crux of the matter is that not every condition is a cause too, in German: "... nicht jede Bedingung ist Ursache ... "(see Bar, Carl Ludwig von, 1871, p. 4). However, and in contrast to a condition, every cause as such is indeed a condition too, in German: "Jede Ursache ist ... auch eine Bedingung ... "(see Bar, Carl Ludwig von, 1871, p. 4). In general, a cause U_t is a necessary condition of an effect W_t . In other words, **without** a cause U_t **no** effect W_t . One consequence of the necessary condition relationship between cause and effect is that "... an effect which had no cause would no longer be an effect." (see Hegel, Georg Wilhelm Friedrich, 1991, p. 151). However, a cause U_t being a necessary condition of an effect W_t is equivalent to an effect W_t being a sufficient condition of the same cause U_t and vice versa too. In our everyday words,

without

 U_t

no

 W_t

is equivalent with

if

 W_t

then

 U_t

and vice versa. As can be seen, there is a kind of strange mirroring between U_t and W_t at the same Bernoulli trial t. Lastly, both are converses of each other too. In other words, U_t 's being a necessary condition of W_t 's is equivalent to W_t 's being a sufficient condition of U_t 's (and vice versa). In general, it is

$$(U_t \vee W_t) \equiv (W_t \vee U_t) \equiv ((U_t \vee W_t) \wedge (W_t \vee U_t)) \equiv +1 \tag{160}$$

	Effect W _t			
		TRUE	FALSE	
Cause	TRUE	a _t	b _t	$\overline{U_t}$
U_{t}	FALSE	$c_t = 0$	d_t	\underline{U}_t
		W _t	W _t	+1

Table 20. Without Ut no Wt

		Cau		
		TRUE	FALSE	
Effect	TRUE	a_t	$c_t = 0$	W_t
\mathbf{W}_{t}	FALSE	b_t	d_t	$\underline{\mathbf{W}}_t$
		Ut	Ut	+1

Table 21. If W_t then U_t

The other side of the causal relation at the same (period of) time / Bernoulli trial t is the fact that a cause U_t is equally a sufficient condition of an effect W_t too or shortly **if** cause U_t **then** effect W_t . One straightforward consequence of this fundamental relationship between a cause and an effect is that "... a cause which had no effect would not be a cause ..." (see Hegel, Georg Wilhelm Friedrich, 1991, p. 151). But even this is not without difficulties, because a cause U_t being a sufficient condition of an effect W_t is equivalent to effect W_t being a necessary condition of the same cause U_t . In different words,

if

 U_t

then

 W_t

is equivalent with

without

 W_t

no

U_t.

		Effect W _t				
		TRUE	FALSE			
Cause	TRUE	a _t	$b_t = 0$	Ut		
Ut	FALSE	c_{t}	d_t	$\underline{\boldsymbol{U}}_t$		
		W_t	$\underline{\mathbf{W}}_{t}$	+1		

Table 22. If U_t then W_t

		Cause U _t				
		TRUE	FALSE			
Effect	TRUE	a _t	c _t	W _t		
\mathbf{W}_{t}	FALSE	$b_t = 0$	d_t	$\underline{\boldsymbol{W}}_t$		
		Ut	$\underline{\mathbf{U}}_{t}$	+1		

Table 23. Without W_t no U_t

To bring it to the point, necessary and sufficient conditions are at the end converses (see Gomes, Gilberto, 2009) of each other and far more than this. In fact, there is a kind of reciprocity or mirroring between cause and effect. Necessary and sufficient conditions are relationships used to describe the relationship between two events at the same Bernoulli trial t. In more detail, if U_t then W_t is equivalent with W_t is necessary for U_t , because the truth of U_t guarantees the truth of W_t . In general, it is

$$(U_{\mathsf{t}} \vee W_{\mathsf{t}}) \equiv (W_{\mathsf{t}} \vee U_{\mathsf{t}}) \equiv ((U_{\mathsf{t}} \vee W_{\mathsf{t}}) \wedge (W_{\mathsf{t}} \vee U_{\mathsf{t}})) \equiv +1 \tag{161}$$

In other words, it is impossible to have U_t without W_t (Bloch, 2011). Similarly, U_t is sufficient for W_t , because U_t being true always implies that W_t is true, but U_t not being true does not always imply that W_t is not true. And we should use this relationships to make our point. In general, **without** gaseous oxygen (U_t) , there is **no** burning wax candle (W_t) ; hence the relationship **if** burning wax candle (W_t) **then** gaseous oxygen (U_t) is equally true and given. This everyday knowledge is known and secured since centuries and might be illustrated as follows.

Wax candle B _t							Gaseous oxygen A _t		
		burning	not burning				present	not present	
Gaseous	present	a _t	b _t	A _t	Wax candle	burning	a _t	$c_t = 0$	B_t
oxygen _t	not present	$c_t = 0$	d_t	$\underline{\mathbf{A}}_{t}$		not burning	b_t	d_t	$\underline{\mathbf{B}}_{t}$
		B _t	$\underline{\mathbf{B}}_{\mathrm{t}}$	+1			A _t	$\underline{\mathbf{A}}_{t}$	+1
Table 24. Without A_t no B_t					Table	e 25. If B _t then	ı A _t		

Nonetheless, and independently of this secured everyday knowledge, a burning wax candle is a sufficient condition of gaseous oxygen but not the cause of gaseous oxygen.

Given all the circumstances, it is at least this simple **counter-example** which provides us with a convincing evidence that **a sufficient condition alone is not enough to describe a cause completely**. In general, a cause as such cannot be reduced to a simple sufficient condition.

In contrast to this obvious fact, other authors prefer another approach to the definition of a cause. "So that, more explicitly, if a given particular event is regarded as having been sufficient to the occurrence of another, it is said to have been its cause; if regarded as having been necessary to the occurrence of another, it is said to have been a condition of it; ..." (see Ducasse, 1926, p. 58). Therefore, in order

to be a cause of oxygen, additional evidence is necessary that a burning wax candle is a necessary condition of gaseous oxygen too. However, even if the relationship **without** gaseous oxygen **no** burning wax candle is given, this relationship is not given vice versa. The relationship **without** burning wax candle **no** gaseous oxygen is not given. Like other fundamental concepts, the concepts of cause and effect can be associated with difficulties too. Under certain conditions, the causal relationship between U_t and W_t , when correctly defined and recognised, is closely allied with the requirement that a certain study or that at least other, different studies provided evidence of a necessary condition between U_t and W_t and of a sufficient condition between U_t and W_t and if possible of **a necessary and sufficient condition** between U_t and W_t too.

Mathematically, a necessary and sufficient condition between U_t and W_t is defined as

$$(U_{\mathsf{t}} \vee \underline{W}_{\mathsf{t}}) \wedge (\underline{U}_{\mathsf{t}} \vee W_{\mathsf{t}}) \equiv +1 \tag{162}$$

However, I think it necessary to make a clear distinction between a necessary and sufficient condition and the converse relationship (Eq. 160) above.

$$((U_{\mathsf{t}} \vee \underline{W}_{\mathsf{t}}) \wedge (\underline{W}_{\mathsf{t}} \vee U_{\mathsf{t}})) \neq (U_{\mathsf{t}} \vee \underline{W}_{\mathsf{t}}) \wedge (\underline{U}_{\mathsf{t}} \vee W_{\mathsf{t}}) \tag{163}$$

2.3.2.2. The direction of causation In general, a cause is related to its own effect in its own way and vice versa (see Mackie, 1966, p. 160) too. The effect (see Black, 1956) of this cause is itself related to its own cause in some way in which the cause is not related to its own effect (see Dummett and Flew, 1954). This can be considered as one of the reasons why the relation between cause and effect is taken to be asymmetrical.

2.3.2.3. The priority of cause to effect Contemporary discussions of causation are greatly influenced by the causal relation that 'an effect W_t is causally dependent upon a cause U_t '. However, under certain conditions (mono-causality), to say that 'an effect W_t is causally dependent upon a cause U_t ' is to say that 'if a cause U_t had not occurred, then an effect W_t would not have occurred too.' (see Lewis, David Kellogg, 1973, 1974). However, what came first, the hen or the egg, the cause or the effect?

2.3.3. Definition causal relationship k

Definition 2.53 (Causal relationship k).

Nonetheless, mathematically, the causal(Barukčić, 2011a,b, 2012) relationship (Barukčić, 1989, 1997, 2005, 2016b, 2017a,c, 2021c) between a cause U_t (German: Ursache) and an effect W_t (German: Wirkung), denoted by $k(U_t, W_t)$, is defined *at each single(Thompson, 2006) Bernoulli trial t* in terms of statistics and probability theory 18 , 19 , 20 as

$$k(U_{t}, W_{t}) \equiv \frac{\sigma(U_{t}, W_{t})}{\sigma(U_{t}) \times \sigma(W_{t})}$$

$$\equiv \frac{p(U_{t} \wedge W_{t}) - p(U_{t}) \times p(W_{t})}{\sqrt[2]{(p(U_{t}) \times (1 - p(U_{t}))) \times (p(W_{t}) \times (1 - p(W_{t})))}}$$
(164)

where σ (U_t, W_t) denotes the co-variance between a cause U_t and an effect W_t at every single Bernoulli trial t, σ (U_t) denotes the standard deviation of a cause U_t at the same single Bernoulli trial t, σ (W_t) denotes the standard deviation of an effect W_t at same single Bernoulli trial t. Table 26 illustrates the theoretically possible relationships between a cause and an effect.

Table 26. Sample space and the causal relationship k

	Effect B _t				
		TRUE	FALSE		
Cause	TRUE	p(a _t)	p(b _t)	$p(U_t)$	
A_{t}	FALSE	$p(c_t)$	$p(d_t)$	$p(\underline{\boldsymbol{U}}_t)$	
		$p(W_t)$	$p(\underline{\mathbf{W}}_t)$	+1	

However, even if one thinks to recognise the trace of Bravais (Bravais, 1846) (1811-1863) - Pearson's (1857-1936) "product-moment coefficient of correlation" (Galton, 1877, Pearson, 1896) inside the causal relationship k (Barukčić, 1989, 1997, 2005, 2016b, 2017a,c) both are completely different. According to Pearson: "The fundamental theorems of correlation were for the first time and almost exhaustively discussed by B r a v a i s ('Analyse mathematique sur les probabilities des erreurs de situation d'un point.' Memoires par divers Savans, T. IX., Paris, 1846, pp. 255-332) nearly half a century ago." (Pearson, 1896) Neither does it make much sense to elaborate once again on the issue causation (Blalock, 1972) and correlation, since both are not identical (Sober, 2001) nor does it make sense to insist on the fact that "Pearson's philosophy discouraged him from looking too far behind phenomena." (Haldane, 1957) Whereas it is essential to consider that the causal relationship k, in contrast to Pearson's product-moment coefficient of correlation (Pearson, 1896) or to Pearson's phi

¹⁸Ilija Barukčić, "The Mathematical Formula of the Causal Relationship k," International Journal of Applied Physics and Mathematics vol. 6, no. 2, pp. 45-65, 2016. https://doi.org/10.17706/ijapm.2016.6.2.45-65

¹⁹Barukčić, Ilija. (2015). The Mathematical Formula Of The Causal Relationship k. https://doi.org/10.5281/zenodo.3944666
²⁰Ilija Barukčić. The causal relationship k. MATEC Web Conf., 336 (2021) 09032 DOI https://doi.org/10.1051/matecconf/202133609032

coefficient(Pearson, 1904b), is defined at every single Bernoulli trial t. This might be a very *small* difference. However, even a small difference might determine a difference. However, in this context and in any case, this small difference *makes*(Barukčić, 2018a) the difference.

2.4. Axioms

Whether science needs new and obviously generally valid statements (axioms) which are able to assure the truth of theorems proved from them may remain an unanswered question. In order to be accepted, a new axiom candidate (see Easwaran, 2008) should be at least as simple as possible and logically consistent to enable advances in our knowledge of nature. The importance of axioms is particularly emphasized by Albert Einstein. "Die wahrhaft großen Fortschritte der Naturerkenntnis sind auf einem der Induktion fast diametral entgegengesetzten Wege entstanden." (see Einstein, 1919, p. 17). In general, lex identitatis, lex contradictionis and lex negationis have the potential to denote the most simple, the most general and the most far-reaching axioms of science, the foundation of our today's and of our future scientific inquiry.

2.4.1. Principium identitatis (Axiom I)

Principium identitatis or **lex identitatis** or axiom I, is closely related to central problems of metaphysics, epistemology and of science as such. It turns out that it is more than rightful to assume that

$$+1 \equiv +1 \tag{165}$$

is true, otherwise there is every good reason to suppose that nothing can be discovered at all.

Identity as the epitome of a self-identical or of self-reference is at the same time different from difference, identity is free from difference, identity is not difference, identity is at the same time the other of itself, identity is non-identity. Identity as simple equality with itself is determined by a non-being, by a non-being of its own other, by a non-being of difference, identity is different from difference. Identity is in its very own nature different and is in its own self the opposite of itself (symmetry). It is equally

$$-1 \equiv -1 \tag{166}$$

In general, +1 and -1 are distinguished, however these distinct are related to one and the same 1. Identity as a vanishing of otherness, therefore, is this distinguishedness in one relation. It is

$$0 \equiv +1 - 1 \equiv 0 \times 1 \equiv 0 \tag{167}$$

Identity, as the unity of something and its own other is in its own self a separation from difference, and as a moment of separation might pass over into an equivalence relation which itself is reflexive, symmetric and transitive. Nonetheless, backed by thousands of years of often bitter human experience, the scientific development has taught us all that human knowledge is relative too. Even if experiments and other suitable proofs are of help to encourage us more and more in our belief of the correctness of a theory, it is difficult to prove the correctness of a theorem or of a theory et cetera once and for all. The challenge for all the science is the need to comply with Einstein's position: "Niemals aber kann die Wahrheit einer Theorie erwiesen werden. Denn niemals weiß man, daß auch in Zukunft eine Erfahrung bekannt werden wird, die Ihren Folgerungen widerspricht..." (Einstein, 1919).

Albert Einstein's position translated into English: 'But the truth of a theory can never be proven. For one never knows if future experience will contradict its conclusion; and furthermore, there are always other conceptual systems imaginable which might coordinate the very same facts.'Our human experience tells us that everything in life is more or less transitory, and that nothing lasts. As a result of our knowledge and experience, several scientific theories have a glorious past to look back on, but all the glory of such scientific theories might remain in the past if scientist don't continue to innovate. In a word, theories can be refuted by time.

"No amount of experimentation can ever prove me right; a single experiment can prove me wrong."

(Albert Einstein according to: Robertson, 1998, p. 114)

In the light of the foregoing, it is clear that appropriate axioms and conclusions derived from the same are a main logical foundation of any 'theory'.

"Grundgesetz (Axiome) und Folgerungen zusammen bilden das was man eine 'Theorie' nennt.

(Einstein, 1919)

However, another point is worth being considered again. One single experiment can be enough to refute a whole theory. Albert Einstein's (1879-1955) message translated into English as: *Basic law (axioms) and conclusions together form what is called a 'theory'* has still to get round. However, an axiom as a free creation of the human mind which precedes all science should be like all other axioms, as simple as possible and as self-evident as possible. Historically, the earliest documented use of **the law of identity** can be found in Plato's dialogue Theaetetus (185a) as "... each of the two is different from the other and the same as itself "21". However, Aristotle (384–322 B.C.E.), Plato's pupil and equally one of the greatest philosophers of all time, elaborated on the law of identity too. In Metaphysica, Aristotle wrote:

"... all things ... have some unity and identity."

(see Aristotle, of Stageira (384-322 B.C.E), 1908, Metaphysica, Chapter IV, 999a, 25-30, p. 66)

²¹Plato's dialogue Theaetetus (185a), p. 104.

In Prior Analytics, ²², ²³ Aristotle, a tutor of Alexander, the thirteen-year-old son of Philip, the king of Macedon, is writing: "When A applies to the whole of B and of C, and is other predicated of nothing else, and B also applies to all C, A and B must be convertible. For since A is stated only of B and C, and B is predicated both of itself and of C, it is evident that B will also be stated of all subjects of which A is stated, except A itself. "²⁴, ²⁵ For the sake of completeness, it should be noted at the outset that Aristotle himself preferred **the law of contradiction** and **the law of excluded middle** as examples of fundamental axioms. Nonetheless, it is worth noting that **lex identitatis** is an axiom too, which possess the potential to serve as the most basic and equally the most simple axiom of science but has been treated by Aristotle in an inadequate manner without having any clear and determined meaning for Aristotle himself. Nonetheless, something which is really just itself is equally different from everything else. In point of fact, is such an equivalence (Degen, 1741) which everything has to itself inherent or must the same be constructed by human mind and consciousness. Can and how can something be **identical with itself** (Förster and Melamed, 2012, Hegel, Georg Wilhelm Friedrich, 1812a, Koch, 1999, Newstadt, 2015) and in the same respect different from itself. An increasingly popular view on identity is the one advocated by Gottfried Wilhelm Leibniz (1646-1716):

"Chaque chose est ce qu'elle est. Et dans autant d'exemples qu'on voudra A est A,
B est B. "
(Leibniz, 1765, p. 327)

or $\mathbf{A} = \mathbf{A}$, $\mathbf{B} = \mathbf{B}$ or $+\mathbf{1} = +\mathbf{1}$. In other words, a thing is what it is (Leibniz, 1765, p. 327). Leibniz' **principium identitatis indiscernibilium** (p.i.i.), the principle of the indistinguishable, occupies a central position in Leibniz' logic and metaphysics and was formulated by Leibniz himself in different ways in different passages (1663, 1686, 1704, 1715/16). All in all, Leibniz writes:

"C'est
le principe des indiscernables,
en vertu duquel
il ne saurait exister dans la nature deux êtres identiques.

Il n'y a point deux individus indiscernables. " (see Leibniz, Gottfried Wilhelm, 1886, p. 45)

Exactly in complete compliance with Leibniz, Johann Gottlieb Fichte (1762 - 1814) elaborates on this subject as follows:

²²Aristotle, Prior Analytics, Book II, Part 22, 68a

²³Kenneth T. Barnes. Aristotle on Identity and Its Problems. Phronesis. Vol. 22, No. 1 (1977), pp. 48-62 (15 pages)

²⁴Aristotle, Prior Analytics, Book II, Part 22, 68a, p. 511.

²⁵Ivo Thomas. On a passage of Aristotle. Notre Dame J. Formal Logic 15(2): 347-348 (April 1974). DOI: 10.1305/ndjfl/1093891315

"Each thing is what it is; it has those realities which are posited when it is posited,

(A = A.) " (Fichte, 1889)

Georg Wilhelm Friedrich Hegel (1770 – 1831) himself objected the Law of Identity by claiming that "A = A is ... an empty tautology. "(see Hegel, Georg Wilhelm Friedrich, 1991, p. 413) provided an example of his own mechanical understanding of the Law of Identity. "the empty tautology: nothing is nothing; ... from nothing only nothing becomes ... nothing remains nothing. "(see Hegel, Georg Wilhelm Friedrich, 1991, p. 84). Nonetheless, Hegel preferred to reformulate an own version of Leibniz principium identitatis indiscernibilium in his own way by writing that "All things are different, or: there are no two things like each other. "(see Hegel, Georg Wilhelm Friedrich, 1991, p. 422). Much of the debate about identity is still a matter of controversy. This issue has attracted the attention of many authors and has been discussed by Hegel too. In this context, it is worth to consider Hegel's radical position on identity.

"The other expression of the law of identity: A cannot at the same time be A and not-A, has a negative form; it is called the law of contradiction."

(Hegel, Georg Wilhelm Friedrich, 1991, p. 416)

We may, usefully (see Barukčić, 2019a), state Russell's position with respect to the identity law as mentioned in his book 'The problems of philosophy' (see Russell, 1912). In particular, according to Russell.

"...principles have been singled out by tradition under the name of 'Laws of Thought.' They are as follows:

(1) **The law of identity**: 'Whatever is,is.

(2) The law of contradiction: 'Nothing can both be and not be.'

(3) The law of excluded middle: 'Everything must either be or not be.'

These three laws are samples of self-evident logical principles, but are not really more fundamental or more self-evident than various other similar principles: for instance, the one we considered just now, which states that what follows from a true premise is true. The name 'laws of thought' is also misleading, for what is important is not the fact that we think in accordance with these laws, but the fact that **things behave in accordance with them**; "

(see Russell, 1912, p. 113)

Russell's critique, that we tend too much to focus only on the formal aspects of the 'Laws of Thoughts' with the consequence that "... we thing in accordance with these laws" (see Russell, 1912, p. 113) is

justified. Judged solely in terms of this aspect, it is of course necessary to think in accordance with the 'Laws of Thoughts'. But this is not the only aspect of the 'Laws of Thoughts'. The other and may be much more important aspect of these 'Laws of Thoughts' is the fact that quantum mechanical objects or that "... things behave in accordance with them" (see Russell, 1912, p. 113).

2.4.2. Principium contradictionis (Axiom II)

Principium contradictionis or **lex contradictionis** ²⁶, ²⁷, ²⁸ or axiom II, the other of lex identitatis, the negative of lex identitatis, the opposite of lex identitatis, a complementary of lex identitatis, can be expressed mathematically as

$$+0 \equiv 0 \times 1 \equiv +1 \tag{168}$$

In addition to the above, from the point of view of mathematics, axiom II (equation 168) is equally the most simple mathematical expression and formulation of a contradiction. However, there is too much practical and theoretical evidence that a lot of 'secured' mathematical knowledge and rules differ too generously from real world processes, and the question may be asked whether mathematical truths can be treated as absolute truths at all. Many of the basic principle of today's mathematics allow every single author defining the real world events and processes et cetera in a way as everyone likes it for himself. Consequentially, a resulting dogmatic epistemological subjectivism and at the end agnosticism too, after all, is one of the reasons why we should rightly heed the following words of wisdom of Albert Einstein.

"I don't believe in mathematics."

(Albert Einstein cited according to Brian, 1996, p. 76)

In the long term, however, the above attitude of mathematics is not sustainable. History has taught us time and time again that objective reality has the potential to correct wrong human thinking slowly but surely, and many more than this. Objective reality has demonstrably corrected wrong human thinking again and again in the past.

²⁶Horn, Laurence R., "Contradiction", The Stanford Encyclopedia of Philosophy (Winter 2018 Edition), Edward N. Zalta (ed.), URL = https://plato.stanford.edu/archives/win2018/entries/contradiction/.

²⁷Barukčić I. Aristotle's law of contradiction and Einstein's special theory of relativity. Journal of Drug Delivery and Therapeutics (JDDT). 15Mar.2019;9(2):125-43. https://jddtonline.info/index.php/jddt/article/view/2389

²⁸Barukčić, Ilija. (2020, December 28). The contradiction is exsiting objectively and real (Version 1). Zenodo. https://doi.org/10.5281/zenodo.4396106

Despite all the adversities, it is necessary and crucial to consider that a self-identical as the opposite of itself is no longer only self-identity but a difference of itself from itself within itself. In other words, "All things are different, or: there are no two things like each other ... is, in fact, opposed to the law of identity ..." (see Hegel, Georg Wilhelm Friedrich, 1991, p. 422) Each on its own and without any respect to the other is distinctive within itself and from itself and not only from another. As the opposite of its own something, is no longer only self-identity, but also a negation of itself out of itself and therefore a difference of itself from itself within itself. In other words, in opposition, a self-identical is able to return into simple unity with itself, with the consequence that even as a self-identical the same self-identical is inherently self-contradictory. A question of fundamental theoretical importance is, however, why should something be itself and at the same time the other of itself, the opposite of itself, not itself? Is something like this even possible at all and if so, why and how? These and similar questions have occupied many thinkers, including Hegel.

"Something is therefore alive only in so far as it contains contradiction within it, and moreover is this power to hold and endure the contradiction within it."

(see Hegel, Georg Wilhelm Friedrich, 1991, p. 440)

However, as directed against identity, contradiction itself is also at the same time a source of self-changes of a self-identical out of itself.

"... contradiction
is the root of all movement and vitality;
it is only in so far as something has a contradiction within it
that it moves, has an urge and activity."

(see Hegel, Georg Wilhelm Friedrich, 1991, p. 439)

The further advance of science will throw any contribution to scientific progress of each of us back into scientific insignificance, as long as principium contradictionis is not given enough and the right attention. The contradiction ²⁹ is existing objectively and real and is the heartbeat of every self-identical. We have reason to be delighted by the fact that very different aspects of principium contradictionis have been examined since centuries from different angles by various authors. According to Aristotle, principium contradictionis applies to everything that is, it is the first and the firmest of all principles of philosophy.

²⁹Barukčić, Ilija. (2020, December 28). The contradiction is existing objectively and real (Version 1). Zenodo. https://doi.org/10.5281/zenodo.4396106

"... the same ... cannot at the same time belong and not belong to the same ... in the same respect ... This, then, is the most certain of all principles"

(see Aristotle, of Stageira (384-322 B.C.E), 1908, Metaph., IV, 3, 1005b, 16–22)

Principium contradictionis or axiom II has many facets. As long as we follow Leibniz in this regard, we should consider that "Le principe de contradiction est en general ... "(Leibniz, 1765, p. 327). Scientist inevitably have false beliefs and make mistakes. In order to prevent scientific results from falling into logical inconsistency or logical absurdity, it is necessary to posses among other the methodological possibility to start a reasoning with a (logical) contradiction too. However and in contrast to the way of reasoning with inconsistent premises as proposed by para-consistent (Carnielli and Marcos, 2001, da Costa, 1974, 1958, Priest, 1998, Priest et al., 1989, Quesada, 1977) and other logic, in the absence of technical and other errors of reasoning, the contradiction itself need to be preserved. In other words, from a contradiction does not anything follows but the contradiction itself while the theoretical question is indeed justified "What is so Bad about Contradictions?" (Priest, 1998). Historically, the principle of (deductive) explosion (Carnielli and Marcos, 2001, Priest, 1998, Priest et al., 1989), coined by 12th-century French philosopher William of Soissons, demand us to accept that anything, including its own negation, can be proven or can be inferred from a contradiction. In short, according to ex falso sequitur quodlibet, a (logical) contradiction implies anything. Respecting the principle of explosion, the existence of a contradiction (or the existence of logical inconsistency) in a scientific theorem, rule et cetera is disastrous. However, the historical development of science shows that scientist inevitably revise the theories, false positions and claims are identified once and again, and we all make different kind of mistakes. In order to avert disproportionately great damage to science and to prevent reducing science into pure subjective belief, a negation of the principle of explosion is required. Nonetheless, a justified negation of the ex contradictione quodlibet principle (Carnielli and Marcos, 2001) does not imply the correctness of para consistent logic (Carnielli and Marcos, 2001, da Costa, 1974, 1958, Priest, 1998, Priest et al., 1989, Quesada, 1977) as such as advocated especially by the Peruvian philosopher Francisco Miró Quesada (Quesada, 1977) and other (Carnielli and Marcos, 2001, da Costa, 1974, 1958, Priest, 1998, Priest et al., 1989). In general, scientific theories appear to progress from lower and simpler to higher and more complex levels. However, high level theories cannot be taken for granted because high level theories are grounded on a lot of assumptions, definitions and other procedures and may rest upon too much erroneous stuff even if still not identified. Therefore, it should be considered to check at lower at simpler levels like with like.

2.4.2.1. Zero power zero

Theorem 2.1. *In general, it is*

$$+0^{+2} \equiv +0 \tag{169}$$

is false.

Proof by direct proof. The premise

$$+0 \equiv +1 \tag{170}$$

is false. In the following, any rearrangement of the premise which is free of (technical) errors, need to end up at a contradiction. In other words, the contradiction will be preserved. We obtain

$$+0 \times +0 \equiv +1 \times +0 \tag{171}$$

Equation 171 becomes

$$+0^{+2} \equiv +0 \tag{172}$$

2.4.2.2. Zero divided by zero

Theorem 2.2. *In general*,

$$\frac{1}{0} \equiv \frac{0}{0} \tag{173}$$

is false.

Proof by direct proof. If the premise

$$+1 \equiv +0 \tag{174}$$

is false, then the relationship

$$\frac{1}{0} \equiv \frac{0}{0} \tag{175}$$

is also false.

2.4.3. Principium negationis (Axiom III)

Lex negationis or axiom III, is often mismatched with simple opposition. However, from the point of view of philosophy and other sciences, identity, contradiction, negation and similar notions are equally mathematical descriptions of the most simple laws of objective reality. What sort of natural process is negation at the end? Mathematically, we define principium negationis or lex negationis or axiom III as

Negation(0)
$$\times 0 \equiv \neg (0) \times 0 \equiv +1$$
 (176)

where ¬ denotes (logical (Boole, 1854) or natural) negation (Ayer, 1952, Förster and Melamed, 2012, Hedwig, 1980, Heinemann, Fritz H., 1943, Horn, 1989, Koch, 1999, Kunen, 1987, Newstadt, 2015, Royce, 1917, Speranza and Horn, 2010, Wedin, 1990b). In this context, there is some evidence that

$$Negation(1) \times 1 \equiv \neg (1) \times 1 = 0 \tag{177}$$

Logically, it follows that

$$Negation(1) \equiv 0 \tag{178}$$

In the following we assume that axiom I is universal. Under this assumption, the following theorem follows inevitably.

Theorem 2.3 (Zero divided by zero). According to classical logic, it is

$$\frac{0}{0} \equiv 1 \tag{179}$$

Proof by direct proof. The premise

$$1 \equiv 1 \tag{180}$$

is true. It follows that

$$0 \equiv 0 \\
\equiv 0 \times 1$$
(181)

In the following, we rearrange the premise (see equation 176, p. 79). We obtain

$$0 \times (\text{Negation}(0) \times 0) \equiv 0 \tag{182}$$

Equation 182 changes slightly (see equation 177, p. 79). It is

$$(Negation(1) \times 1) \times (Negation(0) \times 0) \equiv 0$$
 (183)

Equation 183 demands that

$$(Negation(1)) \times (Negation(0)) \times 0 \equiv 0$$
(184)

Equation 184 is logically possible (see equation 167, p. 71) only if

$$(Negation(1)) \times (Negation(0)) \equiv 1$$
 (185)

(see theorem 2.1, equation 169) whatever the meaning of Negation(1) or of Negation(0) might be, equation 185 demands that

$$Negation(0) \equiv \frac{1}{Negation(1)}$$
 (186)

and that

$$Negation(1) \equiv \frac{1}{Negation(0)}$$
 (187)

Equation 186 simplifies as (see equation 178, p. 79)

Negation(0)
$$\equiv \frac{+1}{\text{Negation}(1)}$$

 $\equiv \frac{+1}{+0}$ (188)

It follows that

$$\neg (0) \times 0 \equiv \frac{1}{0} \times 0 \equiv \frac{0}{0} \equiv 1 \tag{189}$$

To bring it to the point. Classical logic, assumed as generally valid, demands that

$$\frac{0}{0} \equiv 1 \tag{190}$$

Concepts like identity, difference, negation, opposition et cetera engaged the attention of scholars at least over the last twenty-three centuries (see also Horn, 1989, Speranza and Horn, 2010). As long as we first and foremost follow Josiah Royce, negatio or negation "is one of the simplest and most fundamental relations known to the human mind. For the study of logic, no more important and fruitful relation is known." (see also Royce, 1917, p. 265) But, do we really know what, for sure, what negation is? Based on what we know about negation, Aristotle (see also Wedin, 1990a) has been one of the first to present a theory of negation, which can be found in discontinuous chunks in his works the Metaphysics, the Categories, De Interpretatione, and the Prior Analytics (see also Horn, 1989, p. 1). Negation (see also Newstadt, 2015) as a fundamental philosophical concept found its own very special melting point especially in Hegel's dialectic and is more than just a formal logical process or operation which converts true to false or false to true. Negation as such is a natural process too and equally 'an engine of changes of objective reality" (see also Barukčić, 2019a). However, it remains an open question to establish a generally accepted link between this fundamental philosophical concept and an adequate counterpart in physics, mathematics and mathematical statistics et cetera. Especially the relationship between creation and conservation or creatio ex nihilio

also Donnelly, 1970, Ehrhardt, 1950, Ford, 1983), determination and negation (see also Ayer, 1952, Hedwig, 1980, Heinemann, Fritz H., 1943, Kunen, 1987) has been discussed in science since ancient (see also Horn, 1989, Speranza and Horn, 2010) times too. Why and how does an event occur or why and how is an event created (creation), why and how does an event maintain its own existence over time (conservation)? The development of the notion of negation leads from Aristotle to Meister Eckhart (see also Eckhart, 1986) von Hochheim (1260-1328), commonly known as Meister Eckhart (see also Tsopurashvili, 2012) or Eckehart, to Spinoza (1632 – 1677), to Immanuel Kant (1724-1804) and finally to Georg Wilhelm Friedrich Hegel (1770-1831) and other authors too. One point is worth being noted, even if it does not come as a surprise, it was especially Benedict de Spinoza (1632 – 1677) as one of the philosophical founding fathers of the Age of Enlightenment who addressed the relationship between determination and negation in his lost letter of June 2, 1674 to his friend Jarig Jelles (see also Förster and Melamed, 2012) by the discovery of his fundamental insight that "determinatio negatio est" (see also Spinoza, 1674, p. 634). Hegel went even so far as to extended the slogan raised by Spinoza into to "Omnis determinatio est negatio" (see also Hegel, Georg Wilhelm Friedrich, 1812b, 2010, p. 87). Finally, it did not take too long, and the notion of negation entered the world of mathematics and mathematical logic at least with Boole's (see also Boole, 1854) publication in the year 1854. "Let us, for simplicity of conception, give to the symbol x the particular interpretation of men, then 1 - x will represent the class of 'not-men'." (see also Boole, 1854, p. 49). Finally, the philosophical notion negation found its own way into physics by the contributions of authors like Woldemar Voigt (see Voigt, 1887), George Francis FitzGerald (see FitzGerald, 1889), Hendrik Antoon Lorentz (see Lorentz, 1892, 1899), Joseph Larmor (see Larmor, 1897), Jules Henri Poincaré (see Poincaré, 1905) and Albert Einstein (see Einstein, 1905) by contributions to the physical notion "Lorentz factor".

3. Results

3.1. Normalisation of a and c

Theorem 3.1 (Normalisation of a and c). *In general, it is*

$$+1 \equiv 1 - \frac{c}{B} \tag{191}$$

Proof by direct proof. The premise

$$+1 \equiv +1 \tag{192}$$

is true. In the following, we rearrange this premise. We obtain

$$a \equiv a \tag{193}$$

Adding term c, it is

$$a + c \equiv a + c \tag{194}$$

According to our definition, it is $a + c \equiv B$. Eq. 194 becomes

$$\frac{a}{B} + \frac{c}{B} \equiv 1 \tag{195}$$

Rearranging 195, it is

$$\frac{a}{B} \equiv 1 - \frac{c}{B} \tag{196}$$

Assuming a necessary condition relationship between events A and B (conditio sine qua non), it is $\frac{a}{B} = +1$. Eq. 196 becomes

$$1 \equiv 1 - \frac{c}{R} \tag{197}$$

3.2. Normalisation of c and d

Theorem 3.2 (Normalisation of c and d). *In general, it is*

$$+1 \equiv 1 - \frac{c}{\underline{A}} \tag{198}$$

Proof by direct proof. The premise

$$+1 \equiv +1 \tag{199}$$

is true. In the following, we rearrange this premise. We obtain

$$c \equiv c \tag{200}$$

Adding term d, it is

$$c + d \equiv c + d \tag{201}$$

According to our definition, it is $c + d \equiv A$. Eq. 201 becomes

$$\frac{c}{A} + \frac{d}{A} \equiv 1 \tag{202}$$

Rearranging 202, it is

$$\frac{d}{A} \equiv 1 - \frac{c}{A} \tag{203}$$

Assuming a necessary condition relationship between events A and B (conditio sine qua non), it is $\frac{d}{A} = +1$. Eq. 203 becomes

$$1 \equiv 1 - \frac{c}{A} \tag{204}$$

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3.3. Index of unfairness I

Theorem 3.3 (Index of unfairness I). In general, it is

$$IOU \equiv \frac{A+B}{N} - 1 \tag{205}$$

Proof by direct proof. The premise

$$+1 \equiv +1 \tag{206}$$

is true. In the following, we rearrange the premise (see equation 197). From the point of view of an observational study it is

$$1 - \frac{c}{R} \equiv +1 \tag{207}$$

Equation 207 is rearranged further. From the point of view of an experimental study (see equation 204), it is

$$1 - \frac{c}{B} \equiv 1 - \frac{c}{A} \tag{208}$$

The study design of both studies should ensure that the same results can be achieved. Simplifying equation 208 it is

$$\frac{c}{B} \equiv \frac{c}{\underline{A}} \tag{209}$$

Equation 209 is divided by c. Under conditions where c = 0, it is $\frac{0}{0} = 1$ (Barukčić, 2018c, 2019b, 2020e, Barukčić and Ufuoma, 2020b, Barukčić and Barukčić, 2016). We obtain

$$\frac{1}{B} \equiv \frac{1}{A} \tag{210}$$

or

$$\underline{A} \equiv B \tag{211}$$

Equation 211 is rearranged as

$$N - A \equiv B \tag{212}$$

or

$$N \equiv A + B \tag{213}$$

and

$$1 \equiv \frac{A+B}{N} \tag{214}$$

or

$$0 \equiv \frac{A+B}{N} - 1 \tag{215}$$

One objective of the design of a study which investigates a necessary condition relationship between events A and B (conditio sine qua non) should be to ensure as much as possible an index of unfairness (see also Barukčić, 2019c) equal to

$$IOU \equiv \frac{A+B}{N} - 1 \equiv 0 \tag{216}$$

3.4. Normalisation of a and b

Theorem 3.4 (Normalisation of a and b). *In general, it is*

$$1 \equiv 1 - \frac{b}{B} \tag{217}$$

Proof by direct proof. The premise

$$+1 \equiv +1 \tag{218}$$

is true. In the following, we rearrange this premise. We obtain

$$a \equiv a$$
 (219)

Adding term b, it is

$$a + b \equiv a + b \tag{220}$$

According to our definition, it is $a + b \equiv A$. Eq. 220 becomes

$$\frac{a}{A} + \frac{b}{A} \equiv 1 \tag{221}$$

Rearranging 221, it is

$$\frac{a}{A} \equiv 1 - \frac{b}{A} \tag{222}$$

Assuming a sufficient condition relationship between events A and B, it is $\frac{a}{A} = +1$. Eq. 222 becomes

$$1 \equiv 1 - \frac{b}{A} \tag{223}$$

3.5. Normalisation of b and d

Theorem 3.5 (Normalisation of b and d). *In general, it is*

$$1 \equiv 1 - \frac{b}{B} \tag{224}$$

Proof by direct proof. The premise

$$+1 \equiv +1 \tag{225}$$

is true. In the following, we rearrange this premise. We obtain

$$b \equiv b \tag{226}$$

Adding term d, it is

$$b + d \equiv b + d \equiv \underline{B} \tag{227}$$

According to our definition, it is $b + d \equiv \underline{B}$. Equation 227 becomes

$$\frac{b}{B} + \frac{d}{B} \equiv 1 \tag{228}$$

Rearranging 228, it is

$$\frac{d}{B} \equiv 1 - \frac{b}{B} \tag{229}$$

Assuming a sufficient condition relationship (conditio per quam) between events A and B, it is $\frac{d}{\underline{B}} = +1$. Equation 229 becomes

$$1 \equiv 1 - \frac{b}{B} \tag{230}$$

3.6. Index of unfairness II

Theorem 3.6 (Index of unfairness II). *In general, it is*

$$IOU \equiv \frac{A+B}{N} - 1 \equiv 0 \tag{231}$$

CAUSATION *ISSN: 1863-9542*

https://www.doi.org/10.5281/zenodo.6816088

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Proof by direct proof. The premise

$$+1 \equiv +1 \tag{232}$$

is true. In the following, we rearrange the premise (see equation 223). From the point of view of an experimental study it is

$$1 - \frac{b}{A} \equiv +1 \tag{233}$$

Equation 233 is rearranged further. From the point of view of an observational study (see equation 230), it is

$$1 - \frac{b}{A} \equiv 1 - \frac{b}{B} \tag{234}$$

Simplifying equation 234 it is

$$\frac{b}{A} \equiv \frac{b}{B} \tag{235}$$

Equation 235 is divided by b. Under conditions where b = 0, it is $\frac{0}{0} = 1$ (Barukčić, 2018c, 2019b, 2020e, Barukčić and Ufuoma, 2020b, Barukčić and Barukčić, 2016). We obtain

$$\frac{1}{A} \equiv \frac{1}{B} \tag{236}$$

or

$$\underline{B} \equiv A \tag{237}$$

Equation 237 is rearranged as

$$N - B \equiv A \tag{238}$$

or

$$N \equiv A + B \tag{239}$$

and

$$1 \equiv \frac{A+B}{N} \tag{240}$$

or

$$0 \equiv \frac{A+B}{N} - 1 \tag{241}$$

One objective of the design of a study which investigates a sufficient condition relationship (conditio per quam) between events A and B should be to ensure an index of unfairness (see also Barukčić, 2019c) as much as possible equal to

$$IOU \equiv \frac{A+B}{N} - 1 \equiv 0 \tag{242}$$

4. Discussion

In practice, comparability of medical studies of different types or even of the same type is not possible as straightforward as it could be or as it should be. Even an additional simple statement of p-values or confidence intervals is not sufficient to be able to rely upon the results published by a scientific article. Out of the many, different approaches that can be taken towards the investigation of a certain scientific issue, it is necessary to ensure minimum standard conditions thus that different study groups can achieve the same scientific results. A certain relationship between event A and event B given independently and outside of human mind and consciousness, independently of any study and study design, is the way, the same is. From the point of view of an experimental study, the relationship between an event A and another event B should to be the same as seen from the point of view of an observational study. In this respect, it is important to remember that a contradiction is neither possible nor desirable. Unfortunately - since one is theory and the other is practice - it might be necessary to conduct studies out at least in compliance with conditions outlined before. One of these conditions is an index of unfairness (IOU) equal to 0 or very near to 0.

5. Conclusion

Investigation of necessary or sufficient conditions require an index of unfairness of IOU = 0.

Acknowledgments

No funding or any financial support by a third party was received. Thanks to Overleaf, Zotero, Gnuplot, Zenodo, TeX Shop and other.

6. Patient consent for publication

Not required.

Conflict of interest statement

No conflict of interest to declare.

Private note

The definition section of a paper need not and does not necessarily contain new scientific aspects. Above all, it also serves to better understand a scientific publication, to follow every step of the arguments of an author and to explain in greater details the fundamentals on which a publication is based. Therefore, there is no objective need to force authors to reinvent a scientific wheel once and again unless such a need appears obviously factually necessary. The effort to write about a certain subject in an original way in multiple publications does not exclude the necessity simply to cut and paste from an earlier work, and has nothing to do with self-plagiarism. However, such an attitude cannot simply be transferred to the sections' introduction, results, discussion and conclusions et cetera.

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I was born October, 1st 1961 in Novo Selo, Bosnia and Herzegovina, former Yogoslavia. I am of Croatian origin. From 1982-1989 C.E., I studied human medicine at the University of Hamburg, Germany. Meanwhile, I am working as a specialist of internal medicine. My basic field of research since my high school days at the Wirtschaftsgymnasium Bruchsal, Baden Württemberg, Germany is the mathematization of the relationship between a cause and an effect valid without any restriction under any circumstances including the conditions of classical logic, probability theory, quantum mechanics, special and general theory of relativity, human medicine et cetera. I endeavour to investigate positions of quantum mechanics, relativity theory, mathematics et cetera, only insofar as these positions put into question or endanger **the general validity of the principle of causality**.



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