P-values, confidence intervals, and significance testing - navigating the minefield when communicating your findings CBD talk

Stefanie Muff

November 10, 2020

The ongoing controversy around p-values

The ASA's Statement on *p*-Values: Context, Process, and Purpose

Ronald L. Wasserstein^{a*} & Nicole A. Lazar^a pages 129-133

In February 2014, George Cobb, Professor Emeritus of Mathematics and Statistics at Mount Holyoke College, posed these questions to an ASA discussion forum:

Q: Why do so many colleges and grad schools teach p = 0.05?

A: Because that's still what the scientific community and journal editors use.

Q: Why do so many people still use p = 0.05?

A: Because that's what they were taught in college or grad school.

Cobb's concern was a long-worrisome circularity in the sociology of science based on the use of bright lines such as p < 0.05: "We teach it because it's what we do; we do it because it's what we teach." This concern was brought to the attention of the ASA Board.

(Wasserstein and Lazar 2016)

Lots of publications in the past decades...

STATISTICAL ERRORS

P values, the 'gold standard' of statistical validity, are not as reliable as many scientists assume.

BY REGINA NUZZO

COMMENT - 20 MARCH 2019

Scientists rise up against statistical significance

Valentin Amrhein, Sander Greenland, Blake McShane and more than 800 signatories call for an end to hyped claims and the dismissal of possibly crucial effects.

Valentin Amrhein (50), Sander Greenland & Blake McShane

A Dirty Dozen: Twelve P-Value Misconceptions

Steven Goodman

The P value is a massive of statistical evidence that appears in virtually all medical research papers. In insepretables in most extendincy effected because it is not part of any formal system of statistical inference. As a most, the P public inferenced manning is any formal system of statistical process. The public inference is not part of the public inference in the public inference is not part of the public inference in the public inference in the public consequence of the same proper discontinuous control of the minor part of the public interesponders of the minor manner than the public interesponder of the public interesponders of the public interesponders of the minor most of the public interesponders of the public interesponders

Why Most Published Research Findings

Are False



lactors that influence this problem and some concluders thereof.

Modeling the Framework for False Positive findings
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ontiferancian) of research discoveries, is a consequence of the conservieus, yet illiforated strategy of claiming concludes reasonable fluidings shelly on the basis of a single study assessed by formal statistical significance, topically for a poalse loss than 60%. Research is not most appropriately represented and summarized by peaches, but, underturnately, there is a wiferspecial sosien that medical records articles. It can be a proposed to the control of the contr

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expected salates of the 2 s 2 sable as given in Table 1. After a research finding has been claimed based on achieving formed standard significant the postendy probability that it is to to the positive predictive value, FPY. The FPY is also the complementary probability of what Wachedder et al., have called the false positive report.

Ioannidis (2005), Goodman (2008), Nuzzo (2014), Amrhein, Greenland, and McShane (2019), ...

P-values / statistical significance criticism

P-value **criticism** is as **old** as statistical significance testing (1920s!). Issues:

- The sharp line p < 0.05 is arbitrary and significance testing according to it may lead to mindless statistics (Gigerenzer 2004).
- P-hacking / data dredging: Search until you find a result with p < 0.05.
- Publication bias: Studies with p < 0.05 are more likely to be published than "non-significant" results.
- HARKING: Hypothesizing After the Results are Known.
- Model selection using *p*-values \rightarrow model selection bias.

Note: R.A. Fisher, the "inventor" of the p-value (1920s) didn't mean the p-value to be used in the way it is used today, which is: doing a single experiment and use p < 0.05 for a conclusion.

From Goodman (2016):

Fisher used "significance" merely to indicate that an observation was worth following up, with refutation of the null hypothesis justified only if further experiments "rarely failed" to achieve significance. This is in stark contrast to the modern practice of making claims based on a single demonstration of statistical significance.

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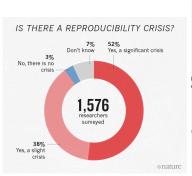
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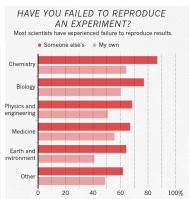
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The misuse of *p*-values is partially responsible for the reproducibility/replicability crisis in science!

A reproducibility crisis?

A survey carried out by Nature in 2016, sheds light on researcher's experiences and thoughts.



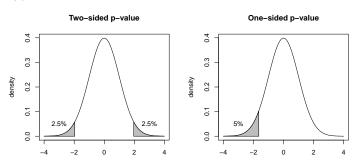


But wait, what is the problem with the *p*-value?

The p-value is not a very intuitive concept...

Formal definition

The p-value is the probability to observe a data summary (e.g., a t-value) that is at least as extreme as the one observed, given that the Null Hypothesis is correct.



Right or wrong?

- 1. The p-value is the probability that the null hypothesis is true.
- 2. p = 0.02 means that the alternative hypothesis is true with 98% probability.
- 3. The *p*-value is the type-1 error rate.
- 4. The p-value is the probability that the result happened by chance.
- 5. If p > 0.05, we can conclude that there is no effect.
- 6. Two studies with p > 0.05 and p < 0.05 are in a conflict.

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Sorry, but this is all bullshit...

Some explanations

The p-value is

$$p = \mathsf{P}(\mathsf{Result} \,|\, H_0)$$
 ,

 $^{^{1}\}mathrm{P}(H_{0}\,|\,\mathrm{Result}) = \tfrac{\mathrm{P}(\mathrm{Result}\,|\,H_{0})\cdot\mathrm{P}(H_{0})}{\mathrm{P}(\mathrm{Result})}$

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The p-value is

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but unfortunately it is not the reverse (why?¹):

$$p \neq \mathrm{P}(H_0 \, | \, \mathrm{Result})$$
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Similarly:

$$1 - p \neq P(H_1 | \text{Result})$$
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Reasons for large p-values:

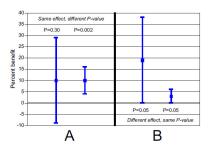
- Low sample size (\rightarrow low power).
- The truth is not far from the null hypothesis.
- Collinear covariates.

Significance vs relevance

Paul D. Ellis in *The Essential Guide to Effect Sizes* (2010, chapter 2):

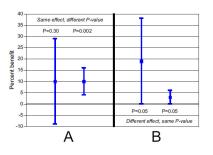
Indeed, statistical significance, which partly reflects sample size, may say nothing at all about the practical significance of a result. [....] To extract meaning from their results [...] scientists need to look beyond p values and effect sizes and make informed judgments about what they see.

- A low p-value does not automatically imply that a variable is "important" and vice versa.
- "Is there an effect?" v.s. 'How much of an effect is there?".



Goodman (2008)

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 $Goodman\ (2008)$

Problem: The *p*-value blands the estimated effect size with its uncertainty.

An example

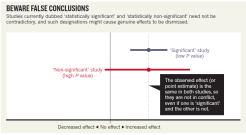
The WHO recommendation concerning smoking and the consumption of processed meat. Both, smoking and meat consumption, are "significantly" increasing the probability to get cancer.

- 50g processed meat per day increases the risk for colon cancer by a factor of 1.18 (+18%).
- Smoking increases the risk to get any type of cancer by a factor of 3.6 (+260%).

Thus: Although both, meat consumption and smoking, are carcinogenic ("significant"), their effect sizes are vastly different!

Are two studies in conflict?

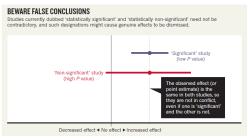
• In the following example, two studies find the same effect size, but one is significant (p = 0.02) and the other one is not (p = 0.09).



Amrhein, Greenland, and McShane (2019)

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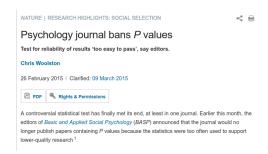
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- This is obviously no conflict, only the uncertainty is different.
- **Again:** The *p*-value blands the effect size with its uncertainty.

Shall we abolish p-values?



Shall we abolish p-values?



- But that throws the baby out with the bath water. It's as if we would forbid trains because they cannot fly to South America...
- p-values are not "good" or "bad". They contain important information, and they have strengths and weaknesses.

What should we do then?



Retire statistical significance
Valentin Amrhein, Sander Greenland, Blake McShane and more than 800 signatories
call for an end to hyped claims and the dismissal of possibly crucial effects.

- In many situations it is not justified to make a strict yes/no decision.²
- **Instead**: accumulating evidence over more and more studies.³

 $^{^2}$ And we are usually not forced to! In contrast to e.g. clinical trials.

³That's why it is so important to publish non-significant results, too! And: the importance of meta-analyses.

But... an (almost) randomly selected example

- Last week I went to the *Evolution* website and just picked the most recent paper (published November 1st, first on the list).
- Guess how many times the word *significant* was used.

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Answer: 55

A snapshot from the paper:

S7), but no clear pattern emerged. We found significant G x E in female size between Control-Dry (P = 2.78E-03). In male size, G x E was significant between Control-Dry (P = 6.39E-03), Dry-Hot (P = 5.39E-03), Dry-Hot-Dry (P = 0.04), and Hot-Hot-Dry (P = 3.93E-03). Genetic correlations between female and male size were close to one (Table 2, Figure 5) in all conditions, suggesting that size cannot evolve independently in both sexes. I_A of female and male size was not significantly different (P = 0.08) considering all conditions. Although differences in Control seemed to be substantial (Figure 4B-D), they were not significant (h^2 : P = 0.26; I_A : P = 0.06). We found that environmental change did not influence I_A and h^2 of female size (P = 0.13; P = 0.14) nor male I_A (P = 0.13), but had a significant effect on male h^2 (P = 0.02).

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How could we do better?

Suggestion 1: Language matters!

Rewrite your results and use a gradual interpretation of the p-value.

For single (observational) studies, the following has been suggested already decades ago (Bland 1986):

Interpreting the P value

Greater than 0.1:

As a rough and ready guide, we can think of P values as indicating the strength of evidence like this:

P value Evidence for a difference or

relationship

Between 0.05 and 0.1: Weak evidence

Between 0.01 and 0.05: Evidence

Less than 0.01: Strong evidence

Less than 0.001: Very strong evidence

Suggestion 2: Report effect sizes and 95% CIs

Ask:

- Is the effect size (biologically, medically, socially...) relevant?
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However

- \bullet The choice of the 95% is again somewhat arbitrary. We could also go for 90% or 99% or any other interval.
- The 95% CI should **not be misused for simple hypothesis testing** in the sense of "Is 0 in the confidence interval or not?" that is just significance testing.

A results table from an example where I was involved (Imo et al. 2018):

Table 4. Evidence for the association with log-transformed mercury values in urine (ug/g creatinine).

n = 164	Variable	Coefficient	95% CI	p-Value
Very strong evidence	Amalgam fillings	0.33	0.24, 0.42	< 0.001
	Last time sea fish	0.32	0.17, 0.47	< 0.001
	Age	-0.04	-0.06, -0.02	< 0.001
	Interaction age × mother	0.05	0.02, 0.08	< 0.001
Strong evidence	Mother (indicator)	-0.97	-1.64, -0.31	0.004
	Smoking	0.30	0.09, 0.50	0.005
	Sea fish	0.08	0.03, 0.13	0.003
Little or no evidence	Log ₁₀ Hg soil	0.02	-0.06, 0.10	0.64
	Limit of quantification	-0.08	-0.25, 0.09	0.37
	Country of birth near the sea	-0.01	-0.16, 0.15	0.93
	Eats vegetables from region	0.07	-0.03, 0.18	0.18

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We found very strong evidence for a positive association of mercury in urine with amalgam fillings (regression coefficient: 0.33; 95% CI: 0.24–0.42; p < 0.001).

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Cl: Confidence interval

We found very strong evidence for a positive association of mercury in urine with amalgam fillings (regression coefficient: 0.33; 95% CI: 0.24–0.42; p < 0.001).

We found no evidence for an association of log-transformed mercury concentrations in soil with log-transformed concentrations in urine (regression coefficient: 0.02; 95% CI: -0.06-0.10; p=0.64).

The interpretation of the *p*-value depends!

- Observational vs experimental study
- Confirmatory vs exploratory analysis

Practice in drug regulation

Clinical trials (CTs) for **drug approval** underlie strict requirements – since decades.

- CTs are randomized controlled trials.
- Study protocols that are published even before any patient is treated.
- Preregistration of study protocols and analysis plans.
- Two Trials Rule:

"at least two adequate and well-controlled studies, each convincing on its own, to establish effectiveness."

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Clinical trials are *experimental* and *confirmatory*, thus it is ok to draw causal conclusions.

Our situation is – for many reasons – not comparable to clinical trials:

- Clinical trials: Always experimental intervention and very strict regulations.
- Ecology: (Often) observational studies, lots of researchers degrees
 of freedom, usually no preregistration, exploratory data analysis,
 no study protocols, model selection,...

But, why not pick the good things from medical statistics?

Final suggestions:

- Use p-values, but no significance testing⁴.
- Always report *effect sizes* and *confidence intervals*.
- Think about the *relevance*, not only the *significance*.
- More gradual interpretation of p-values, rewrite your results sections.
- Do not use *p*-values for model selection.

 $^{^4}$ unless you analyze a randomized controlled trial

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Question: Would you find a "translation guideline" to rewrite your results useful?

Thanks for feedback

⁴unless you analyze a randomized controlled trial

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