

# **Pvalue** (by the ASA statement)



UNIVERSITAT POLITÈCNICA DE CATALUNYA BARCELONATECH

Departament d'Estadística i Investigació Operativa











# Psychology journal bans P values

Test for reliability of results 'too easy to pass', say editors.





# P value ban: small step for a journal, giant leap for science

### False-Positives, p-Hacking, Statistical Power, and Evidential Value

Leif D. Nelson University of California, Berkeley Haas School of Business

> Summer Institute June 2014



# Six Ways to p-Hack

- 1. Stop collecting data once *p*<.05
- 2. Analyze many measures, but report only those with *p*<.05.
- 3. Collect and analyze many conditions, but only report those with *p*<.05.
- 4. Use covariates to get *p*<.05.
- 5. Exclude participants to get *p*<.05.
- 6. Transform the data to get *p*<.05.

# Perspective



### The NEW ENGLAND JOURNAL of MEDICINE

## Improving Patient Safety through Transparency

# Research: increasing value, reducing waste

#### Research: increasing value, reducing waste 1

THE LANCET

How to increase value and reduce waste when research priorities are set

lain Chalmers, Michael B. Bracken, Ben Djulbegovic, Silvio Garattini, Jonathan Grant, A. Metin Gülmezoglu, David W. Howells, John P. A loannids, Sandy Oliver

Research: increasing value, reducing waste 2

Increasing value and reducing waste in research design, conduct, and analysis

Research: increasing value, reducing waste 3

Increasing value and reducing waste in biomedical research regulation and management

Research: increasing value, reducing waste 4

Increasing value and reducing waste: addressing inaccessible research

Research: increasing value, reducing waste 5

Reducing waste from incomplete or unusable reports of biomedical research



#### EDITORIAL

### The scandal of poor medical research

BMJ 1994; 308 doi: http://dx.doi.org/10.1136/bmj.308.6924.283 (Published 29 January 1994) Cite this as: BMJ 1994;308:283

### EDITORIAL

# The scandal of poor epidemiological research

BMJ 2004; 329 doi: http://dx.doi.org/10.1136/bmj.329.7471.868 (Published 14 October 2004 Cite this as: BMJ 2004;329:868

### Is animal research sufficiently evidence based to be a cornerstone of biomedical research?

# NIH plans to enhance reproducibility

Francis S. Collins and Lawrence A. Tabak discuss initiatives that the US National Institutes of Health is exploring to restore the self-correcting nature of preclinical research.



Declaración de transparencia para las publicaciones científicas Declaration of transparency for scientific publications

BMJ

# **Why Most Published Research Findings**

# **Are False**

John P. A. Ioannidis





1-β	R	u	Practical Example	PPV
0.80	1:1	0.10	Adequately powered RCT with little	0.85
			bias and 1:1 pre-study odds	
0.95	2:1	0.30	Confirmatory meta-analysis of good quality RCTs	1- 0.85
0.80	1:3	0.40	Meta-analysis of small inconclusive studies	0.41
0.20	1:5	0.20	Underpowered, but well-performed phase I/II RCT	0.23
0.20	1:5	0.80	Underpowered, poorly performed phase I/II RCT	0.17
0.80	1:10	0.30	Adequately powered exploratory epidemiological study	0.20
0.20	1:10	0.30	Underpowered exploratory epidemiological study	0.12
0.20	1:1,000	0.80	Discovery-oriented exploratory research with massive testing	0.0010
0.20	1:1,000	0.20	As in previous example, but with more limited bias (more standardized)	0.0015

The ASA's statement on p-values: context, process, and purpose



Ronald L. Wasserstein and Nicole A. Lazar

1. P-values can indicate how incompatible the data are with a specified statistical model.

2. P-values do **NOT** measure the probability that the studied hypothesis is true, or the probability that the data were produced by random chance alone.

3. Scientific conclusions and business or policy decisions should **NOT** be based only on whether a p-value passes a specific threshold.

4. Proper inference requires full reporting and transparency

5. A p-value, or statistical significance, does **NOT** measure the size of an effect or the importance of a result.

6. By itself, a p-value does **NOT** provide a good measure of evidence regarding a model or hypothesis.

Declaración de la ASA sobre la significación estadística y los p-valores







1. P-values can indicate how incompatible the data are with a specified statistical model





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# Denominator changed











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2. P-values do NOT measure the probability that the studied hypothesis is true



### Confusion Over Measures of Evidence (p's) Versus Errors ( $\alpha$ 's) in Classical Statistical Testing

Raymond HUBBARD and M. J. BAYARRI

(c) 2003 American Statistical Association DOI: 10.1198/0003130031856

The American Statistician, August 2003, Vol. 57, No. 3 171

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ESSAY

DOI 10.1007/s10654-016-0149-3

### Statistical tests, P values, confidence intervals, and power: a guide to misinterpretations

Sander Greenland<sup>1</sup> · Stephen J. Senn<sup>2</sup> · Kenneth J. Rothman<sup>3</sup> · John B. Carlin<sup>4</sup> · Charles Poole<sup>5</sup> · Steven N. Goodman<sup>6</sup> · Douglas G. Altman<sup>7</sup>



In other words, if you concluded there is a difference in

test performance between the groups, there is less than a 5 percent chance that

you are wrong and more than a 95 percent chance that you are right.

Lesson without

Standard

Deviation = 10

Mean = 90%

Music

100

90

# **Why Most Published Research Findings**

# **Are False**

John P. A. Ioannidis

**Table 4.** PPV of Research Findings for Various Combinations of Power  $(1 - \beta)$ , Ratio of True to Not-True Relationships (*R*), and Bias (*u*)



1-β	R	u	Practical Example	PPV
0.80	1:1	0.10	Adequately powered RCT with little	0.85
			bias and 1:1 pre-study odds	
0.95	2:1	0.30	Confirmatory meta-analysis of good quality RCTs	I- 0.85
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# 3.- P value







### Credibilidad de la ciencia



#### Calculadora Cur

Curva de credibilidad Ayuda

Esta aplicación pretende mostrarle que la posibilidad de repetir un resultado depende de:

- la base científica de la hipótesis (mayor justificación previa, mayor credibilidad);
- la capacidad del estudio para alcanzar sus objetivos según su diseño, tamaño y precisión (mayor potencia, mayor credibilidad);
- el control del riesgo de error por falsas señales positivas (mayor control, mayor credibilidad).

Vd. debe jugar con estos 3 parámetros, y el gráfico adjunto le mostrará cómo cambia el escenario: un efecto real o no, combinado con resultado del análisis significativo o no. Además, también obtendrá la probabilidad de que un resultado "estadísticamente significativo" se repita en el futuro.

#### Expectativa del efecto







### Credibilidad de la ciencia

Calculadora C

Curva de credibilidad Avuda



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#### Expectativa del efecto





$$P(\text{Efecto real}|\text{Signif.}) = \frac{P(\text{Efecto real} \cap \text{Signif.})}{P(\text{Signif.})} = \frac{0.02}{0.06} = 0.333$$



Link to applet: http://shiny-eio.upc.edu/bne/efectos2/





### Adapted Treatment Guided by Interim PET-CT Scan in Advanced Hodgkin's Lymphoma

Peter Johnson, M.D., Massimo Federico, M.D., Amy Kirkwood, M.Sc., Alexander Fosså, M.D., Leanne Berkahn, M.D., Angelo Carella, M.D., Francesco d'Amore, M.D., Gunilla Enblad, M.D., Antonella Franceschetto, M.D., Michael Fulham, M.D., Stefano Luminari, M.D., Michael O'Doherty, M.D., Pip Patrick, Ph.D., Thomas Roberts, B.Sc., Gamal Sidra, M.D., Lindsey Stevens, Paul Smith, M.Sc., Judith Trotman, M.D., Zaid Viney, M.D., John Radford, M.D., and Sally Barrington, M.D.

ABSTRACT

#### BACKGROUND

We tested interim positron-emission tomography-computed tomography (PET-CT) as a measure of early response to chemotherapy in order to guide treatment for patients with advanced Hodgkin's lymphoma.

#### METHODS

Patients with newly diagnosed advanced classic Hodgkin's lymphoma underwent a baseline PET-CT scan, received two cycles of ABVD (doxorubicin, bleomycin, vinblastine, and dacathazine) chemotherapy, and then underwent an interim PET-CT scan. Images were centrally reviewed with the use of a 5-point scale for PET findings. Patients with negative PET findings after two cycles were randomly assigned to continue ABVD (ABVD group) or omit bleomycin (AVD group) in cycles 3 through 6. Those with positive PET findings after two cycles received EEACOFP (bleomycin, etoposide, doxorubicin, cyclophosphamide, vincristine, procarbazine, and prednisone). Radiotherapy was not recommended for patients with negative findings on interim scans. The primary outcome was the difference in the 3-year progression-free survival rate between randomized groups, a noninferiority comparison to exclude a difference of 5 or more percentage points.

#### RESULTS

A total of 1214 patients were registered; 937 of the 1119 patients (83.7%) who underwent an interim PET-CT scan according to protocol had negative findings. With a median follow-up of 41 months, the 3-year progression-free survival rate and overall survival rate in the ABVD group were 85.7% (95% confidence interval [CI], 82.1 to 88.6) and 97.2% (95% CI, 95.1 to 98.4), respectively; the corresponding rates in the AVD group were 84.4% (95% CI, 95.1 to 98.7). The absolute difference in the 3-year progression-free survival rate (ABVD minus AVD) was 1.6 percentage points (95% CI, -3.2 to 5.3). Respiratory adverse events were more severe in the ABVD group than in the AVD group. BEACOPP was given to the 172 patients with positive findings on the interim scan, and 74.4% had negative findings on a third PET-CT scan; the 3-year progression-free survival rate 87.5% and the overall survival rate 87.8%. A total of 62 patients died during the trial (24 from Hodgkin's lymphoma), for a 3-year progression-free survival rate of 82.6% and an overall survival rate of 95.8%.

#### CONCLUSIONS

Although the results fall just short of the specified noninferiority margin, the omission of bleomycin from the ABVD regimen after negative findings on interim PET resulted in a lower incidence of pulmonary toxic effects than with continued ABVD but not significantly lower efficacy. (Funded by Cancer Research UK and Others; ClinicaTrials.gov number, NCT00678327)

University of Southampton, Southampton (P.J.), Cancer Research UK and University College London Cancer Trials Centre (A.K., P.P., T.R., L.S., P.S.) and the PET Imaging Centre, King's College London, King's Health Partners, St. Thomas' Hospital (M.O., Z.V., S.B.), London, the Department of Haematology, Lincoln County Hospital, Lincoln (G.S.), and the Department of Medical Oncology, Christie Hospital, Manchester (J.R.) - all in the United Kingdom; the Department of Diagnostic, Clinical, and Public Health Medicine, University of Modena and Reggio Emilia, Modena (M. Federico, A. Franceschetto, S.L.), the Department of Hematology, San Martino University Hospital, Genoa (A.C.), and Arcispedale Santa Maria Nuova-Istituti di Ricovero e Cura a Carattere Scientifico, Reggio Emilia (S.L.) - all in Italy; the Department of Medical Oncology, Oslo University Hospital, Oslo (A. Fossá); the Department of Haematology, Auckland City Hospital, Auckland, New Zealand (L.B.); the Department of Hematology, Aarhus University Hospital, Aarhus, Denmark (F.A.): the Department of Immunology, Genetics, and Pathology, Uppsala University, Uppsala, Sweden (G.E.); and the Department of Molecular Imaging, Royal Prince Alfred Hospital (M. Fulham), and Concord Repatriation General Hospital, University of Sydney (J.T.), Sydney. Address reprint requests to Dr. Johnson at the Cancer Research UK Centre, University of Southampton, Somers Cancer Research Bldg., Southampton General Hospital, Southampton SO16 6YD, United Kingdom, or at johnsonp@soton.ac.uk.

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# **Thanks**



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