### Surprise, Surprise: It's a false positive

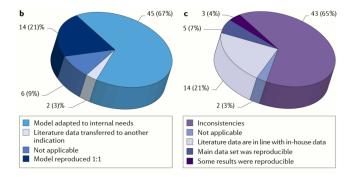
#### Jarrod Hadfield

University of Edinburgh

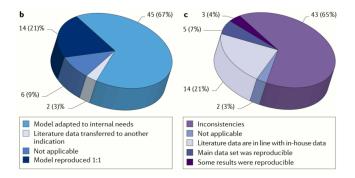
November 24, 2017

• Prinz, F. et al. (2011) Believe it or not: how much can we rely on published data on potential drug targets? Nature Reviews Drug Discovery 10

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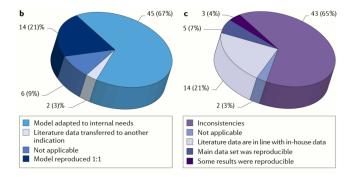


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- Begley, CG. & Ellis, LM. (2012) Drug development: Raise standards for preclinical cancer research. Nature 483
- Open Science Collaboration (2015) Estimating the reproducibility of psychological science. Science 349

• U.S. House of Representatives Hearing on 'Scientific Integrity and Transparency' (2013)

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- 'Budding scientists must be taught technical skills, including statistics, and must be imbued with scepticism towards their own results and those of others. Researchers ought to be judged on the basis of the quality, not the quantity, of their work.'
- 'We need to develop a value system where simply moving on from one's mistakes without publicly acknowledging them severely damages, rather than protects, a scientific reputation.'

## How Science Goes Wrong



# How Science Goes Wrong



'Science still commands enormous - if sometimes bemused - respect. But its privileged status is founded on the capacity to be right most of the time and to correct its mistakes when it gets things wrong. And it is not as if the universe is short of genuine mysteries to keep generations of scientists hard at work. The false trails laid down by shoddy research are an unforgivable barrier to understanding.'

- Background
- Problems
- Under-graduate Project
- Solutions
- Unacknowledged Issues

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  - Wacholder, S. et al. (2004) Assessing the Probability That a Positive Report is False: An Approach for Molecular Epidemiology Studies. Journal of the National Cancer Institute 96, 434-442
  - Ioannidis, JPA. (2005) Why Most Published Research Findings Are False PLoS Medicine 2 e124

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Experiment (n=30)	Small	0.08	0.61
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Average	Medium	0.50	0.91

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Average	Small	0.80	
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#### False Positive Report Probability

$$1 - FPRP = \frac{(1 - \beta)\pi}{(1 - \beta)\pi + \alpha(1 - \pi)}$$

Scenario	Effect Size	$\pi = 0.5$	$\pi = 0.1$	
Ideal		0.94	0.64	
Experiment (n=30)	Small	0.61	0.15	
Experiment (n=30)	Medium	0.84	0.37	
Average	Small	0.80	0.31	
Average	Medium	0.91	0.53	

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$$1 - FPRP = rac{(1-eta)\pi}{(1-eta)\pi+lpha(1-\pi)}$$

Scenario	Effect Size	$\pi = 0.5$	$\pi = 0.1$	$\pi = 0.01$	
Ideal		0.94	0.64	0.14	
Experiment (n=30)	Small	0.61	0.15	0.02	
Experiment (n=30)	Medium	0.84	0.37	0.05	
Average	Small	0.80	0.31	0.04	
Average	Medium	0.91	0.53	0.09	

#### False Positive Report Probability

$$1 - FPRP = \frac{(1 - \beta)\pi}{(1 - \beta)\pi + \alpha(1 - \pi)}$$

Scenario	Effect Size	$\pi = 0.5$	$\pi = 0.1$	$\pi = 0.01$	$\alpha = 0.23$
Ideal		0.94	0.64	0.14	0.034
Experiment (n=30)	Small	0.61	0.15	0.02	0.003
Experiment (n=30)	Medium	0.84	0.37	0.05	0.011
Average	Small	0.80	0.31	0.04	0.009
Average	Medium	0.91	0.53	0.09	0.021

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- Simmons, JP. (2011) False-Positive Psychology: Undisclosed Flexibility in Data Collection and Analysis Allows Presenting Anything as Significant Psychological Science 22 1359-1366
- *Researcher degrees of freedom*: how many decisions were made during the course of data collection, analysis, presentation and publication.

Are qualitative assessments of  $\pi$  useful?

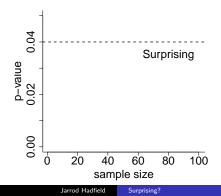
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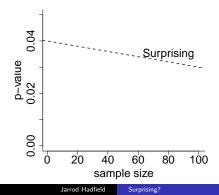
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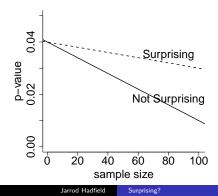
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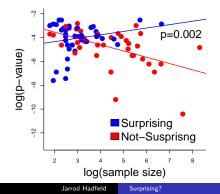
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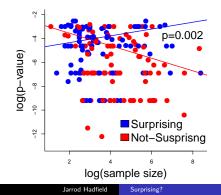
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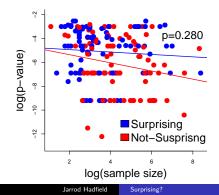
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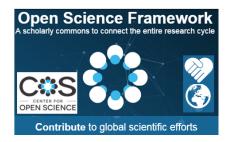


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• Digitally pre-register study hypotheses, data collection and analysis plans as a record of intent.

Should we bring (qualitative) priors into it?

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