# Open Science and reproducible research

Thursday 12th December 2019

Brendan Palmer,

Clinical Research Facility - Cork & School of Public Health



@B\_A\_Palmer





# Can we believe what we see in the literature?









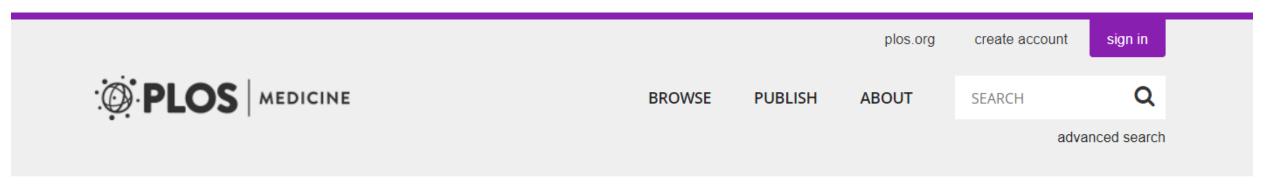
### The Prevalence of Inappropriate Image Duplication in Biomedical **Research Publications**

Elisabeth M. Bik,<sup>a</sup> Arturo Casadevall,<sup>b,c</sup> Ferric C. Fang<sup>d</sup>

Department of Medicine, Division of Infectious Diseases, Stanford School of Medicine, Stanford, California, USAa; Department of Molecular Microbiology and Immunology, Johns Hopkins Bloomberg School of Public Health, Baltimore, Maryland, USAb; Department of Medicine, Johns Hopkins School of Medicine, Baltimore, Maryland, USA<sup>c</sup>; Departments of Laboratory Medicine and Microbiology, University of Washington School of Medicine, Seattle, Washington, USA<sup>d</sup>

Inaccurate data in scientific papers can result from honest error or intentional falsification. This study attempted to **ABSTRACT** determine the percentage of published papers that contain inappropriate image duplication, a specific type of inaccurate data. The images from a total of 20,621 papers published in 40 scientific journals from 1995 to 2014 were visually screened. Overall, 3.8% of published papers contained problematic figures, with at least half exhibiting features suggestive of deliberate manipulation. The prevalence of papers with problematic images has risen markedly during the past decade. Additional papers written by authors of papers with problematic images had an increased likelihood of containing problematic images as well. As this analysis focused only on one type of data, it is likely that the actual prevalence of inaccurate data in the published literature is higher. The marked variation in the frequency of problematic images among journals suggests that journal practices, such as prepublication image screening, influence the quality of the scientific literature.

### Can we believe what we read in the literature?



■ OPEN ACCESS

**ESSAY** 

### Why Most Published Research Findings Are False

John P. A. Ioannidis

Published: August 30, 2005 • https://doi.org/10.1371/journal.pmed.0020124

| 68,836    | 2,931    |
|-----------|----------|
| Save      | Citation |
| 2,768,586 | 10,482   |
| View      | Share    |

### And publication bias is very real



Follow

Elsevier editor Spada acknowledging that null results are not even considered for Addictive Behaviors, seemingly not realizing how problematic that is. Offering a lower prestige alternative journal doesn't make that right.



Professor M. M. Spada said:

"Articles that may not traditionally be considered by Addictive Behaviors, including negative/null data papers, studies using smaller samples and cross-sectional designs, replication studies, cross-cultural research, and case reports will be welcome by its sister journal Addictive Behaviors Reports."

Editor-in-Chief Professor M. M. Spada London South Bank University Journal Metrics

> CiteScore: 3.10 ①
Impact Factor: 2.686 ①

Journal Metrics

> CiteScore: 2.11 ①



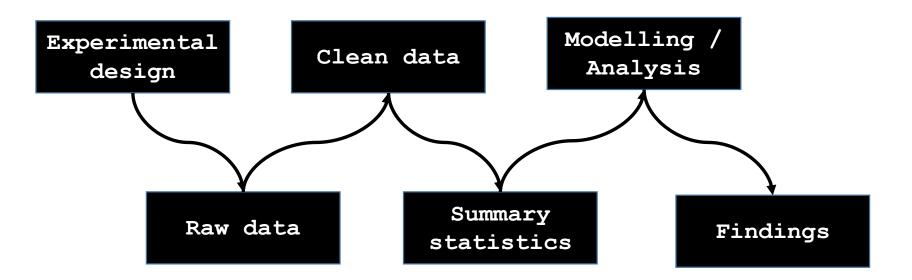
### p-values do not define a study

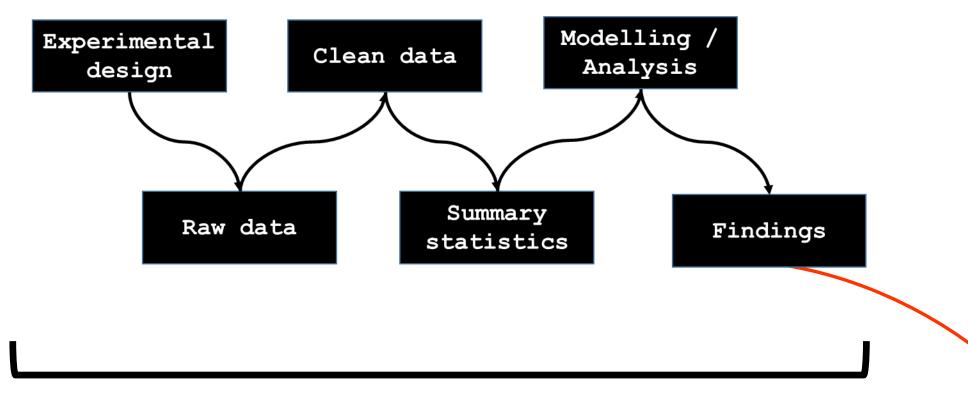


Statistics: P values are just the tip of the iceberg

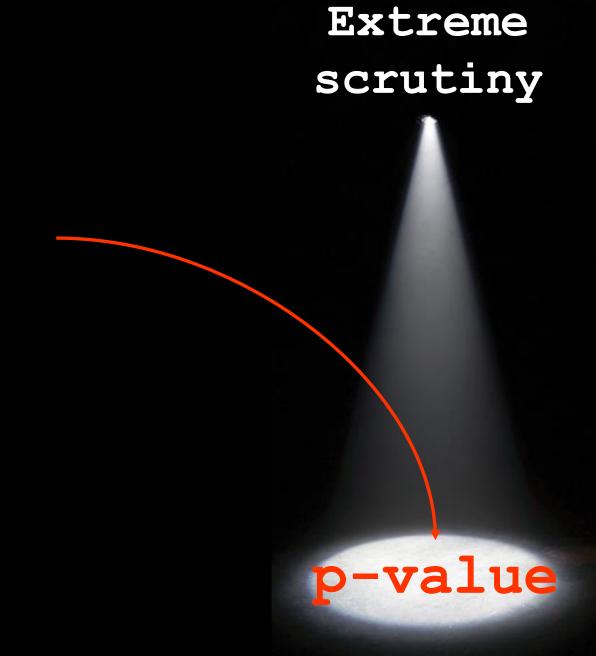
Jeffrey T. Leek & Roger D. Peng

28 April 2015





Little debate



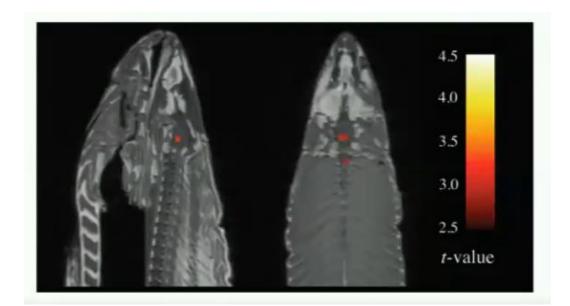
### Ig Noble research - 2012 Neuroscience winner

### Major Journal of Serendipitous and Unexpected Results

#### Neural Correlates of Interspecies Perspective Taking in the Post-Mortem Atlantic Salmon: An Argument For Proper Multiple Comparisons Correction

Craig M. Bennett 1\*, Abigail A. Baird 2, Michael B. Miller 1 and George L. Wolford 3

<sup>&</sup>lt;sup>3</sup>Department of Psychological and Brain Sciences, Moore Hall, Dartmouth College, Hanover, NH 03755



One mature Atlantic Salmon (Salmo salar) participated in the fMRI study. The salmon measured approximately 18 inches long, weighed 3.8 lbs, and was not alive at the time of scanning. It is not known if the salmon was male or female, but given the post-mortem state of the subject this was not thought to be a critical variable.

The task administered to the salmon involved completing an open-ended mentalizing task. The salmon was shown a series of photographs depicting human individuals in social situations with a specified emotional valence, either socially inclusive or socially exclusive. The salmon was asked to determine which emotion the individual in the photo must have been experiencing.

Several active voxels were observed in a cluster located within the salmon's brain cavity (see Fig. 1). The size of this cluster was  $81 \text{ mm}^3$  with a cluster-level significance of p = 0.001.

Either we have stumbled onto a rather amazing discovery in terms of post-mortem ichthyological cognition, or there is something a bit off with regard to our uncorrected statistical approach.

<sup>&</sup>lt;sup>1</sup>Department of Psychology, University of California at Santa Barbara, Santa Barbara, CA 93106

<sup>&</sup>lt;sup>2</sup>Department of Psychology, Blodgett Hall, Vassar College, Poughkeepsie, NY 12604

### What is your research hypothesis?

Article

# Too True to be Bad: When Sets of Studies With Significant and Nonsignificant Findings Are Probably True

Social Psychological and Personality Science 2017, Vol. 8(8) 875-881 © The Author(s) 2017 Reprints and permission: sagepub.com/journalsPermissions.nav DOI: 10.1177/1948550617693058 journals.sagepub.com/home/spp





Daniël Lakens<sup>1</sup> and Alexander J. Etz<sup>2</sup>

### Consider the following example:

- Starting out,  $H_0$  and  $H_1$  are equally likely
  - α is controlled for 0.05
  - The study has 80% power
  - What is the most likely outcome?
  - A True positive, B True negative,
  - C False Positive, D False negative

### Result

 What is the most likely outcome? A - True positive - 40% B - True negative - 47.5% C - False Positive - 2.5% D - False negative - 10% What is the best strategy to improve this outcome? Pick a better hypothesis If our alternate hypothesis is more likely... •  $H_0 = 40\%$  and  $H_1 60\%$ A - True positive - 48% B - True negative - 38% C - False Positive - 2%

D - False negative - 12%

# Today



### Don't do what Donny Dont does!



"In short, peer review misses all the hard stuff, and a worrying amount of the easy stuff"

James Heathers, Northwestern University

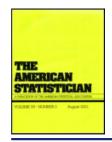
### #datathugs



### Brian Wansink: The grad student who never said no

"Every day we would scratch our heads, ask "Why," and come up with another way to reanalyze the data with yet another set of plausible hypotheses. Eventually we started discovering solutions"

# The winds of change



#### The American Statistician



ISSN: 0003-1305 (Print) 1537-2731 (Online) Journal homepage: http://amstat.tandfonline.com/loi/utas20

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

Ronald L. Wasserstein & Nicole A. Lazar

Eur J Epidemiol (2016) 31:337–350 DOI 10.1007/s10654-016-0149-3

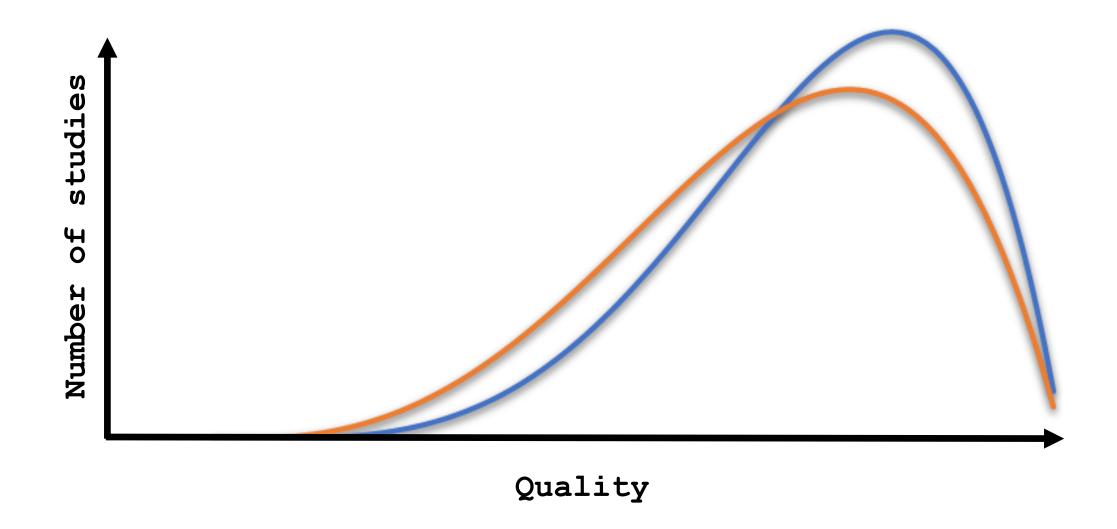


#### ESSAY

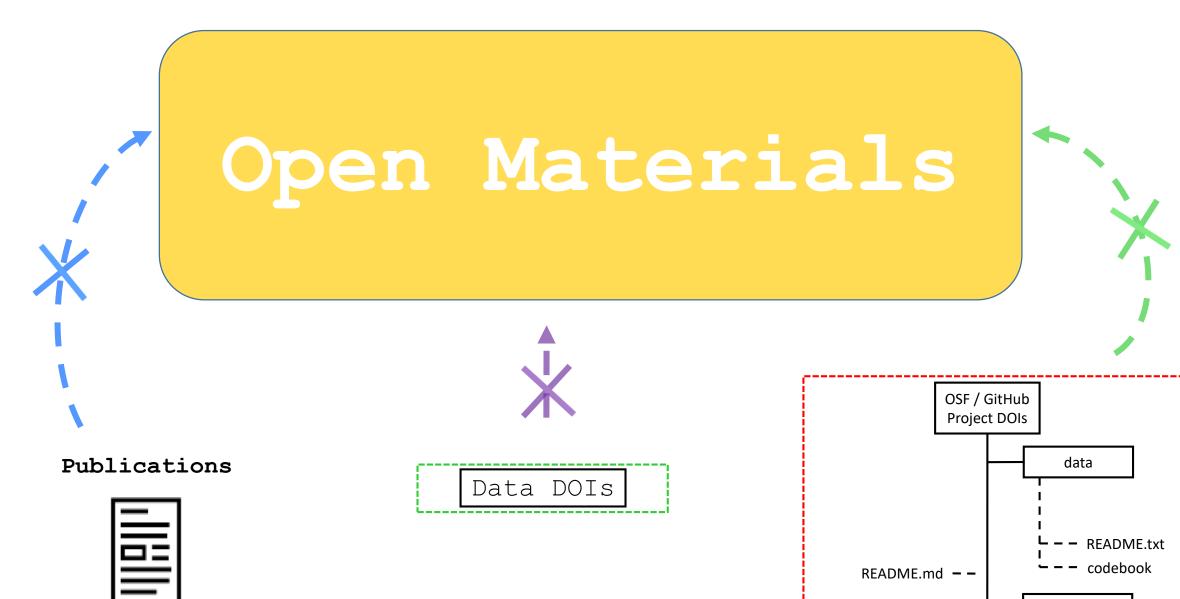
### Statistical tests, *P* values, confidence intervals, and power: a guide to misinterpretations

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>

### Can we shift the distribution?



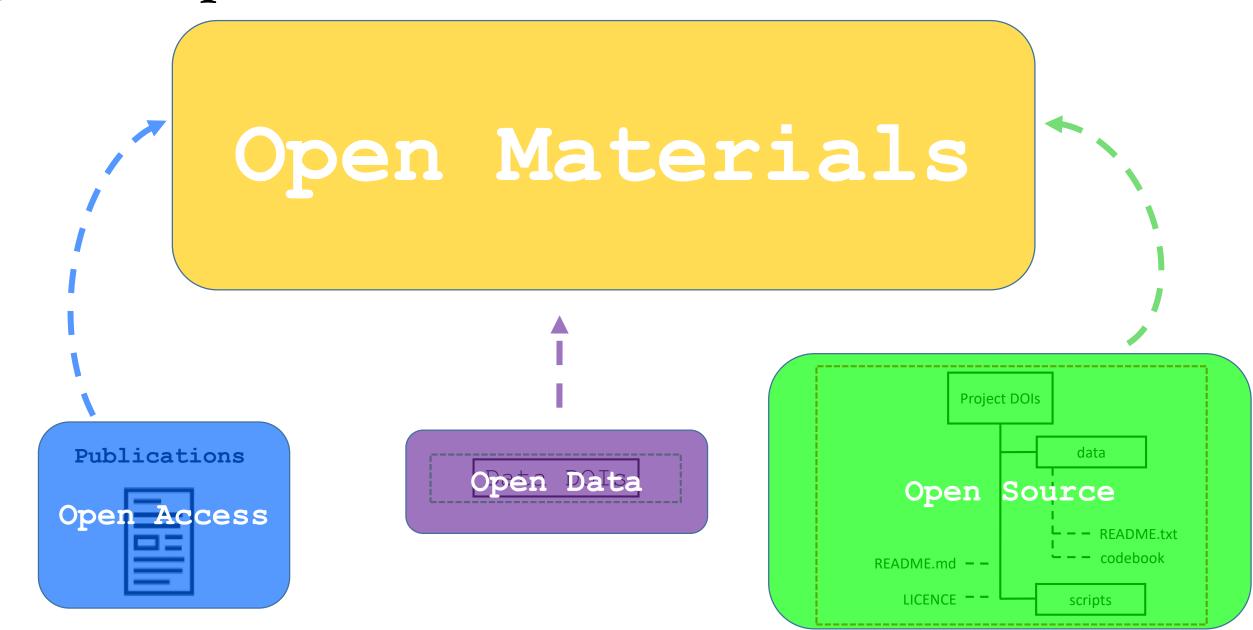
### The 'Closed Science Picture'



LICENCE

scripts

### The 'Open Science Picture'



### Can I see your data and code?

#### 1989

\* Corresponding author.

#### 1999

\* Corresponding author. Mailing address: Institute of Human Virology, 725 West Lombard St., Rm. N649, University of Maryland, Baltimore, MD 21201. Phone: (410) 706-4680. Fax: (410) 706-4694. E-mail: devico@umbi.umd.edu.

#### 2009

\* Corresponding author. Mailing address: Department of Biochemistry and Molecular Biology, The Pennsylvania State University, 201 Althouse Laboratory, University Park, PA 16802. Phone: (814) 863-8705. Fax: (814) 865-7927. E-mail: cec9@psu.edu.

#### Also 2019



### Single-Cell Virus Sequencing of Influenza Infections That Trigger Innate Immunity

Finally, we process the annotated cell-gene matrix in R to generate the plots shown in this paper. This analysis utilized a variety of R and Bioconductor (90) packages, including Monocle (91, 92) and ggplot2. A Jupyter notebook that performs these analyses is at <a href="https://github.com/jbloomlab/IFNsorted\_flu\_single\_cell/blob/master/monocle\_analysis.ipynb">https://github.com/jbloomlab/IFNsorted\_flu\_single\_cell/blob/master/monocle\_analysis.ipynb</a>,

2019

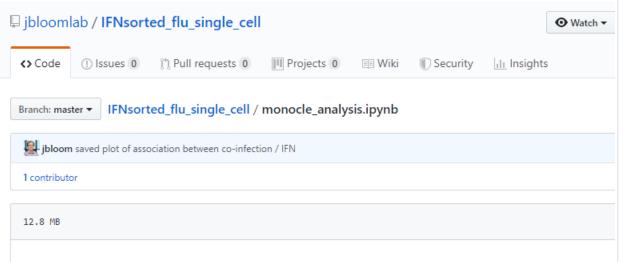


"I'm the 38th author..."
"Wow, that sucks."



#### Guidelines for the use and interpretation of assays for monitoring autophagy (3rd edition).

Klionsky DJ, Abdelmohsen K, Abe A, Abedin MJ, Abeliovich H, Acevedo Arozena A, Adachi H, Adams CM, Adams PD, Adeli K, Adhihetty PJ, Adler SG, Agam G, Agarwal R, Aghi MK, Agnello M, Agostinis P, Aguirre-Ghiso J, Airoldi EM, Ait-Si-Ali S, Akematsu T, Akporiaye ET, Al-Rubeai M, Albaiceta GM, Albanese C, Albani D, Albert ML, Aldudo J, Algül H, Alirezaei M, Alloza I, Almasan A, Almonte-Beceril M, Alnemri ES, Alonso C, Altan-Bonnet N, Altieri DC, Alvarez S, Alvarez-Erviti L, Alves S, Amadoro G, Amano A, Amantini C, Ambrosio S, Amelio I, Amer AO, Amessou M, Amon A, An Z, Anania FA, Andersen SU Andley UP, Andreadi CK, Andrieu-Abadie N, Anel A, Ann DK, Anoopkumar-Dukie S, Antonioli M, Aoki H, Apostolova N, Aguila S, Aguilano K, Araki K, Arama E, Aranda A, Araya J, Arcaro A, Arias E, Arimoto H, Ariosa AR, Armstrong JL, Arnould T, Arsov I, Asanuma K, Askanas V, Asselin E, Atarashi R, Atherton SS Atkin JD, Attardi LD, Auberger P, Auburger G, Aurelian L, Avalli R, Avagliano L, Avantaggiati ML, Avrahami L, Awale S, Azad N, Bachetti T, Backer JM, Bae DH, Bae JS, Bae ON, Bae SH, Baehrecke EH, Baek SH, Baghdiguian S, Bagniewska-Zadworna A, Bai H, Bai J, Bai XY, Bailly Y, Balaji KN, Balduini W, Ballabio A, Balzan R, Banerjee R, Bánhegyi G, Bao H, Barbeau B, Barrachina MD, Barreiro E, Bartel B, Bartolomé A, Bassham DC, Bassi MT, Bast RC Jr Basu A, Batista MT, Batoko H, Battino M, Bauckman K, Baumgarner BL, Bayer KU, Beale R, Beaulieu JF, Beck GR Jr, Becker C, Beckham JD, Bédard PA Bednarski PJ, Begley TJ, Behl C, Behrends C, Behrends CM, Behrns KE, Bejarano E, Belaid A, Belleudi F, Bénard G, Berchem G, Bergamaschi D, Bergami M, Berkhout B, Berliocchi L, Bernard A, Bernard M, Bernassola F, Bertolotti A, Bess AS, Besteiro S, Bettuzzi S, Bhatlacharyya S, Bhutla SK, Biagosch C, Bianchi MW, Biard-Piechaczyk M, Billes V, Bincoletto C, Bingol B, Bird SW, Bitoun M, Bjedov I, Blackstone C, Blanc L, Blanco GA, Blomhoff HK, Boada-Romero E, Böckler S, Boes M, Boesze-Battaglia K, Boise LH, Bolino A, Boman A, Bonaldo P, Bordi M, Bosch J, Botana LM, Botti J, Bou G, Bouché M. Bouchecareilh M, Boucher MJ, Boulton ME, Bouret SG, Boya P, Boyer-Guittaut M, Bozhkov PV, Brady N, Braga VM, Brancolini C, Braus GH, Bravo-San Pedro JM, Brennan LA, Bresnick EH, Brest P, Bridges D, Bringer MA, Brini M, Brito GC, Brodin B, Brookes PS, Brown EJ, Brown K, Broxmeyer HE, Bruhat A, Brum PC, Brumell JH, Brunetti-Pierri N, Bryson-Richardson RJ, Buch S, Buchan AM, Budak H, Bulavin DV, Bultman SJ, Bultynck G, Bumbasirevic V, Burelle Y, Burke RE, Burmeister M, Bütikofer P, Caberlotto L, Cadwell K, Cahova M, Cai D, Cai J, Cai Q, Calatayud S, Camougrand N, Campanella M, Campbell GR, Campbell M, Campello S, Candau R, Caniggia I, Cantoni L, Cao L, Caplan AB, Caraglia M, Cardinali C, Cardoso SM, Carew JS, Carleton LA, Carlin CR, Carloni S, Carlsson SR, Carmona-Gutierrez D, Carneiro LA, Carnevali O, Carra S, Carrier A, Carroll B, Casas C, Casas J, Cassinelli G, Castets P, Castro-Obregon S, Cavallini G, Ceccherini I, Cecconi F, Cederbaum Al, Ceña V, Cenci S, Cerella C, Cervia D, Cetrullo S, Chaachouay H, Chae HJ, Chagin AS, Chai CY, Chakrabarti G, Chamilos G, Chan EY, Chan MT, Chandra D, Chandra P, Chang CP, Chang RC, Chang TY, Chatham JC, Chatteriee S, Chauhan S, Che



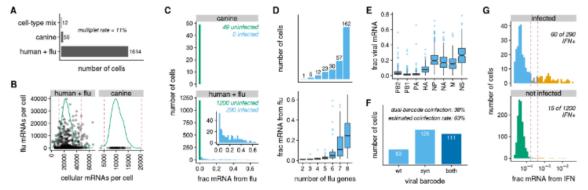
#### Table of Contents

- · Analyze viral features associated with IFN induction
  - Setup for analysis
    - Load / install packages
    - · Notebook-wide variables / functions
  - Get cell-gene matrices
    - Specify cell types
    - Load cell-gene matrix
  - Count cells and annotate multiplets
    - Annotate cross-celltype multiplets
    - Number of cells and multiplet frequency
    - Plot summarizing cell counts and multiplets
  - Filter multiplets and low-quality cells
    - Remove cross-celltype multiplets
    - Number of cellular and flu mRNAs, bounds for filtering
    - · Plot cellular / flu mRNAs with filters
    - · Filter cells with extreme mRNA amounts
  - Call infection / gene presence from canine cell thresholds
    - · Constant fraction or number of mRNAs from flu?
    - · Confirm equal mix of flu barcodes in canine cells
    - Look at segment frequencies
    - Get human cells for infection-status calling
    - · Compute P-value flu is above background
    - · Call infected cells by amount of total flu
    - Call gene presence/absence

#### Figures for paper

We have made all of the plots above, and saved some of them to the figures directory already by using the isfig=TRUE argument to saveShowPlot. However, there are others that we want to assemble into multi-panel figures. We do that here.

First, we assemble a figure that shows the calling of cells, infected cells, and IFN+ cells:



Now a supplementary figure to the one above with the single-cell transcriptomic data:



### FAIR is a part of our life now!

#### **Data Guidelines**

Wellcome

- 1. Background
  - 1.1 Open Data Policy
  - 1.2 Fair Data Principles
- 2. Share Your Data in 3 Steps
  - 2.1 Prepare Your Data for Sharing
  - 2.2 Select a Repository
  - 2.3 Add a Data Availability Statement to Your Manuscript
  - 2.4 Linking your datasets to your article

Some types of data benefit from visualization within the article. Wellcome Open Research welcomes the submission of manuscripts featuring <u>Plot.ly interactive figures</u> and <u>Code Ocean compute capsules</u>. For further detail, please <u>contact us</u>.



#### Social Research Ethics Committee (SREC)

#### ETHICS APPROVAL FORM

https://www.ucc.ie/en/research/about/ethics/

https://www.ucc.ie/en/media/research/research/researchdocuments/UCCCodeofResearchConduct.pdf. SREC advises against storing research data on non UCC approved cloud-based storage services. Physical data must be stored in a locked cabinet and you must specify who has permission to access this data.

#### Research Data Management



Good data governance and stewardship are key components of good research practice. In this regard, Science Foundation Ireland supports that research data should be Findable, Accessible, Interoperable and Reusable (FAIR)\*. Appropriate data management and data sharing are fundamental to all stages of the research process and support high quality, reproducible research. As such, access to research data arising in whole or in part from SFI funding should be as open as possible.



#### **FAIR Data Management**

Describe the approach to data management that will be taken during and after the project, including who will be responsible for data management and data stewardship. The word limit is <u>500 words</u>.



A set of Digital Object Compliance principles that describes the properties of digital objects that enables them to be findable, accessible, interoperable and reproducible (FAIR).

<sup>&</sup>lt;sup>4</sup> Data management should follow the FAIR guiding principles (Findability, Accessibility, Interoperability & Reusability). See, for example, Wilkinson, M. D. et al. (2016) The FAIR Guiding Principles for Scientific Data Management and Stewardship. Full text: <a href="http://www.nature.com/articles/sdata201618">http://www.nature.com/articles/sdata201618</a>. It is required that all staff and student researchers store those data which are required to replicate research findings, and the information required to enable re-use of data. Details of the UCC policy on research data storage can be found in section 8 of the Code of Research Conduct (2016):

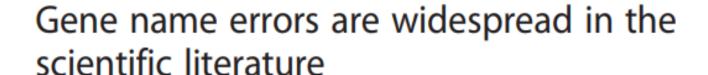
### Beware of default settings

Ziemann et al. Genome Biology (2016) 17:177 DOI 10.1186/s13059-016-1044-7

Genome Biology

#### COMMENT

**Open Access** 





Mark Ziemann<sup>1</sup>, Yotam Eren<sup>1,2</sup> and Assam El-Osta<sup>1,3\*</sup>

#### Abstract

The spreadsheet software Microsoft Excel, when used with default settings, is known to convert gene names to dates and floating-point numbers. A programmatic scan of leading genomics journals reveals that approximately one-fifth of papers with supplementary Excel gene lists contain erroneous gene name conversions.

frequently reused. Our aim here is to raise awareness of the problem.

We downloaded and screened supplementary files from 18 journals published between 2005 and 2015 using a suite of shell scripts. Excel files (.xls and.xlsx suffixes) were converted to tabular separated files (tsv) with ssconvert (v1.12.9). Each sheet within the Excel file was converted to a separate tsv file. Each column of data in the tsv file was screened for the presence of gene sym-

- Why did it take us until 2016 to discover this?

# \*cough\* We've known for a long time \*cough\*

### **BMC Bioinformatics**



Correspondence



Mistaken Identifiers: Gene name errors can be introduced inadvertently when using Excel in bioinformatics

Barry R Zeeberg<sup>†1</sup>, Joseph Riss<sup>†2</sup>, David W Kane<sup>3</sup>, Kimberly J Bussey<sup>1</sup>, Edward Uchio<sup>4</sup>, W Marston Linehan<sup>4</sup>, J Carl Barrett<sup>2</sup> and John N Weinstein<sup>\*1</sup>

2004

### But it doesn't end there

### Date and time expressed according to ISO 8601 [refresh]

**Date** 2019-10-15

Date and time in 2019-10-15T19:49:52+00:00

UTC

2019-10-15T19:49:52Z

20191015T194952Z

Week 2019-W42

Date with week 2019-W42-2

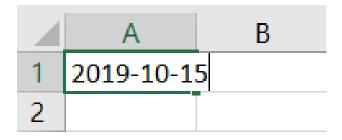
number

**Date without** --10-15<sup>[1]</sup>

year

Ordinal date 2019-288

- YYYY-MM-DD or YYYYMMDD
- Type this into Excel



- And hit return

| 1 | А          | В |
|---|------------|---|
| 1 | 15/10/2019 |   |
| 2 |            |   |

- DD/MM/YYYY

### Beware of default settings

Print



Data and computer code should be made publicly available at an early stage - or else ... esarastudillo

Last week we learned a famous 2010 academic paper, relied on by political big-hitters to

bolster arguments for austerity cuts, contained significant errors; and that those errors came
down to misuse of an Excel spreadsheet.

in LinkedIn

Sadly, these are not the first mistakes of this size and nature when handling data. So what on Earth went wrong, and can we fix it?

Harvard's <u>Carmen Reinhart</u> and <u>Kenneth Rogoff</u> are two of the most respected and influential academic economists active today.

## What can you do?



Reproducibility is important because the you of 3 months ago is terrible at answering email! - @tracykteal at #2016dssummit

**Follow** 

1:17 PM - 26 Oct 2016 from Manhattan, NY



# Get organised

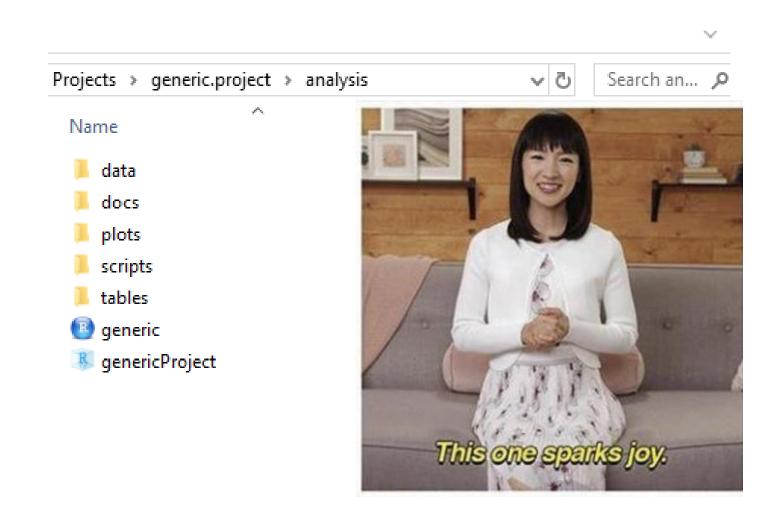
- Help your future-self

| B Palmer Medicine Files > 4a Project            | Pyrosequencing analysis  | > Pyrosequencing Paner | Draft Paner incl Figs > Submission    | > IVI Resubmission     | > JVI_resubmission_files > Final Final version |
|---|--------------------------|------------------------|---------------------------------------|------------------------|--|
| b_r diffici_tviculciffc_r files * 4a r toject * | i yrosequenenig_unuiysis | * Tyrosequencing_raper | Prair_i apei_inci_i igs P Sabinission | * JVI_ICC3GDITII331011 | 7 7 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1        |

| Name                                   | Date modified    |
|--|------------------|
| Cover_letter_B_A_Palmer_Sept_2014      | 10/09/2014 17:05 |
| 🔓 Fig_1_Sept_14                        | 11/09/2014 10:31 |
| Fig_1_Sept_14                          | 10/09/2014 23:07 |
| 🔓 Fig_2_Sept_14                        | 11/09/2014 10:31 |
| Fig_2_Sept_14                          | 10/09/2014 23:07 |
| Fig_3_Sept_14                          | 11/09/2014 10:31 |
| Fig_3_Sept_14                          | 10/09/2014 23:07 |
| 🔓 Fig_4_Sept_14                        | 11/09/2014 10:31 |
| Fig_4_Sept_14                          | 10/09/2014 23:07 |
| 🔓 Fig_5_Sept_14                        | 11/09/2014 10:33 |
| Fig_5_Sept_14                          | 10/09/2014 23:07 |
| HCV_UDPS_B_A_Palmer_Sept_14            | 17/09/2014 12:21 |
| Response_to_Reviewer_Sept_14           | 10/09/2014 22:42 |
| Supplementry_Figure_B_A_Palmer_Sept_14 | 29/08/2014 13:21 |
| Supplementry_Figure_B_A_Palmer_Sept_14 | 10/09/2014 22:31 |
| Tables_B_A_Palmer_Sept_2014            | 10/09/2014 22:09 |



# Define a generic project structure



### Give your files and folders informative names

- Make your file names:
  - Machine readable
  - 2. Human readable
  - 3. Work with default ordering

### NO

#### Name

- All unique 4a amino acid Sequences (B-N).fas
- All unique 4a amino acid Sequences (B-N).meg
- All\_AA\_haplotypes.meg
- All\_AA\_haplotypes\_with\_clonal\_sequences.meg
- BS100\_AA\_with\_clones
- BS100\_AA\_with\_clones.nwk
- BS1000\_AA\_pyro&clones
- BS1000\_AA\_pyro&clones.nwk
- BS1000\_AA\_pyro\_only
- BS1000\_AA\_pyro\_only.nwk
- BS1000\_Unique\_Clonal\_AA

### Yes

Projects > 2016-08-08\_RespPCT > analysis > scripts

Name

R 01\_clean\_data
R 02\_plots
R 03\_tables
R 04\_stats\_analysis
R 05\_post\_hoc\_stats
R functions
R randomization
R tables

# Putting the pieces together

A: Define a project structure

B: Set a naming convention

C: Use scripted workflows

D: Digital notebooks

Reproducible research

E: Version control

F: Data packaging

# Coming to a UCC near you in 2020



BIG NEWS: We've just received approval for a new postgraduate module where we'll teach reproducible scripted workflows through #rstats!

Thanks to all who helped get it to this point, but special mention to @statsepi who set the ball rolling 2

@UCC > Modules > PG6030 😸

**Module Content:** The module will introduce fundamental concepts of reproducible research alongside hands-on training in the R programming language. Students will be instructed on data collation, curation and management techniques that will serve as a foundation towards downstream visualisation, analysis and reporting via scripted, reproducible workflows.

- Email b.palmer@ucc.ie to register your interest
- First class likely to run in April 2020

### Questions...



There is no learning without having to pose a question. And a question requires doubt.

