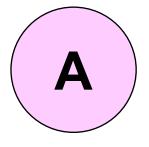
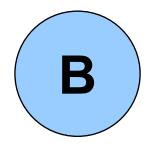


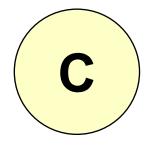
# Research Integrity or How to be a Good Scientist

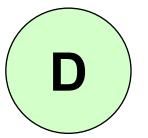
Jo Montgomery, Sarah Inglesfield, Anne Segonds-Pichon, Laura Biggins and Simon Andrews





















# **An illustration**



# **Research Integrity**

MMR = Measles, Mumps and Rubella





**Autism** 



#### 1998 Link MMR autism

Wakefield et al.

n=12 children

https://pubmed.ncbi.nlm.nih.gov/9500320/

#### 2010

Retraction of the Wakefield et al. paper

https://www.ncbi.nlm.nih.gov/pmc/articles/PMC2831678/

#### 2010

Andrew Wakefield found guilty of serious professional misconduct

https://www.bmj.com/content/340/bmj.c593

1999 No link MMR autism Taylor *et al.* 

n=498 children

https://pubmed.ncbi.nlm.nih.gov/10376617

2002-2005: No link MMR autism

Many more studies

https://pubmed.ncbi.nlm.nih.gov/15366972/ https://pubmed.ncbi.nlm.nih.gov/15877763/ 2001: No link MMR autism Dales et al.

n=600-1900 children each year over 14 years

https://pubmed.ncbi.nlm.nih.gov/11231748/

# **Research Integrity**





Lancet. 1998 Feb 28;351(9103):637-41. doi: 10.1016/s0140-6736(97)11096-0.

Ileal-lymphoid-nodular hyperplasia, non-specific colitis, and pervasive developmental disorder in children

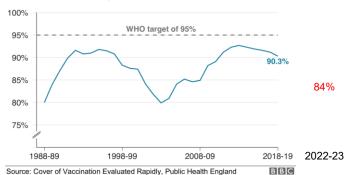
A J Wakefield <sup>11</sup>, S H Murch, A Anthony, J Linnell, D M Casson, M Malik, M Berelowitz, A P Dhillon, M A Thomson, P Harvey, A Valentine, S E Davies, J A Walker-Smith

Affiliations + expand
PMID: 9500320 DOI: 10.1016/s0140-6736(97)11096-0

- Scientists and organisations across the world spent a great deal of time and money refuting the results of a minor paper in the Lancet.
- MMR vaccination: 1995: 95% to 2002: 81 %
- Measles outbreaks in the UK in 2008 and 2009
- 2020: Uptake of MMR vaccine: 91% (still below herd immunity)

#### MMR first dose coverage is falling in England

% of children who completed first dose of vaccine at 24 months



# **Research Integrity**



Rise in measles cases prompts vaccination campaign in England

UKHSA declares national incident as figures suggest more than 3.4 million children have not had MMR jab



- **2016**: UK declared measles free Now lost this status
- Increases in number of measles cases
- 2022-23: 84% children in England
- (74% in London, vs 90% South West)
- Need vaccination rate of 95%
- Current decline due to:

Vaccine misinformation
Not serious?
Difficulty accessing appointments
Impact of covid pandemic

# Obvious examples are obvious, but...





# Outline of the course







# What is Research Integrity?



# Research Integrity a.k.a. Scientific Integrity

- Scientific integrity (From Wikipedia): Scientific integrity deals with 'best practices' or rules of professional practice of researchers.
- Organisation for Economic Co-operation and Development (OECD) report, 2007 replication (or reproducibility) crisis and the fight against scientific misconduct.

# Research Integrity a.k.a. Scientific Integrity

 The replication crisis - scientific studies are difficult or impossible to replicate or reproduce.

## **Psychology:**

Open Science Collaboration
(100 papers from 2012)
36% of the replications yielded significant findings vs 97% in the original studies.

## **Cancer Research:**

Reproducibility Project: Cancer Biology (53 papers from 2010 to 2012)
25% experiments could be reproduced.
Replication effect sizes were 85% smaller on average than the original findings.

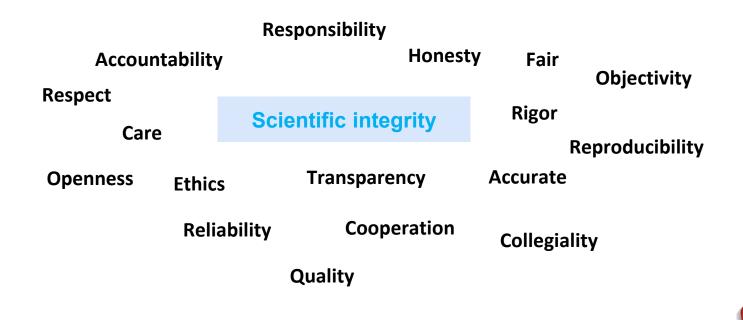


# Which words are most important/synonymous with integrity? Exercise





# Research Integrity Many words



Accurate

Care

Collegiality

Cooperation

**Ethics** 

Fair

Honesty

Objectivity

**Openness** 

Quality

Reliability

Reproducibility

Respect

Responsibility

Rigor

Transparency

# Research Integrity More than words

**Scientific integrity** 



# **Research Integrity**

Research Integrity means conducting, reviewing and reporting research in a responsible way that allows ourselves and others to have confidence that our research is trustworthy and of the highest quality. It also sets the culture of the Institute as an inclusive, respectful, open and outstanding place to work.

**BI-RES-005 RESEARCH INTEGRITY POLICY** 





# Research Integrity in practice



# Research Integrity: In Practice

Accountability Accurate Care Collegiality Cooperation **Ethics** Fair Honesty Objectivity **Openness** Quality Reliability Reproducibility

Respect

Rigor

Responsibility

Transparency

Question Results **Experimental Design Data Analysis** Sample Size **Data Exploration Experiment** 



# Research Integrity in practice Experimental design



Accurate

Care

**Collegiality** 

Cooperation

**Ethics** 

Fair

Honesty

**Objectivity** 

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Quality

**Reliability** 

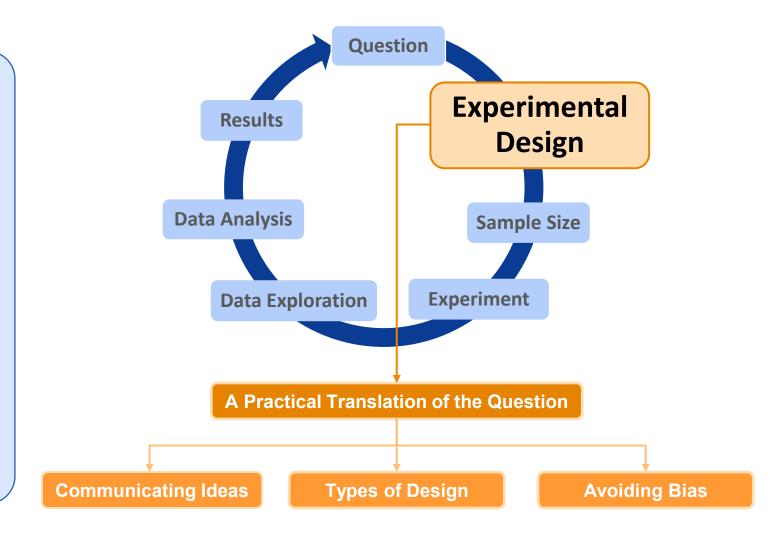
Reproducibility

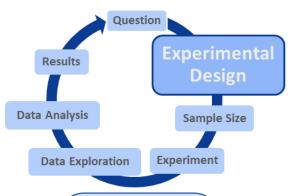
Respect

Responsibility

Rigor

Transparency





Accurate

Care

Collegiality

#### Cooperation

Ethics

Fair

Honesty

Objectivity

Openness

Quality

Reliability

Reproducibility

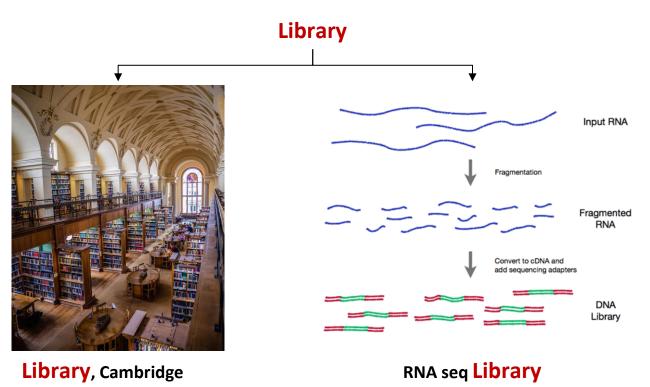
Respect

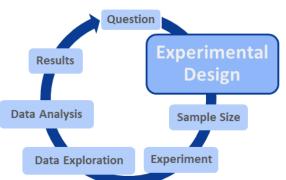
Responsibility

Rigor

Transparency

## **Clear Communication**





Accurate

Care

**Collegiality Cooperation** 

Ethics

Fair

Honesty

Objectivity

Openness

Quality

Reliability

Reproducibility

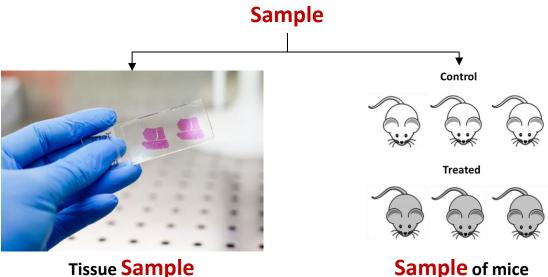
Respect

Responsibility

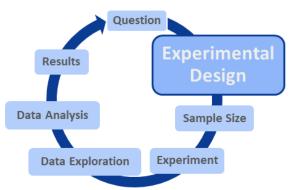
Rigor

Transparency

## **Clear Communication**



- Different words to describe the same data/graphs
- Different traditions in different labs, areas of science



## **Appropriate Type of Design**

Distinguish real differences from experimental artefacts...

...But experiments can be big and complicated

Accountability

Accurate

Care

Collegiality

#### Cooperation

**Ethics** 

Fair

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Honesty Objectivity

Openness

Quality

#### Reliability

Reproducibility

Respect

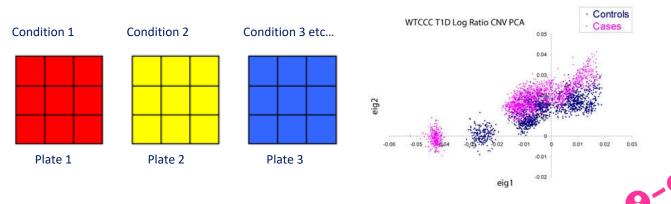
Responsibility

**Rigor** 

Transparency

#### Seminal Wellcome Trust GWAS Study:

- 14000 cases of 7 diseases & 3000 shared controls
- Each processed at different sites and genotyped on distinct series of plates



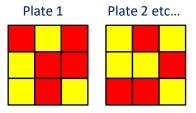
Differences between control and cases are confounded by plate

# Results Experimental Design Data Analysis Sample Size Data Exploration Experiment

## **Appropriate Type of Design**

#### GenADA multi-site collaborative study:

- 875 Alzheimers patients, 850 controls, 9 sites
- Randomised Block Design



#### Accountability

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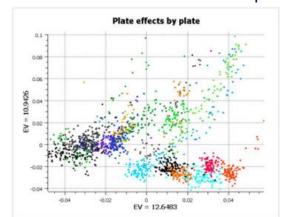
Respect

Responsibility

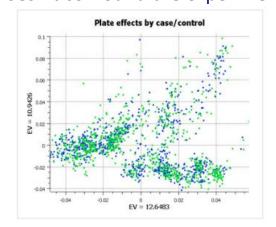
Rigor

Transparency

## Still have differences between plates



### Doesn't confound the experiment



https://blog.goldenhelix.com/stop-ignoring-experimental-design-or-my-head-will-explode/



## **Avoiding Bias**

Humans are not always good at remaining objective...



**Cognitive Bias = Pareidolia** 

Biases can also impact our experiments



# **Confirmation bias Exercise**





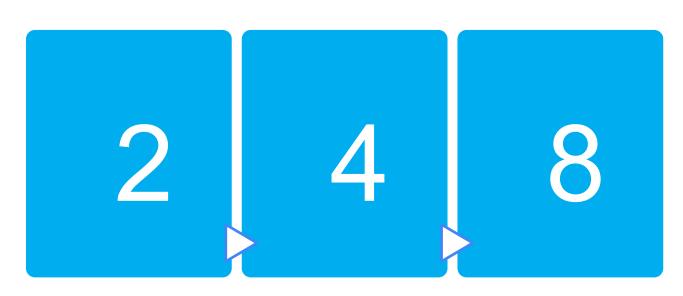
# A Quick Exercise on Confirmation Bias and Hypothesis Testing

You will be presented with 3 numbers in a sequence.

You need to guess the **rule** that governs the sequence.

You can suggest any 3 numbers you like, and we will tell you whether or not your sequence follows the rule.

# A Quick Exercise on Confirmation Bias and Hypothesis Testing





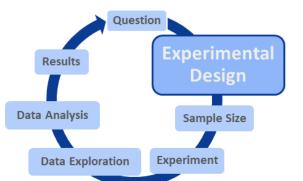
# A Quick Exercise on Confirmation Bias and Hypothesis Testing

What's the rule?



# **Confirmation bias**

What **Objective** confirms facts your beliefs What you see



# **Experimenter Bias**

Accountability

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Cooperation

**Ethics** 

Fair

Honesty

#### **Objectivity**

Openness

Quality

Reliability

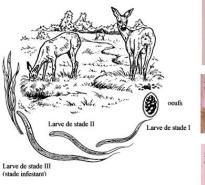
#### Reproducibility

Respect

Responsibility

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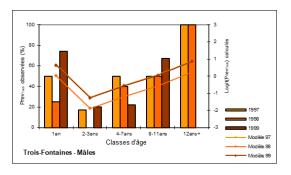




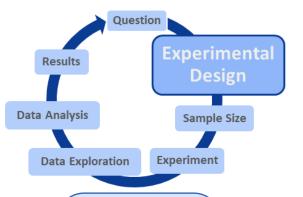








Scientists should be **blind** ... if they want to be **objective**!



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Openness

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Reliability

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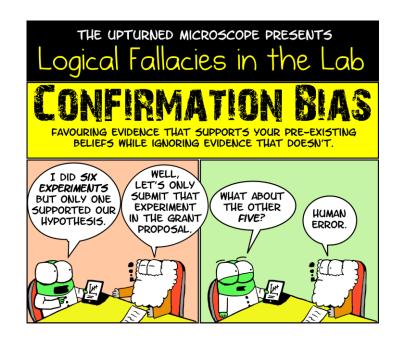
Rigor

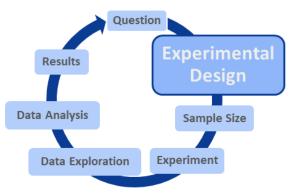
Transparency

## **Blinding**

is important to avoid

## **Confirmation bias**





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#### Reproducibility

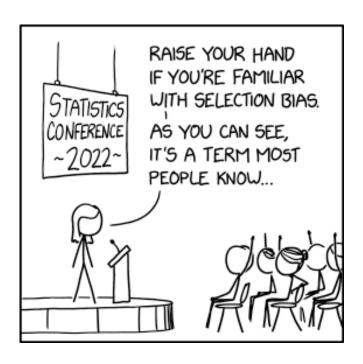
Respect

Responsibility

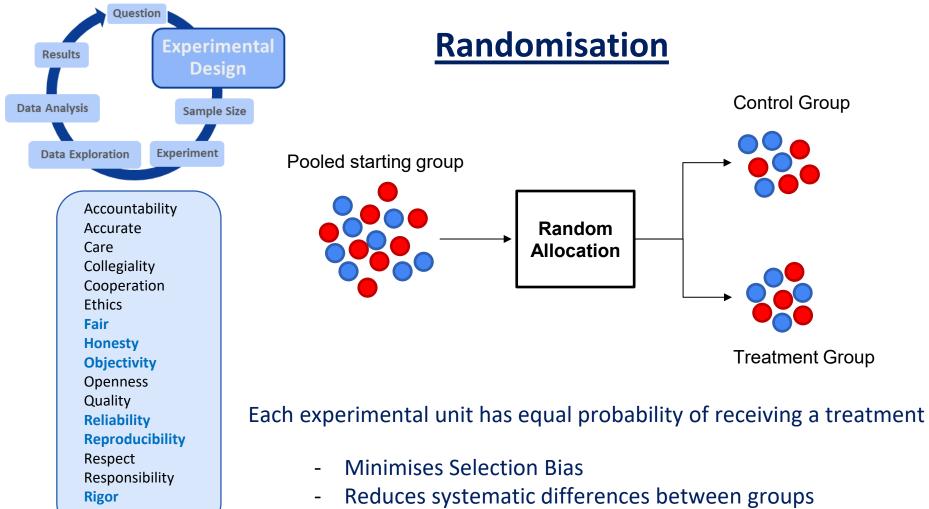
Rigor

Transparency

# **Selection Bias**

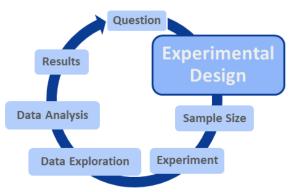


Biases in our sample populations can impact on our conclusions too



Transparency

Reduces systematic differences between groups



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**Ethics** 

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Honesty

### **Objectivity**

Openness

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#### Reproducibility

Respect

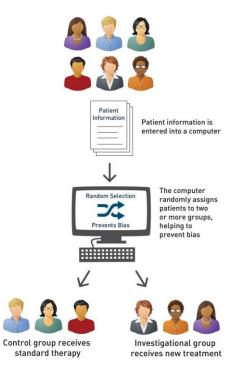
Responsibility

Rigor

Transparency

# **Random Allocation**

#### **CLINICAL TRIALS RANDOMIZATION**







#### **Random Allocation**

Data Analysis

Sample Size

Data Exploration

Experiment

Accountability Accurate

Care

Collegiality

Cooperation

**Ethics** 

Fair

**Honesty** 

**Objectivity** 

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Pick out a mouse at random, first 3 get the treatment

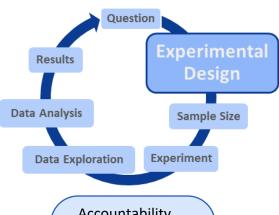
Is this random?

Does it have the potential to introduce bias?









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**Ethics** 

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Openness

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Reliability

#### Reproducibility

Respect

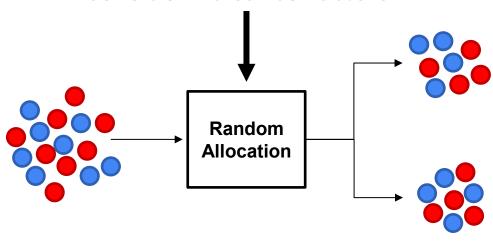
Responsibility

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Transparency

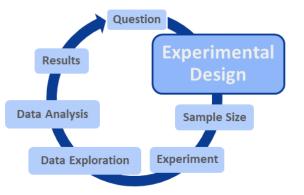
#### **Randomisation More Broadly**





Also consider randomisation **throughout** the experiment

https://eda.nc3rs.org.uk/experimental-design-allocation#randomisationstrategy



#### In a nutshell

#### Accountability

Accurate

Care

Collegiality

Cooperation

Ethics

Fair Honesty

#### **Objectivity**

Openness

Quality

Reliability

Reproducibility

Respect

Responsibility

Rigor

Transparency

#### Good experimental design...

- Translates the scientific question into lab work
- Prevents subjectivity
- Reduces effects of nuisance variables

...Is a fair way to do science



# Research Integrity in practice Sample size



Accountability

Accurate

#### Care

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Fair

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Reliability

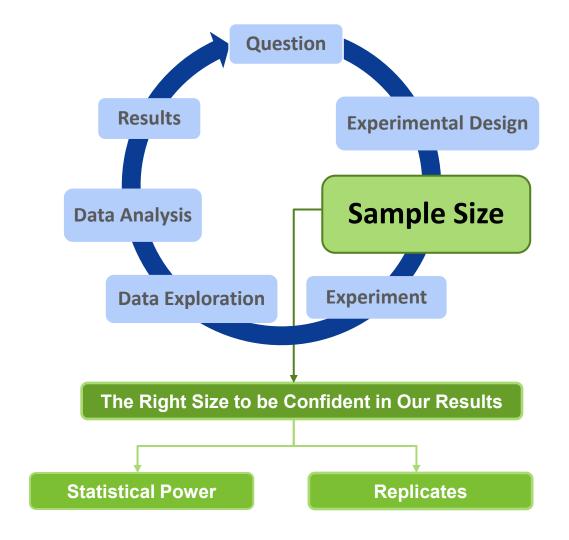
Reproducibility

Respect

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#### **Statistical power: an analogy**

Accountability

Accurate

Care

Collegiality

Cooperation

**Ethics** 

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Reliability

Reproducibility

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Responsibility

**Rigor** 

Transparency

You send your child into the basement to find a tool. The child comes back and says "it isn't there". What do you conclude?

#### In the house

"If the tool really is in the basement, what are the chances that your child would have found it?"

How long did the child spend looking?



How big is the tool?



r



How messy is the basement?



0



#### In the lab

"If there is a difference between 2 conditions, what are the chances that your experiment will pick it up (p<0.05)?"

How many mice do you look at? Sample size



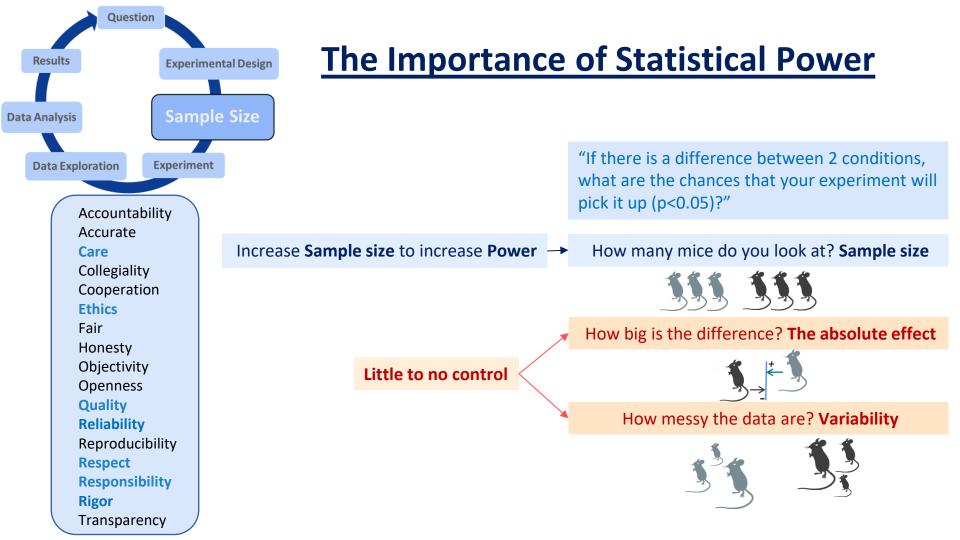
How big is the difference? **The absolute effect** 

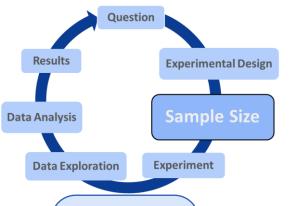


How messy the data are? Variability









#### The Importance of Statistical Power

Low Powered Studies have a greater chance of failing to detect a real effect

BUT that's all probability...

So some underpowered studies will detect a real effect

Are these results trustworthy?





Accountability

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Care

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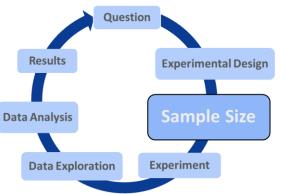
Reproducibility

Respect

Responsibility

Rigor

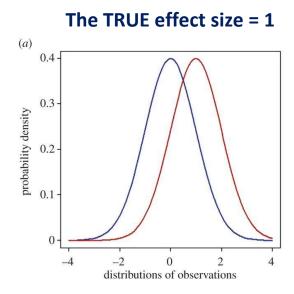
Transparency

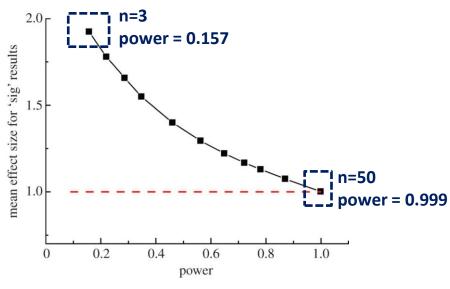


#### **The Importance of Statistical Power**

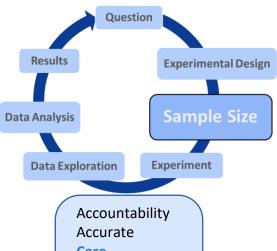
The Problem of the "Inflation Effect"...

Accountability Accurate Care Collegiality Cooperation **Ethics** Fair Honesty Objectivity **Openness** Quality Reliability Reproducibility Respect Responsibility Rigor Transparency





https://royalsocietypublishing.org/doi/10.1098/rsos.140216



Care

Collegiality

Cooperation

**Ethics** 

Fair

Honesty

Objectivity

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Quality

Reliability

Reproducibility

Respect

Responsibility

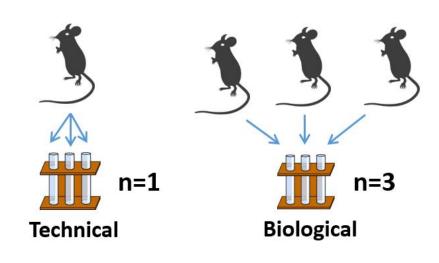
Rigor

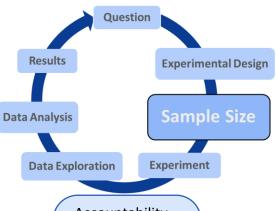
**Transparency** 

#### **Defining Replicates**

By the way: **replicates** = repeat = sample = library

#### **Technical** versus biological replicates





#### What is a Biological Replicate in Practice?



1 biological replicate: ( A

 $\mathbf{A}$ ) 1 mouse

**B**) 1 cage

Accountability
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Reliability

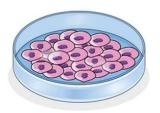
Reproducibility

Respect

Responsibility

Rigor

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1 biological replicate:



**B** ) 1 petri dish

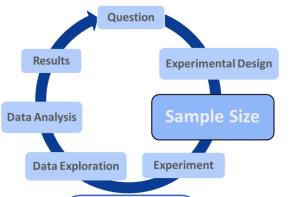


1 biological replicate:



**B** ) 1 petri dish





#### In a nutshell

Accountability

Accurate Care

Collegiality

Cooperation **Ethics** 

Fair

Honesty Objectivity

Openness

Quality

Reliability

Reproducibility

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Transparency

Sample Size & Power are key to confident results

Underpowered Studies are more likely to:

- Fail to detect real effects
- Overestimate the effect size of detected effects

More biological replicates increase our evidence

How Many? Formalise with power calculations....









# Research Integrity in practice Data Exploration & Data Analysis



Accountability

**Accurate** 

Care

**Collegiality** 

Cooperation

**Ethics** 

Fair

Honesty

Objectivity

**Openness** 

Quality

Reliability

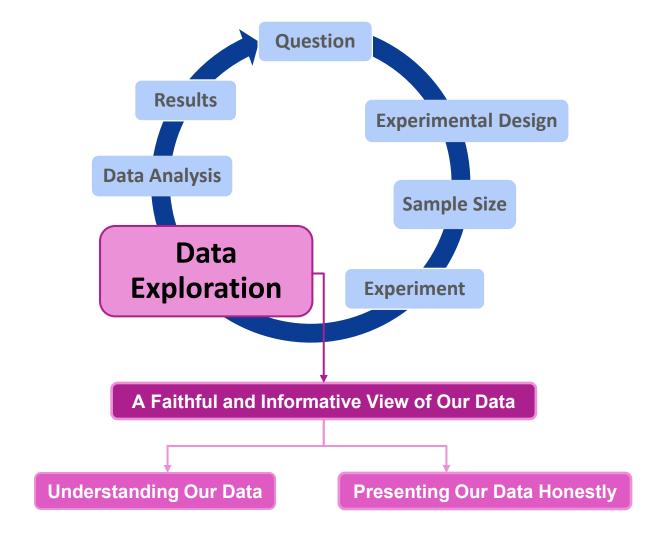
Reproducibility

Respect

Responsibility

Rigor

**Transparency** 



Accountability Accurate Care Collegiality Cooperation **Ethics** Fair Honesty **Objectivity Openness** Quality Reliability Reproducibility Respect Responsibility Rigor

**Transparency** 



**Builds Our Understanding and Confidence** 

# **Data Exploration: Understanding Our Data**

Accountability

Accurate

Care

Collegiality

Cooperation

**Ethics** 

Fair

**Honesty** 

**Objectivity** 

**Openness** 

Quality

Reliability

Reproducibility

Respect

Responsibility

**Rigor** 

**Transparency** 

#### Often we summarise our data to key values

N	182
X Mean	54.26
Y Mean	47.83
X SD	16.76
Y SD	26.93
Correlation	-0.06

Can be really useful...

...And also really not!

## **Data Exploration: Anscombe's Quartet**

Accountability

Accurate

Care

Collegiality

Cooperation

**Ethics** 

Fair

**Honesty** 

**Objectivity** 

**Openness** 

Quality

Reliability

Reproducibility

Respect

Responsibility

Rigor

**Transparency** 

"A computer should make both calculations and graphs"

Dataset 1 Dataset 2		Dataset 3		Dataset 4			
X1	Y1	X2	Y2	Х3	Y3	X4	Y4
10.00	8.04	10.00	9.14	10.00	7.46	8.00	6.58
8.00	6.95	8.00	8.14	8.00	6.77	8.00	5.76
13.00	7.58	13.00	8.74	13.00	12.74	8.00	7.71
9.00	8.81	9.00	8.77	9.00	7.11	8.00	8.84
11.00	8.33	11.00	9.26	11.00	7.81	8.00	8.47
14.00	9.96	14.00	8.10	14.00	8.84	8.00	7.04
6.00	7.24	6.00	6.13	6.00	6.08	8.00	5.25
4.00	4.26	4.00	3.10	4.00	5.39	19.00	12.50
12.00	10.84	12.00	9.13	12.00	8.15	8.00	5.56
7.00	4.82	7.00	7.26	7.00	6.42	8.00	7.91
5.00	5.68	5.00	4.74	5.00	5.73	8.00	6.89



Francis Anscombe

4 datasets
Each consisting of X and Y variable

All 4 datasets have the same summary statistics...

#### Accountability Accurate Care Collegiality Cooperation **Ethics** Fair **Honesty Objectivity**

Openness Quality

Reliability

Reproducibility

Respect

Responsibility

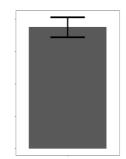
Rigor

**Transparency** 

#### Data Exploration: Anscombe's Quartet

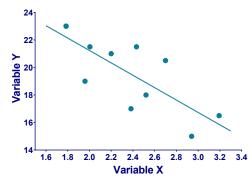
The average and spread of the conditions appears the same

	Data	aset 1	Dataset 2		Dataset 3		Dataset 4	
	<b>X1</b>	Y1	X2	Y2	Х3	Y3	Х4	Y4
N	11	11	11	11	11	11	11	11
Mean	9.00	7.50	9.00	7.50	9.00	7.50	9.00	7.50
STD	3.31	2.03	3.31	2.03	3.31	2.03	3.31	2.03
SEM	1.00	0.612	1.00	0.612	1.00	0.612	1.00	0.612



#### The relationship between X & Y can be described the same

	Correlation (r)	Line of Best Fit
x1 vs. y1	r = 0.8164	Y = 0.5001*X + 3.000
x2 vs. y2	r = 0.8162	Y = 0.5000*X + 3.001
x3 vs. y3	r = 0.8163	Y = 0.4997*X + 3.002
x4 vs. y4	r = 0.8165	Y = 0.4999*X + 3.002



#### Accountability

Accurate

Care

Collegiality

Cooperation

**Ethics** 

Fair

**Honesty** 

**Objectivity** 

**Openness** 

Quality

**Reliability** 

Reproducibility

Respect

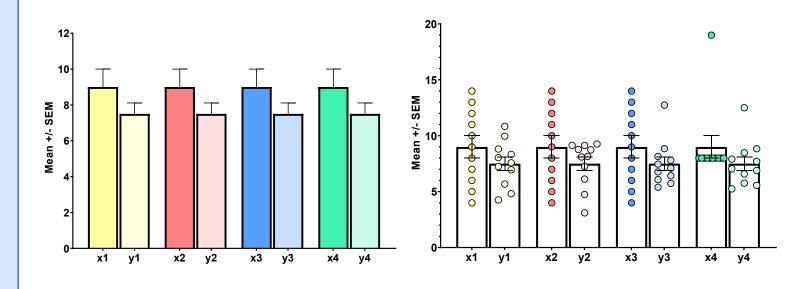
Responsibility

Rigor

**Transparency** 

## Data Exploration: Anscombe's Quartet

How does the data behave within the groups?

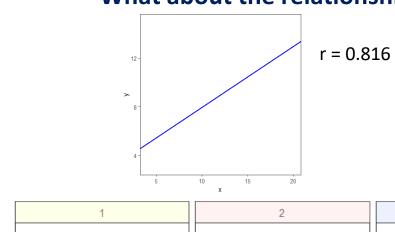


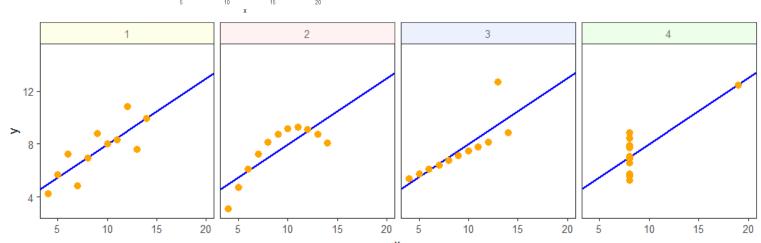
We're still not getting a complete view of the data What about the relationship between X & Y?

### **Data Exploration: Anscombe's Quartet**

What about the relationship between X & Y?

Y = 0.500\*X + 3.00





Accountability
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Responsibility

**Rigor** 

**Transparency** 

#### Can You Predict The Data Structure?

We already know what the data will look like...

Accountability
Accurate
Care
Collegiality
Cooperation
Ethics
Fair

Honesty

**Objectivity** 

**Openness** 

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**Reliability** 

Reproducibility

Respect

Responsibility

**Rigor** 

**Transparency** 

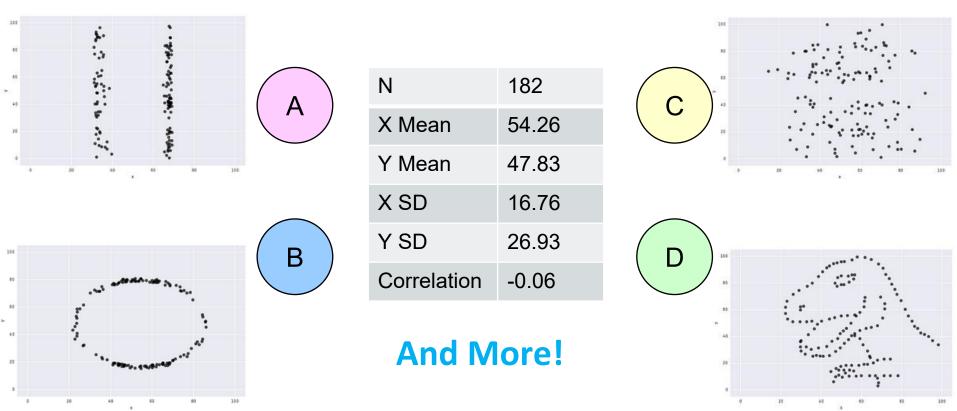
	$\bigwedge$	N	182	
ine A	A	X Mean	54.26	
		Y Mean	47.83	
_	_	X SD	16.76	
Circle	В	Y SD	26.93	
Circle B		Correlation	-0.06	







# The Datasaurus Dozen



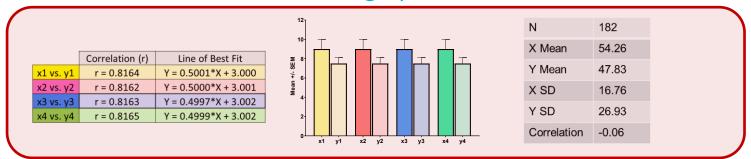
https://www.research.autodesk.com/publications/same-stats-different-graphs/

#### Accountability Accurate Care Collegiality Cooperation **Ethics** Fair Honesty **Objectivity Openness** Quality Reliability Reproducibility Respect Responsibility Rigor

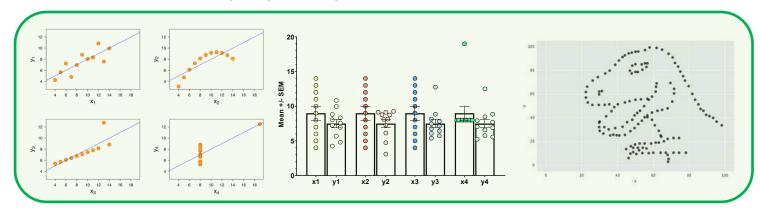
**Transparency** 

# **Data Exploration: Seeing is Believing!**

Beware of statistical or graphical summaries...



#### ...without proper exploration & visualisation!



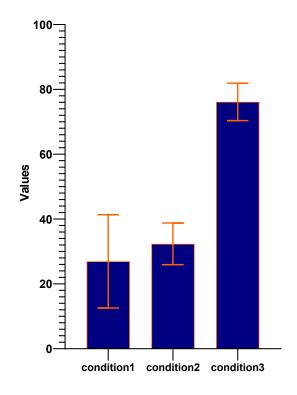


# Data Exploration Exercise





## What do you think of this graph?

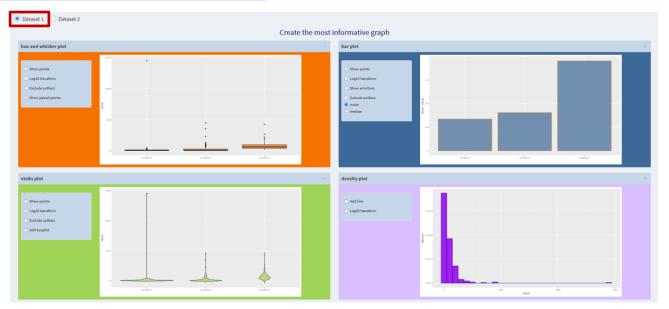




Values from 3 conditions from one experiment

# Representing Dataset 1: Create the most informative graph

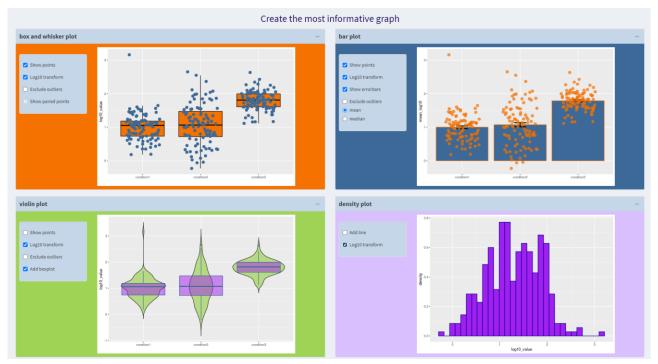
https://tinyurl.com/RIdataExp Make sure you are looking at Dataset 1



Plot for initial exploration of the data? Plot for presentation/ publication?



# Representing Dataset 1: Create the most informative graph

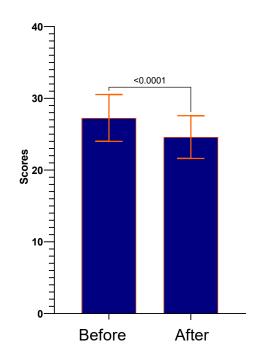


Which plots did we choose and why?

Any differences between initial exploration and presentation?



# What do you think of this graph?

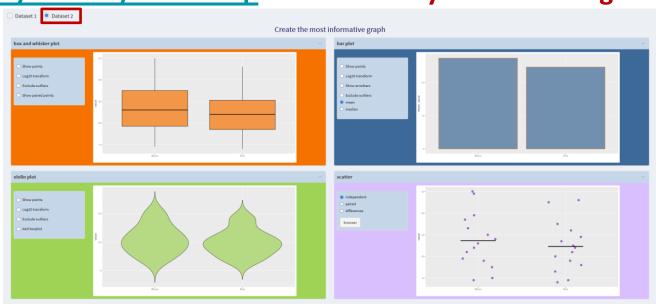




Values for before-after treatment from 4 experiments.

# Representing Dataset 2: Create the most informative graph

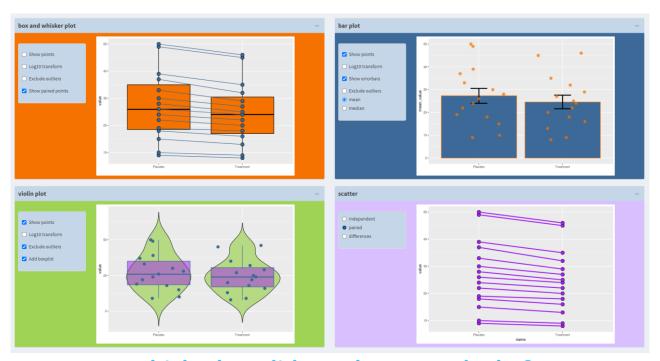
https://tinyurl.com/RIDataExp Make sure you are looking at Dataset 2







# Representing Dataset 2: Create the most informative graph



Which plots did we choose and why? What do you think about the stats now?

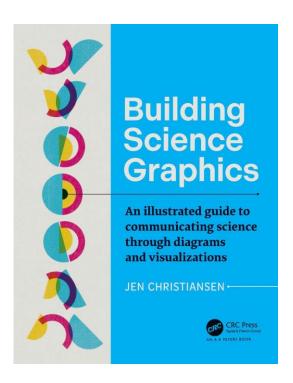


#### An Aside to Help with Better Figure Design...



Scientific Figure Design



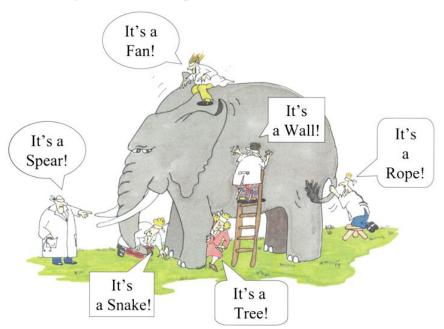




# Data exploration What can go wrong if we don't do it?!



# Data exploration Less Exploring more Assuming!



Risk missing the actual story the data is telling...

# **Data Exploration...**



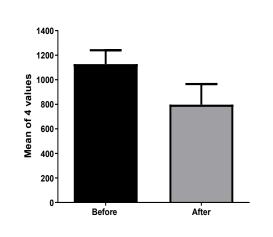
...Suspicious Summaries

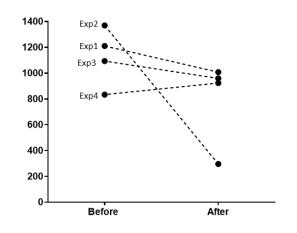
# Question Results Experimental Design Data Analysis Sample Size Data Exploration Experiment

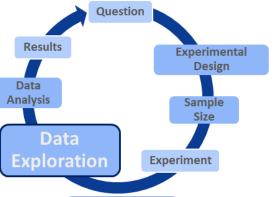
## **Example 1**

- Four experiments: Before-After treatment effect on a variable of interest.
- <u>Hypothesis</u>: Treatment will decrease the levels of the variable of interest



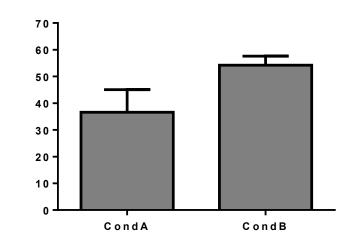


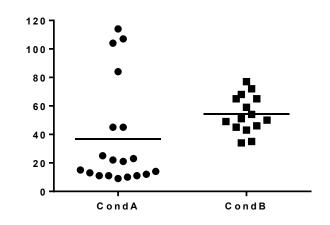


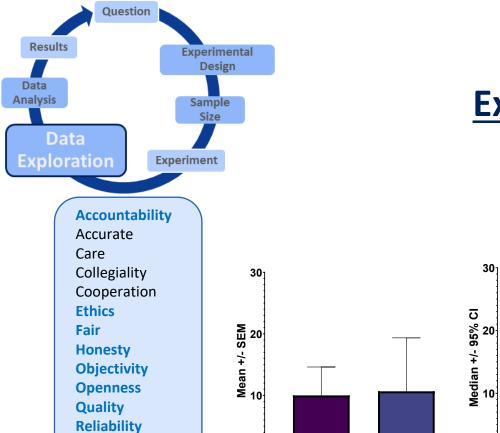


#### **Example 2**









Condition 1

**Condition 2** 

Reproducibility

Responsibility

**Transparency** 

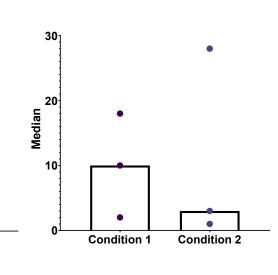
Respect

**Rigor** 

#### **Example 3**

Condition 1

**Condition 2** 



### **Data Exploration...**



#### ...Dubious Datasets

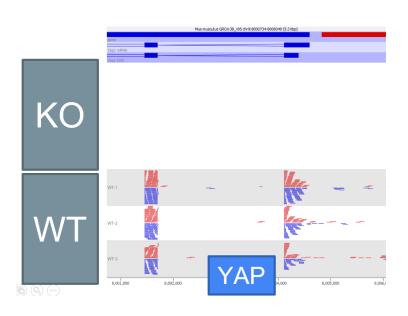
# Question Results Experimental Design Data Analysis Sample Size Data Exploration Experiment

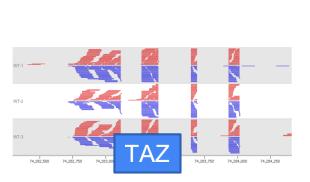
#### **Example 1: A Knockout?**

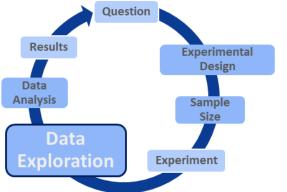
#### nature neuroscience

YAP and TAZ control peripheral myelination and the expression of laminin receptors in Schwann cells









#### **Example 2: A Case of Mistaken Identity**

OPEN & ACCESS Freely available online

PLOS GENETICS

In Vivo Effects of Histone H3 Depletion on Nucleosome Occupancy and Position in Saccharomyces cerevisiae

#### **Accountability**

Accurate

Care

Collegiality

Cooperation

**Ethics** 

Fair

Honesty

**Objectivity** 

**Openness** 

Quality

Reliability

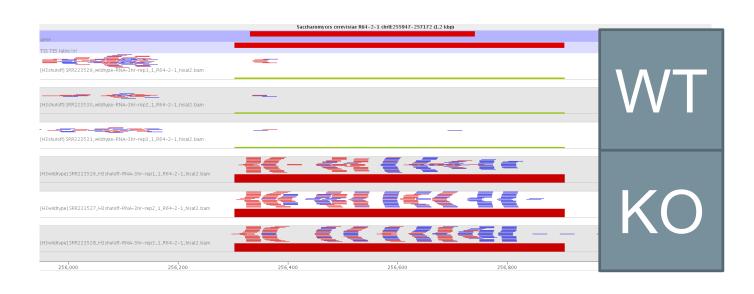
Reproducibility

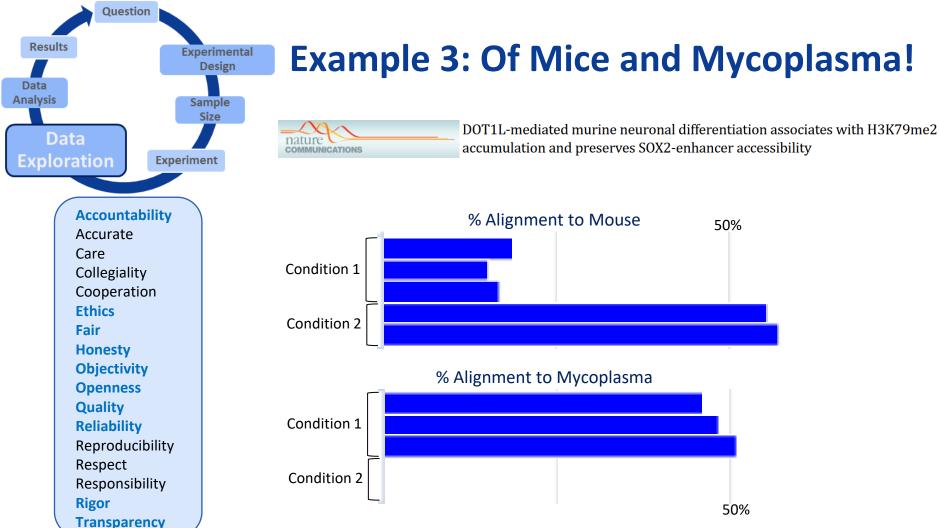
Respect

Responsibility

Rigor

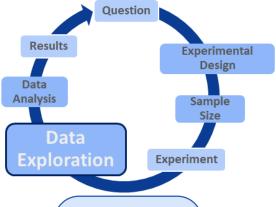
**Transparency** 





50%

50%



Accurate

Care

Collegiality Cooperation

Ethics

Fair

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Reliability

Reproducibility

Respect

Responsibility

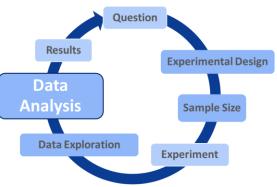
Rigor

**Transparency** 

### All highlight the perils of assuming and not exploring....



...Is it okay if you don't know there's a problem?



Accurate

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Respect

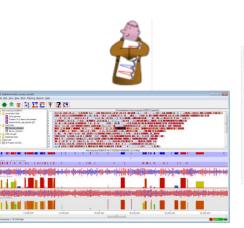
Responsibility

Rigor

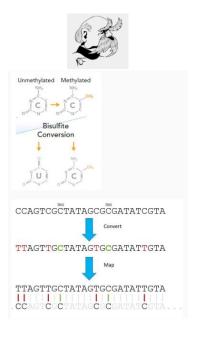
Transparency

#### **Analytical Tools**

The right tools for the job!







- Know what's out there
- Learn how to use it



Accurate

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Collegiality

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Reproducibility

Respect

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Rigor

Transparency

#### **Help With Analytical Tools**



### Core Skills Courses

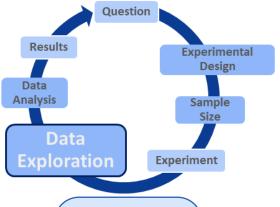
e.g. R Programming



## Application Specific Courses

e.g. NGS Analysis

https://www.bioinformatics.babraham.ac.uk/training.html



Accurate

Care

Collegiality Cooperation

Ethics

Fair

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Objectivity

**Openness** 

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**Rigor** 

**Transparency** 

#### In a nutshell

#### **Data exploration** is CRITICAL to:

- Understand our data
- Be confident in our findings

Ensure our results are quality and reliable

... Good for everyone!

Accurate Care

Collegiality

Cooperation

**Ethics** 

Fair

#### **Honesty**

Objectivity

Openness

Quality

Reliability

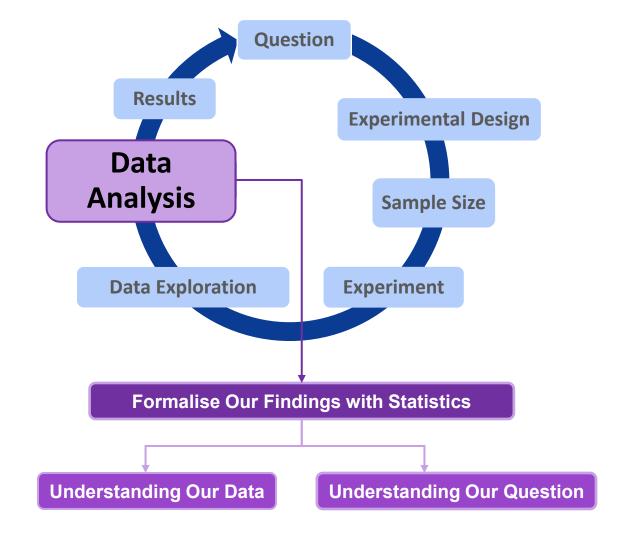
Reproducibility

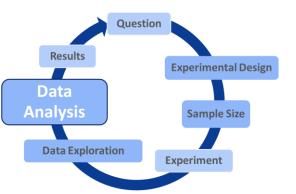
Respect

Responsibility

**Rigor** 

Transparency





#### **Statistical Analysis**

Translate the hypothesis/question into statistical questions

By choosing the right test!

Accountability

Accurate

Care

Collegiality

Cooperation

**Ethics** 

Fair

Honesty

Objectivity

Openness

Quality

Reliability

Reproducibility

Respect

Responsibility

Rigor

Transparency

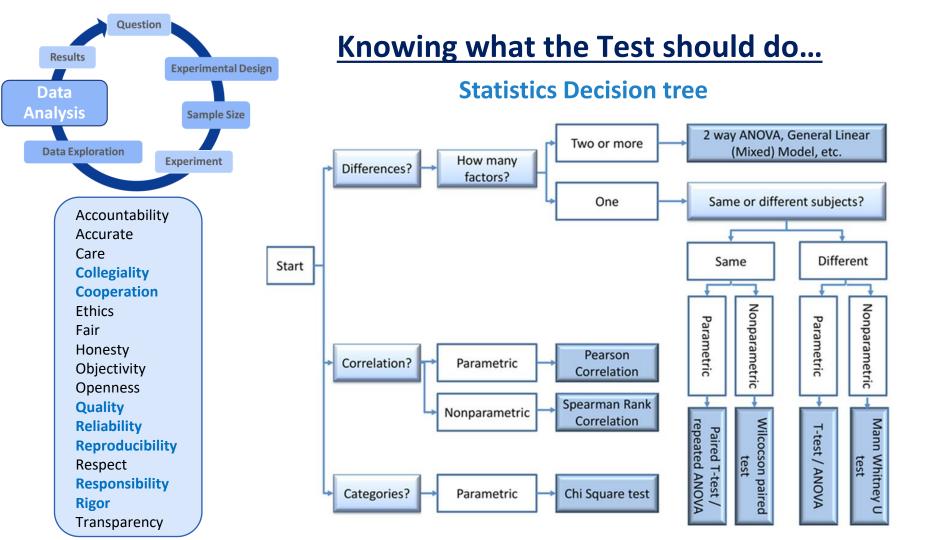
#### **Healthy approach:**

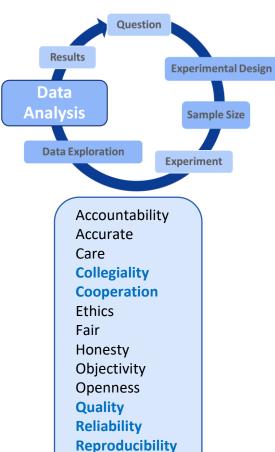
"It's not about knowing the name of the test...

...It's knowing what the test should do"



**Anne Segonds-Pichon** 





Respect

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Responsibility

Transparency

#### **Knowing what the Test should do...**



### Statistics Using R



Babraham Significant Significa

Using GraphPad Prism

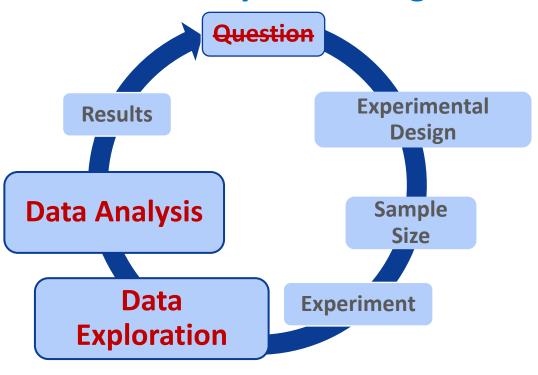
https://www.bioinformatics.babraham.ac.uk/training.html

Accountability Accurate Care Collegiality Cooperation **Fthics** Fair Honesty Objectivity **Openness** Quality Reliability Reproducibility Respect Responsibility Rigor

Transparency

### **Research Integrity**

More than 1 way to Investigate!



**Exploratory data analysis** 

Accurate

Care

Collegiality

Cooperation

**Ethics** 

Fair

**Honesty** 

**Objectivity** 

**Openness** 

Quality

Reliability

Reproducibility

Respect

Responsibility

Rigor

**Transparency** 

#### In a nutshell



Data exploration should always be a pivotal step of analysis

Stats helps us formalise our findings











## Research Integrity What does Ethics mean?



Accurate

#### Care

Collegiality Cooperation

#### **Ethics**

Fair Honesty Objectivity Openness

Quality Reliability

Reproducibility

Respect

Responsibility

Rigor

Transparency

### Research Integrity Ethics and Animal Welfare

- If we need **biological replicates** to be confident in our results, why not have **as many as we can?**
- And what does it have to do with integrity?
- Time to talk about the Home Office, ASPA and the 3 Rs



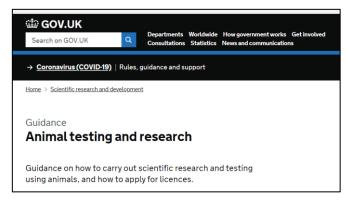




### Research Integrity Home Office

**The Home Office** (HO) is a ministerial department of the Government of the UK, responsible for immigration, security and law and order.

- But more importantly: animal welfare





By Steph Gray, CC BY-SA 2.0, https://commons.wikimedia.org/w/index.php?curid=31387711

Accurate

#### Care

Collegiality

Cooperation

#### **Ethics**

Fair

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**Openness** 

Quality

Reliability

Reproducibility

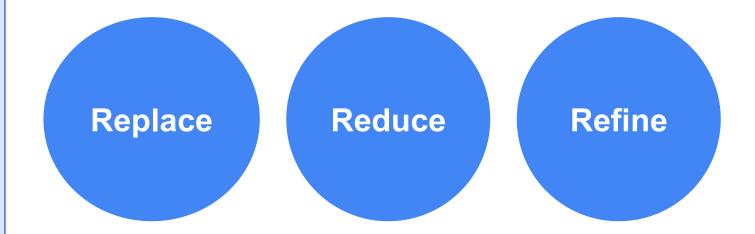
Respect

Responsibility

Rigor

Transparency

### Research Integrity The 3 Rs



https://wellcome.org/sites/default/files/wtp057673 0.pdf

### Research Integrity

The 3Rs: about animal welfare

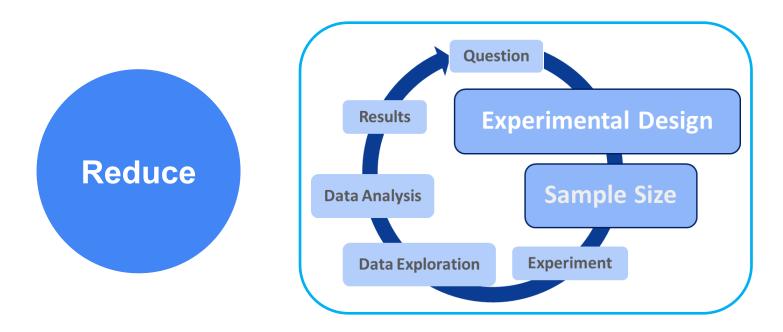




Avoiding or replacing the use of animals in areas where they otherwise would have been used.

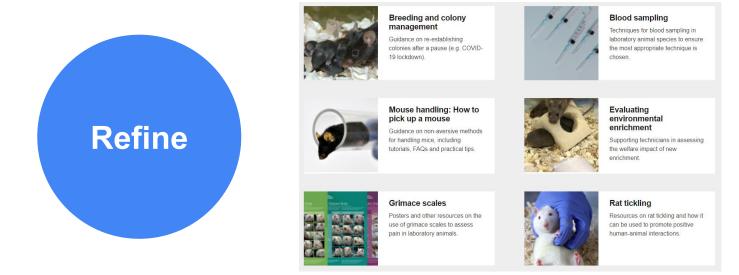
### **Research Integrity**

The 3Rs: about animal welfare



Minimising the number of animals used consistent with scientific aims.

## Research Integrity The 3Rs: about animal welfare



Minimising the pain, suffering, distress or lasting harm that research animals might experience.

### Research Integrity The 3 Rs at Babraham: AWERB

The Hub People Committees V Institute V Science Facilities V



#### Animal Welfare & Ethical Review Body

#### Remit

To provide the campus with independent ethical advice on the balance of harms to benefits within scientific projects using animals. To monitor standards of animal care and welfare, to support and advise named persons and licensees working under the Animals (Scientific Procedures) Act and to advise the Establishment Licence Holder on the suitability of Project Licence applications. To develop initiatives and guidelines leading to the widest possible application of the 3Rs (refinement, reduction and replacement) both on the campus and amongst the wider scientific community. In accordance with our commitments to the Concordat on Openness in animal research, any staff member with a concern that falls within the AWERB remit is encouraged to speak to the AWERB chair or any other committee member.



### **Good Research in Practice**



Accurate

Care

Collegiality

Cooperation

**Fthics** 

**Fair** 

**Honesty** 

Objectivity

**Openness** 

Quality

Reliability

Reproducibility

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Responsibility

Rigor

**Transparency** 

#### **Good Research in Practice**

#### The Research Process

- Responsibilities
- Competence
- Project planning
- Quality Control

#### **Laboratory Practice**

- Health and safety
- Handling of samples and materials
- Facilities and equipment
- Documentation of procedures and methods
- Research/work records



Get a GRiP!

'Good Research in Practice' (GRIP) supports and measures the more practical aspects of Research Integrity standards compliance on the ground in the lab.





## Keeping track of the research How?





Accuracy

Care

**Collegiality** 

Cooperation

**Ethics** 

Fair

**Honesty** 

Objectivity

**Openness** 

Quality

Reliability

Reproducibility

Respect

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Rigor

Transparency

## Research Integrity Laboratory Notebooks



### Accountability Accuracy

Care

Collegiality

Cooperation

**Ethics** 

Fair

**Honesty** 

Objectivity

**Openness** 

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Reproducibility

Respect

Responsibility

Rigor

**Transparency** 

## Research Integrity Laboratory Notebooks





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Accuracy

Care

**Collegiality** 

Cooperation

**Ethics** 

Fair

**Honesty** 

Objectivity

**Openness** 

Quality

Reliability

Reproducibility

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**Transparency** 

## Research Integrity Laboratory Notebooks



Accuracy

Care

**Collegiality** 

Cooperation

**Ethics** 

Fair

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**Transparency** 

## Research Data Data Storage



Legal Requirements

**Practicalities** 

Accuracy

Care

**Collegiality** 

Cooperation

**Ethics** 

Fair

**Honesty** 

Objectivity

**Openness** 

Quality

Reliability

Reproducibility

Respect

Responsibility

Rigor

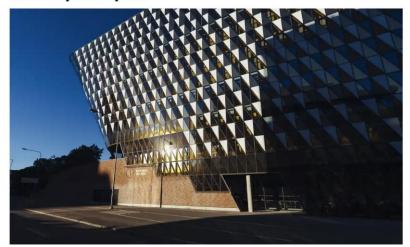
**Transparency** 

## Research Data Data Storage

#### Sweden

Decades of research destroyed after freezer fails at Swedish university

Estimated value of the samples thought to be in the millions as incident reported to police



An internal investigation has been launched at the Karolinska Institutet despite no indication of sabotage. Photograph: Bloomberg/Getty Images



- Interruption in the supply of liquid nitrogen leading to the destruction of samples from multiple institutions.
- Valued in the millions.
- "Those worst affected are those researching leukaemia, they have gathered samples from patients over as much as 30 years,"

## What should we be doing? Expectations and Responsibilities







Created by Anggara Putra from Noun Project



Created by shashank singh from Noun Project



Created by Good Wife from Noun Project

Accuracy

Care

**Collegiality** 

Cooperation

**Ethics** 

Fair

**Honesty** 

Objectivity

**Openness** 

Quality

Reliability

Reproducibility

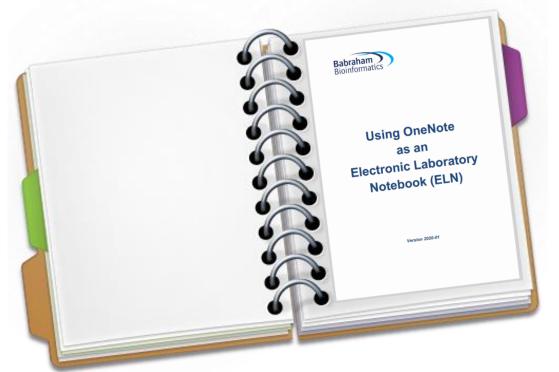
Respect

Responsibility

Rigor

**Transparency** 

### Research Data OneNote





### Responsibility



Accountability Accurate Care Collegiality Cooperation **Ethics** Fair Honesty Objectivity **Openness** Quality Reliability

Reproducibility

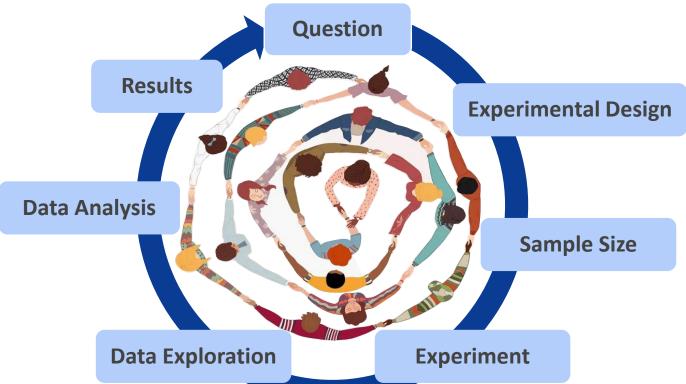
Responsibility

Transparency

Respect

Rigor

# **Research Integrity Wider Responsibility** Question Results



Accuracy

Care

**Collegiality** 

Cooperation

**Ethics** 

Fair

**Honesty** 

Objectivity

**Openness** 

Quality

Reliability

Reproducibility

Respect

Responsibility

Rigor

**Transparency** 

## **Research Integrity**



Accuracy

Care

**Collegiality** 

**Cooperation** 

**Ethics** 

Fair

**Honesty** 

Objectivity

**Openness** 

Quality

Reliability

Reproducibility

Respect

Responsibility

Rigor

**Transparency** 

## **Research Integrity**

**Questioning** 



Accuracy

Care

**Collegiality** 

Cooperation

**Ethics** 

Fair

Honesty

Objectivity

**Openness** 

Quality

Reliability

Reproducibility

Respect

Responsibility

**Rigor** 

**Transparency** 

# **Research Integrity**

**Collaboration and Competition** 



Accuracy

Care

**Collegiality** 

Cooperation

**Ethics** 

Fair

**Honesty** 

Objectivity

**Openness** 

Quality

Reliability

Reproducibility

Respect

Responsibility

Rigor

**Transparency** 

## **Research Integrity**

**The Game** 



### **Accountability** Accuracy Care **Collegiality** Cooperation **Fthics** Fair Honesty Objectivity **Openness** Quality Reliability Reproducibility Respect Responsibility Rigor **Transparency**

# Research Integrity Wider Responsibility and Scientific Community





### **Research Integrity**

#### Review

#### This means both

- Having your work reviewed by others peer review journals, papers,
- conferences esc
- Reviewing other people's work.

Reviewing also neems more locally. Within your group, department, institute.

Officiam dissalfile positive, not combative.

Your group have a vected interest in you and your crience being nucreodal and getting the many right. Listen to them.

Actively usek input and questioning Don't take it personally Be positive in your criticism of cithens' science

It makes you and your science better. Other people knowlets of things. It works both ways. You do it for them, they do it for you.

As pour science progresses, you poss through these barriers, pick up issues, find alternative viewpoints.

You don't vision to be taken done in a public harm (see hower of an example above a ) who closed to take the south to an interval and underworked and underworked and their findings were agriced the common understanding of excipace distance. They were purchased and disproved uponly and enthormodicity. We district note this to bupper to you it. List meeting per time a to take in the purchased and office the district of the distr

Lab meetings et are a training ground, An apportunity to conditing ratio mistaker, have the ctory, get things right before you go to conference/review etc.

It will save you from future emboraconvent, time and effort to go through this.

#### fillhot does reviewing the odersce mean to you?

All Further tolising on the peer review processic available.



R to B conjugately seed a



#### FAIR

#### Friedblin

#### Accessible

#### Interspecible

(which could never replicate the spirit of the analysis, rather than just jump.

through exactly the came loops)







Papraducib

financing your advance is SABR communication the value of your salence and your data.

Your data has sake above and beyond what you created it for, especially with big datasets:

Make your data as useful as possible (you're in work for you and others). Others can pull things out of your data. This is good for stations, and with involvy adds solve to your research. Also good for the planet -not repeating what lact already level date.

This is about chite-disclosure—so we full little do this, to wake data accessible.

BUT there is a real celling paint here to make it as accessible as possible—it

At It we re-process more jacquescing data than we green the new data. And we green the a Life.

It's also important that YMU make use of public data as much as possible.

Also, notice sure —if your work is bosed on an existing paperyldata, that you do a RNDRIR sheek on the actual data.

This is also a learning opportunity (practice viscolining and analysing the type of data you will likely to collecting) and also a good described as an exercise after all time and showly and also a few and an exercise and a service are also as a service and a servi

How does this link to research integrity?



E in E origination with



#### Pre-prints

The positives and regatives to pre-prints are a bit more number.

The problem is that published data is often out of date. By the time a paper cores out (6 months +, mostly more), takes time to get published, findings may have moved on.

A sway to circumsent the delay in getting research out BUT it's not peer reviewed.

It looks like a paper, but it's not:

Not peer reviewed

Not fact checked

A High proportion on bloarchise never reakes it into a paper.

If we consider that about half of all that is published (and peer reviewed) is a rong, this pushes that likelihood up even morewith bioarchive.

However, it is worth thinking about if your data is time critical. You may also need to check with your preferred journal – will they penalise you/not accept if you've published on bloomchive first?

#### What are the pros and coss?





How does this link to research integrity?



E in England 1984 annual State of the Company State (1984) from



#### Open Access

#### This is a bit of a game.

Open access can have benefits to you, and others.

There are open access journals, or some which you can pay a

fee to make open access.

This reeans your paper is more accessible.

Also accessible to automatic test mining, so even more can be gleaned from your data (see FAIR).

Something like PubMed central will also make papers open access – with about a 6 month delay – on loads of journals, not just open access ones.

Wears you can pull out information, including API etc (autoprogram).

Remember: Just because something is in a paper, doesn't mean it's actually true!

#### What are the pros and cons?



How does this link to research integrity?



The Resignation of the Committee of the



Accuracy

Care

**Collegiality** 

Cooperation

**Ethics** 

Fair

Honesty

Objectivity

**Openness** 

Quality

Reliability

Reproducibility

Respect

Responsibility

Rigor

**Transparency** 

## **Research Integrity**

"Academia is at its best a meritocracy and at its worst a pit of vipers. A grown-up version of the children's game snakes and ladders. The quest for knowledge and the pushing back of frontiers has created a hierarchal system with super demanding and, at times, conflicting metrics (postdocs must publish, publish, publish but to apply for a lectureship position you need to have a teaching track record)."

Emma Williams

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# Research Integrity In a nutshell

Applying research integrity principles is our

responsibility as scientists



Accountability
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# Research Integrity In the lab



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# Research Integrity Work culture



Accurate

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# Research Integrity PhD Students



It doesn't matter if it 'doesn't work'

Accurate

#### Care

Collegiality

Cooperation

#### **Ethics**

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## **Research Integrity**



...but what if it is too hard?

Accurate

#### Care

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Reproducibility

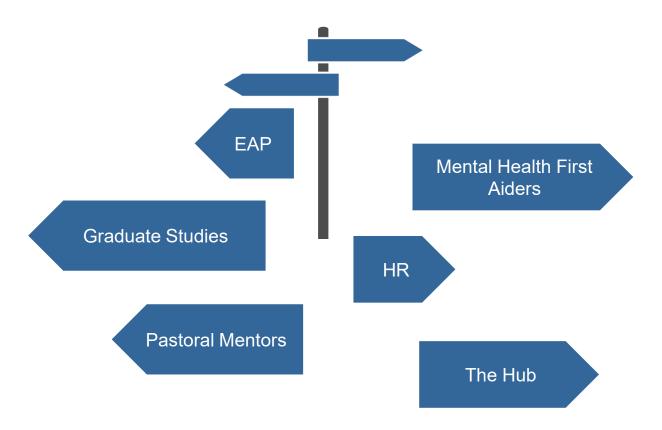
Respect

Responsibility

Rigor

Transparency

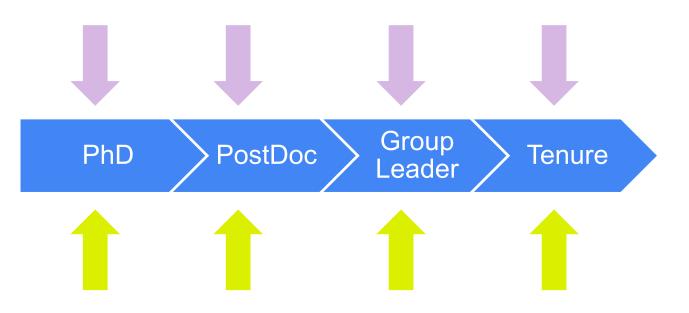
## **Research Integrity**



Accountability Accurate Care Collegiality Cooperation **Ethics** Fair Honesty Objectivity **Openness** Quality Reliability Reproducibility Respect Responsibility Rigor

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## **Research Integrity**



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## **Research Integrity**

Which roles are most important or responsible for Research Integrity?



Director







Technician







Group Leader

Facilities





Visiting Scientist

**Tenure** 



### **Generative Al**



### Have You Used Generative AI?

**Accountability** Accurate

Care

**Collegiality** 

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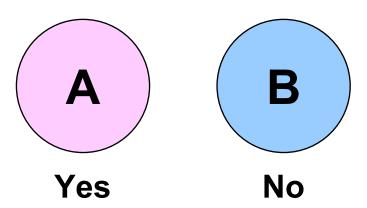
Transparency











### Accountability Accurate

Care

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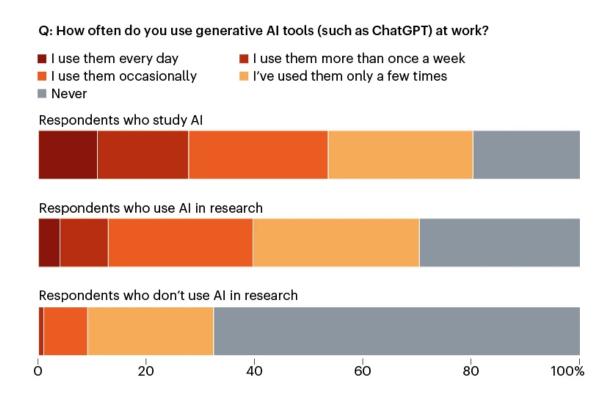
Respect

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### **Have You Used Generative AI?**



Al and science: what 1,600 researchers think

https://www.nature.com/articles/d41586-023-02980-0#correction-0

## Why Use Generative AI?

Accountability
Accurate

Care

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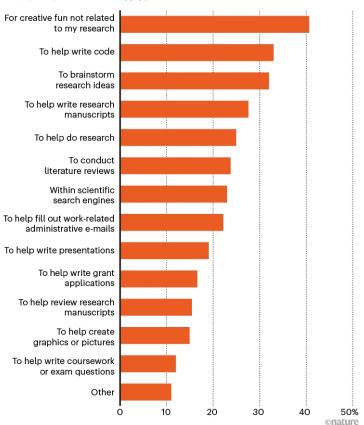
Respect

Responsibility

Rigor

**Transparency** 

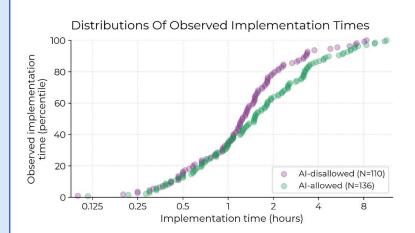
Q: What do you use generative AI tools (such as ChatGPT and other large language models) for? (Choose all that apply.)



### **Considerations For Generative Al**

### Will it Actually Help?

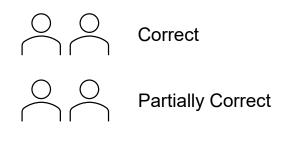
### **Software Developers**

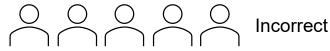


Measuring the Impact of Early-2025 AI on Experienced Open-Source Developer Productivity, Becker *et al* 2025

#### **Scientists**

From 9 Self-Reported "Correct" Answers





Attendees of the Joint International Bioinformatics Hackathon 2025.

# Accountability Accurate Care

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### **Considerations For Generative Al**

### **Accountability**

Accurate

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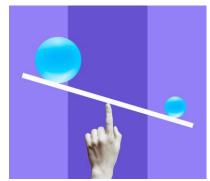
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### Will it Harm?

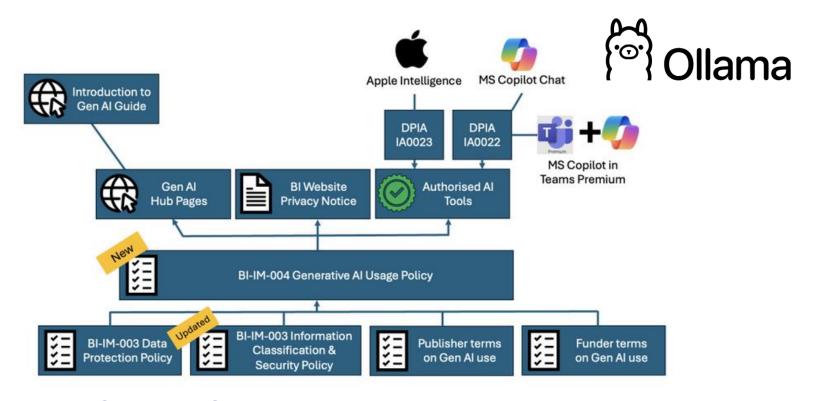






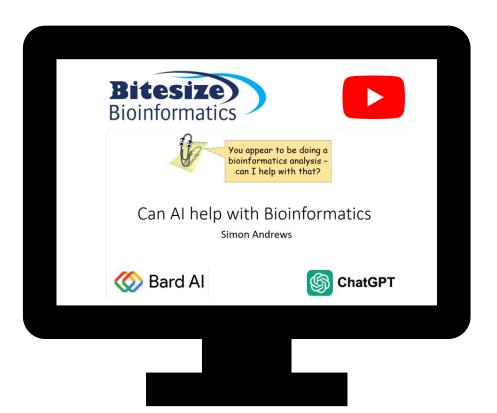
Ultimately it is our responsibility

### **Generative AI at Babraham**



See The Generative AI Pages on The Hub

### **Generative AI and Bioinformatics**







# How our integrity may be tested





### **Publications**





### **Publish or Perish?**



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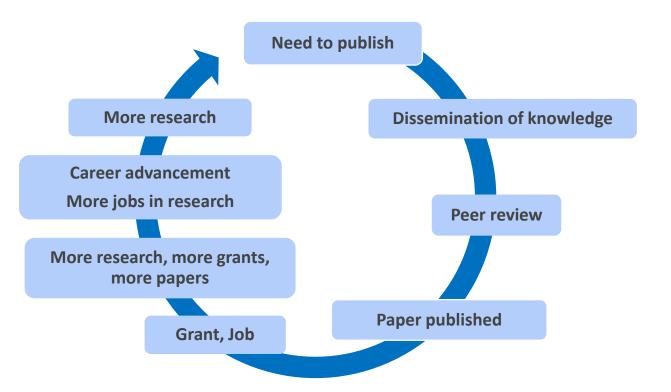
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## **Publications: the good**



Accurate

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Reproducibility

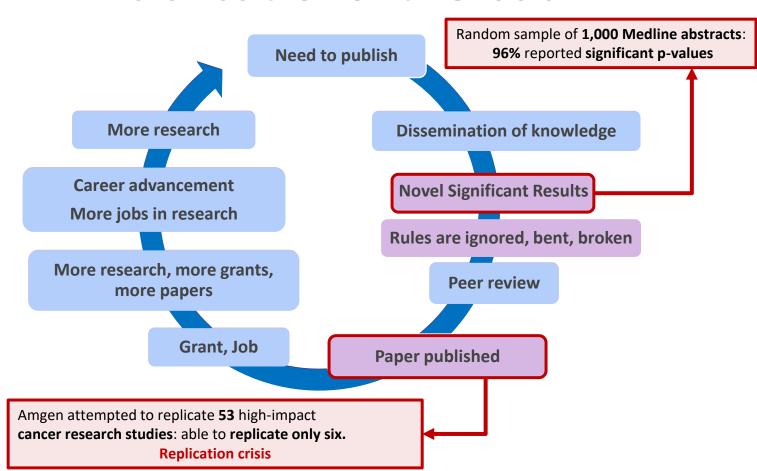
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### **Publications: the bad**



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### Publications: the bad

### The most cited paper to be retracted

### Pluripotency of mesenchymal stem cells derived from adult marrow

Nature. 2002 Jul 4;418(6893):41-9. doi: 10.1038/nature00870. Epub 2002 Jun 20.

June 2024

**Retraction** of the paper



December 2019

New concerns raised by **Elisabeth Bik** 

https://pubpeer.com/publications/DF95 522E3585E37663CAD1972E70BD#

June 2002

Paper published in Nature

22 years!

June 2007 **Erratum published** 

Concerns about reproducibility and duplicated images

**Primarily by New Scientist** 

**Investigation by University** 

**Retraction of related paper** 

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	Published	Accesses	Citations
Original Article	20 June 2002	27k	4492
Erratum Note	14 June 2007	6289	39
Retraction Note	17 June 2024	19k	1

(Numbers as of Oct. 2024)

Most authors agree with the retraction\*

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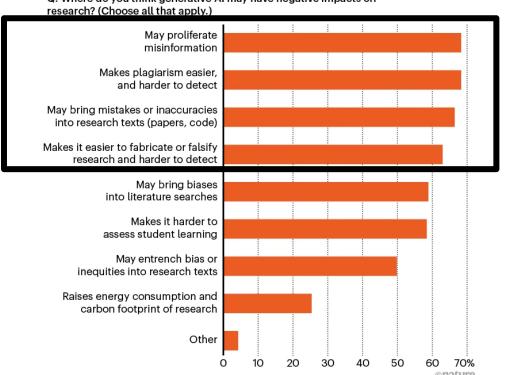
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# Publications: the ugly Generative Al Concerns

Q: Where do you think generative AI may have negative impacts on



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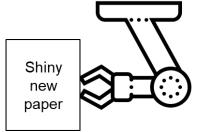
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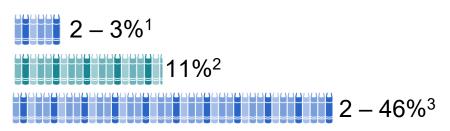
**Transparency** 

# Publications: the ugly Paper Mills





### **Estimated Prevalence**



- 1. https://www.nature.com/articles/d41586-023-03464-x
- 2. https://www.medrxiv.org/content/10.1101/2023.05.06.23289563v2
- 3. https://publicationethics.org/node/55256



In 2023 Hindawi closed 4 journals because they were "heavily compromised by paper mills"

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### **Publications: the balance**





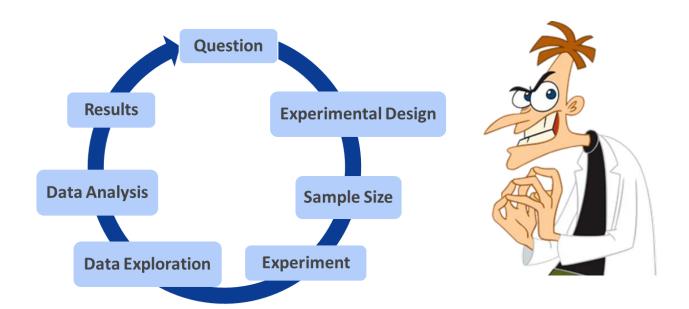
### Misconduct



### **Accountability** Accurate Care **Collegiality** Cooperation **Ethics** Fair **Honesty Objectivity Openness** Quality Reliability Reproducibility Respect Responsibility Rigor

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# Research Integrity Misconduct

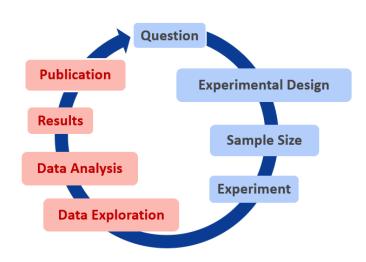


### **Misconduct: How?**



There are many ways to misbehave when it comes to research

Plagiarism
Fabrication and falsification
Inappropriate image manipulation
Non-publication of data
Faulty data-gathering procedures
Poor data storage and retention
Misleading authorship
Sneaky publication practices



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### Misconduct: Why?

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Tina Gunsalus





## **Misconduct: Spot the TRAGEDIES**



**Group and authority pressure** 

**Entitlement** 

**Deception** 

**Ambition** 

Rationalisation

A Simple Request from their Professor

Verify the numbers in a figure matched the spreadsheet

**A Questionable Outcome** 

Paper published with the new student as a co-author

Temptation They accept the credit

Why?

Incrementalism

**Embarrassment** 

Stupid systems

A real case from Tina Gunsalus
https://www.nature.com/articles/d41586-018-05145-6

The New Student



### Misconduct: In a nutshell



"I never met anyone who said, yeah, you know, that was the day I woke up and decided to you know, put my career at risk, potentially go to jail, embarrass my family, lose my job."



Tina Gunsalus

Accountability

Cooperation **Fthics** Fair

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### **Honesty**

Objectivity Openness Quality Reliability Reproducibility Respect Responsibility Rigor

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We need to be aware of our feelings and circumstances And acknowledge how they might influence us

## The Only Acceptable Tragedy!\*

### STEPS Tragedy • The Dance - How To...



STEP 1 - Put your hands up beside your ears in 'Tragedy' shock!



STEP 2a - Raise your right arm.



STEP 2b - Raise your left arm so both arms are up.



STEP 3 - Clasp both hands together on your heart.



STEP 4 - Stretch both arms out straight in front with your palms up.



STEP 5a - Put your hands to your head as in STEP 1 and then step to your right.



STEP 5b - Repeat the same move to your left. Repeat the same two moves again.



STEP 6a - Step to the left and swing your left arm out with your right hand on your hip.



STEP 6b - Step to the left and swing your right arm out with your left hand on your hip.



STEP 7 - Hold your right arm out and stop the traffic!



STEP 8 - Keep your right arm out, turn to your right and roll your left shoulder three times.

https://www.youtube.com/watch?v=OiwDHHcHPh0

\*To be fully transparent we should note the Bee Gees did it first!



### The bottom line



## Research Integrity It's about being a good scientist



**Research integrity** 

is about owning every step of our research, and benefits everyone.





# What do you think now? Which words are most important/synonymous with integrity?





## Research Integrity More than words





### The Hub

## Research Integrity

### **Useful resources**





BI-HR-001 Code of Conduct
BI-HR-005 Disciplinary Policy
BI-COR-010 Whistleblowing Policy
BI-RES-001 Authorship Policy
BI-RES-004 Research Misconduct Policy
BI-RES-005 Research Integrity Policy
Record Retention Policy
Data Management Guidance

## Research Integrity Useful resources



#### **Training Courses**



www.bioinformatics.babraham.ac.uk



MS Teams channel and mailing list. Contact

Richard.Acton@babraham.ac.uk

### **Research Integrity**

**Useful resources** 

## nature









"Integrity is doing the right thing even when no one is watching."

— C.S. Lewis

"Integrity is doing the right thing when you don't have to-when no one else is looking or will ever know—when there will be no congratulations or recognition for having done so."

Excerpted from <u>Shattering the Glass Slipper: Destroying Fairy-tale Thinking Before It Destroys You.</u>
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