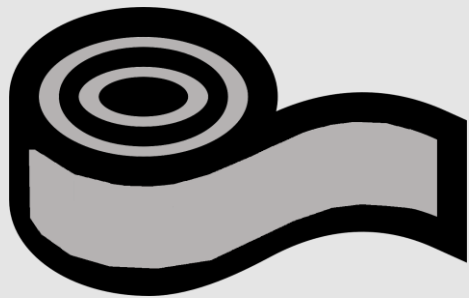


Statistical Rules to Tape to Your Forehead

BERDC Special Topics Talk 6



DaCCoTA
DAKOTA CANCER COLLABORATIVE
ON TRANSLATIONAL ACTIVITY

Dr. Mark Williamson
Biostatistics, Epidemiology,
and Research Design Core

Opening

Goal: present a clear list of rules for running a statistical analysis that are *useful* and *memorable*

- Sources: personal experience and the expertise of others
- Not exhaustive, inviolable, or strictly linear
- Example using shark data

Before Moving On:

Pre-test: https://und.qualtrics.com/jfe/form/SV_ebTk64vnD3yrrPo



Sources

Zuur & Ieno, 2016

A protocol for conducting and presenting results of regression-type analyses

Protocol for conducting and presenting results of regression-type analyses

1. State appropriate questions
2. Visualize the experimental design
3. Conduct data exploration
4. Identify the dependency structure in the data
5. Present the statistical model
6. Fit the model
7. Validate the model
8. Interpret and present the numerical output of the model
9. Create a visual representation of the model
10. Simulate from the model

Kass et al, 2016

Ten Simple Rules for Effective Statistical Practice

- Rule 1 Statistical Methods Should Enable Data to Answer Scientific Questions
- Rule 2 Signals Always Come with Noise
- Rule 3 Plan Ahead, Really Ahead
- Rule 4 Worry about Data Quality
- Rule 5 Statistical Analysis Is More Than a Set of Computations
- Rule 6 Keep it Simple
- Rule 7 Provide Assessments of Variability
- Rule 8 Check Your Assumptions
- Rule 9 When Possible, Replicate!
- Rule 10 Make Your Analysis Reproducible

Goodman et al, 2014

Ten Simple Rules for the Care and Feeding of Scientific Data

ASA, 2018

Ethical Guidelines for Statistical Practice

The Rules

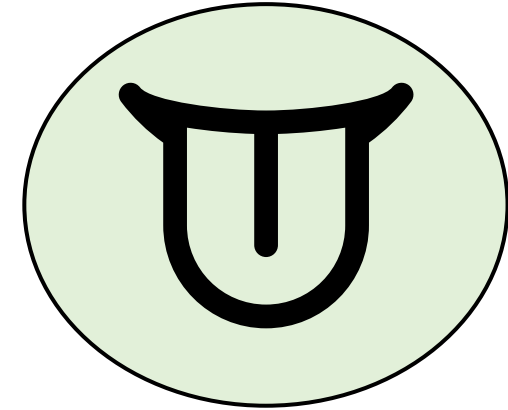
- 1) Annunciation 🏹
- 2) Exploration 🦶
- 3) Assumptions 🦴
- 4) Building 💪
- 5) Testing 🖐️
- 6) Justifying 🦶
- 7) Presentation 👁️
- 8) Interpretation 🧠



Rule 1: Annunciation

Clearly state your research questions.

- **Write one or two sentences**
- **Be painfully explicit**
- **Keep identity of variables in mind**



Abstract Examples:

Here, we use genome-wide CRISPR–Cas9 screening to establish that a T cell receptor (TCR) recognized and killed most human cancer types via the monomorphic MHC class I-related protein, MR1, while remaining inert to noncancerous cells (Crowther et al. 2020)

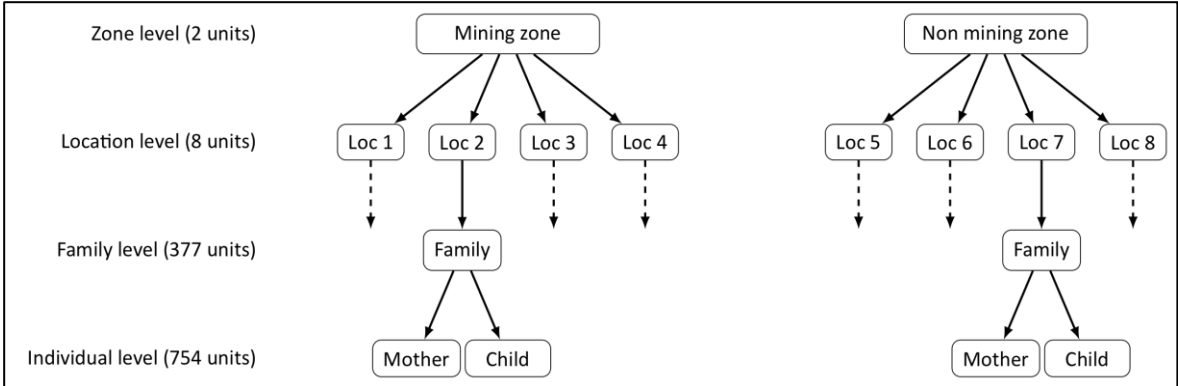
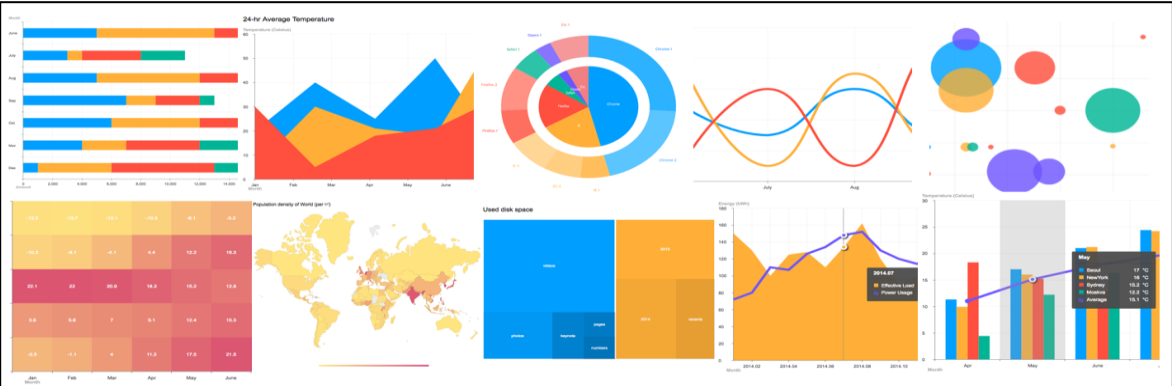
Using a combination of adrenalectomy, denervation, chemogenetics, cell ablation and knockout of the adrenergic receptor specifically in melanocyte stem cells, we find that the stress-induced loss of melanocyte stem cells is independent of immune attack or adrenal stress hormones (Zhang et al. 2020)

Here we sequenced whole genomes of 632 colonies derived from single bronchial epithelial cells across 16 subjects. Tobacco smoking was the major influence on mutational burden, typically adding from 1,000 to 10,000 mutations per cell; massively increasing the variance both within and between subjects; and generating several distinct mutational signatures of substitutions and of insertions and deletions (Yoshida et al. 2020)

Rule 2: Exploration

Get to know your data in and out.

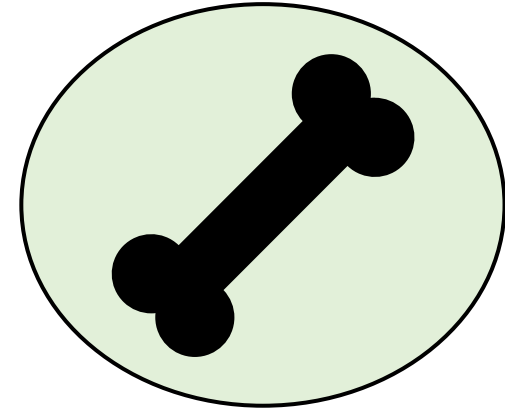
- Summary statistics are your friend
- Graphs, graphs, graphs!
- Determine design and dependencies



Rule 3: Assumptions

Ensure that your data meets test assumptions.

- **Check during design, before analysis, after analysis**
- **Be ready to try different types of tests**
- **Use many piece of information**



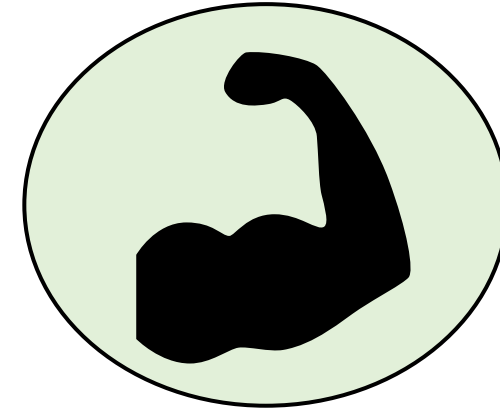
Rule 4: Building

Build one or more statistical models for your data.

- **Model type**
- **Model equation**
- **Model selection**

$$\begin{aligned}
 RD_{ijk} &\sim \text{Beta}(\pi_{ijk}) \\
 E(RD_{ijk}) &= \pi_{ijk} \\
 \text{var}(RD_{ijk}) &= \pi_{ijk} \times (1 - \pi_{ijk}) / (1 + \theta) \\
 \text{logit}(\pi_{ijk}) &= \text{Time}_{ijk} + \text{Relatedness}_{ijk} + \text{GroupSize}_{ijk} \\
 &\quad + \text{Time}_{ijk} \times \text{Relatedness}_{ijk} \\
 &\quad + \text{Relatedness}_{ijk} \times \text{GroupSize}_{ijk} + \text{Groomer}_l \\
 &\quad + \text{Hour}_j + \text{Receiver}_l \\
 \text{Groomer}_l &\sim N(0, \sigma_{\text{Groomer}}^2) \\
 \text{Hour}_j &\sim N(0, \sigma_{\text{Hour}}^2) \\
 \text{Receiver}_l &\sim N(0, \sigma_{\text{Receiver}}^2)
 \end{aligned}$$

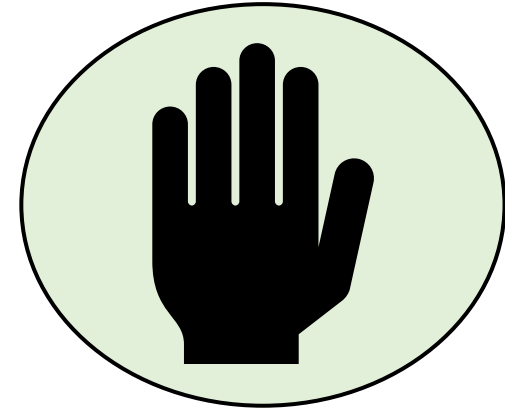
$$\begin{aligned}
 \text{NCalls}_{ij} &\sim \text{Poisson}(\mu_{ij}) \\
 E(\text{NCalls}_{ij}) &= \mu_{ij} \\
 \text{log}(\mu_{ij}) &= \text{SexParent}_{ij} + \text{FoodTreatment}_{ij} \\
 &\quad + \text{ArrivalTime}_{ij} + \text{SexParent}_{ij} \times \text{FoodTreatment}_{ij} \\
 &\quad + \text{SexParent}_{ij} \times \text{ArrivalTime}_{ij} + \text{Nest}_i \\
 \text{Nest}_i &\sim N(0, \sigma^2)
 \end{aligned}$$



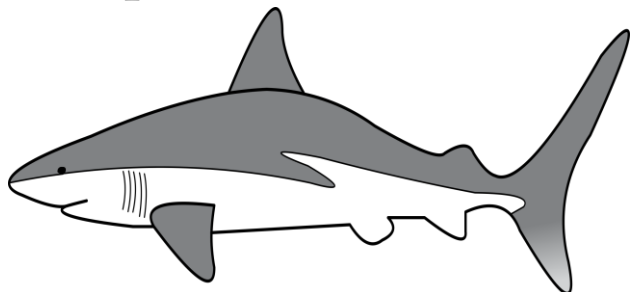
Rule 5: Testing

Test your statistical models on your data.

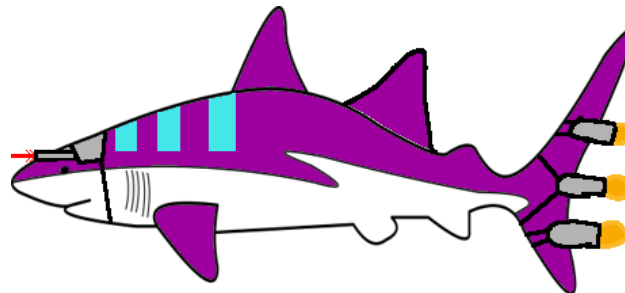
- **Start with the simplest model**
- **Different methods of tweaking**
- **Keep careful notes**



'Simple Shark Model'



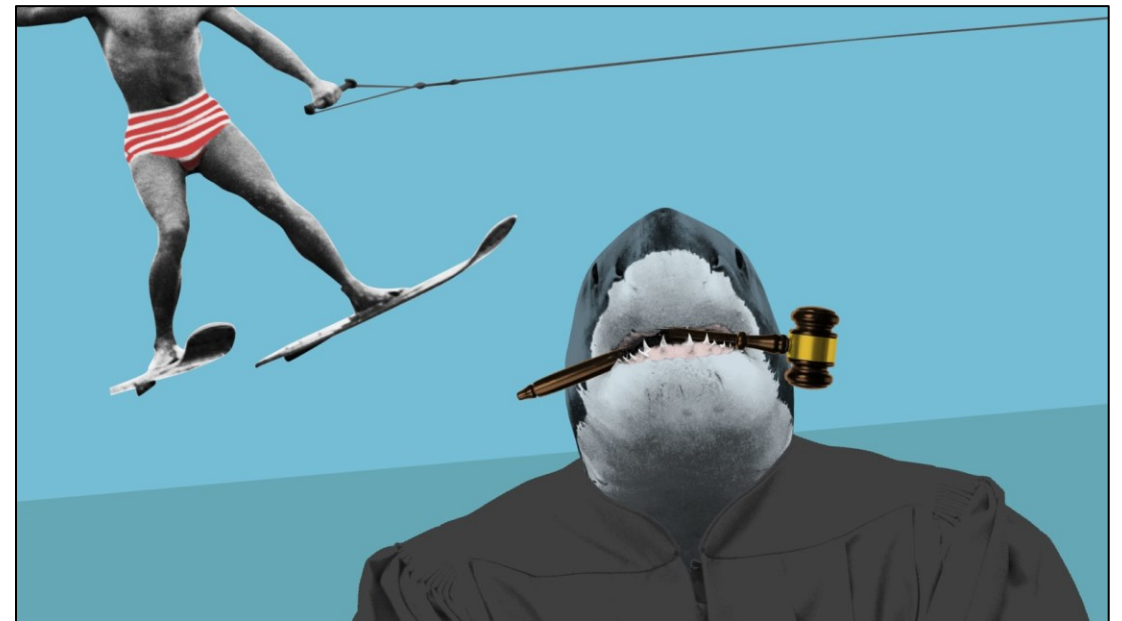
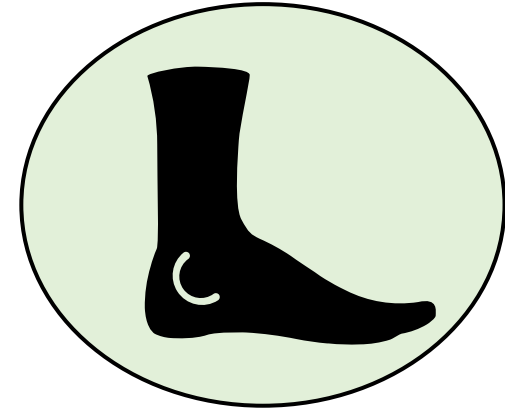
'Advanced Shark Model'



Rule 6: Justifying

Justify your model results.

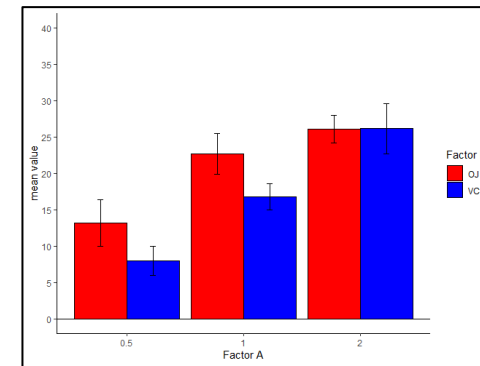
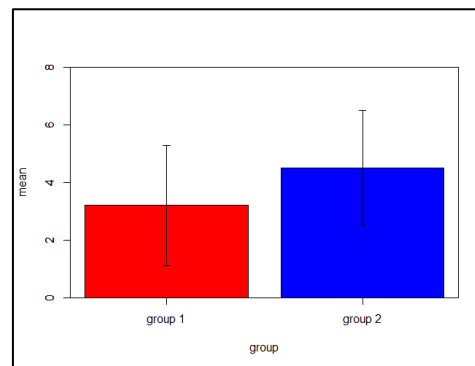
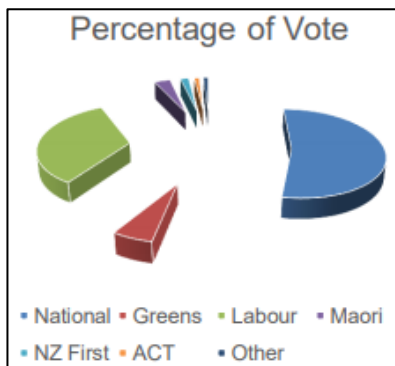
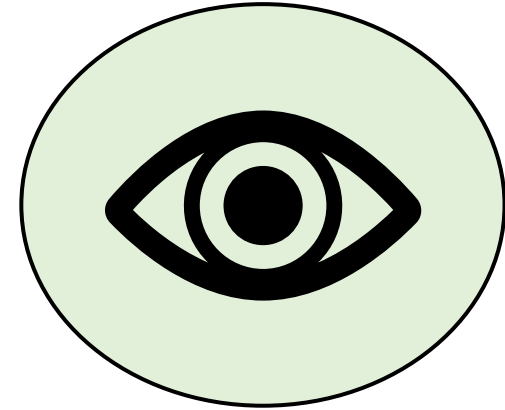
- **Fit statistics**
- **Link to summary stats and research background**
- **Have a 'why' for each piece of the statistical process**



Rule 7: Presentation

Present your results through text, tables, and figures.

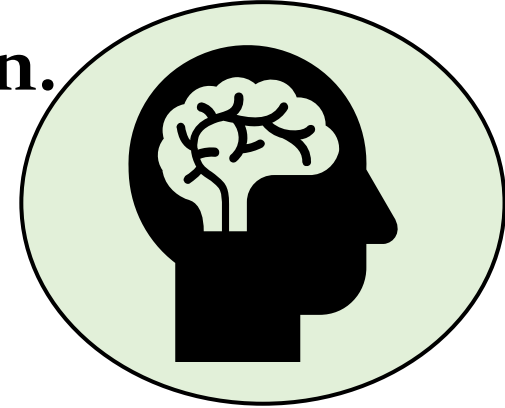
- **Written results should include estimates, test statistics, and significance values**
- **Tables and figures should add clarity, not banality**
- **Make uncertainty a key feature**






Rule 8: Interpretation

Clearly explain what your results are and what they mean.

- **Follow numerical results with clear wording**
- **Relate back to research question**
- **Project outside the study**



The Rules (again)

- 1) Annunciation 
- 2) Exploration 
- 3) Assumptions 
- 4) Building 
- 5) Testing 
- 6) Justifying 
- 7) Presentation 
- 8) Interpretation 

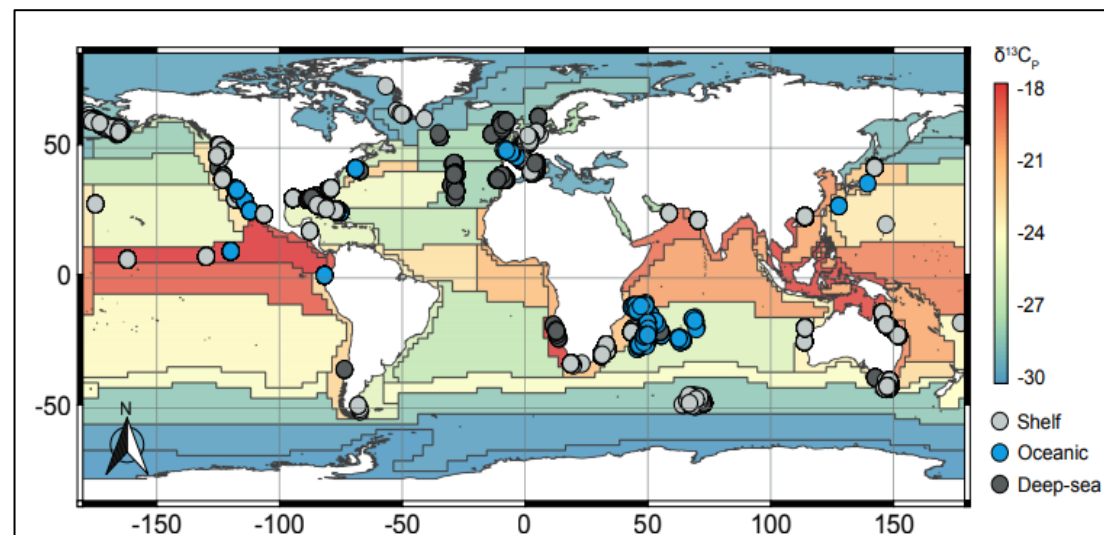
Shark Example

A Global Perspective on the Trophic Geography of Sharks

Abstract

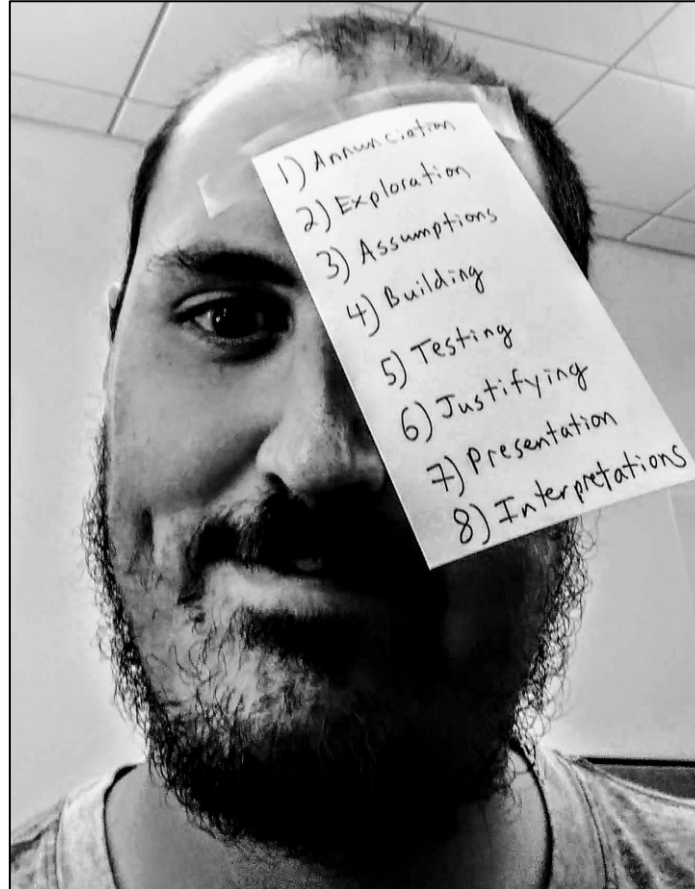
Sharks are a diverse group of mobile predators that forage across varied spatial scales and have the potential to influence food web dynamics. The ecological consequences of recent declines in shark biomass may extend across broader geographic ranges if shark taxa display common behavioural traits. By tracking the original site of photosynthetic fixation of carbon atoms that were ultimately assimilated into muscle tissues of 5,394 sharks from 114 species, we identify globally consistent biogeographic traits in trophic interactions between sharks found in different habitats. We show that populations of shelf-dwelling sharks derive a substantial proportion of their carbon from regional pelagic sources, but contain individuals that forage within additional isotopically diverse local food webs, such as those supported by terrestrial plant sources, benthic production and macrophytes. In contrast, oceanic sharks seem to use carbon derived from between 30° and 50° of latitude. Global-scale compilations of stable isotope data combined with biogeochemical modelling generate hypotheses regarding animal behaviours that can be tested with other methodological approaches.

- 1) **Annunciation**
- 2) **Exploration**
- 3) **Assumptions**
- 4) **Building**
- 5) **Testing**
- 6) **Justifying**
- 7) **Presentation**
- 8) **Interpretation**



Wrap-up

**May these rules
'stick' with you on
your statistical
journey**



**Please take the
post-test and
survey:**

Post-test: https://und.qualtrics.com/jfe/form/SV_4Tol5Rj6GcoGkv4

Survey: https://und.qualtrics.com/jfe/form/SV_5iHQvVd200BpoSa

References

Rule Reference Materials:

- <https://besjournals.onlinelibrary.wiley.com/doi/epdf/10.1111/2041-210X.12577>
- <https://journals.plos.org/ploscompbiol/article?id=10.1371%2Fjournal.pcbi.1004961>
- <https://www.ncbi.nlm.nih.gov/pmc/articles/PMC3998871/pdf/pcb.1003542.pdf>
- <https://www.amstat.org/asa/files/pdfs/EthicalGuidelines.pdf>

Example Abstracts:

- <https://www.nature.com/articles/s41590-019-0578-8>
- <https://www.nature.com/articles/s41586-020-1935-3>
- <https://www.nature.com/articles/s41586-020-1961-1>

Other:

- <https://www.theanalysisfactor.com/three-rules-statistical-analysis-unlearn/>
- https://www.jmp.com/en_be/statistics-knowledge-portal/what-is-regression/simple-linear-regression-assumptions.html
- https://www.iase-web.org/islp/apps/gov_stats_graphing/GoodBad/GoodBadGraphs.pdf
- <https://www.thedailybeast.com/democrats-have-already-jumped-the-shark-on-judge-kavanaugh>

Shark Data:

- https://nsuworks.nova.edu/cgi/viewcontent.cgi?article=1916&context=occ_facarticles
- <https://datadryad.org/stash/dataset/doi:10.5061/dryad.d1f0d>

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