

Introduction to Reproducible Research

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How Do You Know if a Data Analysis is Successful?

Parable

ARTICLES



Genomic signatures to guide the use of chemotherapeutics

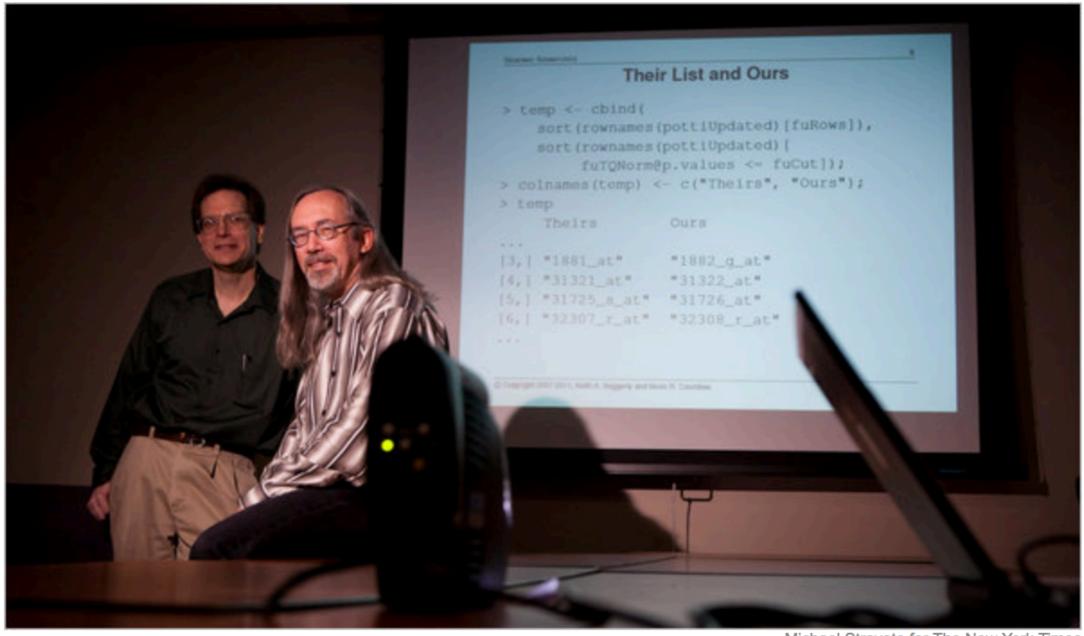
Anil Potti^{1,2}, Holly K Dressman^{1,3}, Andrea Bild^{1,3}, Richard F Riedel^{1,2}, Gina Chan⁴, Robyn Sayer⁴, Janiel Cragun⁴, Hope Cottrill⁴, Michael J Kelley², Rebecca Petersen⁵, David Harpole⁵, Jeffrey Marks⁵, Andrew Berchuck^{1,6}, Geoffrey S Ginsburg^{1,2}, Phillip Febbo^{1–3}, Johnathan Lancaster⁴ & Joseph R Nevins^{1–3}

Deception at Duke



"Rock Star" Statisticians

How Bright Promise in Cancer Testing Fell Apart

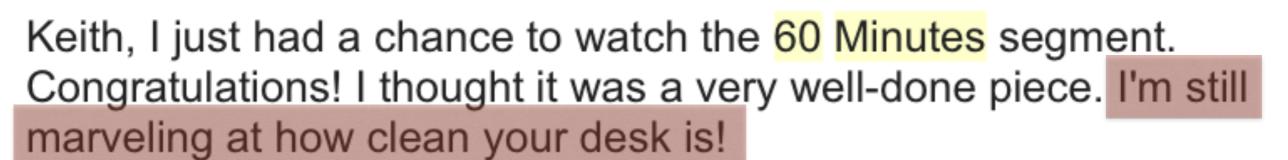


Michael Stravato for The New York Times

"Deception" at MDACC

Roger D. Peng <rpeng@jhsph.edu>

to Keith 🖃



Best, -roger



Follow-up Discussion

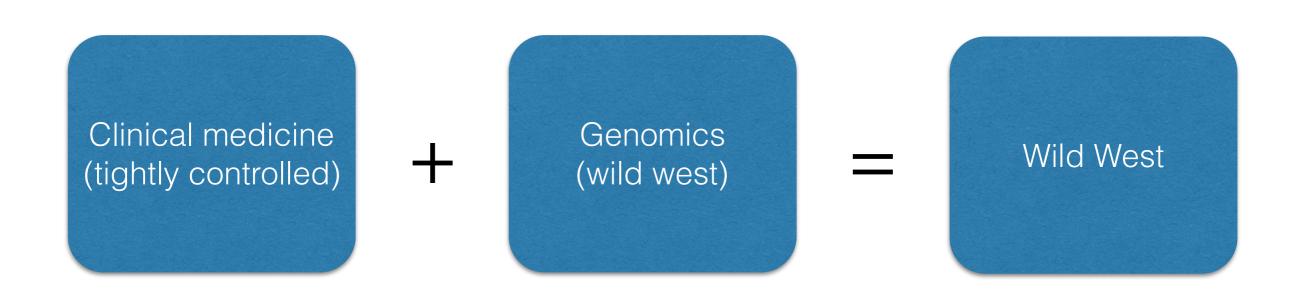
Steve Goodman sgoodman@jhmi.edu via googlegroups.com Unsubscribe



to reproducible-r. 💌

BTW, I felt that Keith and Kevin's 45 seconds was akin to listening to "Ride of the Valkyries"in a TV commercial instead of hearing the whole of Die Walkure. There ain't nothin' better than the full Die Baggerly, as long as Keith is singing!

Lessons?



Institute of Medicine Committee

REPORT BRIEF SMARCH 2012

INSTITUTE OF MEDICINE

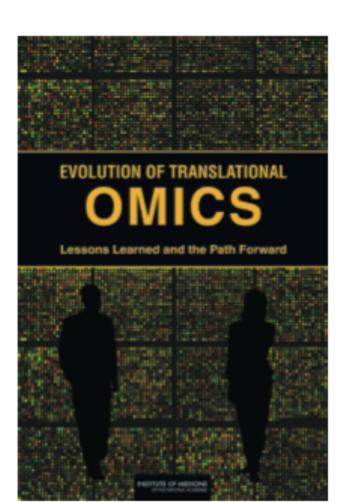
OF THE NATIONAL ACADEMIES

Advising the nation • Improving health

For more information visit www.iom.edu/translationalomics

Evolution of Translational Omics

Lessons Learned and the Path Forward



The IOM Report

- Data/metadata used to develop test should be made publicly available
- The computer code and fully specified computational procedures used or development of the omics-based test should be made available
- Ideally, the computer code that is released will encompass all of the steps of computational analysis, including all data preprocessing steps

Replication and Reproducibility

Replication

- Focuses on the validity of the scientific claim
- "Is this claim true?"
- Ultimate standard for scientific evidence
- New investigators, data, analytic methods, labs, instruments, etc.
- Important in studies that can impact policy or regulation

Reproducibility

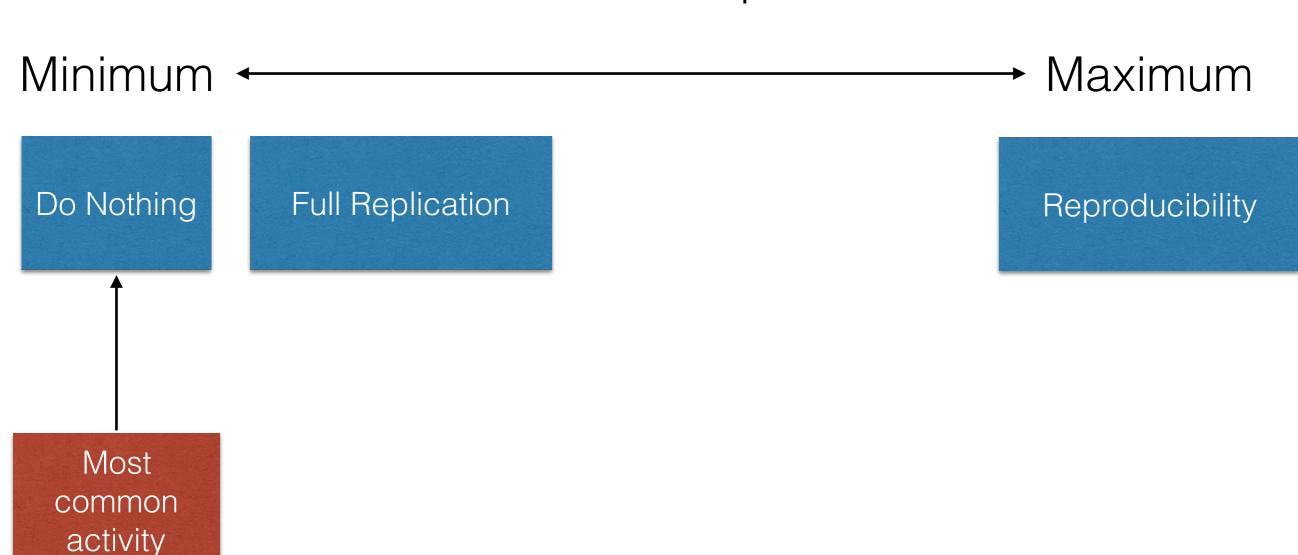
- Focuses on the validity of the data analysis
- "Can we trust this analysis?"
- A minimum standard
- New investigators, same data, same methods
- Important when replication is impossible

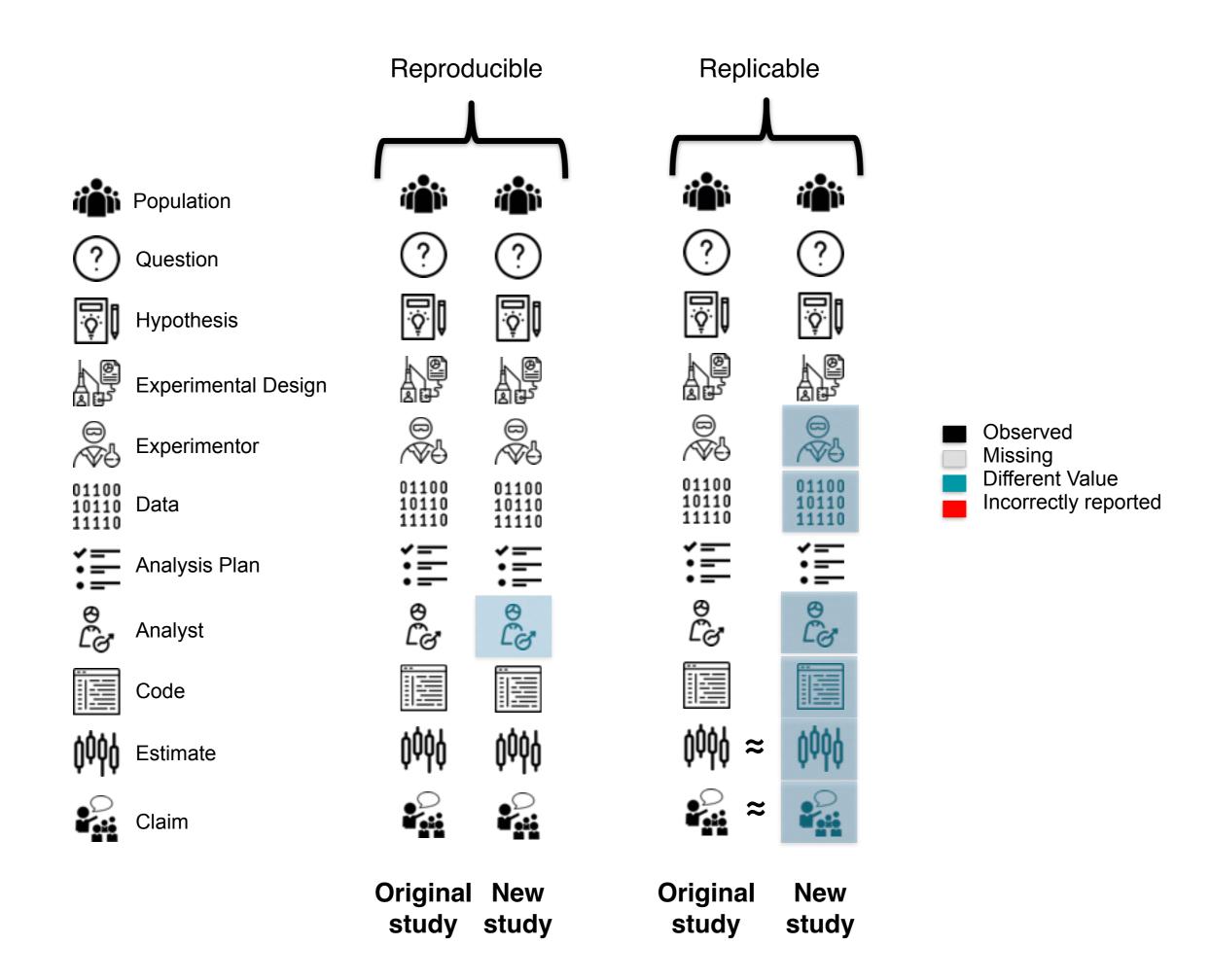
What's Wrong with Replication?

- Nothing, but...
- Some studies cannot be replicated
 - No time, opportunistic
 - No money
 - Unique
- Reproducible Research: Make analytic data and code available so that others may reproduce findings

Upon Seeing Your Work...

Information Required



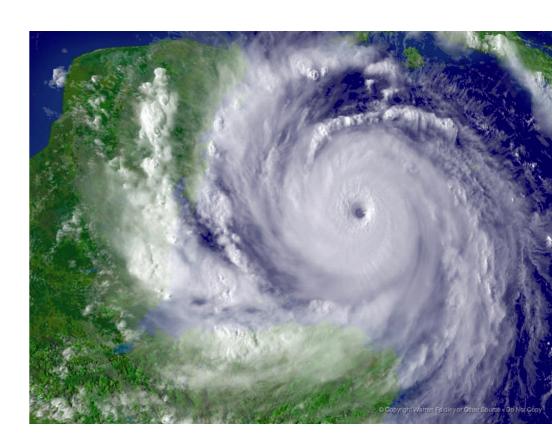


Why Do We Need Reproducible Research?

- New technologies increasing data collection throughput
- Data are more complex and high dimensional
- Existing databases can be merged into new and bigger databases
- Computing power is greatly increased, allowing more sophisticated/complicated analyses
- For every field "X" there is a field "Computational X"

Air Pollution and Health: A Perfect Storm?

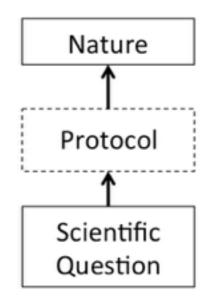
- Estimating small health effects in the presence of much stronger signals
- Results inform substantial policy decisions and affect many stakeholders
- EPA regulations can cost billions of dollars
- Complex statistical methods are needed and subjected to intense scrutiny



The End Result

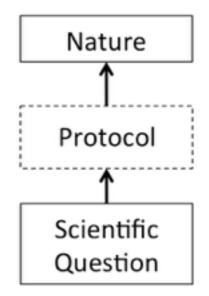
- Basic analyses can be difficult to describe
- Heavy computational requirements are thrust upon people without adequate training in statistics and computing
- Errors are more easily introduced into long and complex analysis pipelines
- Knowledge transfer is limited
- Complicated analyses cannot be trusted

Published Article



Author

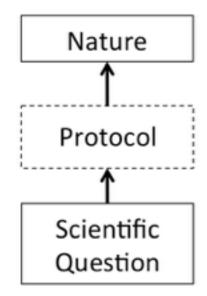
Published Article



Reader

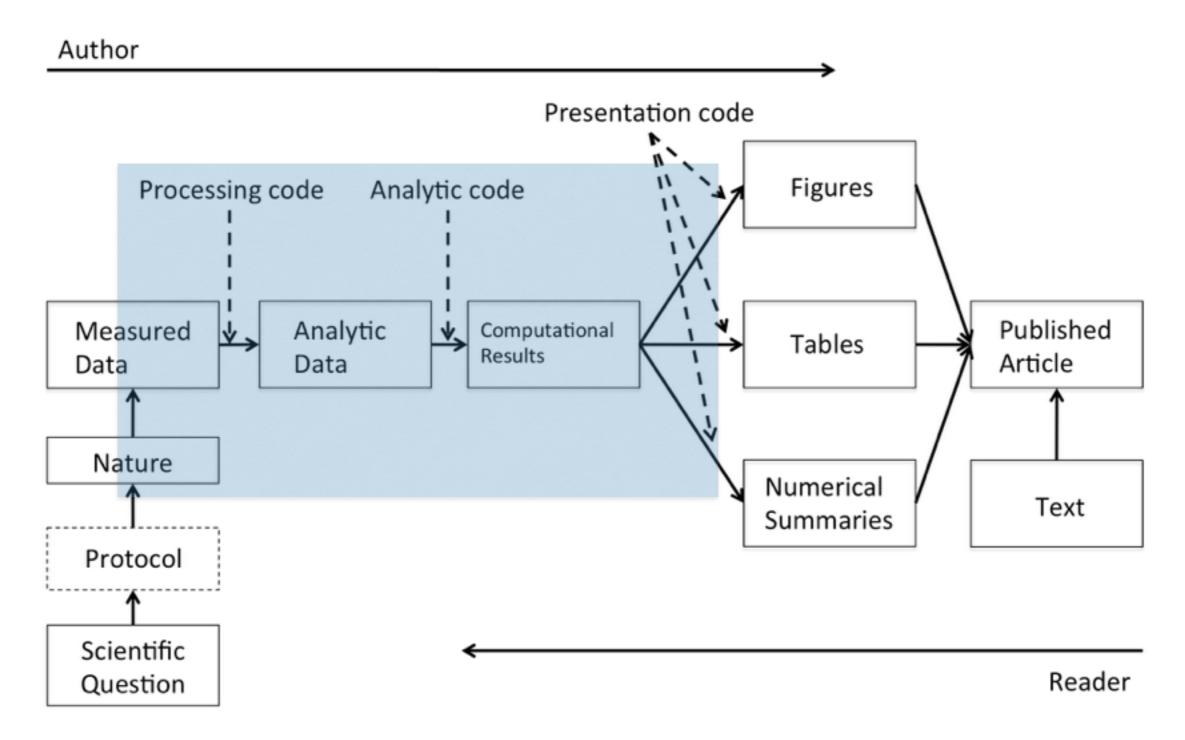
Author

Published Article



Express train to nature

Reader



- Analytic data are available
- Analytic (and preprocessing) code are available
- Documentation of code and data
- Standard means of distribution

- Authors
 - Want to make their research reproducible
 - Want tools for RR to make their lives easier (or at least not much harder)
- Readers
 - Want to reproduce (and perhaps expand upon) interesting findings
 - Want tools for RR to make their lives easier

Challenges

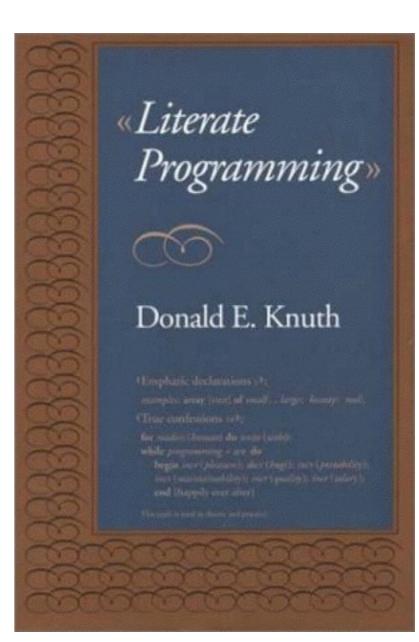
- Authors must undertake considerable effort to put data and results on the web (may not have resources like a web server)
- Readers must download data/results individually and piece together which data go with which code sections, etc.
- Readers may not have the same resources as authors
- Few tools to help authors/readers (although toolbox is growing!)

Recent Developments

- Software: iPython Notebooks, knitr, markdown, LONI, Galaxy
- Repositories: GitHub, NCBI, ICPSR, Dataverse
- Policy: Science, Nature, PLOS ONE, OSTP, NIH

Literate Statistical Programming

- An article/report is a stream of text and code
- Analysis code is divided into text and code "chunks"
- Each code chunk loads data and computes results
- Presentation code formats results (tables, figures, etc.)
- Article text explains what is going on
- Literate programs can be weaved to produce humanreadable documents and tangled to produce machine-readable documents
- See Literate Programming by Donald Knuth



Literate Statistical Programming

- Literate programming is a general concept that requires
 - A documentation language (human readable)
 - A programming language (machine readable)
- Sweave uses LaTeX and R as the documentation and programming languages
- Sweave was developed by Friedrich Leisch (member of the R Core) and is maintained by R core
- Main web site: http://www.statistik.lmu.de/~leisch/Sweave

Literate Statistical Programming

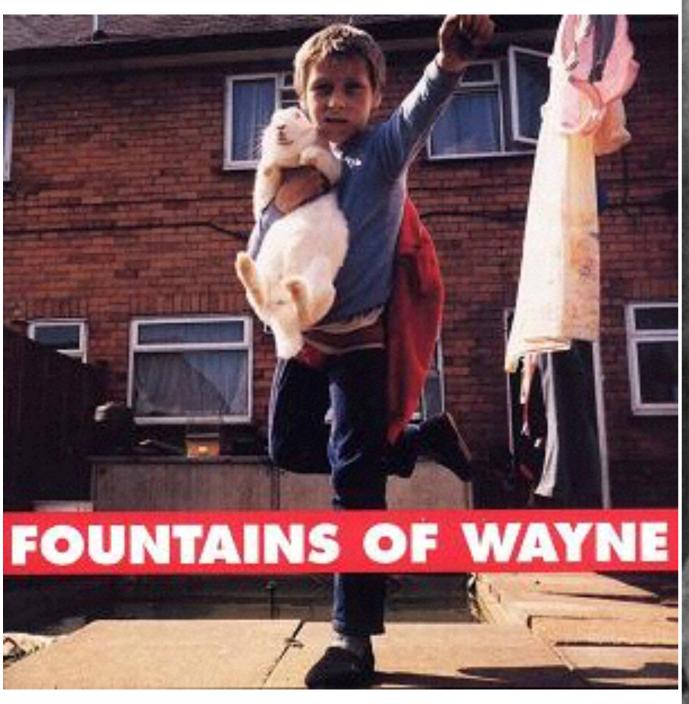
- knitr is package that brings together many features added on to Sweave to address limitations
- knitr uses R as the programming language knitr was developed by Yihui Xie (while a graduate student in statistics at Iowa State, now at RStudio)
- knitr uses the R programming language (although others are allowed) and variety of documentation languages
 - LaTeX, Markdown, HTML
- Built into RStudio pipeline
- See http://yihui.name/knitr/



What Problem Does Reproducibility Solve?

- What we get
 - Transparency / Improved knowledge transfer
 - Data availability
 - Software / Methods
- What we do NOT get
 - Validity / Correctness of the analysis

Computational Research has a Communication Problem





```
Verse 1:
                     Bb
It's you and me on a beach/ In nineteen-ninety eight
Leaning into the breeze/ From the willows and rivermen grace
Are reborn in this place/ I'm assured the procedure is painless
The taxicab with no brakes/ around the mountain pass
                                         Bb
Keep your head in your hands/ if anybody askes what you mean when
         Αm
You were picking a fight/ you were only complimenting the waitress
Chorus 1:
                                Bb
Give us a room, with a mountain view:
                                                   Bb
A tiny cabana by the water, yeah by the water and I,
Got a rental for an hour or two, for a ride up the coast
And a dip in the ocean.
(Repeat Intro)
Verse 2:
The waterfront is alight/ with Citronella flame
Tourists flash in the night/ from the grottoes and
                               Bb
                     Am
Gathering now on the heel-worn planks for a drunken
Form another, a mumble (?)
And lovers paddle a boat/ on the molten bay
                     Bb
Peering into the the reeds/ on a ripple
And playing it cool/ in a bar by the pool
With a Caribbean Kiss Amaretto
```

```
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With a Caribbean Kiss Amaretto

I. TEIL.

Aufführungsrecht vorbehalten Hymnus: Veni, ereator spiritus. Droits desicution réservés. Bas-Klartnette in B 1.2.8.4. Pagott. Kontra-Fagott. 1.2.3.4.Trompete in P. 1.2.3.4. Pressure. Pauken. Orgel. 1.2. Sopran. 1.2.Alt. Tepor. Bariton. Baß. Knabenchor. Sopran. Alt. Tenor. Sopran. Alt. Tenor. Be5. 1. Violine. 2. Violine. Bratsche, Violoncell.

The Central Problem

Data Analysis = ???

What's Next?

- Reproducibility is critical for communicating a data analysis
- One cannot sufficiently describe an analysis in words
- General consensus about its importance
- Infrastructure for making all research reproducible is not there yet, but things are ever improving

How Do You Know if a Data Analysis is Successful?

- Reproducible
- Uses the best available statistical methods of analysis

"There ain't nothin' better than the full Die Baggerly, as long as Keith is singing!"

-Steve Goodman