Steps toward reproducible research

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> kbroman.org github.com/kbroman @kwbroman Slides: bit.ly/steps2rr



Karl -- this is very interesting, however you used an old version of the data (n=143 rather than n=226).

I'm really sorry you did all that work on the incomplete dataset.

Bruce

The results in Table 1 don't seem to correspond to those in Figure 2.

Where did we get this data file?

Why did I omit those samples?

Which image goes with which experiment?

How did I make that figure?

In what order do I run these scripts?

"Your script is now giving an error."

"The attached is similar to the code we used."

Reproducible

VS.

Replicable

Reproducible

VS.

Correct

kbroman.org/steps2rr

A little bit reproducible is better than not reproducible.

A little bit open is better than not open.

Strive to make each project a bit better organized than the last.

File organization and naming are powerful weapons against chaos.

Jenny Bryan

Your closest collaborator is you six months ago, but you don't reply to emails.

(paraphrasing Mark Holder)

Have sympathy for your future self.

```
RawData/
                        Notes/
DerivedData/
                        Refs/
Python/
                        ReadMe.txt
R/
                        ToDo.txt
Ruby/
                        Makefile
Analysis/
Figures/
```

Chaos

```
AimeeNullSims/
                  Deuterium/
                                         Ping/
AimeeResults/
                  ExtractData4Gary/
                                         Ping2/
                  FromAimee/
                                         Ping3/
AnnotationFiles/
Brian/
                  GoldStandard/
                                         Ping4/
Chr6_extrageno/
                  HumanGWAS/
                                         Plav/
Chr6 segdis/
                  Insulin/
                                         Prdm9/
ChrisPlaisier/
                  Int2_for_Mark/
                                         RBM_PlasmaUrine_2012-03-08/
Code4Aimee/
                  Islet 2011-05/
                                         Slco1a6/
                  MappingProbes/
CompAnnot/
                                         StudyLineupMethods/
CondScans/
                  MultiProbes/
                                         kidney_chr6.R
D20 2012-02-14/
                  NewMap/
                                         pck2 sucla2.R
D20 cellcvcle/
                  Notes/
                                         penalties.txt
D2Ocorr/
                  NullSims/
                                         transeQTL4Lude/
Data4Aimee/
                  NullSims 2009-09-10/
Data4Tram/
                  PepIns_2012-02-09/
```

```
betw tissue corr.R
                       expr scatterplot allprobes.R
                                                       gve similarity alltissues.R
coatcolor lod.R
                       expr_scatterplots_dup.R
                                                       gve_similarity.R
colors.R
                       expr_scatterplots_mix.R
                                                       gve_supp.R
cover_fig.R
                       expr_scatterplots_swap.R
                                                       insulin_lod.R
eqtl_counts_10.R
                       expr_swaps.R
                                                       local_eqtl_locations.R
eatl counts.R
                       func.R
                                                       my plot map.R
eve hist.R
                       genotype plates.R
                                                       my plot scanone.R
eve scheme.R
                       gve_hist.R
                                                       sex_vs_X.R
eve similarity.R
                       gve new.R
                                                       xchr fig.R
eve_similarity_supp.R
                                                       xist_and_y.R
                       gve.R
expr_corr_dup.R
                       gve_scheme.R
expr corr mix.R
                       gve similarity 2ndbest.R
```

```
fig1.png fig5.png
fig10.png fig6.png
fig2.png fig7.png
fig3.png fig8.png
fig4.png fig9.png
```

- Machine readable
 - No spaces
 - No special characters except _ and -
- ► Human readable
 - Explain the contents
- Consistent
 - Name similar files in a similar way
- Make use of computer's sorting
 - pad numbers with 0's (e.g., 01, 02, ...)
 - start with general grouping, then more specific
 - dates like 2019-05-14

PUBLIC SERVICE ANNOUNCEMENT:

OUR DIFFERENT WAYS OF WRITING DATES AS NUMBERS CAN LEAD TO ONLINE CONFUSION, THAT'S WHY IN 1988 ISO SET A GLOBAL STANDARD NUMERIC DATE FORMAT.

THIS IS THE CORRECT WAY TO WRITE NUMERIC DATES:

2013-02-27

THE FOLLOWING FORMATS ARE THEREFORE DISCOURAGED:

02/27/2013 02/27/13 27/02/2013 27/02/13 20130227 2013.02.27 27.02.13 27-02-13 27.2.13 2013.15.8904109 MMXIII-II-XXVII MMXIII $\frac{LVII}{CCCLXV}$ 1330300800 ((3+3)×(111+1)-1)×3/3-1/3³ 2014 115555 10/11011/1101 02/27/20/13 $\frac{2}{9}$ 1 $\frac{1}{2}$ 3 $\frac{7}{2}$

```
0 vcf2db.R
1 prep geno.R
2 prep pheno clin.R
2 prep pheno otu.R
3 prep covar.R
4 prep analysis pheno clin.R
4 prep analysis pheno otu.R
5 scans.R
6 grab peaks.R
7 find nearby peaks.R
```

No "final" in file names



No "final" in file names

Deprecated/ ReadMe.txt adipose int1 final.RData adipose int2 final.RData adipose mlratio final.RData adipose mlratio ngrank final.RData adipose prcomp.RData aligned geno with pmap.RData batches final.RData batches_raw_final.RData cpl final.RData d2o final.RData gastroc int1 final.RData gastroc int2 final.RData gastroc mlratio final.RData gastroc mlratio ngrank final.RData gastroc prcomp.RData hypo int1 final.RData hypo int2 final.RData hypo mlratio final.RData hypo mlratio final old.RData hypo mlratio ngrank final.RData hypo mlratio ngrank final old.RData hypo omit.RData

hypo prcomp.RData islet int1 final.RData islet int2 final.RData islet mlratio final.RData islet mlratio ngrank final.RData islet prcomp.RData kidney int1 final.RData kidney int2 final.RData kidney mlratio final.RData kidney mlratio ngrank final.RData kidney prcomp.RData lipomics final rev2.RData liverTG final.RData liver int1 final.RData liver int2 final.RData liver mlratio final.RData liver mlratio ngrank final.RData liver prcomp.RData mirna final.RData necropsy_final_rev2.RData plasmaurine final rev.RData pmark.RData rbm final.RData

No "final" in file names

Deprecated/ ReadMe.txt adipose int1 final.RData adipose int2 final.RData adipose mlratio final.RData adipose mlratio ngrank final.RData adipose prcomp.RData aligned geno with pmap.RData batches final.RData batches_raw_final.RData cpl final.RData d2o final.RData gastroc int1 final.RData gastroc int2 final.RData gastroc mlratio final.RData gastroc mlratio ngrank final.RData gastroc prcomp.RData hypo int1 final.RData hypo int2 final.RData hypo mlratio final.RData hypo mlratio ngrank final.RData hypo mlratio ngrank final old.RData hypo omit.RData

hypo prcomp.RData islet int1 final.RData islet int2 final.RData islet mlratio final.RData islet mlratio ngrank final.RData islet prcomp.RData kidney int1 final.RData kidney int2 final.RData kidney mlratio final.RData kidney mlratio ngrank final.RData kidney prcomp.RData liverTG final.RData liver int1 final.RData liver int2 final.RData liver mlratio final.RData liver mlratio ngrank final.RData liver prcomp.RData mirna final.RData pmark.RData rbm final.RData

batches raw v1.rds batches v1.rds clinical cpl v2.rds clinical d2o v2.rds clinical lipomics v4.rds clinical liverTG v2.rds clinical mirna v2.rds clinical necropsy v4.rds clinical plasmaurine v3.rds clinical rbm v2.rds Deprecated/ geneexpr int1 adipose v2.rds geneexpr int1 gastroc v2.rds geneexpr_int1_hypo_v2.rds geneexpr int1 islet v2.rds geneexpr int1 kidney v2.rds geneexpr_int1_liver_v2.rds geneexpr_int2_adipose_v2.rds geneexpr int2 gastroc v2.rds geneexpr_int2_hypo_v2.rds geneexpr_int2_islet_v2.rds geneexpr int2 kidney v2.rds geneexpr int2 liver v2.rds geneexpr mlratio adipose v2.rds geneexpr mlratio gastroc v2.rds geneexpr_mlratio_hypo_v1.rds geneexpr mlratio hypo v2.rds geneexpr mlratio islet v2.rds geneexpr mlratio kidney v2.rds geneexpr mlratio liver v2.rds geneexpr mlratio ngrank adipose v2.rds geneexpr_mlratio_nqrank_gastroc_v2.rds geneexpr mlratio ngrank hypo v1.rds geneexpr mlratio ngrank hypo v2.rds geneexpr mlratio ngrank islet v2.rds geneexpr mlratio ngrank kidney v2.rds geneexpr mlratio ngrank liver v2.rds geneexpr_omit_hypo.rds geneexpr prcomp adipose v2.rds geneexpr_prcomp_gastroc_v2.rds geneexpr prcomp hypo v2.rds geneexpr_prcomp_islet_v2.rds geneexpr prcomp kidney v2.rds geneexpr_prcomp_liver_v2.rds geno_aligned_w_pmap.rds geno pmark.rds ReadMe.txt

Document your work

- What is all of this stuff?
- ► What was your analysis process?

ightarrow ReadMe files

Organizing data in spreadsheets

	А	В	С	D	E	F	G
1	1MIN						
2			Normal			Mutant	
3	В6	146.6	138.6	155.6	166	179.3	186.9
4	BTBR	245.7	240	243.1	177.8	171.6	188.1
5							
6	5MIN						
7			Normal			Mutant	
8	В6	333.6	353.6	408.8	450.6	474.4	423.8
9	BTBR	514.4	610.6	597.9	412.1	447.4	446.5

Organizing data in spreadsheets

	Α	В	С	D
1	ttt_min	strain	mutation	response
2	1	B6	normal	146.6
3	1	B6	normal	138.6
4	1	В6	normal	155.6
5	1	В6	mutant	166
6	1	B6	mutant	179.3
7	1	B6	mutant	186.9
8	1	BTBR	normal	245.7
9	1	BTBR	normal	240
10	1	BTBR	normal	243.1
11	1	BTBR	mutant	177.8
12	1	BTBR	mutant	171.6
13	1	BTBR	mutant	188.1
14	5	B6	normal	333.6
15	5	В6	normal	353.6

Organizing data in spreadsheets

- Make it a rectangle
- Individual measurements as rows; variables as columns
- Single header row
- One item per cell
- No empty cells
- No calculations in the raw data
- No highlighting or coloring as data

"What the heck is 'FAD_NAD SI 8.3_3.3G'?"

Metadata

Create a data dictionary

- Explain each column
- Include different versions of the variable names (compact vs descriptive)
- Units
- Allowable values

▶ The metadata are data

Make it a rectangle

Data dictionary

	А	В	С	D
1	name	plot_name	group	description
2	mouse	Mouse	demographic	Animal identifier
3	sex	Sex	demographic	Male (M) or Female (F)
4	sac_date	Date of sac	demographic	Date mouse was sacrificed
5	partial_inflation	Partial inflation	clinical	Indicates if mouse showed partial pancreatic inflation
6	coat_color	Coat color	demographic	Coat color, by visual inspection
7	crumblers	Crumblers	clinical	Indicates if mouse stored food in their bedding
8	diet_days	Days on diet	clinical	Number of days on high–fat diet

Everything with a script

If you do something once, you'll do it 1000 times.

Reproducible reports

Gough project diagnostics

Karl Broman, 3 March 2014

Combine genotypes and phenotypes

I've combined the initial genotypes (using the re-clustered genotypes for plates 14-16) with the well-behaved portion of the re-run genotypes. I'm focusing on 36813 markers that are informative (though, as we'll see, there are still a lot of badly behaved and basically non-informative markers that need to be removed). I've combined data on replicate samples, to give one set of genotype calls for each sample.

There are 1497 genotyped mice and 1464 phenotyped mice. All of the mice in the phenotype data have genotypes, but there are 33 genotyped mice with no phenotypes, including 3 Gough mice and 30 F2 progeny.

Reproducible reports

Gough project diagnostics

```
25 I've combined the initial genotypes (using the re-clustered genotypes
Karl
     26 for plates 14-16) with the well-behaved portion of the re-run
     27 genotypes. I'm focusing on 'r totmar(g)' markers that are informative
Col
     28 (though, as we'll see, there are still a lot of badly behaved and
I've
     29 basically non-informative markers that need to be removed).
the v
     30 I've combined data on replicate samples, to give one set of genotype
infor
     31 calls for each sample.
info
     32
give
     33 There are 'r nind(g)' genotyped mice and 'r nrow(phe)' phenotyped
     34 mice. All of the mice in the phenotype data have genotypes, but there
Ther
     35 are 'r sum(is.na(match(gid, pid)))' genotyped mice with no phenotypes.
data
        including `r sum(g$pheno$gen[which(is.na(match(gid, pid)))]==0)`
mice
        Gough mice and `r sum(g$pheno$gen[which(is.na(match(gid, pid)))]==2)`
     38 F2 progeny.
```

```
R/analysis.html: R/analysis.Rmd Data/cleandata.csv
    cd R;R -e "rmarkdown::render('analysis.Rmd')"

Data/cleandata.csv: R/prepData.R RawData/rawdata.csv
    cd R;R CMD BATCH prepData.R

RawData/rawdata.csv: Python/xls2csv.py RawData/rawdata.xls
    Python/xls2csv.py RawData/rawdata.xls > RawData/rawdata.csv
```

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R/analysis.html: R/analysis.Rmd Data/cleandata.csv
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```

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    Python/xls2csv.py RawData/rawdata.xls > RawData/rawdata.csv
```

Write modular code

- Modular code is easier to understand, maintain, and reuse.
- ► Turn repeated code into functions
- Combine useful functions into a package or module

Keeping track of versions

- ► Google drive / Dropbox / Box
- Version numbers in file names
- Formal version control (e.g., git/GitHub)
 - Browse changes
 - Try new things without fear of breaking what works
 - Jump to the state of the project at any time point
 - Merge simultaneous changes from multiple people

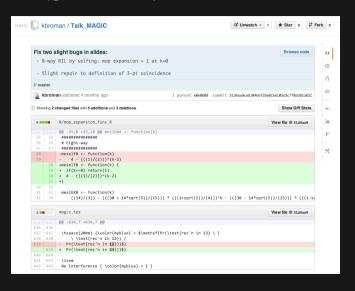




Greatly simplify the public domain stuff in the ReadMe			
kbroman authored 15 days ago		latest commit	f1777ef192 🔁
Figs	Add crazy table from preCC paper		4 months ago
Perl	Add lines_of_code_by_version.csv to repository		4 months ago
m R	Another fix regarding map expansion in 8-way RIL by selfing at k=0		4 months ago
gitignore	Add lines_of_code_by_version.csv to repository		4 months ago
Makefile	Revise Readme to link to version for web		4 months ago
ReadMe.md	Greatly simplify the public domain stuff in the ReadMe		15 days ago
magic.tex	Fix two slight bugs in slides:		4 months ago

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```
28
          -meself8 <- function(k)</pre>
               4 - (((1)/(2)))^{(k-2)}
     28
         +meself8 <- function(k) {
          + if(k==0) return(1)
     30
          + 4 - (((1)/(2)))^{(k-2)}
          +}
30
     32
31
     33
           mesibX8 <- function(k)</pre>
32
                ((14)/(3)) - (((30 + 14*sqrt(5))/(15)))
     34
```

Backups

- Multiple places, including off-site
- ► Automatic

License your software

Pick a license, any license

- Jeff Atwood

Share your stuff

▶ Code

- GitHub / BitBucket
- Zenodo (archival, with DOIs)

▶ Data

- Domain-specific repository (e.g., dbGAP)
- General repository (e.g., github, figshare, zenodo, datadryad)
- Institutional repository

Summary

- 1. Organize your project
- 2. Choose good names for things
- 3. Document what's what
- 4. Organize data as a rectangle
- 5. Metadata is data
- 6. Everything with a script
- 7. Even better: reproducible reports
- 8. Automate the process (GNU Make)
- 9. Write modular code (functions and packages)
- 10. Use version control (git/GitHub)
- 11. License your software
- 12. Share your data and code

Other considerations

Testing

are you getting the right answers?

Software versions

will your stuff work when dependencies change?

Large-scale computations

computation time + dependence on cluster environment

Collaborations

coordinating who does what and where things live

Collaboration

- Do more, by working in parallel
- Do more, through diversity of ideas and skills
- Reproducible pipelines have immediate advantages
- Tests of reproducibility
- Code review

Challenges in collaborations

- ► Shared vision?
- ► Compromise
- Coordination
- Communication
- Sharing code and data
- Synchronization

Challenges in collaborations

- ➤ Shared vision?
- ▶ Compromise
- Coordination
- Communication
- ► Sharing code and data
- ▶ Synchronization
- ▶ Weakest link?

Challenges

(totally hypothetical)

"Could we meet to talk about the data file structure?"

"Could we meet to talk about the data file structure?"

"No."

"Wait, these results seem to be based on the older SNP map."

"Could you write the methods section?"

"But I didn't do the work, and we don't have the code that was used."

"My data analyst has taken a job at Google."

"Could you do these analyses? X said they would, but they're not responding to my emails."

Shared vision

- ▶ Publication
- ▶ Code & data sharing
- ▶ Who will do what
- ▶ Timeline
- ► Ongoing sharing of methods, results

Shared workspace

- Project structure
- Data and metadata formats
- Software environment
- Automated sync (or it won't happen)

Technology for sharing

- Data
 - figshare
 - dropbox / box / google drive
- Code
 - github / bitbucket
- Pipeline / workflow
 - make / drake / snakemake / rake
- Full environment
 - docker containers
 - mybinder.org / wholetale.org

The most important tool is the mindset, when starting, that the end product will be reproducible.

Keith Baggerly

The second-most important tool is training.

- me

Slides: bit.ly/steps2rr



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