STAT 545A Class meeting #10 Wednesday, October 10, 2012

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Review of last class

Develop an R coding style. Keep your projects organized.

Use names. Avoid Magic Numbers. Break tasks and scripts in small pieces. Avoid hand-hacking, e.g. by writing functions.

Take advantage of .Rprofile and .Renviron to make persistent changes to your R life.

Manage your R installation so that upgrading once or twice a year is relatively painless.

<u>Source</u> is real. Other things ... not so much.

Make sure you can reproduce your analytical results and your figures from the input data without breaking much of a sweat.

Review of last class

Keep your eyes/ears/mind open about professional tools for reproducible research, collaboration and open science, and version control. You will probably need to address this as your training and career goes on.

Look at Sweave and knitr for creating dynamic, integrated documents of a data analysis. JB favors knitr, based on R Markdown, since markdown comes up in so many other places.

Consider git (JB's choice), mercurial, and Subversion (SVN) for version control.

If you choose git, then github is a great way to share your work with others and/or to collaborate.

Review of last class

(Semi-?) automate getting non-figure stuff out of R and ready for inclusion into, e.g. manuscripts, webpages.

Use write.table for spreadsheet-y stuff. dput / save / sink useful in special cases. Use packages like xtable / Hmisc / R2HTML for writing HTML and LaTeX tables.

Get started on your final project!

Do not underestimate the time and effort required to capture and tame a "wild" dataset (versus those found in captivity, i.e. data analyzed ad nauseum in textbooks, in previous classes you took, in R documentation, etc.). There will be weird stuff you must [1] detect and [2] fix.

How to ask a question to maximize chance of getting an answer.

The R Inferno

Patrick Burns¹

30th April 2011

Read about the 9th circle of R hell in <u>The R Inferno</u>.

Circle 9

Homework reading

Unhelpfully Seeking Help

Here live the thieves, guarded by the centaur Cacus. The inhabitants are bitten by lizards and snakes.

There's a special place for those who—not being content with one of the 8 Circles we've already visited—feel compelled to drag the rest of us into hell.

The road to writing a mail message should include at least the following stops:

9.1 Read the appropriate documentation.

"RTFM" in the jargon. There is a large amount of documentation about R, both official and contributed, and in various formats. A large amount of documentation means that it is often nontrivial to find what you are looking for—especially when frustration is setting in and blood pressure is rising.

Breathe.

There are various searches that you can do. R functions for searching include <code>help.search</code>, <code>RSiteSearch</code> and <code>apropos</code>.

If you are looking for particular functionality, then check the Task Views (found on the left-side menu of CRAN).

If you have an error, then look in rather than out—debug the problem. One way of debugging is to set the **error** option, and then use the **debugger** function:

options(error=dump.frames)
command that causes the error
debugger()

The **debugger** function then provides a menu of the stack of functions that have been called at the point of the error. You can inspect the state of the objects inside these functions, and hopefully understand what the problem is.

The R Inferno

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"If someone has the wit and knowledge to answer your question, they probably have other things they would like to do. Making your message clear, concise and user-friendly gives you the best hope of at least one of those strangers diverting their attention away from their life towards your problem." How to ask a question to maximize chance of getting an answer.

Homework reading

"<u>How To Ask Questions The Smart Way</u>" by Eric Raymond and Rick Moen

It's OK to be ignorant; it's not OK to play stupid.

So, while it isn't necessary to already be technically competent to get attention from us, it is necessary to **demonstrate the kind of attitude that leads to competence: alert, thoughtful, observant, willing to be an active partner in developing a solution**.

The best way to get a rapid and responsive answer is to ask it like a person with smarts, confidence, and clues who just happens to need help on one particular problem.

<u>R-help</u> (you probably shouldn't be posting here!) has a good posting guide

000		R: Mailing Lists Posting Guide	x		
IP + @http://www.R-pro	ject.org/posting-guide.html		•	C Q- R-help	0
🛱 🇰 myCourseWebStuff 🕶 myWebStu	ff * myWorkRSS (55) * myFunRSS (1) *	Pin It StorCenter ▼ UBC ▼ NYT	Merriam-Webster OnLine Jim Bryan	The R Project Google Maps	>
dai-a03 Bryan Lab	Home Bryan Lab	R: Mailing Lists Posting Guide			+

Posting Guide: How to ask good questions that prompt useful answers

This guide is intended to help you get the most out of the R mailing lists, and to avoid embarrassment. Like many responses posted on the list, it is written in a concise manner. This is not intended to be unfriendly - it is more a consequence of allocating the limited available time and space to technical issues rather than to social niceties.

The list: Remember that R is free software, constructed and maintained by volunteers. They have various reasons for contributing software and participating on the mailing lists, but often have limited time.

Homework reading

Do your homework before posting: If it is clear that you have done basic background research, you are far more likely to get an informative response. See also Further Resources further down this page.

- Do help.search("keyword") and apropos("keyword") with different keywords (type this at the R prompt).
- Do RSiteSearch("keyword") with different keywords (at the R prompt) to search R functions, contributed packages and R-Help postings. See ? RSiteSearch for further options and to restrict searches.
- Read the online help for relevant functions (type ?functionname, e.g., ?prod, at the R prompt)
- If something seems to have changed in R, look in the latest NEWS file on CRAN for information about it.
- Search the R-faq and the R-windows-faq if it might be relevant (<u>http://cran.r-project.org/faqs.html</u>)
- Read at least the relevant section in <u>An Introduction to R</u>
- · If the function is from a package accompanying a book, e.g., the MASS package, consult the book before posting
- The R Wiki has a section on finding functions and documentation

Technical details of posting: See General Instructions for more details of the following:

- No HTML posting (harder to detect spam) (note that this is the default in some mail clients you may have to turn it off). Note that chances have become relatively high for 'HTMLified' e-mails to be completely intercepted (without notice to the sender).
- No binary attachments except for PS, PDF, and some image and archive formats (others are automatically stripped off because they can contain
 malicious software). Files in other formats and larger ones should rather be put on the web and have only their URLs posted. This way a reader has the
 option to download them or not.
- Use an informative subject line (not something like `question')
- For new subjects, compose a new message and include the 'r-help@R-project.org' (or 'r-devel@R-project.org') address specifically. (Replying to an
 existing post and then changing the subject messes up the threading in the archives and in many people's mail readers.)
- If you can't send from an email address that simply accepts replies, then say so in your posting so that people are not inconvenienced when they try to
 respond to your message
- · Some consider it good manners to include a concise signature specifying affiliation

Where we go from here

Today: Two group tests. "Scaling up": different tests, many two group comparisons. Using figures to convey bulk statistical results instead of big tables of numbers. Use two group testing problem as way to introduce the bootstrap approach.

Next Monday: Continuing to develop and illustrate the bootstrap. Robust regression.

Next Wednesday: Smooth regression and cross validation.



Rationale for growth studies yeast deletion mutants

- Analogy: flipping circuit breakers in a house to determine which lights and outlets are controlled by each circuit
- If the deletion mutant for gene g is defective at some biological activity, that suggests that gene g contributes to that activity.
- Growth studies are the 'entry-level' study. In real life, we often measure more complicated phenotypes and subject the mutant to additional challenges, e.g. treatment with drugs or deletion/mutation of additional genes. Also, this type of data is often integrated with other types of studies.

Y = a quantitative measure of growth

e.g. growth rate or # cells at study end

Z = the specific yeast gene that was deleted e.g.YDLI33W

Yeast genome has 16 chromosomes.

Each gene lives somewhere on one of these chromosomes.

Therefore, each deletion mutant is also associated with one yeast chromosome.



Y = a quantitative measure of growth

e.g. growth rate or # cells at study end

Z = the specific yeast gene that was deleted e.g.YDLI33W

X = the chromosome on which the gene is

e.g. "chromosome D"

So, I quantitative variable Y and 2 categorical variables X, Z. Kind of like Y = lifeExp and Z = country and X = continent in Gapminder data.

Minimum you need to know

- Measuring growth -- an example of a phenotype -- on thousands of different strains of yeast. This is the response.
- Each strain is characterized by lacking the DNA for one gene.
- Each gene is found on one of the sixteen chromosomes in the yeast genome. This is the (potential) explanatory variable.
- It is interesting, for biological and experimental reasons, to ask whether the distribution of 'growth after gene deletion' differs across the chromosomes.
- Gapminder analology: interesting to assess whether the distribution of life expectancy differs across the continents.

the typical "two groups" testing problem

$$Y \mid X = x_i \sim F_i$$
$$Y \mid X = x_j \sim F_j$$
$$F_i = F_j ?$$

Y = measure of growth for mutant lacking a specific gene

X = chromosome on which this gene is found

Do you think there are systematic differences in the phenotype distribution across the 16 chromosomes?



 Data source: Giaever G, Flaherty P, Kumm J, Proctor M, Nislow C, et al. (2004) Chemogenomic profiling: identifying the functional interactions of small molecules in yeast. Proc Natl Acad Sci U S A 101: 793-798. <u>Pubmed</u>. DOI: <u>10.1073/pnas.</u> <u>0307490100</u>

All of my code, data, figures, and results for this example: <u>yeastDeletionGrowth</u>

I have done some preprocessing and simplification of a subset of the data associated with the reference above. Specifically, we are looking at growth data on yeast deletion mutants growing in control conditions. Underlying dataset available supplementary information from the article above.

> peek(gDat)

	geneDel	chromo	chromoPretty	pheno
844	YDR008C	4	D / IV	9.686326
2691	YIL134W	9	I / IX	9.188266
3034	YJR091C	10	J / X	10.167940
3322	YKR024C	11	K / XI	10.530350
4270	YMR260C	13	M / XIII	10.338061
4359	YNL029C	14	N / XIV	9.855063
4973	YOR123C	15	0 / XV	7.098627

Each row consists of

- geneDel = name of the gene that was deleted
- chromo = the associated chromosome (an integer between I and I6)
- chromoPretty = a prettier version of the chromosome (more suitable for labeling in tables and figures)
- pheno = a growth phenotype (due to experimental realities and preprocessing, the units are meaningless, i.e. don't expect to see a cell count here)

How many observations? How many genes? How many observations per gene?

> str(gDat) # 5666 observations
'data.frame': 5666 obs. of 4 variables:
\$ geneDel : Factor w/ 5521 levels "YAL001C", "YAL002W",...: | 2 3 4 5 6 7 8 9 10 ...
\$ chromo : int | | | | | | | | | ...
\$ chromoPretty: Factor w/ 16 levels "A / I", "B / II",...: | | | | | | | | | | ...
\$ pheno : num 9.39 9.40 10.38 10.54 8.65 ...

```
> table(table(gDat$geneDel))
```

```
This is puzzling.

5666 observations

5521 unique genes

5376 genes contribute 1 observation, 145 contribute 2

5376 + 145 = 5521 \checkmark

5376 + 2*145 = 5666 \checkmark
```

```
table() is a useful function
sometimes makes sense: table(table(...))
```

"Exhaustion and checking"

- The principle of exhaustion reassures the reader (and analyst!) that all cases are accounted for.
- GOOD: "Of the 84 patients with myocardial infarction, 22 had subendocardial infarction and 62 transmural infarction, with 78 discharged alive -- a mortality in the hospital of 7 percent."
- BAD: "... 78 patients were discharged alive, leading to an in-hospital mortality rate of 7 percent.

GOOD: 22 myo + 62 subendo = $84 \checkmark$ 84 - 78 = 6 died $6/84 = 0.07 \checkmark$ $6/84 = 0.07 \checkmark$ 6 = 0.07n n = 6/0.07n = 84

"Exhaustion and checking"

- Much easier for a reader to accept / confirm results that are complete and transparent.
 - Don't make the reader do irritating arithmetic.
 - Don't (seem to) contradict yourself!
 - Don't seem like you're hiding something!
- Build your credibility with your audience, i.e. that you are careful, thorough, and honest.
- Presents a golden opportunity for you to catch errors and confusion, before you start the analysis.
- Reference: "Writing About Numbers" by Frederick Mosteller in Medical Uses of Statistics, 2nd edition, Bailer and Mosteller (eds), NEJM Books, 1992.

How many observations per chromosome? How many unique genes?

Chromosome	# observations	# unique genes	# replicated genes	gFre dat
A / I	87	87	0	## \ ++++
B / II	370	370	0	##
C / III	135	132	3	σFre
D / IV	835	754	81	cbi
E/V	248	248	0	
F / VI	105	105	0	## 1
G / VII	561	512	49	writ
H / VIII	221	221	0	
I / IX	203	202	1	
J / X	326	325	1	
K / XI	302	302	0	
L/XII	500	499	1	
M / XIII	442	437	5	
N / XIV	373	372	1	
0 / XV	506	503	3	
P / XVI	452	452	0	

gFreqTable <-

data.frame(obsCount = table(gDat\$chromoPretty))

what if we want to count the number of unique genes ## on each chromosome?

'tapply' applies a function to a ragged array

gFreqTable\$uniqGeneCount <cbind(tapply(gDat\$geneDel, gDat\$chromoPretty, function(x) {return(length(unique(x)))}))

facilitate putting this table in a report
write.table(gFreqTable,

file = paste(whereAml, "chromoFreq.txt", sep = ""), quote = FALSE, row.names = FALSE, sep = "\t")

(easily copied and pasted from "chromoFreq.txt")

Getting to the bottom of the duplicated gene deletions

- See code for details
- Short version:
 - When there are 2 rows for a gene deletion, such as YCL038C, it turns out the observed phenotypes are exactly the same
 - So, it's a not a legitimate, independent observation of the growth phenotype
 - Immediate implication: I've eliminated these duplicates.
- hDat nows holds the clean, duplicate-free data

```
> str(hDat)
'data.frame': 5521 obs. of 4 variables:
 $ geneDel
                : Factor w/ 5521 levels "YAL001C", "YAL002W",..: 1 2 3 4 5 6 7 8..
 $ chromo
                : int 1 1 1 1 1 1 1 1 1 ...
 $ chromoPretty: Factor w/ 16 levels "A / I","B / II",...: 1 1 1 1 1 1 1 1 1 1 ...
 $ pheno
                : num 9.39 9.4 10.38 10.54 8.65 ...
> peek(hDat)
     geneDel chromo chromoPretty
                                       pheno
    YBL102W
                            B / II 9.285750
                    2
190
     YDR089W
                            D / IV 9.528659
917
                    4
                                                                    P/XVI
1040 YDR185C
                    4
                            D / IV 7.079669
1969 YGL201C
                           G / VII 9.754082
                    7
                                                                     O/XV
2118 YGR046W
                    7
                           G / VII 9.262812
                            K / XI 9.479903
3175 YKL085W
                   11
                                                                    N / XIV
3622 YLR176C
                   12
                           L / XII 7.359638
                                                                    M / XIII
> write.table(hDat,
                                                                     L / XII
               jPaste(whereAmI, "data/hDat.txt"),
+
               quote = FALSE, row.names = FALSE, sep = "\t")
                                                                     K / XI
+
                                                                      J/X
> dotplot(table(hDat$chromoPretty),
           origin = 0, type = c("p", "h"),
+
                                                                      1 / IX
           xlab = "# genes")
+
                                                                    H / VIII
> dev.print(pdf,
                                                                     G / VII
             jPaste(whereAmI, "figs/genesPerChromoDotplot.pdf"),
+
                                                                     F / VI
             width = 6, height = 8)
+
quartz
                                                                      E/V
     2
                                                                     D/IV
                                                                     C / III
                                                                      B/II
                                                                      A / I
                                                                           0
                                                                                    200
                                                                                             400
                                                                                                       600
                                                                                           # genes
```

sanity check of # genes on each chromosome





```
> densityplot( ~ pheno | chromoPretty, hDat,
+ plot.points = FALSE, layout = c(4,4),
+ xlab = "Growth phenotype")
> dev.print(pdf,
+ jPaste(whereAmI, "figs/phenoDensityplotByChromo4by4.pdf"),
+ width = 8, height = 6)
quartz
2
```



Real differences in distribution? I'll focus on comparing two chromosomes: 4 vs. 5.

0.0

6

> iDat <- subset(hDat, chromo %in% 4:5)</pre>

> str(iDat) # 'data.frame': 1002 obs. of 4 variables 'data.frame': 1002 obs. of 4 variables: : Factor w/ 5521 levels "YAL001C", "YAL002W",...: 590 591 592 593.. \$ geneDel \$ chromo : int 4 4 4 4 4 4 4 4 4 4 ... \$ chromoPretty: Factor w/ 16 levels "A / I", "B / II",..: 4 4 4 4 4 4 4 4 4 ... : num 9.78 8.23 9.82 7.62 8.24 ... \$ pheno > table(iDat\$chromo) A / I B/II 4 5 C / III 754 248 D/IV > ## highlighting issues around unused factor levels G / VII H / VIII | / IX > table(iDat\$chromoPretty) J / X D / IV A / I B/II C/III E / V F / VI G / VII H / VIII 0 0 754 248 0 0 0 n 0 / XV N / XIV I / IX J / X L / XII M / XIII P / XVI K / XI 0 / XV P / XVI 0 0 0 0 0 0 0 0 > densityplot(~ pheno, iDat, groups = chromoPretty, plot.points = FALSE, auto.key = TRUE, + 0.6 xlab = "Growth phenotype") + Density 0.4 0.2

Growth phenotype

10

8

Real differences in distribution? I'll focus on comparing two chromosomes: 4 vs. 5.



Real differences in distribution?



How would you assess?

Tests that address our question

- t test
- Wilcoxon test, aka Mann-Whitney here
- Kolmogorov-Smirnov test, 2 sample version
- Chi-square test of homogeneity
- I'm sure there are others





t test and Wilcoxon test

- t.test ... I assume all are familiar
 - default in R is to NOT assume common variance, i.e. to perform Welch's t test
 - suitability?
- Wilcoxon test is based on ranks, therefore is nonparametric
 - Rank all the data, ignoring the grouping variable
 - Compute the sum of the ranks for one group
 - Null distribution of this can be worked out / approximated
 - Also called Mann-Whitney test

Kolmogorov-Smirnov test (two sample)

- Hypothesis: $F_i = F_j$, i.e. distributions are same
- Estimate each CDF with the empirical CDF (ECDF)

$$\hat{F}_i(x) = \frac{1}{n} \sum_k I[x_{i,k} \le x]$$

 Test statistic is the maximum of the absolute difference between the ECDFs

$$\max \left| \hat{F}_i(x) - \hat{F}_j(x) \right|$$

- Null distribution does not depend on F_i , F_j (!)
- (I'm suppressing detail here.)

Kolmogorov-Smirnov test (two sample)

```
> ## sadly, Kolmogorov-Smirnov test has no formula interface
> jDat <- split(iDat$pheno, iDat$chromoPretty)</pre>
> str(jDat)
List of 2
 $ D / IV: num [1:754] 9.78 8.23 9.82 7.62 8.24 ...
 $ E / V : num [1:248] 10.57 8.62 10.6 10.57 8.93 ...
> ks.test(x = jDat[[1]], y = jDat[[2]])
    Two-sample Kolmogorov-Smirnov test
                                                                   D/IV
E/V
data:
       jDat[[1]] and jDat[[2]]
D = 0.3349, p-value < 2.2e-16
alternative hypothesis: two-sided
                                           0.6
                                         Density
                                           0.4
                                           0.2
                                           0.0
```

Growth phenotype

10

8

6

Chi-square test of homogeneity

- If F_i = F_j, then ... if we divide the support into bins, the observed relative frequency of these bins should be approximately the same in the two samples
- Obviously, choice of the bin boundaries is a big issue. Certainly true in terms of the theory. In practice, this should work fairly well most of the time.
- A 'pooled' estimate of the expected relative frequencies is obtained by ignoring the grouping variable.
- Conventional chi square test statistic arises from summing the usual terms:
 - (observed expected)² / expected
Could the observed bin counts (top) be random draws from one common underlying distribution (bottom)?

chr4 chr5

12

9

5

9

3

13

27

55

100

14

248

22

96

55

45

37

58

60

99

156

116

754

10

> addmargins(jCounts)

5.5

6 –

6.5

7.5

8.5

9.5

Sum

_

7

8

9

- 6

7

- 8

9

10

9.5

6.5

7.5

- 8.5

_

_

10 - 10.5

10.5 - 11



6

7

8

9

10

11

```
> (jBreaks <- hist(iDat$pheno)$breaks)  # plot is nice by-product!
[1] 5.5 6.0 6.5 7.0 7.5 8.0 8.5 9.0 9.5 10.0 10.5 11.0
> jCounts <- sapply(jDat, function(jPheno) {
+ hist(jPheno, breaks = jBreaks, plot = FALSE)$counts
+ })</pre>
```

```
> rownames(jCounts) <-
+ paste(jBreaks[-length(jBreaks)], jBreaks[-1], sep = " - ")</pre>
```

>	ac	ldn	narg	ins	(j(Count	S)
				ch	r4	chr5	Sum
5	. 5	_	6		22	1	23
6	_	6.	. 5		96	12	108
6	• 5	_	7		55	9	64
7	_	7.	. 5		45	5	50
7.	. 5	_	8		37	9	46
8	_	8.	. 5		58	3	61
8	. 5	-	9		60	13	73
9	_	9.	. 5		99	27	126
9.	. 5	_	10	1	56	55	211
1() -	- 1	L0.5	1	16	100	216
1().5	5 -	- 11		10	14	24
Sι	ım			7	54	248	1002

cut() is another useful function for making a quantitative variable into a factor

addmargins(), prop.table(), etc. helpful for working with contingency tables

> chisq.test(jCounts)

Pearson's Chi-squared test

data: jCounts
X-squared = 110.4444, df = 10, p-value < 2.2e-16</pre>

OK, we just conducted 4 different hypothesis tests. It would be nice to gather them together for comparison, no?

Especially when we scale up to other pairs of chromosomes, where the results might not be so unanimous.

Get to know the result objects

> tTestResult <- t.test(pheno ~ chromo, iDat)</pre>

```
> class(tTestResult)
[1] "htest"

> names(tTestResult)
[1] "statistic" "parameter" "p.value" "conf.int" "estimate"
[6] "null.value" "alternative" "method" "data.name"
```

```
> tTestResult$statistic
    t
-10.05494
```

```
> tTestResult$p.value
[1] 7.906832e-22
```

```
> tTestResult$conf.int
[1] -1.1101599 -0.7472465
attr(,"conf.level")
[1] 0.95
```

In addition to str(), class() and names() will help you understand the basics about most objects.

Once you understand the structure, you're ready to extract the good stuff.

Get to know the result objects

```
> ksTestResult <- ks.test(x = jDat[[1]], y = jDat[[2]])</pre>
```

```
> class(ksTestResult)
[1] "htest"
                                          # quite different from t test
> names(ksTestResult)
[1] "statistic" "p.value" "alternative" "method"
                                                            "data.name"
> ksTestResult$statistic
        D
0.3348592
> ksTestResult$p.value
[1] 0
> ksTestResult$estimate
                                          # not present
NULL
> ksTestResult$conf.int
                                          # not present
NULL
```

Contrast Kolmogorov-Smirnov output with the t test.

make a list and extract from it



- <u>lists</u> can hold just about anything. Here, the objects are hypothesis test results. All have class "htest".
- You don't even really need to understand what that means, as long as you realize you can use sapply() to pull out the pvalues of each component of the list.
- Sometimes key info from non-rectangular objects can be extracted and stored more easily than the entire object.

Now, consider all possible pairwise comparisons of chromosomes.

16 choose 2 = 16 * 15 / 2 = 120

Wouldn't it be cool to conduct all possible two group comparisons? For any one of the tests we're considering?

In exploratory data analysis, you want to find conclusions that hold up under various choices of methodology!

0.6

I / IX

12

10

8

6

Growth phenotype

6

8

N / XIV

J/X

O/XV

K / XI

10

12

P / XVI

L / XII

0.6

0.4 0.2 0.0

0.6 0.4 0.2

0.0

All possible pairs -- disclaimer

- This isn't so elegant statistically -- and unfortunately I see similar things done frequently.
- Remember, we have better ways to systematically answer the question "Is the distribution of phenotype common to all chromosomes?"
 - ANOVA, for starters (although not a great idea here)
 - bootstrap approaches
 - random or mixed effects models
- BUT, maybe this is sometimes a good idea. Here, it still presents a good learning opportunity.

I enumerate all possible pairs of chromosomes and apply all 4 "two groups" tests and gather all of the p-values.

```
> str(tqtRes)
'data.frame':
                120 obs. of 6 variables:
 $ chromoA: int 1 1 1 1 1 1 1 1 1 ...
                2 3 4 5 6 7 8 9 10 11 ...
 $ chromoB: int
                1.04e-05 4.86e-03 2.67e-11 8.66e-01 6.26e-05 ...
 $ t
          : num
 $ wilcox : num
                7.29e-07 2.10e-03 5.39e-11 7.77e-01 1.17e-05 ...
               2.53e-06 1.87e-02 1.26e-08 7.68e-01 2.98e-05 ...
 $ ks
      : num
 $ chisq : num 2.19e-07 5.55e-02 3.27e-09 6.26e-03 3.54e-04 ...
> peek(tqtRes)
                                                                  chisq
    chromoA chromoB
                                       wilcox
                                                        ks
                               t
                 10 5.077492e-01 1.260279e-01 4.265804e-02 3.055263e-02
23
          2
                 15 2.749797e-02 1.080776e-04 3.271894e-06 1.146343e-06
28
          2
43
          4
                  5 7.906832e-22 1.154823e-23 0.000000e+00 4.340871e-19
65
          5
                 16 7.659286e-08 7.746156e-07 7.007464e-06 2.444194e-04
          6
                 15 3.601006e-02 2.847043e-03 6.857203e-03 3.464897e-02
74
                 14 1.344403e-02 1.278636e-01 4.284638e-02 5.906893e-03
90
          8
119
         14
                 16 4.117154e-01 9.092607e-01 2.626741e-01 9.915321e-02
```

How to do this with apply() instead of an explicit loop -- or, even worse, a double loop? See code, if you are interested!

Let's make a scatterplot matrix of the pvalues from the various tests.

How many points will be in each panel?

What do you expect these scatterplots to look like? Why?

> round(cor(subset(tgtRes, select = c(t, wilcox, ks, chisq))), 2)

t wilcox ks chisq t 1.00 0.82 0.79 0.46 wilcox 0.82 1.00 0.83 0.47 ks 0.79 0.83 1.00 0.66

chisa	0.46	0.47	0.66	1.00
Curad	0.10	0.1/	0.00	T • 00



p-values from two groups tests

<pre>> round(cor(foo), 2)</pre>								
	t	wilcox	ks	chisq				
t	1.00	0.97	0.93	0.90				
wilcox	0.97	1.00	0.98	0.93				
ks	0.93	0.98	1.00	0.95				
chisq	0.90	0.93	0.95	1.00				



log10 p-values from two groups tests



p-values from two groups tests



log10 p-values from two groups tests



120 points in each panel, one per chromosome pair = two group comparison

strong agreement in 'statistically significant' region, elsewhere resembles indep. (?) U[0,1]

Now want to drill down to specific chromosome pairs.

Which ones are "different"?

Let's try to avoid the dreaded huge table of numbers



> head	(tgtRes)
--------	----------

	chromoA	chromoB	t	wilcox	ks	chisq
1	1	2	1.038025e-05	7.290265e-07	2.526472e-06	2.189111e-07
2	1	3	4.863497e-03	2.096621e-03	1.869599e-02	5.551262e-02
3	1	4	2.667323e-11	5.387564e-11	1.258664e-08	3.274657e-09
4	1	5	8.663625e-01	7.766479e-01	7.677386e-01	6.262497e-03
5	1	6	6.258230e-05	1.170805e-05	2.981307e-05	3.542402e-04
6	1	7	6.327104e-11	1.857242e-10	3.887342e-08	1.262049e-09

Consider one particular test, e.g. the Wilcoxon test, and focus just on those p-values.

Let's make a heatmap of these p-values.

First, reshape the p-value data.



```
> str(jP$wilcox)
num [1:16, 1:16] 1.00 7.29e-07 2.10e-03 5.39e-11 7.77e-01 ...
 - attr(*, "dimnames")=List of 2
  ..$ : chr [1:16] "A / I" "B / II" "C / III" "D / IV" ...
  ..$ : chr [1:16] "A / I" "B / II" "C / III" "D / IV" ...
> jP$wilcox[1:5, 1:5]
                                       C / III
                                                                   E / V
               A / I
                           B / II
                                                     D / TV
        1.000000e+00 7.290265e-07 2.096621e-03 5.387564e-11 7.766479e-01
A / I
   II 7.290265e-07 1.000000e+00 9.003812e-02 1.123049e-03 2.429079e-13
B /
    III 2.096621e-03 9.003812e-02 1.000000e+00 2.590718e-04 9.181813e-05
С
      5.387564e-11 1.123049e-03 2.590718e-04 1.000000e+00 1.154823e-23
    ΤV
D
       7.766479e-01 2.429079e-13 9.181813e-05 1.154823e-23 1.000000e+00
E / V
```

jP\$wilcox is a 16 x 16 symmetric matrix of p-values.

See Ch. 6 Trivariate displays in lattice 'Trivariate Displays' in <u>Sarkar</u> (2008).

- cloud() gives 3D scatterplots
- wireframe(), contourplot(), and levelplot() deal with surfaces and two-way tables
- Our matrix of p-values is a two-way table
 - Envision a square divided in to 16 * 16 cells
 - Each cell addresses a specific pair of chromosomes
 - Color the cell according to the evidence for a difference in distribution



row

- > ## graphs depicting the pairwise comparisons of phenotype dist'n
- > ## shade the squares according to p-value
- > levelplot(jP\$t, scales = list(x = list(rot = 45))) # wow, that's ugly

RColorBrewer palette 'GnBu'



GnBu (sequential)



Colors based on 'GnBu' palette from RColorBrewer

Key idea: associate a p-value of 1 with white and a p-value of 0 with an actual color.



Show data and modelling / inference results TOGETHER whenever you can.



Ordered by median phenotype

Use a rationale, deliberate ordering.

Ordered by clustering result



Ordered by Jenny



> print(lPlot, pos = c(0.15, 0, 1, 1), more = FALSE)

Ordered by Jenny



Problem: we're using most of our color space to depict distinctions we don't care about, i.e. p-value = 0.3 vs. 0.8.



This transformation will help emphasize differences in small to modest p-values and de-emphasize difference in medium to large p-values.



t-test, JB order, probit transformed p-values

```
## transforming the p-values to optimize mapping into the color ramp
## I want to de-emphasize differences between teeny p-values
## and between very large p-values
## and increase the visual impact of differences in (0.01, 0.10)
pvalueTicks <- c(0.0001, 0.001, 0.01, 0.05, 0.1, 0.5, 1)</pre>
qnScale <- 0.6
colorkeyTicks <- qnorm(pvalueTicks * qnScale)</pre>
jTitle <- "t-test, JB order, probit transformed p-values"
lPlot <-
  levelplot(qnorm$t(jP[jOrd, jOrd] * qnScale), col.regions = myCols,
            scales = list(x = list(rot = 45)),
            xlab = "", ylab = "", main = jTitle,
            colorkey = list(col = myCols,
              labels = list(labels = pvalueTicks, at = colorkeyTicks)))
print(dPlot, pos = c(0, 0.1, 0.2, 0.96), more = TRUE)
print(lPlot, pos = c(0.15, 0, 1, 1), more = FALSE)
```



t-test, JB order, probit transformed p-values

Results are similar, although there is 'more statistical significance' found by

KS test, JB order, probit transformed p-values

1

0.5

0.1

0.05

0.01

0.001

1e-0-

PII

, +*1*11



Colors based on 'GnBu' palette from RColorBrewer



Ordered by Jenny, probit transformed p-values

		<pre>> head(tgtRes[order(tgtRes\$t),], 15)</pre>						
Ì	- 1		chromoA	chromoB	t	wilcox		
	- 0.5	43	4	5	7.906832e-22	1.154823e-23	4.440	
		56	5	7	2.299423e-19	9.751576e-22	7.771	
	- 0.1	51	4	13	4.627917e-19	5.634388e-19	7.138	
ł	- 0.05	81	7	13	7.149913e-16	1.722451e-16	5.147	
	- 0.01	49	4	11	2.159276e-11	6.787809e-15	4.389	
	- 0.001	3	1	4	2.667323e-11	5.387564e-11	1.258	
	10.001	6	1	7	6.327104e-11	1.857242e-10	3.887	
	- 10-04	53	4	15	1.507109e-10	1.018281e-13	3.461	
		79	7	11	3.333448e-10	1.508345e-13	4.924	
		57	5	8	4.496093e-09	1.057575e-07	7.152	
		83	7	15	4.855781e-09	4.288748e-12	5.441	
		18	2	5	6.627043e-09	2.429079e-13	8.360	
		65	5	16	7.659286e-08	7.746156e-07	7.007	
		50	4	12	1.783292e-07	1.877395e-07	2.964	
		61	5	12	2.634741e-07	2.634340e-09	6.877	

With this figure, it easy to recover which pairs are most different and what the actual p-value is (approximately).

There's always room for improvement

- Better size for density plots ... make levelplot rectangular not square? How to make sizing more compatible?
- Mapping of p-values to colors, i.e. can I improve on the probit?
- Color ramp ... can I, should I make it easier to read off stat. sig. pairs at a <alpha> level?
- Put the density plot in those wasted blank diagonal squares ... would be awesome but a pain to do?

My goals in this and all plots

- Don't chicken out on "scaling up".
- Try to replace a table of numbers with a picture.
- Facilitate comparisons
- Facilitate the identification of trends
- Recommended reference: Gelman A, Pasarica C, Dodhia R."Let's Practice What We Preach: Turning Tables into Graphs". The American Statistician, Volume 56, Number 2, I May 2002, pp. 121-130(10). via JSTOR

Statistical Computing and Graphics

Let's Practice What We Preach: Turning Tables into Graphs

Andrew GELMAN, Cristian PASARICA, and Rahul DODHIA

Statisticians recommend graphical displays but often use tables to present their own research results. Could graphs do better? We study the question by going through the tables in a recent issue of the *Journal of the American Statistical Association*. We show how it is possible to improve the presentations using graphs that actually take up less space than the original tables. We find a particularly effective tool to be multiple repeated line plots, with comparisons of interest connected by lines and separate comparisons isolated on different plots.

KEY WORDS: Data reduction; Graph; Table; Visual display.

plays. Our advice follows well-known principles of data display (see, e.g., Tufte 1983; Cleveland 1985) but applied to the presentation or research results as well as raw data.

2. DISPLAYING NUMERICAL RESULTS

Statistical research requires the display of many different kinds of numerical results, including raw numbers, data reductions, inferences, and—for research in theory and methodology—summaries of probability distributions, most notably frequency properties of statistical procedures. Unfortunately, it is still standard to display these latter summaries as tables rather than graphs. In this section of our article, we briefly review the research on data display and perception, and then we

2.2 The Goals and Principles of Statistical Data Display

Constructing the graphs shown in this article actually took <u>a lot of time and several iterations</u>, with each step provoking further thought on the motivation behind various techniques of graphical display. Here, we attempt to identify some of the key ideas, with the hope of making the task easier for future researchers.

Tables are more effective if the goal is to read off exact numbers. However, the interest in a statistical paper typically lies in <u>comparisons</u>, not absolute numbers. For example, there is no reason for the reader to care that the standard error for a particular parameter estimate is 0.029 as computed under a certain method and 0.054 under another method. Rather, the point is that the second number is almost twice as high. When dozens of such potential comparisons are possible, they can be seen much more clearly in a well-chosen graphical display than in a table (see Figure 3). The idea of comparisons provides a theoretical framework for statistical graphics (as is implicit in Tukey 1972, 1977).

from Gelman et al (2002)

Raters' characterization of sentences (absolute score range)	Percent		Frequency of	1996 total employed	Relative
Negative (1-1.9)	23.9	Profession	recent citations	(1,000)	frequency
Neutral (2-2.9)	52.2	Lawyers	8101	880	9.2
Positive (\geq 3)	23.9	Economists	1201	148	8.1
		Architects	1097	160	6.9
Total	100.0	Physicians	3989	667	6.0
		Statisticians	34	14	2.4
Chamatarization	Characterization		479	245	2.0
Gharacterization		Dentists	165	137	1.2
		Teachers			
Negative ()	N	(not university)	3938	4724	0.8
Neutral		Engineers	934	1960	0.5
Positive (-)		Accountants	628	1538	0.4
0% 20% 4	0% 60%	Computer programmers	91	561	0.2

Total

Figure 1. Top panel: Table from Ellenberg (2000) shows the relative frequencies of three categories in a set of 46 ratings of sentences. Bottom panel: Graphical display allows direct comparison without distractions of irrelevant decimal places. Parentheses show ± 1 standard error bounds based on the implicit binomial distribution with n = 46.



20,657

11,034

1.9

from Gelman et al (2002)

A common first reaction to rants about data display is: Yeah, yeah, I know that already. But, as we have seen, we—the statistical profession—don't know it yet! If the world's leading statistical journal doesn't do it right, there is obviously still room for progress. And the first steps are recognizing why the problem exists and forming a step-by-step plan to fix it. The steps we recommend, for most data displays in statistical research, are: (1) identify the key comparisons of interest; (2) display these on small individual plots with comparison lines or axes where appropriate; and (3) establish enough control over the graphical display so that small legible plots can be juxtaposed as necessary.

The graphs in this article are extremely simple, and serious exploratory data analysis benefits from many elaborations, including <u>color, multiple linked (trellis) plots</u>, and <u>dynamic displays</u> (see, for a start, Chambers et al. 1983; Cleveland 1985, 1993). We

from Gelman et al (2002)

Something to think about in your project and future work ... can you complement or replace a big table of numbers with a figure? Can you employ multiple methodologies and assess how same/different the results are?

Now ... moving beyond the "classical" tests and introducing the bootstrap.

CLASS ENDED HERE.