Evolutionary Genetics

LV 25600-01 | Lecture with exercises | 4KP



Jean-Claude Walser

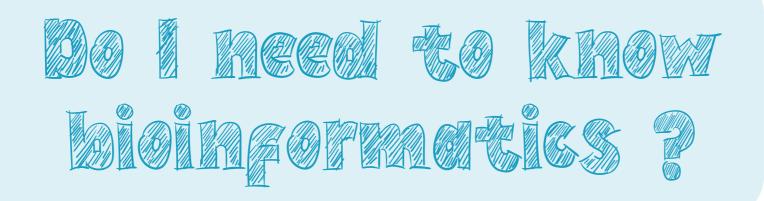
jean-claude.walser [at] usys.ethz.ch





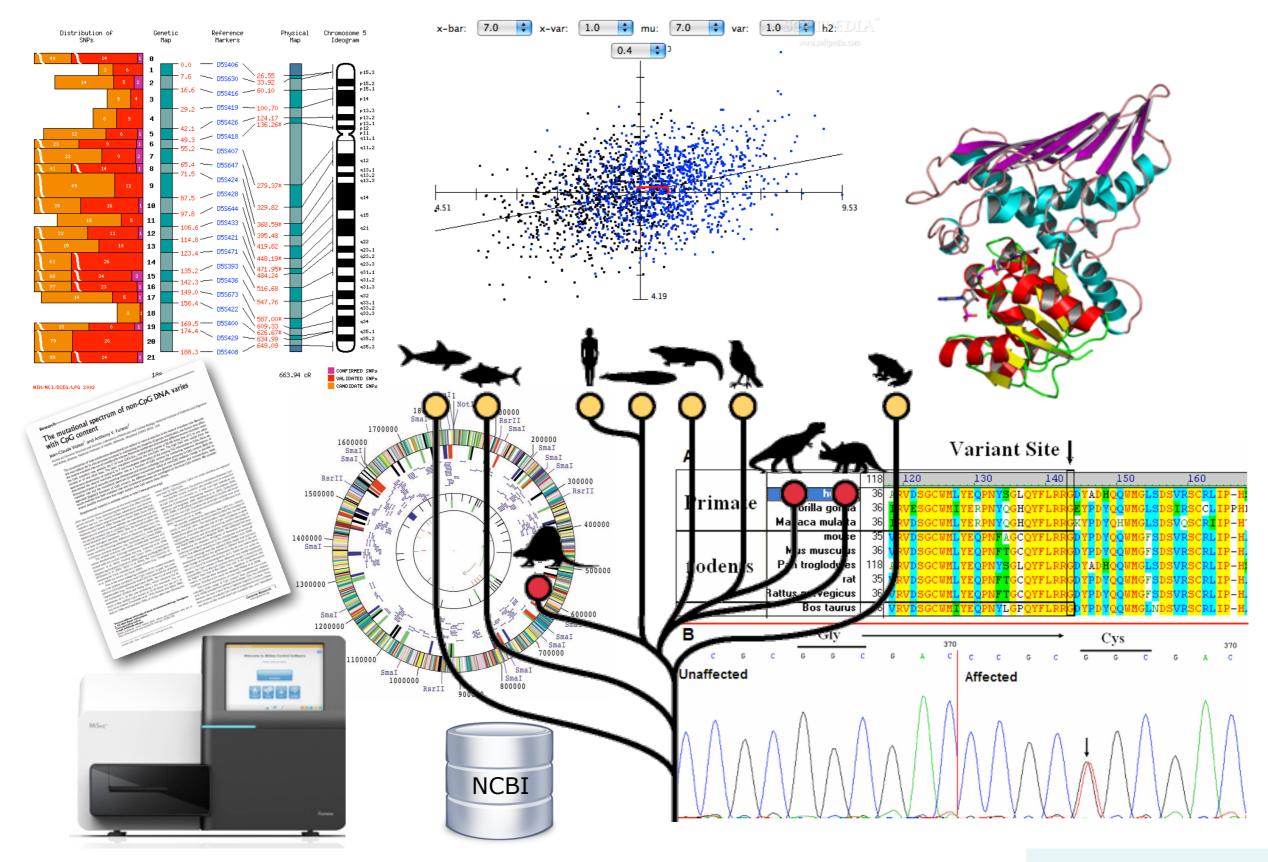


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Yes, absolutely. You don't have to be an expert (or geek) but you should know the basics.

"An **expert** is a person who has made all the mistakes that can be made in a very narrow field." -Niels Bohr



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Imagine, you are part of the **Covid broadcast team** and you are responsible for the **daily statistics**. All laboratories conducting Covid testing have to send daily reports. It is your duty to collect these reports. The labs are using standardised **CSV files**. At the end of a day you have **hundreds of CSV files** and you need to add them to your existing table. Your team-leader recommends to use the **spreadsheet editor** Microsoft Excel to store the data because he already has create a summary report template. Any questions?

The Guardian

CovidNews: how Excel may have caused loss of 16,000 test results in England

Public Health England data error blamed on limitations of Microsoft spreadsheet

One lab had sent its daily test report to PHE (Public Health England) in the form of a **CSV file** – the simplest possible database format, just a list of values separated by commas. That report was then **loaded into Microsoft Excel**, and the new tests at the bottom were added to the main database.

But while CSV files can be any size, Microsoft Excel files can only be 1,048,576 rows long – or, in older versions which PHE may have still been using, a mere 65,536. When a CSV file longer than that is opened, the bottom rows get cut off and are no longer displayed. That means that, once the lab had performed more than a million tests, it was only a matter of time before its reports failed to be read by PHE.

Alex Hern UK technology editor

@alexhern Tue 6 Oct 2020 08.21 BST

Microsoft's spreadsheet software is one of the world's most popular business tools, but it is regularly implicated in errors which can be costly, or even dangerous, because of the ease with which it can be used in situations it was not designed for.

In 2013, an Excel error at JPMorgan masked the loss of almost \$6bn (£4.6bn), after a cell mistakenly divided by the sum of two interest rates, rather than the average. The news led James Kwak, a professor of law at the University of Connecticut, to warn that Excel is "incredibly fragile".

"There is no way to trace where your data comes from, there's no audit trail (so you can overtype numbers and not know it), and there's no easy way to test spreadsheets, for starters. The biggest problem is that anyone can create Excel spreadsheets – badly. Because it's so easy to use, the creation of even important spreadsheets is not restricted to people who understand programming and do it in a methodical, well-documented way," Kwak wrote.

Errors from the spreadsheet software have even changed the very foundations of human genetics. The names of 27 genes have been changed over the past year by the Human Gene Nomenclature Committee, after Microsoft's program continually misformatted them. The genes SEPT1 and MARCH1, for instance, have been changed to SEPTIN1 and MARCHF1 after they were repeatedly turned into dates, while symbols that were common words have been altered so that grammar tools didn't autocorrect them: WARS is now WARS1, for instance.

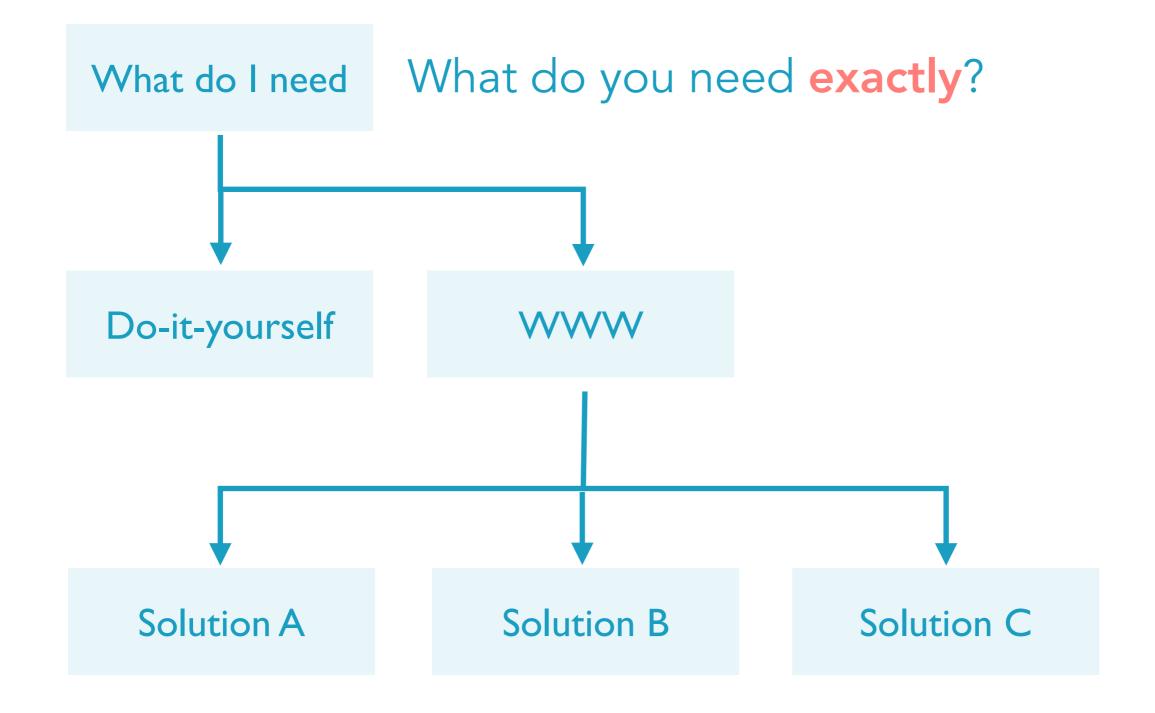


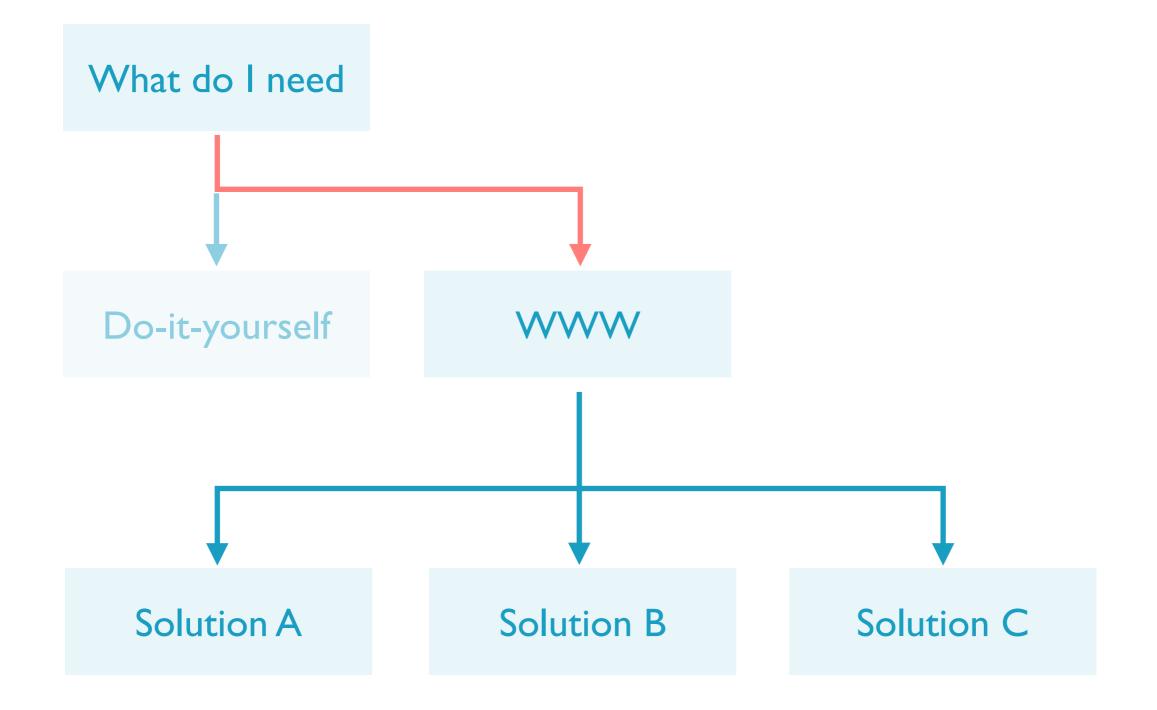
Alex Hern UK technology editor

♥ @alexhern Tue 6 Oct 2020 08.21 BST "A new scientific truth does not triumph by convincing its opponents and making them see the light, but rather because its opponents eventually die, and a new generation grows up that is familiar with it."

-Max Planck







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- What should you know before you download?
- What should you do before you start analysing your data?
- Where can you get help?
- What do you need for the publication / report?

What should you know **before** you download?

- support / history / usage
- requirements
- limitations
- type (e.g. freeware)
- in- and output file format
- last update / update history
- literature search
- what language is used

What should you do **before** you start analysing your data?

- avoid installation errors
- understand installation warnings
- run "simple" tests and/or examples
- explore limitations
- check in- and output format
- speed and resource managment

Where can you get **help**?

- manual(s) r RFM!
- help options (e.g. help button, -help, --h)
- README file(s)
- log file(s)
- user forum
- author(s)

What do you need for the **publication / report** ?

- literature reference or web link
- version
- parameters (better: command line)

Freeware (from "free" and "software") is computer software that is available for use at no cost or for an optional fee (donation).

The **GNU General Public License** (GNU GPL or simply GPL) is the most widely used free software license.

Commercial software, or less commonly, **payware**, is computer software that is produced for sale or that serves commercial purposes.

The term **shareware** (also known as trialware or demoware) refers to proprietary software that is provided to users without payment on a trial basis and is often limited by any combination of functionality, availability or convenience.

Input sequence(s) (or upload file):	?
Upload File	
Run Sequence length:	

Input sequence(s) (or upload file):	?
>Seq1	
ATGGC ATGCG	
>Seq2	
TGACA TTNNN	
Upload File	
Save Sequence length: 11,11	

OPEN CACCESS Freely available online

PLOS COMPUTATIONAL BIOLOGY

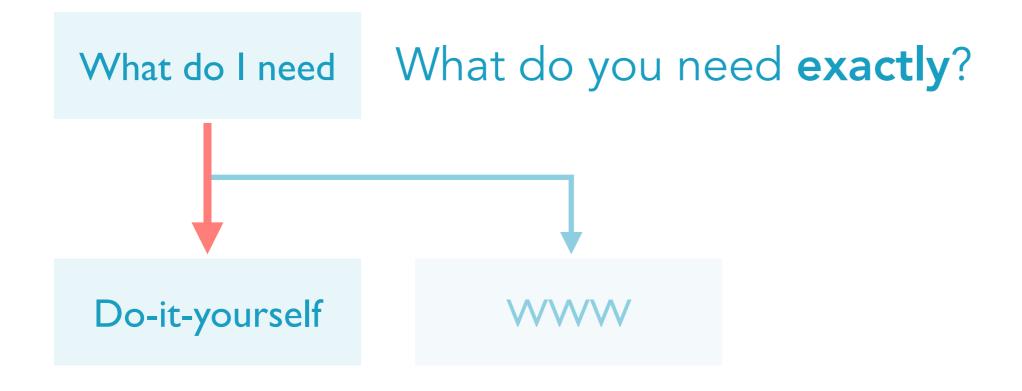
Editorial

Ten Simple Rules for Getting Help from Online Scientific Communities

Giovanni M. Dall'Olio¹*, Jacopo Marino², Michael Schubert³, Kevin L. Keys¹, Melanie I. Stefan⁴, Colin S. Gillespie⁵, Pierre Poulain^{6,7,8}, Khader Shameer^{9,10}, Robert Sugar³, Brandon M. Invergo¹, Lars J. Jensen¹¹, Jaume Bertranpetit¹, Hafid Laayouni¹

Netiquette

- Rule 1. Do Not Be Afraid to Ask a Question
- Rule 2. State the Question Clearly
- Rule 3. New to a Mailing List? Learn the Established Customs before Posting
- Rule 4. Do Not Ask What Has Already Been Answered
- Rule 5. Always Use a Good Title
- Rule 6. Do Your Homework before Posting
- Rule 7. Proofread your Post and Write in Correct English
- Rule 8. Be Courteous to Other Forum Members
- Rule 9. Remember That the Archive of Your Discussion Can Be Useful to Other People
- Rule 10. Give Back to the Community



PROGRAMMING / SCRIPTING IN BIOINFORMATICS

- **R**
- Python (BioPython)
- Perl (BioPerl)
- SQL
- C and C++ (Bio++)
- Ruby (BioRuby)
- PHP and JavaScript
- Java (BioJava)
- Go (BioGo)
- Shell Scripting

entry

Euclid's algorithm for the greatest

An **algorithm** is a **step-by-step procedure** for solving a problem or accomplishing some end especially by a computer. Procedure that produces the answer to a question or the solution to a problem in **a finite number of steps**. An algorithm that produces a yes or no answer is called a decision procedure; one that leads to a solution is a computation procedure. A mathematical formula and the instructions in a computer program are examples of algorithms.

Dynamic Programming is a method for solving complex problems by breaking it down into a collection of simpler subproblems. In order to solve a given problem using a dynamic programming approach, we need to solve different parts of the problem (subproblems), then combine the solutions of the subproblems to reach an overall solution.



Pseudocode is an informal high-level description of the operating principle of a computer program or other algorithm. It uses the structural conventions of a programming language, but is **intended for human reading** rather than machine reading.

1.. If student's grade is greater than or equal to 20

Print "passed"

else

Print "failed"

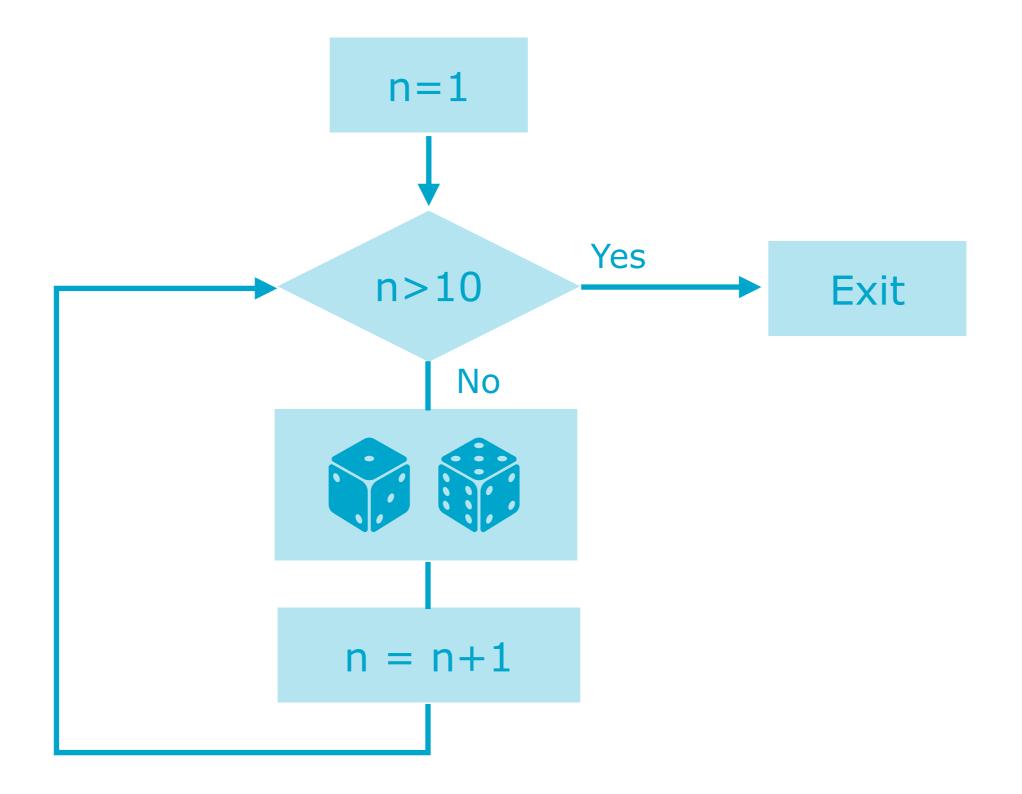
void function fizzbuzz	
<pre>for (i = 1; i<=100; i++) {</pre>	
set print_number to true;	
if i is divisible by 3	
print "Fizz";	
set print_number to false;	
if i is divisible by 5	
print "Buzz";	
set print_number to false;	
<pre>if print_number, print i;</pre>	
print a newline;	
}	

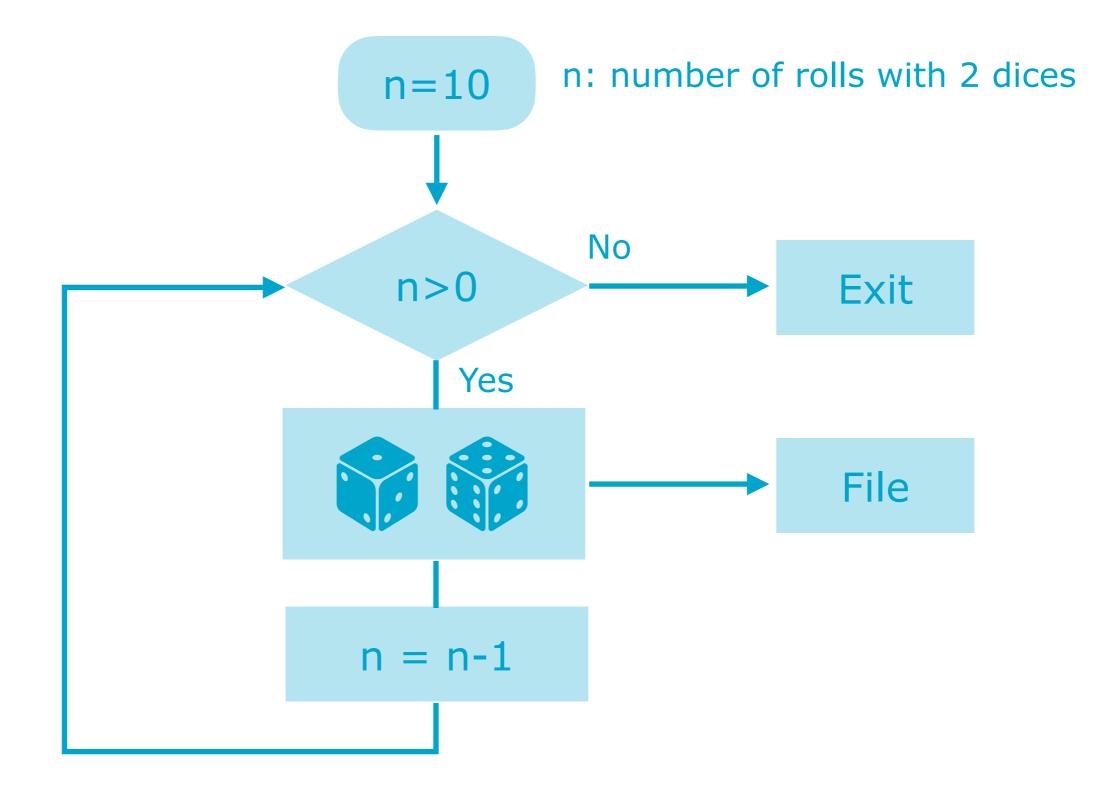
Write some pseudocode for a robot vacuum cleaner.

Example:

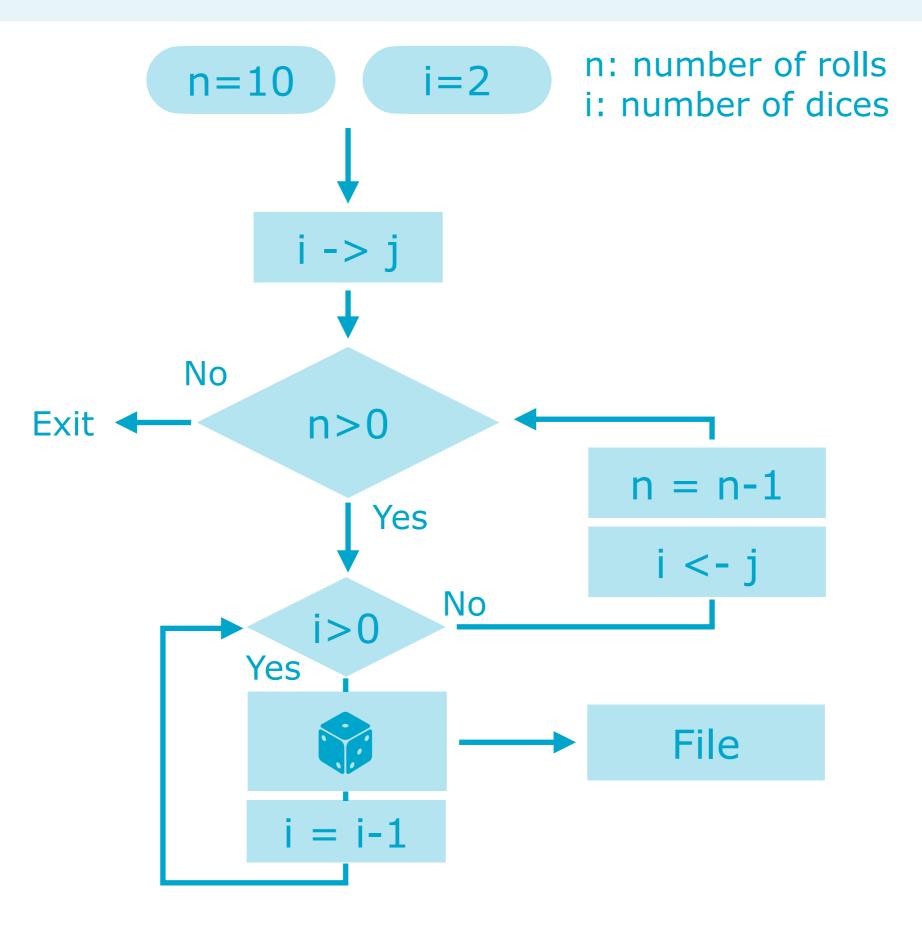
- 1. go straight until you hit something stop
- 2. turn around 45° start again
- 3. go to step 1







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Education

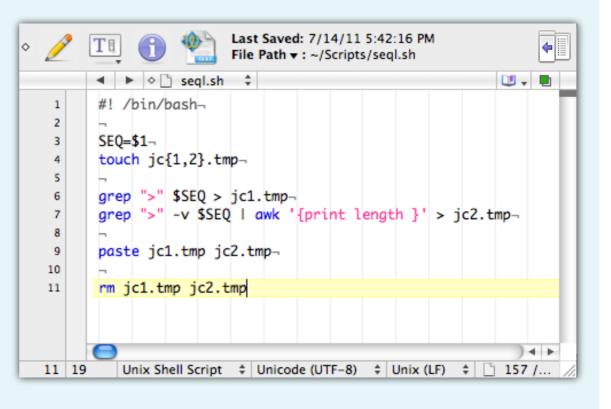
A Quick Guide to Organizing Computational Biology Projects

William Stafford Noble^{1,2}*

1 Department of Genome Sciences, School of Medicine, University of Washington, Seattle, Washington, United States of America, 2 Department of Computer Science and Engineering, University of Washington, Seattle, Washington, United States of America

Most bioinformatics coursework focuses on algorithms, with perhaps some components devoted to learning programming skills and learning how to use existing bioinformatics software. Unfortunately, for students who are preparing for a research career, this type of curriculum fails to address many of the day-to-day organisational challenges associated with performing computational experiments. In practice, the principles behind organising and documenting computational experiments are often learned on the fly, and this learning is strongly influenced by personal predilections as well as by chance interactions with collaborators or colleagues.

Specific help (e.g. complition) Syntax colouring (e.g. trouble shooting) Line numbers (e.g. debuging) Find and replace (e.g. RegEx) Charater Encoding (e.g. line breaks) Special Features (e.g. code folding)



>Most Fonts 1234567890 WWWWWWWWWW IIIIIIIII

AXAXAXAXAX

>Courier
1234567890
WWWWWWWWW
IIIIIIIII
AXAXAXAXAX

Specific help (e.g. complition)

Example: RStudio

> test

◆ testInheritedMethods◆ testVirtual	{methods} {methods}	<pre>testInheritedMethods(f, signatures, test = TRUE, virtual = FALSE, groupMethods = TRUE, where = .GlobalEnv)</pre>
<pre>testthat::</pre>		A set of distinct inherited signatures is generated to test inheritance for all the methods of a specified generic function. If method selection is ambiguous for some of these, a summary of the ambiguities is attached to the returned object. This test should be performed by package authors <i>before</i> releasing a package.
		Press F1 for additional help

Syntax colouring (e.g. trouble shooting)

Example: RStudio
<u>obj</u> ·%>%·¬
filter_taxa(grepl(pattern = "^[a-zA-Z]-(\$, taxon_names)) %>% # remove "odd" taxa-
filter_taxa(taxon_ranks == "o", supertaxa = TRUE) %>% # subset to the order rank
heat_tree(node_label = gsub(pattern = "\\[\\]", replacement = "", taxon_names),
<pre>node_size = n_obs,-</pre>
<pre>node_color = n_obs,-</pre>
node_color_axis_label = "OTU count",
layout = "davidson-harel", initial_layout = "reingold-tilford")

Syntax colouring (e.g. trouble shooting)

<u>obj</u>·%>%·-

- ···filter_taxa(grepl(pattern·=·"^[a-zA-Z]+(")·taxon_names))·%>%·#·remove·"odd"·taxa-
- · filter_taxa(taxon_ranks == "o", supertaxa = TRUE) %>% # subset to the order rank-
- heat_tree(node_label = gsub(pattern = "\\[|\\]", replacement = "", taxon_names),-
-node_size = n_obs,-
- node_color_axis_label = "OTU count",-
- layout = "davidson-harel", initial_layout = "reingold-tilford")¶

<u>obj</u>·%>%·-

filter_taxa(grepl(pattern = "^[a-zA-Z]+\$, taxon_names)) %>% # remove "odd" taxafilter_taxa(taxon_ranks == "o", supertaxa = TRUE) %>% # subset to the order rank heat_tree(node_label = gsub(pattern = "\\[[\\]", replacement = "", taxon_names), node_size = n_obs,node_color = n_obs,layout = "davidson-harel", initial_layout = "reingold-tilford") **Code Folding** - Code folding allows you to easily **show and hide blocks of code** to make it easier to navigate your source file and focus on the coding task at hand.

Example RStudio: To insert a new code section you can use the **Code > Insert Section** command. Alternatively, any comment line which includes at least four trailing dashes (-), equal signs (=), or pound signs (#) automatically creates a code section.

Code Folding in RStudio

```
1 ### ------
                                                        ###
 2 ### Ordination
                                                        ###
  ### ------
3
                                                    --- ###
 4
 5 - ## Libraries ----
 6
7 #install.packages("ggpubr")
8 library("ggpubr")
9
10 - ## Help ----
11
12 # DCA - detrended correspondence analysis using decorana
13 # CCA - correspondence analysis / constrained correspondence analysis (a.k.a. canonical correspondence analysis), via cca
14 # RDA - redundancy analysis, or optionally principal components analysis, via rda
15 # CAP - [Partial] constrained analysis of principal coordinates or distance-based RDA, via capscale
16 # DPCoA - double principle coordinate analysis using a (corrected, if necessary) phylogenetic/patristic distance between species.
17 # NMDS - non-metric multidimenstional scaling of a sample-wise ecological distance matrix onto a user-specified number of axes
18
  # MDS/PCoA - principal coordinate analysis (also called principle coordinate decomposition, multidimensional scaling (MDS), or class
19 #
20 # Syntax
21 # plot_ordination(phyloseq, ordinate(phyloseq, "DCA"), type="samples", color="index")
22
23 - ## Combined ordination plots ----
24 op1 <- plot_ordination(d, ordinate(d, "DCA"), type="samples", color="incubator")</pre>
   op2 <- plot_ordination(d, ordinate(d, "DCA"), type="biplot", shape="group", color="incubator")
25
26
27
   ggarrange(op1, op2
Z8
             labels = c("A", "B"),
29
             ncol = 1, nrow = 2)
30
```

Code Folding in RStudio

1	### ###
2	### Ordination ###
3	### ###
4	
5 ⊦	## Libraries 📾
10)	## Help G
23 -	## Combined ordination plots
24	<pre>op1 <- plot_ordination(d, ordinate(d, "DCA"), type="samples", color="incubator")</pre>
25	<pre>op2 <- plot_ordination(d, ordinate(d, "DCA"), type="biplot", shape="group", color="incubator")</pre>
26	
27	ggarrange(op1, op2
Z8	labels = c("A", "B"),
29	ncol = 1, nrow = 2

Built-in Help: Example for RStudio

- 1 MV<-get_manifests(Data,blocks)¬
- 2 check_MV<-test_manifest_scaling(MV,specs\$scaling)¬</pre>
- 3 gens<-get_generals(MV,path_matrix)¬</pre>
- 4 names(blocks)<-gens\$lvs_names¬</pre>
- 5 block_sizes<-lengths(blocks)-</pre>
- 6 blockinds<-indexify(blocks)-</pre>

Show syntax highlighting in console input



- 2 check_MV<-test_manifest_scaling(MV,specs\$scaling)</pre>
- 3 gens<-get_generals(MV,path_matrix)</pre>
- 4 names(blocks)<-gens\$lvs_names</pre>
- 5 block_sizes<-lengths(blocks)</pre>
- 6 blockinds<-indexify(blocks)</pre>

Reformat Code 企業A

1 MV <- get_manifests(Data, blocks)
2 check_MV <- test_manifest_scaling(MV, specs\$scaling)</pre>

- 3 gens <- get_generals(MV, path_matrix)
 4 names(blocks) <- gens\$lvs_names</pre>
- 5 block_sizes <- lengths(blocks)
- 6 blockinds <- indexify(blocks)</pre>

Preparing data and blocks indexification 3 -4 # building data matrix 'MV' 5 <- get_manifests(Data, blocks) 6 MV check_MV <- test_manifest_scaling(MV, specs\$scaling)</pre> 7 8 # generals about obs, mvs, lvs 9 gens <- get_generals(MV, path_matrix)</pre> 10 11 12 # indexing blocks 13 names(blocks) <- gens\$lvs_names</pre> 14 block_sizes <- lengths(blocks)</pre> 15 blockinds <- indexify(blocks)

Code Style



What is a **coding style**?

Language is a tool that allows people to interact and communicate with each other. The clearer we express ourselves, the better the idea is transferred from our minds to others. The same applies to programming languages: **concise**, **clear and consistent code is easier to read, more fun to work with and ultimately easier to reproduce**.

ggplot2isanrpackageforproducingstatisticalordatagraphicsbutitisunli kemostothergraphicspackagesbecauseithasadeepunderlyinggramm arthisgrammarbasedonthegrammarofgraphicswilkinson2005iscompo sedofasetofindependentcomponentsthatcanbecomposedinmanydiff erentwaysthismakesggplot2verypowerfulbecauseyouarenotlimitedto asetofprespecifiedgraphicsbutyoucancreatenewgraphicsthatarepreci selytailoredforyourproblemthismaysoundoverwhelmingbutbecauseth ereisasimplesetofcoreprinciplesandveryfewspecialcasesggplot2isalso easytolearnalthoughitmaytakealittletimetoforgetyourpreconceptions fromothergraphicstools

Source: Wickham (2009) ggplot2 - Elegant Graphics for Data Analysis

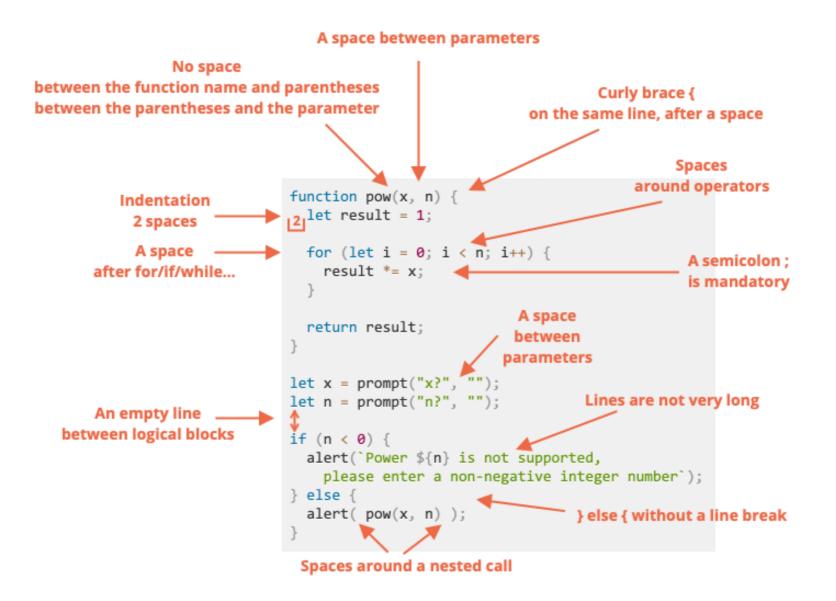
ggplot2 is an r package for producing statistical or data graphics but it is unlike most other graphics packages because it has a deep underlying grammar this grammar based on the grammar of graphics wilkinson 2005 is composed of a set of independent components that can be composed in many different ways this makes ggplot2 very powerful because you are not limited to a set of pre-specified graphics but you can create new graphics that are precisely tailored for your problem this may sound overwhelming but because there is a simple set of core principles and very few special cases ggplot2 is also easy to learn although it may take a little time to forget your preconceptions from other graphics tools

Source: Wickham (2009) ggplot2 - Elegant Graphics for Data Analysis

ggplot2 is an R package for producing statistical, or data, graphics, but it is unlike most other graphics packages because it has a deep underlying grammar. This grammar, based on the Grammar of Graphics (Wilkinson, 2005), is composed of a set of independent components that can be composed in many different ways. This makes ggplot2 very powerful, because you are not limited to a set of pre-specified graphics, but you can create new graphics that are precisely tailored for your problem. This may sound overwhelming, but because there is a simple set of core principles and very few special cases, ggplot2 is also easy to learn (although it may take a little time to forget your preconceptions from other graphics tools).

Source: Wickham (2009) ggplot2 - Elegant Graphics for Data Analysis

Code Style



javascript example / source: https://javascript.info/coding-style



Google's R Style Guide

tidyverse style guide examples

Object names

Good
day_one
day_1
Bad
DayOne
dayone

Parentheses

Good
<u>mean(x, na.rm = TRUE)</u>

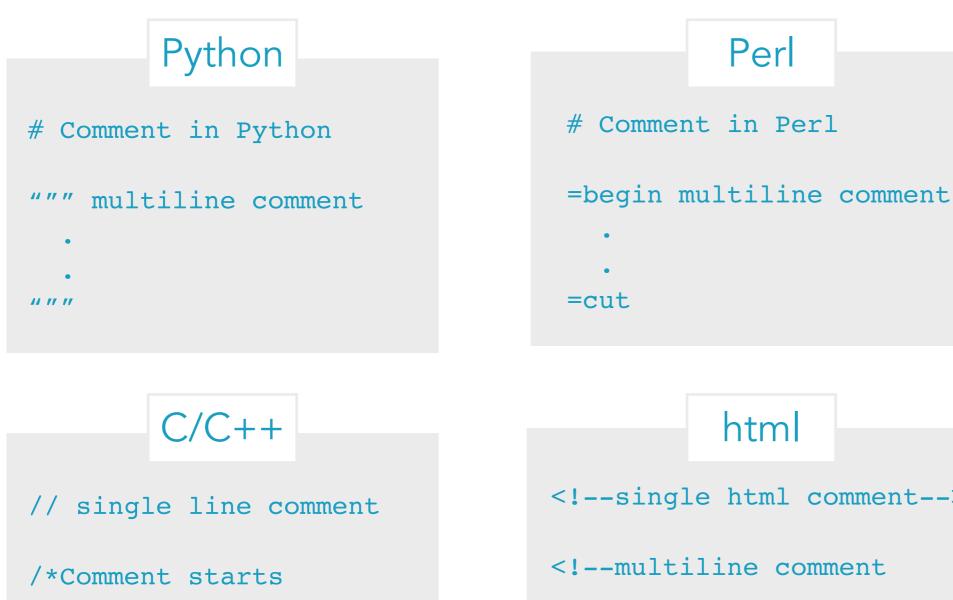
Bad

mean (x, na.rm = TRUE)
mean(x, na.rm = TRUE)

Infix operators

```
# Good
height <- (feet * 12) + inches
mean(x, na.rm = TRUE)
# Bad
height<-feet*12+inches
mean(x, na.rm=TRUE)
```

In computer programming, a comment is a programmer-readable explanation or annotation in the source code of a computer program. They are added with the purpose of making the source code easier for humans to understand, and are generally ignored by compilers and interpreters. The syntax of comments in various programming languages varies considerably.



•

Comment ends*/

Similar to Java

<!--single html comment--> //-->

Type of Comments

- Header / Titles
- Documentation
- Clarification

Headers are important structural elements.

Titles help to give the code structure.

Documentation comments are needed to explain your code.

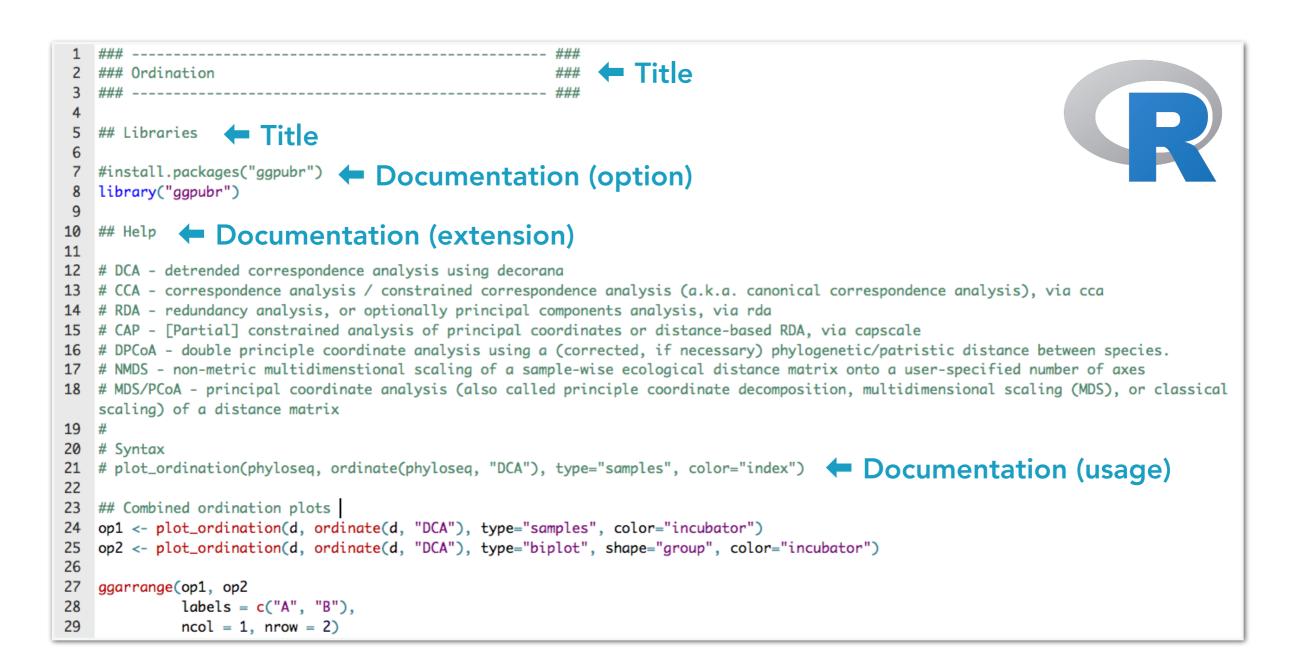
Clarification comments are intended for anyone (including your future self) who may need to maintain, refactor, or extend your code.

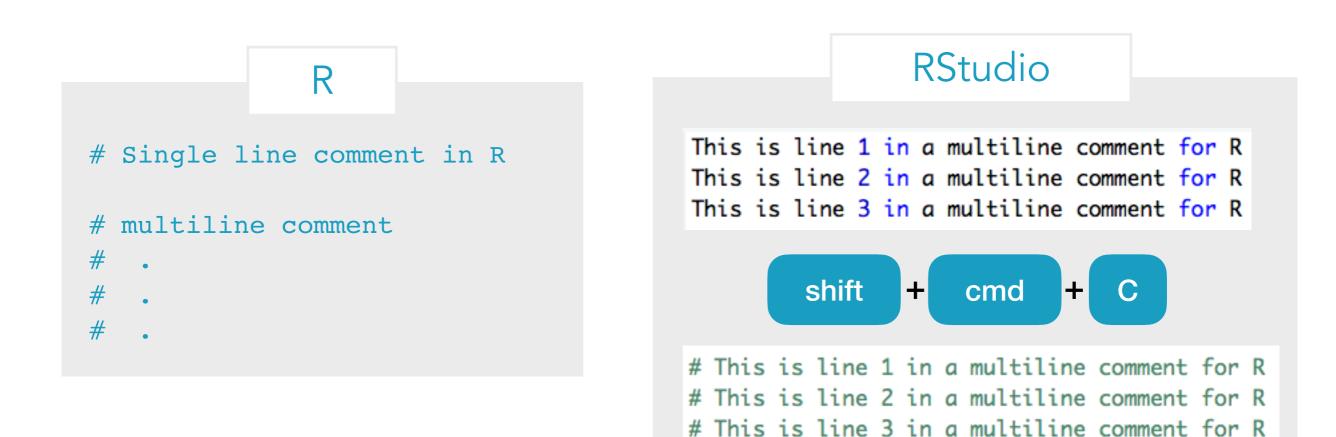
1	##•
2	##··P·R·O·J·E·C·T·-·H·E·L·P·-·F·I·L·E¬
3	## •
4	##·Project·:·Structural·variance·in·Brachypodium·distachyon-
5	##·Data····:Sequel·System·-·Pacific·Biosciences¬
6	## · Run · · · · : · Run 160517-
7	##·Date····: 22.05.17-
8	##•
9	
10	##
11	##·A· ·General·Help-
12	## A Generative the tp
13	
14	##·Project·Aim¬
15	# PacBio · Sequel · reads · should · be · mappen · to · the · reference · genome · of · Brachypodium · distachyon
16	#·(Version·2.4) ·and · structure · differences · identified.¬
17	
18	##·Working·directory¬
19	cd /home/project_Bdist/GV/~
20	##·Raw·data¬
21	
22	ll /home/project_Bdist/GV/data/raw/s231. <u>subreads</u> .bam
23	## •
24	
25	##·B· ·Quality·control-
26	## •
27	
28	## • QC • with • fastqp¬
29	fastqp -a s231 -n 100000 -k 7 count-duplicates data/raw/s231.subreads.bam
30	
31	## C Alienzest
32	##·C· ·Alignment¬
33	## •
34	The second state and state
35	##•Using•the•raw•BAM•and•align•to•reference-
36	pbalign s231.subreads.bam ref.fa s231.aligned.bam⊸
37	
38	## D L Eight Differences
39	##·D· ·Find·Differences-
40	## •
41	"" Field differences between oligende mede ood offerences
42	##·Find·difference·between·aligned·reads·and·reference¬
43	python jscreen.py -v -t 4 -r ref.fa -a ref.ggf3 -bam s231.aligned.bam -out s231_var.txt-
44	

1	##
2	##··P·R·O·J·E·C·T·-·H·E·L·P·-·F·I·L·E¬
3	##.====================================
4	##·Project·:·Structural·variance·in·Brachypodium·distachyon¬
5	##·Data····:·Sequel·System·-·Pacific·Biosciences¬
6	##·Run····:·Run160517¬
7	##·Date····:·22.05.17¬
8	## • ==================================
9	
10	##
11	##·A· ·General·Help-
12	##.
13	
14	##·Project·Aim¬
15	#·PacBio·Sequel·reads·should·be·mappen·to·the·reference·genome·of· <u>Brachypodium</u> ·distachyon¬
16	#•(Version•2.4)•and•structure•differences•identified.¬
17	
18	##·Working·directory¬
19	<pre>cd /home/project_Bdist/GV/¬</pre>
20	
21	##·Raw·data¬
22	ll /home/project_Bdist/GV/data/raw/s231.subreads.bam-
23	
24	## •
25	##·B· ·Quality·control¬
26	##
27	
28	##·QC·with·fastgp-
29	fastgp -a s231 -n 100000 -k 7 count-duplicates data/raw/s231.subreads.bam
30	
31	##
32	##·C· ·Alignment-
33	## •
34	
35	##•Using•the•raw•BAM•and•align•to•reference-
36	pbalign s231.subreads.bam ref.fa s231.aligned.bam
37	
38	## •
39	##·D· ·Find·Differences-
40	## •
41	
42	##·Find·difference·between·aligned·reads·and·reference¬
43	python jscreen.py -v -t 4 -r ref.fa -a ref.ggf3 -bam s231.aligned.bam -out s231_var.txt-
44	

Comments (**#**) Spaces → Structure

Code







	◄ ► (no symbol selected) ↓
3743 3744	<pre>push ax ;ilevel argument for RestoreDC</pre>
3745	; if (vfPrvwDisp)
3746	; InflateRect((LPRECT) rcwClip, 4, 4);
3747	; Assemble Note: coded inline because we're gods
3748	
3749	<pre>lea si, [rcwClipXpLeftRc]</pre>
3750	<pre>cmp [vfPrvwDisp],fFalse</pre>

Why not, if you must

F@!+& piece of R code - it drives me avocados

What about a more constructive way ...

#! I can't figure this out.
#! I need to extract only the records where
#! t < 12 but not > 18.

Good code is self-documenting.

Code tells you how, comments tell you why.

Good developers write good code; great ones also write good comments.



get-cpu-and-memory-usage_v190812_sz.sh.archive
get-cpu-and-memory-usage_v190823_nz.sh.archive
get-cpu-and-memory-usage_v190903_jcw.sh.archive
get-cpu-and-memory-usage_v200303_jcw.sh.archive
get-cpu-and-memory-usage_v200530_nz.sh.archive
get-cpu-and-memory-usage_v210117_jcw.sh

git --distributed-is-the-new-centralized

Git is a free and open source distributed version control system designed to handle everything from small to very large projects with speed and efficiency.

Git is easy to learn and has a tiny footprint with lightning fast performance. It outclasses SCM tools like Subversion, CVS, Perforce, and ClearCase with features like cheap local branching, convenient staging areas, and multiple workflows.



GitLab is a web-based Git repository manager with wiki and issue tracking features, using an open source license, developed by GitLab Inc.

