

# Evolutionary Genetics

LV 25600-01 | Lecture with exercises | 4KP

# Bioinformatics

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# REPRODUCIBLE RESEARCH

Recap

# Bioinformatics - Reproducible Research

**Repeatability** is a measure of the likelihood that, having produced one result from an experiment, you can try the same experiment, with the same setup, and produce that same result. It is a way for researchers to verify that their own results are true and are not just **chance artefacts**.

The **reproducibility** of data is a measure of whether a **different research team can attain results published in a paper using the same methods**. This shows that the results are **not artefacts of the unique setup in one research lab**. It is easy to see why reproducibility is desirable, as it reinforces findings and protects against rare cases of fraud, or less rare cases of human error, in the production of significant results.

**Replicability** - Different team, different experimental setup. If an observation is replicable it should be able to be made by a different team, using a different measuring system and dataset, in a different location, on multiple trials. This would therefore involve collecting data anew.

Source: <https://www.technologynetworks.com/informatics/articles/repeatability-vs-reproducibility>

## Material and Methods

In a first step, all paired-end raw reads were successfully merged using FLASH (version 1.2.9, Magoc and Salzberg 2011) with minimum overlap of 5nt and maximal mismatch ration of 0.8.

## Supplementary Data

```
## (a) Merging overlapping paired-end reads
# -v Version (1.2.9)
# -m minimum overlap (default 10bp)
# -x max mismatch ration (default 0.25)

flash -m 5 -x 0.8 random_1000_R1.fq random_1000_R2.fq -o
merged | tee flash.log
```



CODE / SCRIPTS

Recap

## A Quick Recap

### Code Style

Learn and use a common style.



#

Comment your code and do it generously.

### Code Editor

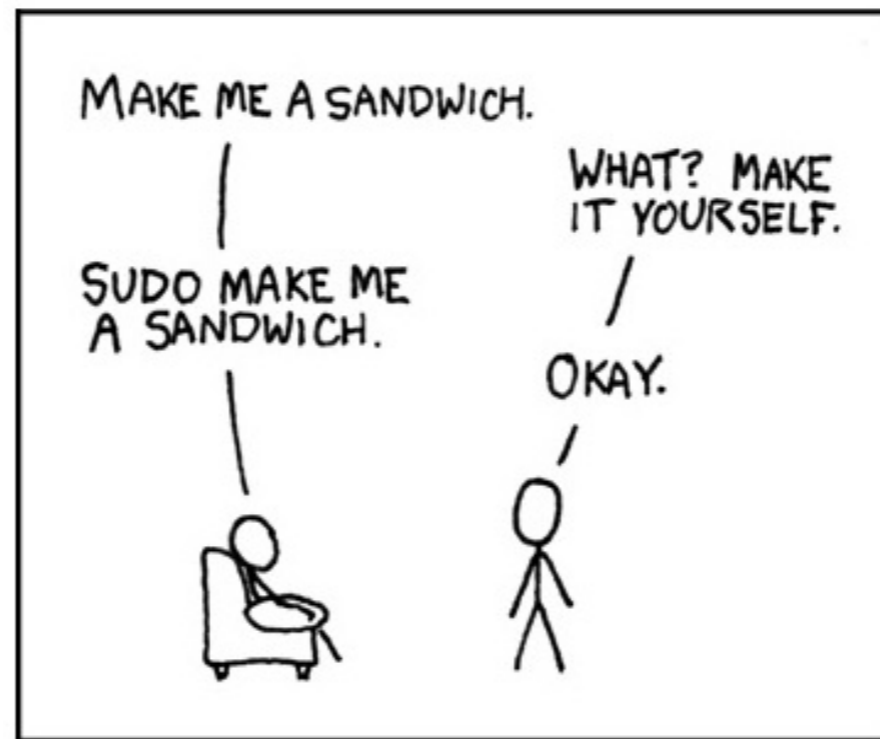
Make use of the different features like syntax highlighting, code folding or auto-completion.



### Version Control

For bigger or collaborative projects use version control and /or cloud based solutions.

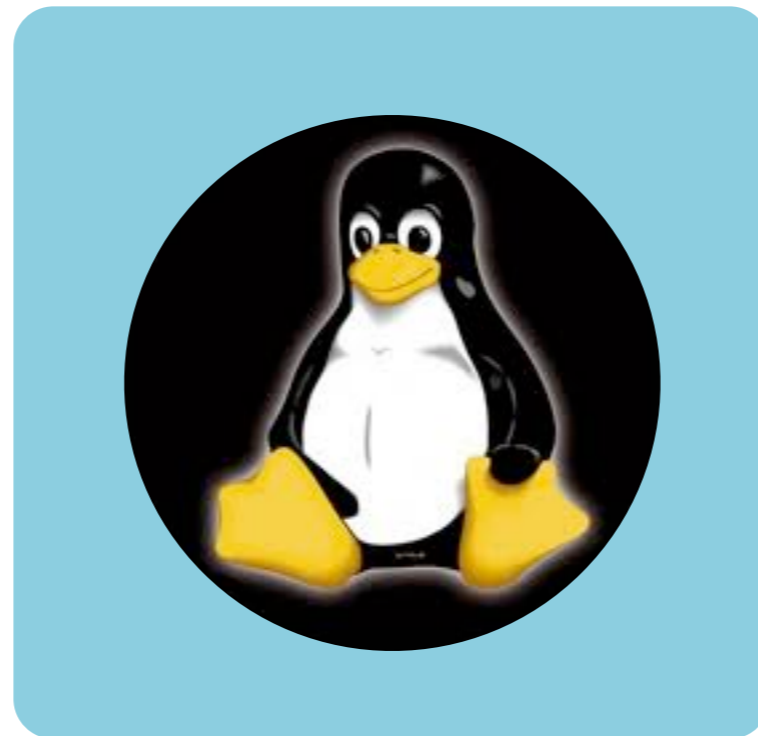
# TERMINAL



## The war of the OS and the conflict of the V



Mac

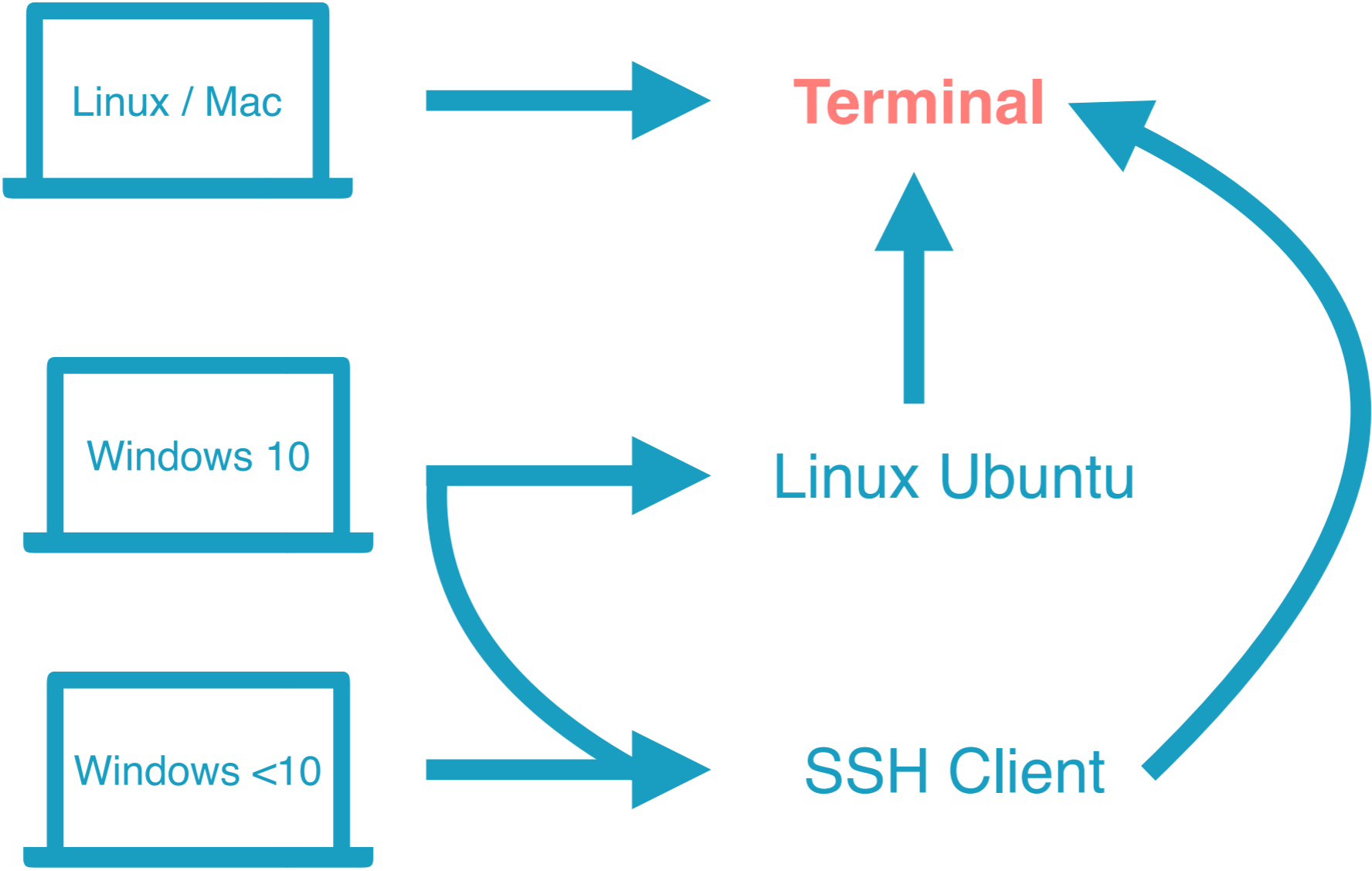


Linux



Windows

# Bioinformatics - Digital Safety



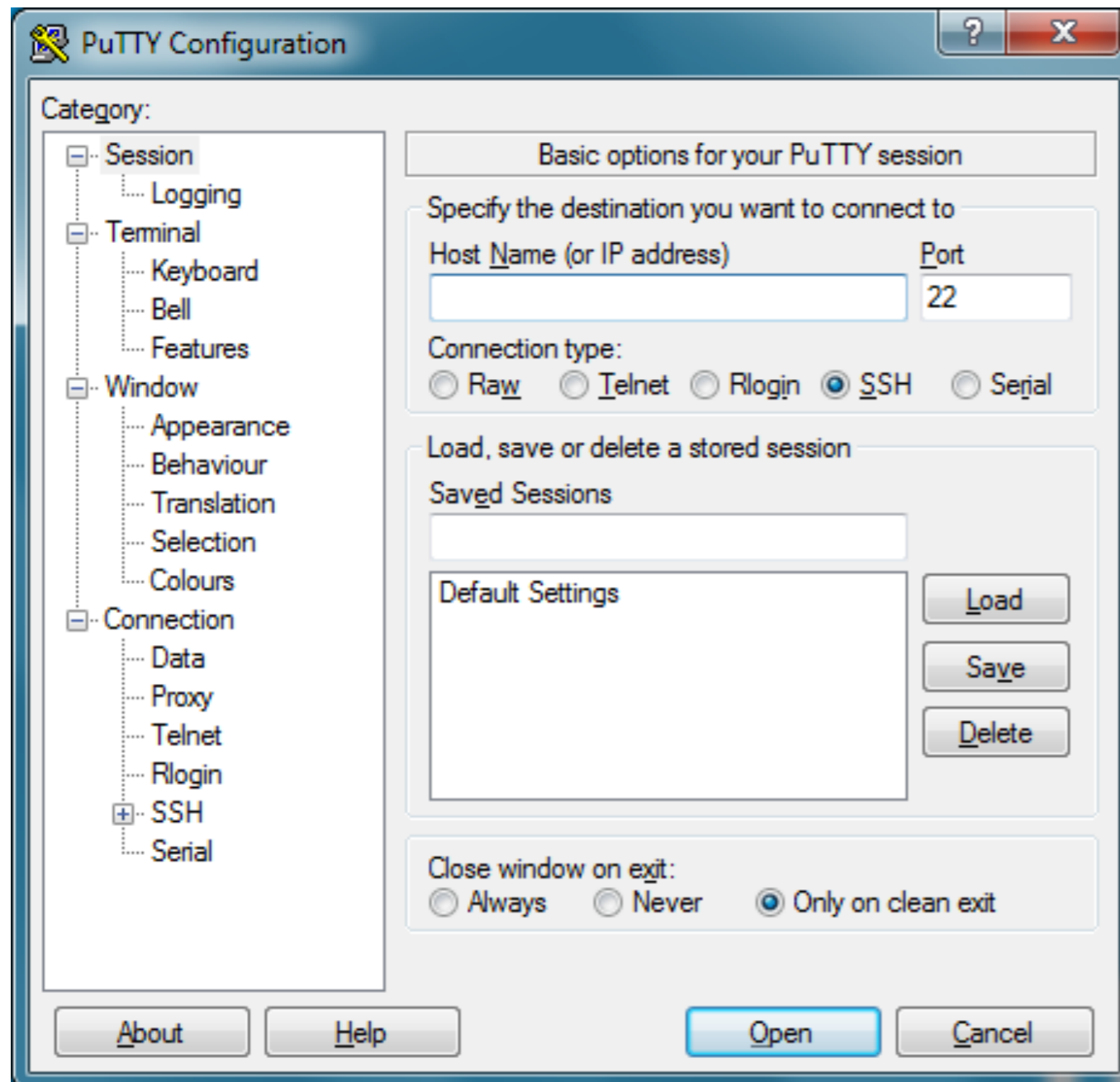
## Putty



PuTTY is an SSH and telnet client, developed originally by Simon Tatham for the Windows platform. PuTTY is open source software that is available with source code and is developed and supported by a group of volunteers.

<http://www.putty.org/>

# Bioinformatics - Introduction



Host Name:  
**student0X@gdcsrv2.ethz.ch**

Note: Change X accordingly.

# Bioinformatics - Introduction

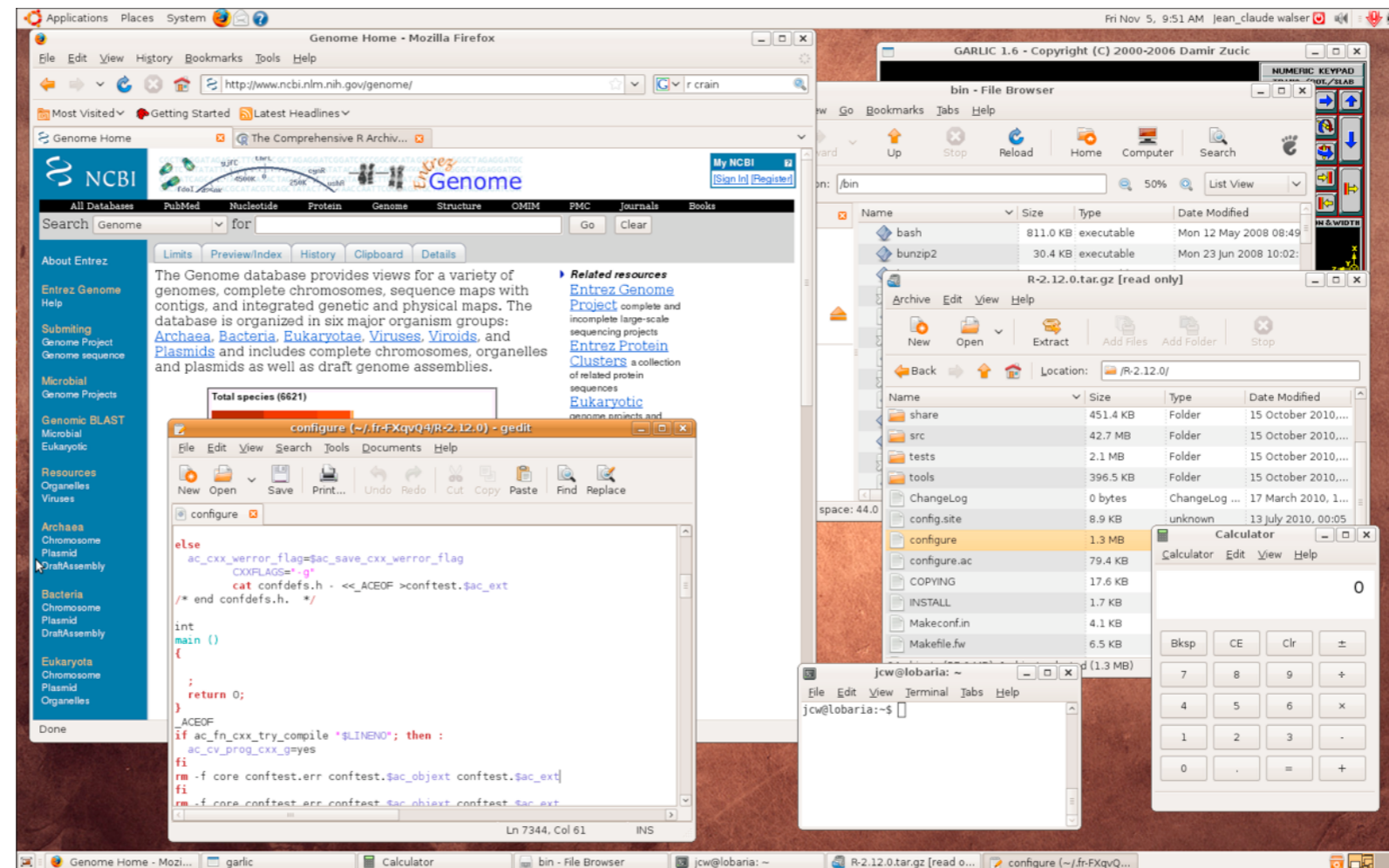


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# Bioinformatics - Terminal

A **graphical user interface (GUI)** - often pronounced goeey - an interface that allows the user (you) to interact with programs in more ways than typing.



GUIs are nice but have limitations. It is also difficult to describe what you exactly did and therefore it is difficult to reproduce.

GUIs were introduced in reaction to the steep learning curve of **command-line interfaces (CLI)**, which require commands to be typed on the keyboard. Since the commands available in command line interfaces can be numerous, complicated operations can be completed using a short sequence of words and symbols. This allows for greater efficiency and productivity once many commands are learned.



UNIX

Where there is a shell, there is a way.

## Shell - Terminal



Shell is a UNIX term for the interactive user interface with an operating system. The shell is the layer of programming that understands and executes the commands a user enters.

Bourne-Shell (sh)

Korn-Shell (ksh)

C-Shell (csh)

TC-Shell (tcsh)

**Bourne-Again-Shell (bash)**

**Debian Almquist Shell (dash)**

Z-Shell (zsh)

A-Shell (ash)

PowerShell / cmd.exe

What do I have?  
\$> echo \${SHELL}

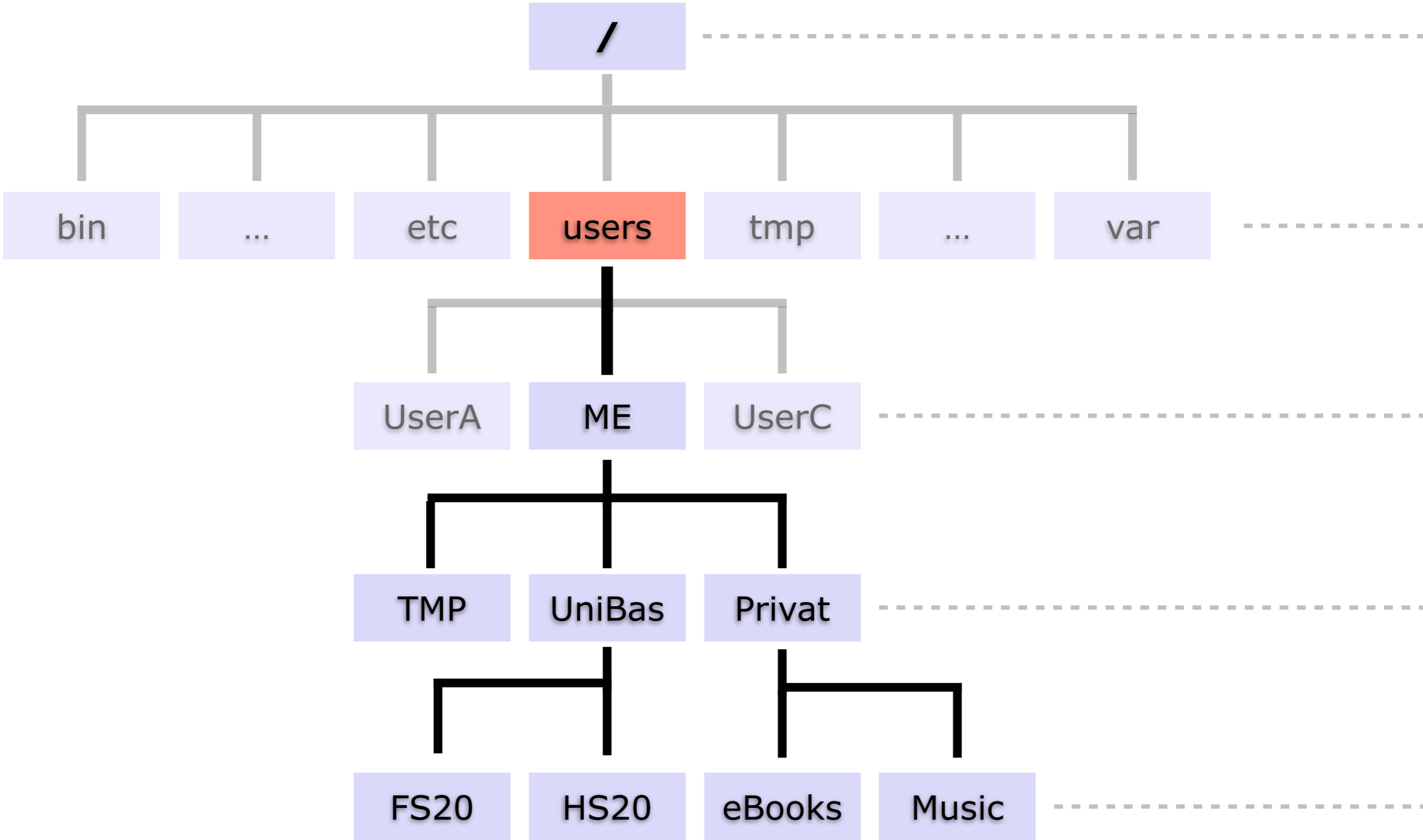
# Bioinformatics - Terminal

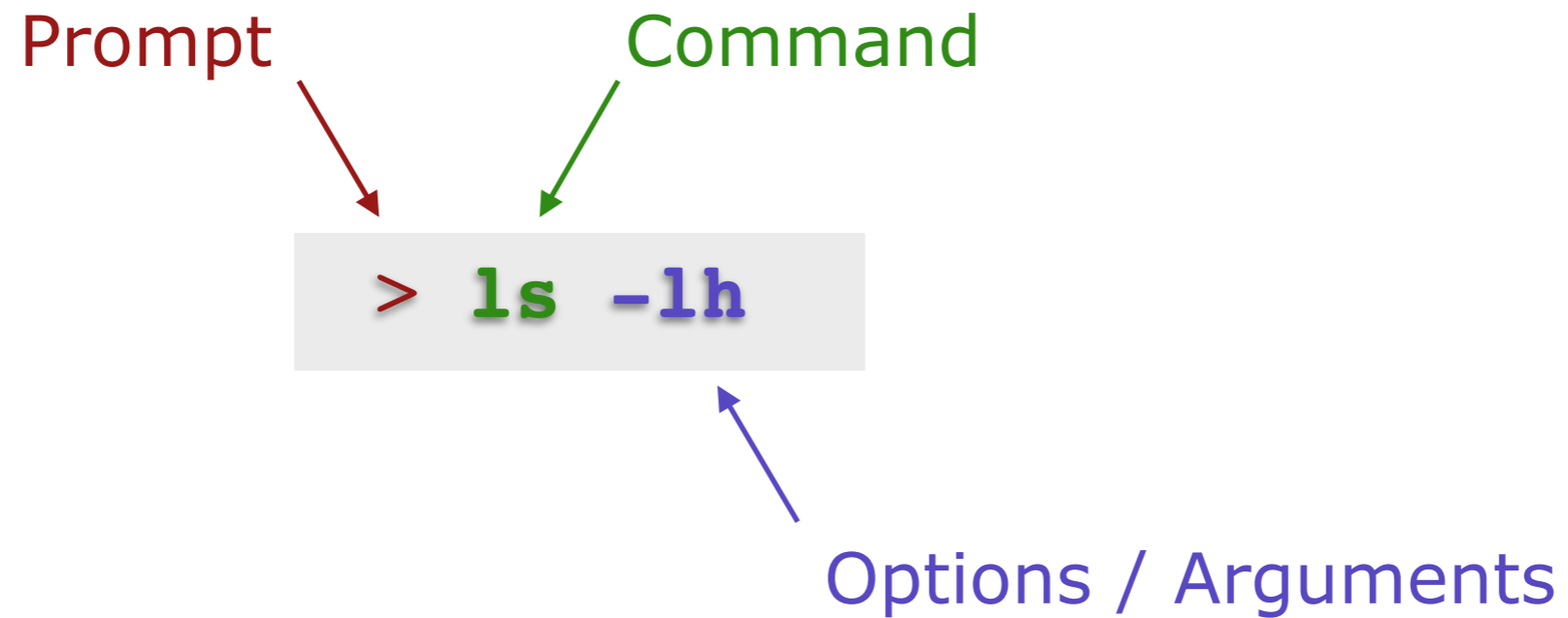


→ Open your (local) terminal

```
/Users/jean-claude — bash — 80x34
[jean-claude]: ll
total 17704
drwxr-xr-x@ 44 jean-claude  staff   1.5K Oct 29 09:40 .
drwxr-xr-x   7 root         admin   238B Mar 11 2010 ..
-rw-----   1 jean-claude  staff    3B Mar 29 2010 .CFUserTextEncoding
-rw-r--r--@   1 jean-claude  staff   15K Nov  5 08:17 .DS_Store
-rw-r--r--   1 jean-claude  staff   2.7K Aug  3 19:01 .RData
-rw-r--r--   1 jean-claude  staff   25K Nov  4 12:56 .Rhistory
drwx-----  11 jean-claude  staff   374B Nov  5 09:27 .Trash
-rw-----   1 jean-claude  staff    0B Nov 10 2009 .Xauthority
drwxr-xr-x   2 jean-claude  staff    68B Apr 20 2009 .Xcode
-rw-----   1 jean-claude  staff   24K Oct 28 17:10 .bash_history
drwx-----   3 jean-claude  staff   102B Aug 23 17:38 .bfglKingsLegacy
drwx-----   3 jean-claude  staff   102B Nov 10 2009 .config
drwx-----   3 jean-claude  staff   102B Apr 20 2009 .cups
drwxr-xr-x  21 jean-claude  staff   714B Nov 10 2009 .fontconfig
-rw-r--r--   1 jean-claude  staff   1.0K Jul  1 12:46 .jalview_properties
-rw-----   1 jean-claude  staff    35B Oct 28 17:10 .lessht
drwx-----   2 jean-claude  staff    68B Apr 20 2009 .macports
```

# Bioinformatics - Terminal





## Get Help

```
> info <command>
```

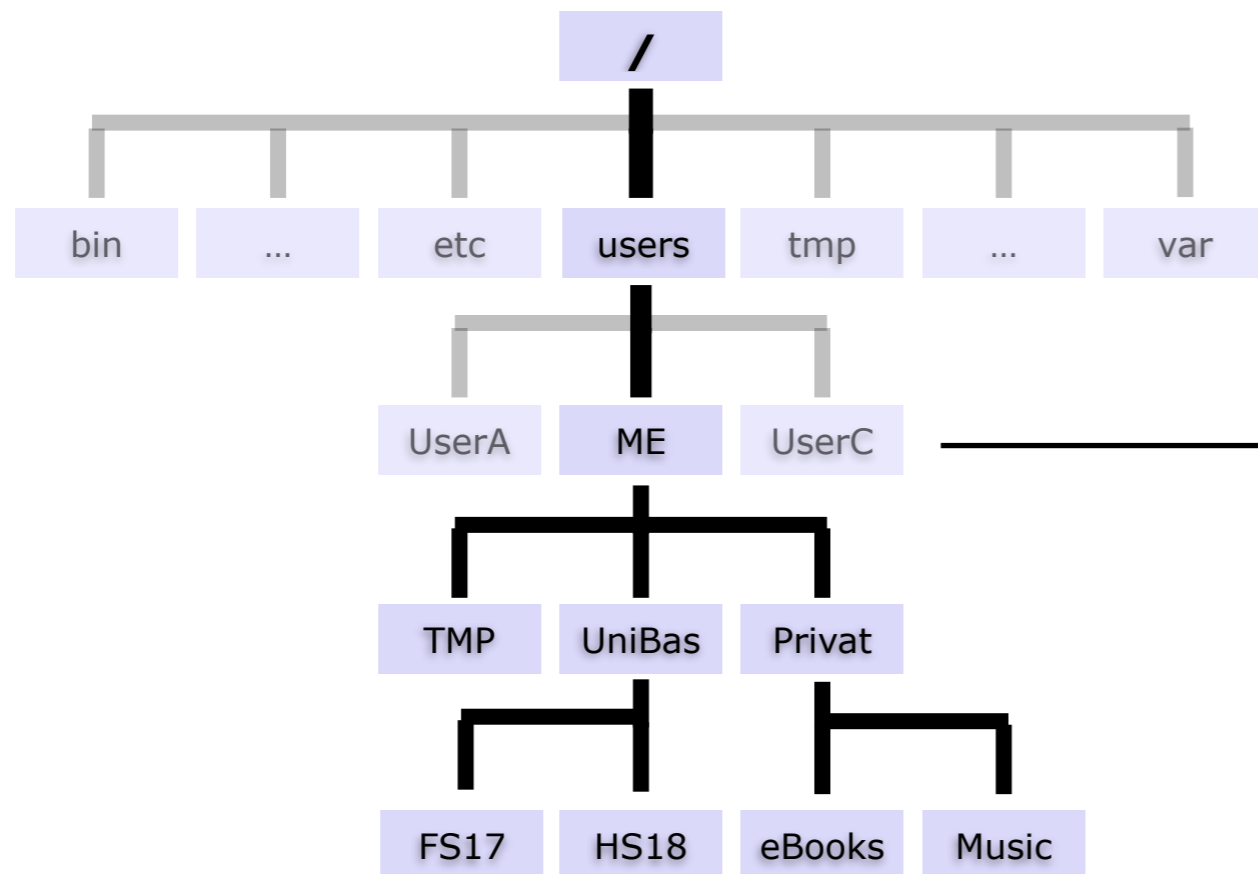
```
> info ls
```

```
> man <command>
```

```
> man ls
```

\* press [Q] key to leave info or help

`pwd` - print name of current/working directory



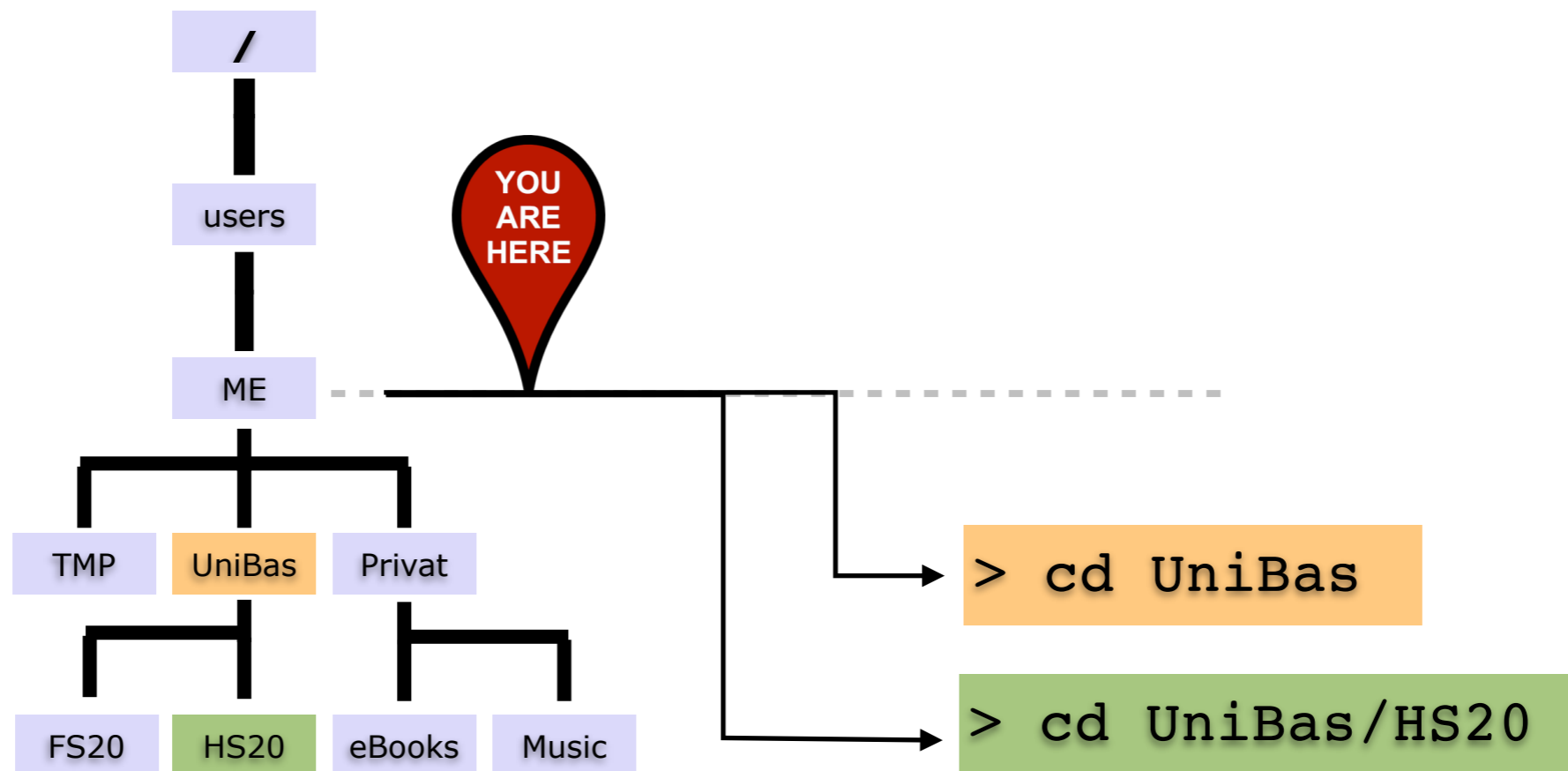
directory with users homes

```
> pwd  
/home/ME
```

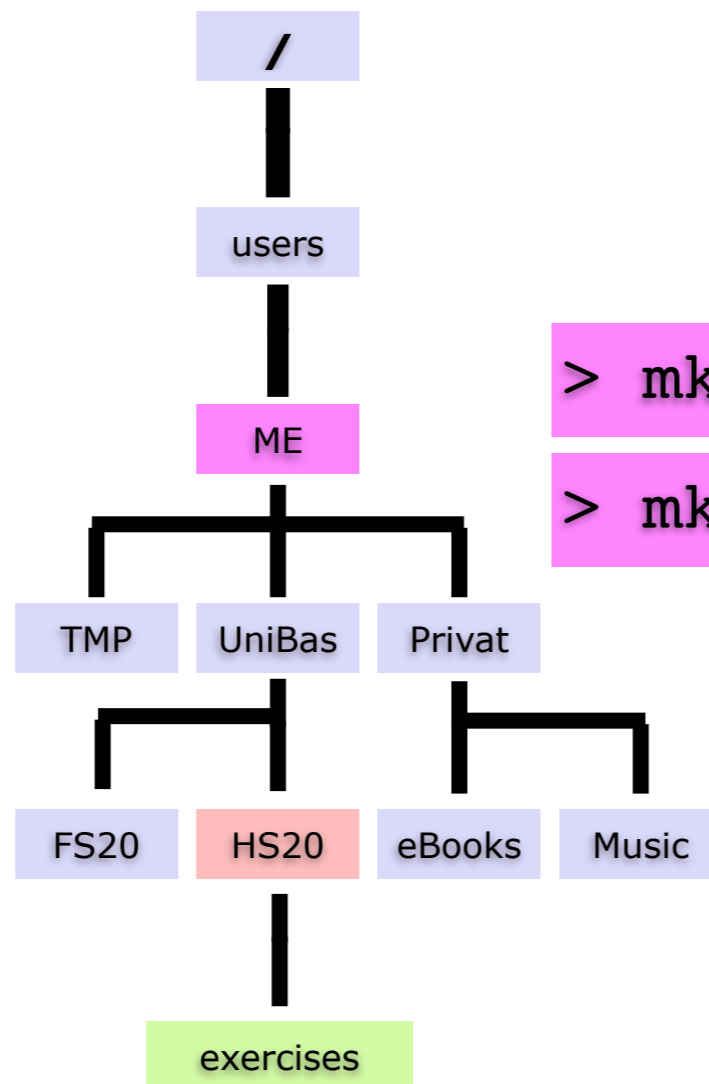
user homes (~)



## cd - change directory



## mkdir - creating directories

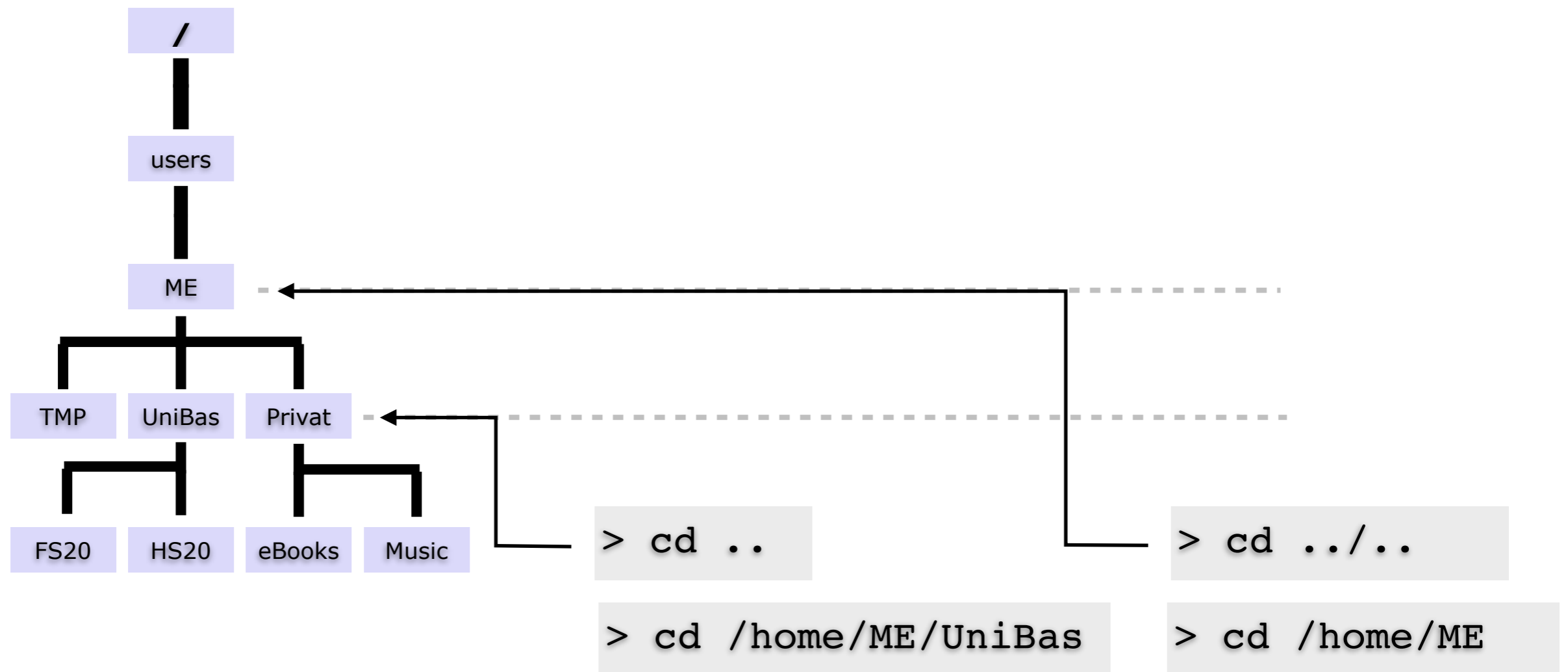


```
> mkdir ME/UniBas/HS20/exercises
```

```
> mkdir -p ME/UniBas/HS20/exercises
```

```
> mkdir exercises
```

`cd - change directory (going up)`

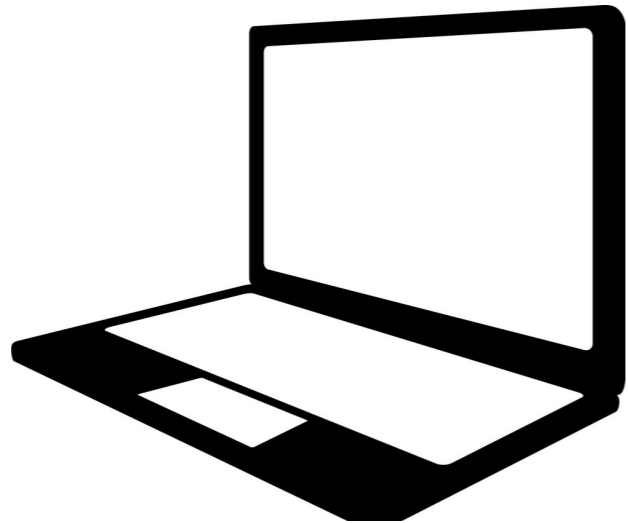


# Open Application

> open /Volumes/Mac\*/Applications/Firefox.app

# Open Text File with Editor

> edit \${HOME}/Documents/TMP/test.txt



Local  
Software  
Management

Software  
Dependencies

Parallel  
Versions

## Version Control



Application **A**

Requirments: Python 2.7

Application **B**

Requirments: Python >3.0



Package, **dependency and environment management** for any language—Python, R, Ruby, Lua, Scala, Java, JavaScript, C/ C++, FORTRAN. Conda as a package manager helps you find and install packages. If you need a package that requires a different version of Python, you do not need to switch to a different environment manager, because conda is also an environment manager. With just a few commands, you can set up a totally separate environment to run that different version of Python, while continuing to run your usual version of Python in your normal environment.

# BIOCONDA<sup>®</sup>

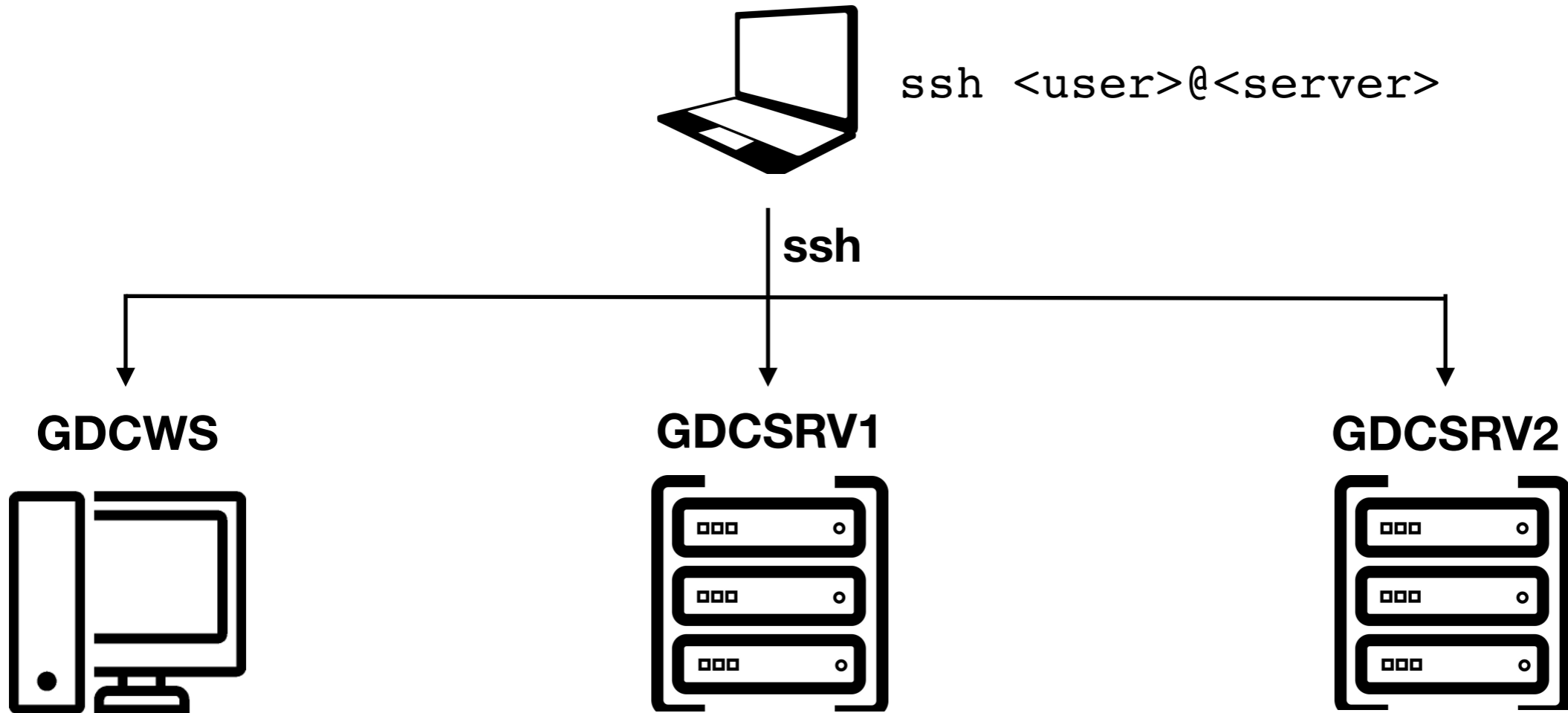
*Bioconda is a channel for the conda package manager specializing in bioinformatics software. The conda package manager makes installing software a vastly more streamlined process. Conda is a combination of other package managers you may have encountered, such as pip, CPAN, CRAN, Bioconductor, apt-get, and homebrew. Conda is both language- and OS-agnostic, and can be used to install C/C++, Fortran, Go, R, Python, Java etc programs on Linux, Mac OSX, and Windows.*



```
python --version
# Python 2.7.15
bwa
# -bash: bwa: command not found
blast -help
# -bash: blast: command not found

conda info --envs
source activate aligners
conda info --envs
python --version
# Python 3.6.7
bwa
blastn -help
```





```
lscpu; free -m
# Architecture: x86_64
# Model name: AMD Opteron(TM)
# CPU(s): 21
# Mem: 161'008k
# NUMA node0 CPU(s): 0-5
# NUMA node1 CPU(s): 6-11
# NUMA node2 CPU(s): 12-17
# NUMA node3 CPU(s): 18-23
```

```
lscpu; free -m
# Architecture: x86_64
# Model name: Intel(R) Xeon
# CPU(s): 160
# Mem: 926'346'512k
# NUMA node0 CPU(s): 0-9,80-89
# NUMA node1 CPU(s): 10-19,90-99
# NUMA node2 CPU(s): 20-29,100-109
# NUMA node3 CPU(s): 30-39,110-119
# NUMA node4 CPU(s): 40-49,120-129
# NUMA node5 CPU(s): 50-59,130-139
# NUMA node6 CPU(s): 60-69,140-149
# NUMA node7 CPU(s): 70-79,150-159
```

```
lscpu; free -m
# Architecture: x86_64
# Model name: Intel(R) Xeon
# CPU(s): 48
# Mem: 775'363'000k
# NUMA node0 CPU(s): 0-11
# NUMA node1 CPU(s): 12-23
# NUMA node2 CPU(s): 24-35
# NUMA node3 CPU(s): 36-47
```

## Local:

```
$ df -h
```

Filesystem	Size	Used	Avail	Capacity
/dev/disk1s1	<b>932Gi</b>	253Gi	676Gi	28%

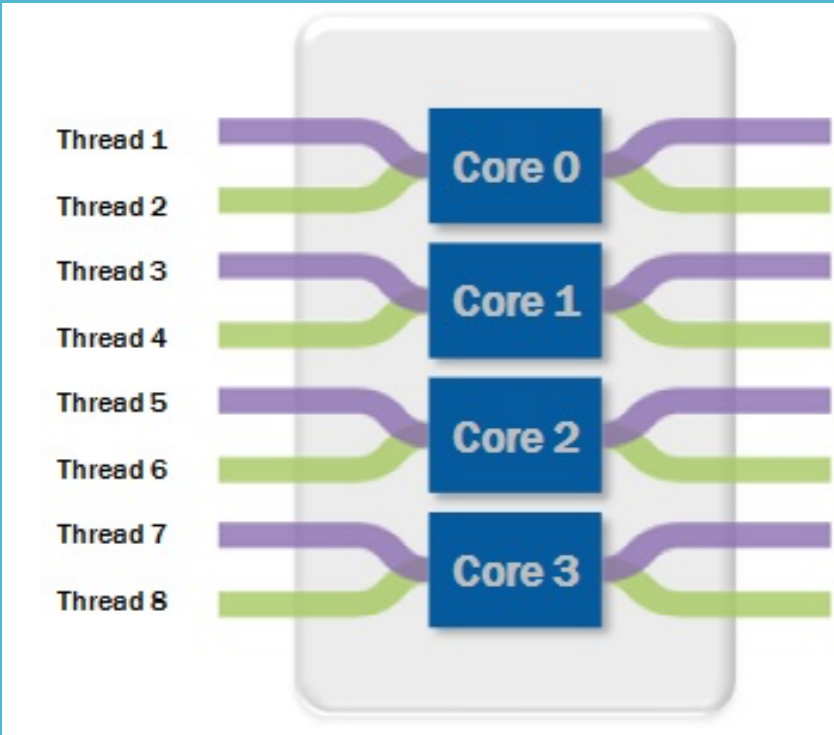
## Remote:

```
$ df -h
```

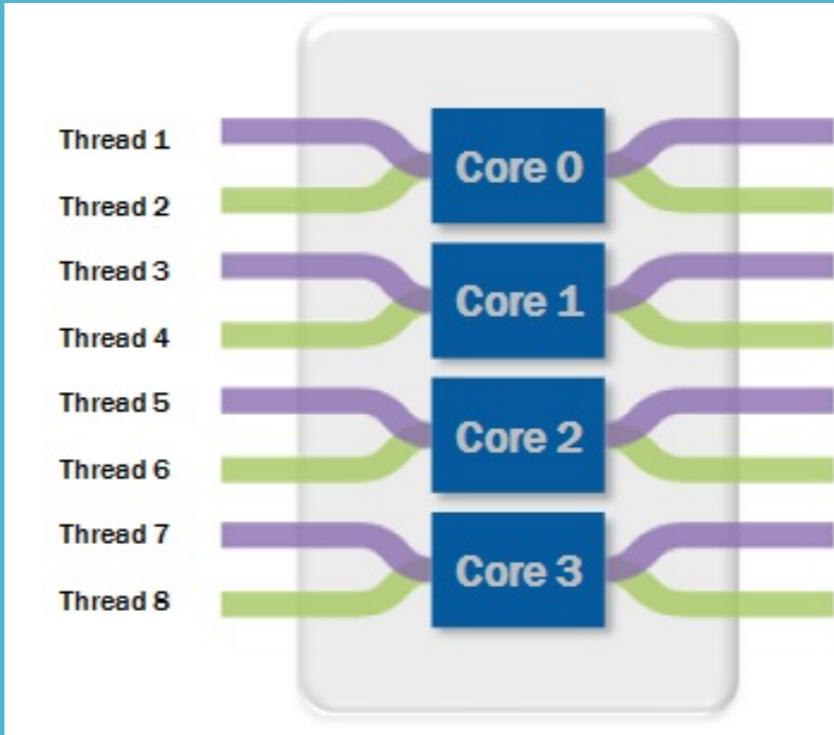
# Filesystem	Size	Used	Avail	Use%	Mounted on
# /dev/sdb	<b>5.3T</b>	2.7T	2.6T	51%	/data/local
# /data/gdc_home	<b>11T</b>	4.6T	6.1T	43%	/gdc_home
# /data2/gdc_home2	<b>22T</b>	7.4T	15T	34%	/gdc_home2
# /data3/gdc_home3	<b>28T</b>	5.2T	23T	19%	/gdc_home3
# /data4/gdc_home4	<b>37T</b>	25T	13T	67%	/gdc_home4
# /data5/gdc_home5	<b>50T</b>	27T	23T	55%	/gdc_home5

## Node

### CPU



### CPU



- **Compute node:** Currently most compute nodes have two sockets, each with a single CPU, volatile working memory (RAM), a hard drive, typically small, and only used to store temporary files, and a network card.
- **CPU:** Central Processing Unit, the chip that performs the actual computation in a compute node. A modern CPU is composed of numerous cores, typically 8 or 10. It has also several cache levels that help in data reuse.
- **Core:** part of a modern CPU. A core is capable of running processes, and has its own processing logic and floating point unit. Each core has its own level 1 and level 2 cache for data and instructions. Cores share last level cache.
- **Threads:** a process can perform multiple computations, i.e., program flows, concurrently. In scientific applications, threads typically process their own subset of data, or a subset of loop iterations.

```
top - 15:31:08 up 198 days, 5:44, 11 users, load average: 2.28, 2.95, 2.74
Tasks: 4418 total, 3 running, 4414 sleeping, 1 stopped, 0 zombie
Cpu(s): 1.4%us, 0.1%sy, 0.0%ni, 98.5%id, 0.0%wa, 0.0%hi, 0.0%si, 0.0%st
Mem: 926346512k total, 915660416k used, 10686096k free, 1139248k buffers
Swap: 41943036k total, 3202716k used, 38740320k free, 858950540k cached
```

PID	USER	PR	NI	VIRT	RES	SHR	S	%CPU	%MEM	TIME+	COMMAND
101909	cstritt	20	0	145m	12m	2216	R	99.5	0.0	1670:03	python2
101912	cstritt	20	0	145m	12m	2216	R	99.1	0.0	1670:03	python2
47027	smrtanal	20	0	85.2g	1.8g	7224	S	39.5	0.2	77856:07	java
61899	jwalser	20	0	18404	4652	948	R	4.7	0.0	0:01.10	top
45866	smrtanal	20	0	65.9g	8.3g	8548	S	0.6	0.9	1050:29	java
525	root	20	0	0	0	0	S	0.3	0.0	1:33.86	ksoftirqd/130
649	root	20	0	0	0	0	S	0.3	0.0	49:55.41	events/6
683	root	20	0	0	0	0	S	0.3	0.0	123:26.83	events/40
8717	root	20	0	0	0	0	S	0.3	0.0	525:50.39	kondemand/41
8826	root	20	0	0	0	0	S	0.3	0.0	389:55.32	kondemand/150
61910	root	20	0	98.4m	3908	2944	S	0.3	0.0	0:00.02	sshd
1	root	20	0	19368	1136	916	S	0.0	0.0	66:56.88	init
2	root	20	0	0	0	0	S	0.0	0.0	0:18.69	kthreadd
3	root	RT	0	0	0	0	S	0.0	0.0	3613:39	migration/0
4	root	20	0	0	0	0	S	0.0	0.0	3:51.22	ksoftirqd/0
5	root	RT	0	0	0	0	S	0.0	0.0	0:00.00	stopper/0
6	root	RT	0	0	0	0	S	0.0	0.0	128:44.45	watchdog/0
7	root	RT	0	0	0	0	S	0.0	0.0	2585:53	migration/1
8	root	RT	0	0	0	0	S	0.0	0.0	0:00.00	stopper/1
9	root	20	0	0	0	0	S	0.0	0.0	2:22.84	ksoftirqd/1
10	root	RT	0	0	0	0	S	0.0	0.0	105:29.73	watchdog/1
11	root	RT	0	0	0	0	S	0.0	0.0	2263:28	migration/2
12	root	RT	0	0	0	0	S	0.0	0.0	0:00.00	stopper/2

## CPU state percentages

us: user

sy: system

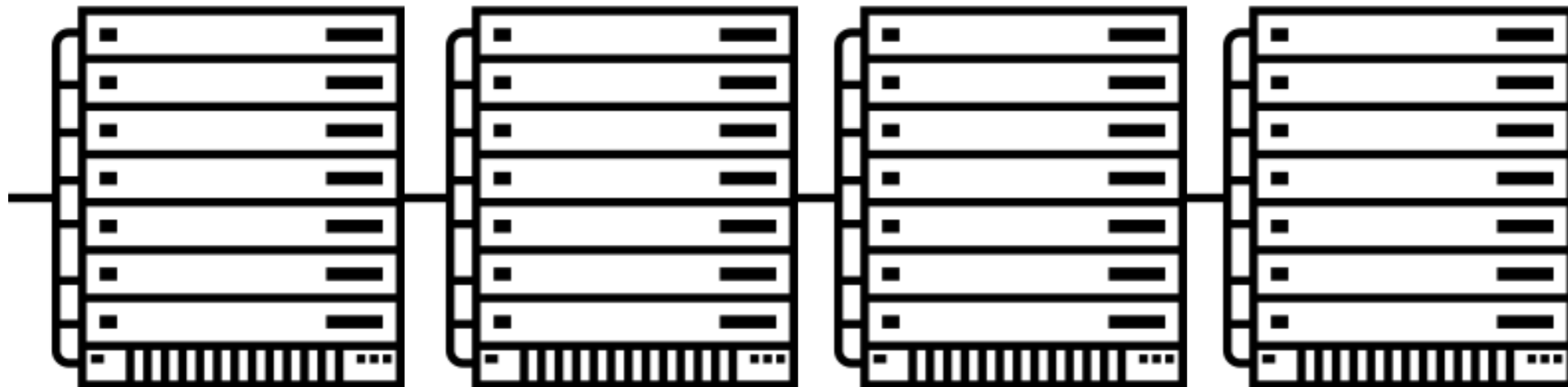
ni: nice

wa: IO-wait

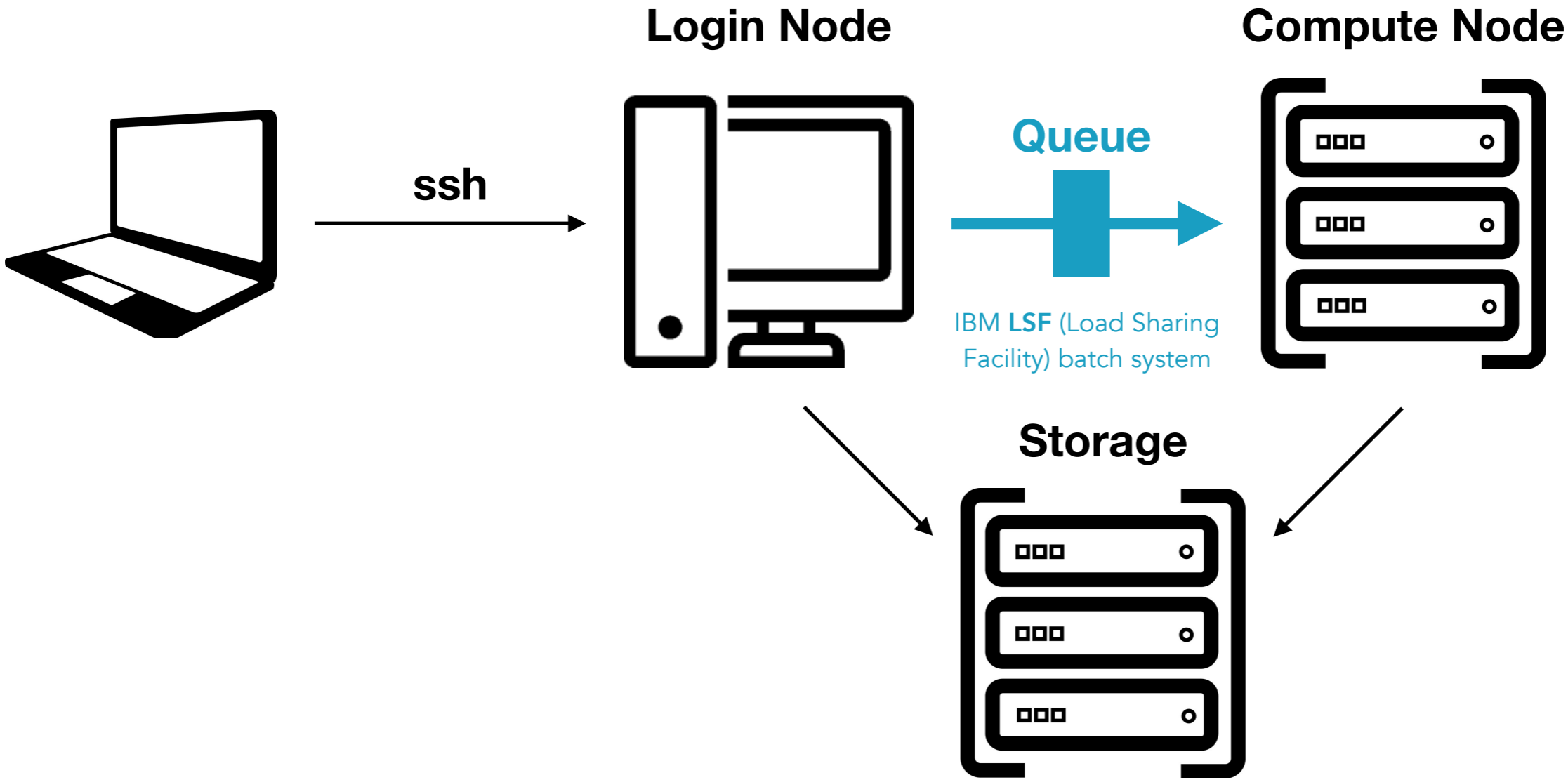
hi: hardware interrupts

si: software interrupts

## High Performance Cluster







- **HPC cluster:** relatively tightly coupled collection of compute nodes. Access to the cluster is provided through a login node. A resource manager and scheduler provide the logic to schedule jobs efficiently on the cluster.
- **Compute node:** an individual computer, part of an HPC cluster. Currently most compute nodes have two sockets, each with a single CPU, volatile working memory (RAM), a hard drive, typically small, and only used to store temporary files, and a network card.
- **CPU:** Central Processing Unit, the chip that performs the actual computation in a compute node. A modern CPU is composed of numerous cores, typically 8 or 10. It has also several cache levels that help in data reuse.
- **Core:** part of a modern CPU. A core is capable of running processes, and has its own processing logic and floating point unit. Each core has its own level 1 and level 2 cache for data and instructions. Cores share last level cache.
- **Threads:** a process can perform multiple computations, i.e., program flows, concurrently. In scientific applications, threads typically process their own subset of data, or a subset of loop iterations.

# Bioinformatics - SSH





## Euler II

Euler II contains **768** compute nodes of a newer generation — [BL460c Gen9](#) —, each equipped with:

- Two 12-core [Intel Xeon E5-2680v3](#) processors (2.5-3.3 GHz)
- Between 64 and 512 GB of DDR4 memory clocked at 2133 MHz (32 × 512 GB; 32 × 256 GB; 32 × 128 GB; 672 × 64 GB)

Euler II also contains **4** very large memory nodes — Hewlett-Packard [DL580 Gen9](#) —, each equipped with:

- Four 16-core [Intel Xeon E7-8867v3](#) processors (2.5 GHz)
- **3072** GB of DDR4 memory clocked at 2133 MHz

## Euler III

Euler III contains **1215** compute nodes — [Hewlett-Packard m710x](#) —, each equipped with:

- A quad-core [Intel Xeon E3-1585Lv5](#) processor (3.0-3.7 GHz)
- 32 GB of DDR4 memory clocked at 2133 MHz
- A 256 GB [NVMe](#) flash drive

All these nodes are connected to the rest of the cluster via 10G/40G Ethernet.

## Euler IV

Euler IV contains **288** high-performance nodes — [Hewlett-Packard XL230k Gen10](#) —, each equipped with:

- Two **18-core** [Intel Xeon Gold 6150](#) processors (2.7-3.7 GHz)
- 192 GB of DDR4 memory clocked at 2666 MHz

All these nodes are connected together via a new 100Gb/s InfiniBand EDR network.

## Euler V

Euler V contains **352** compute nodes — Hewlett-Packard BL460c Gen10 —, each equipped with:

- Two **12-core** [Intel Xeon Gold 5118](#) processors (2.3 GHz nominal, 3.2 GHz peak)
- 96 GB of DDR4 memory clocked at 2400 MHz

## Basic job submission

```
bsub -W 2:00 -n number_of_procs -R "rusage[mem=2048,scratch=5000]" <command>  
<parameters>  
  
# -n request multiple cores (or threads)  
# -R mem default the batch system allocates 1024 MB (1 GB) of memory per  
processor core  
# -R scratch for temporary data
```

## Submission script

```
#!/bin/bash  
#BSUB -J "MyScript"          ## Job Title  
#BSUB -n 10                  ## Number of Cores  
#BSUB -R "rusage[mem=2048]"  ## Memory Request  
#BSUB -W 2:00                ## Running Time  
  
## Load environment  
module load gcc/4.8.2 gdc perl/5.18.4  
  
## ...
```

## Job monitoring













```
[leonhard@euler08 ~]$ bjobs 31989961
Job information
  Job ID           : 31989961
  Status          : RUNNING
  Running on node  : e1268
  User            : leonhard
  Queue          : normal.4h
  Command        : compute_pq.py
  Working directory : $HOME/testruns
Requested resources
  Requested cores   : 1
  Requested memory  : 1024 MB per core
  Requested scratch : not specified
  Dependency       : -
Job history
  Submitted at     : 08:45 2016-11-15
  Started at      : 08:48 2016-11-15
  Queue wait time  : 140 sec
Resource usage
  Updated at      : 08:48 2016-11-15
  Wall-clock      : 34 sec
  Tasks          : 4
  Total CPU time   : 5 sec
  CPU utilization  : 80.0 %
  Sys/Kernel time : 0.0 %
  Total resident memory : 2 MB
  Resident memory utilization : 0.2 %
```





## 2-minute warmup

BY DAREBEE © [darebee.com](http://darebee.com)  
Repeat each exercise for 10 seconds.

 hops on the spot	 side-to-side hops single leg	 hops on the spot	 side-to-side hops feet together
 alt back expansions	 chest expansions	 arm circles (wide)	 arm circles
 hops on the spot	 side-to-side hops single leg	 hip rotations	 torso rotations





# Bioinformatics - Example Questions

Q1

What is the outcome of the following terminal command?

```
head -n 2 seq.fa | grep "^>" -c
```

Q2

What does the following command line do?

```
grep ">" seq.fa | grep "Daphnia" | wc -l seq.fa
```

Q3

What does verbose mean and why would it be important to use?

```
curl -verbose -O https://server.ch/file.zip
```

Q4

With both commands, a new file called seq.fasta is created. What is the difference?

```
cp seq.fa seq.fasta  
mv seq.fa seq.fasta
```



**Q5**

What is the connection between comments and reproducibility?

**Q6**

What is missing in the description below to make it reproducible?

"USEARCH (Edgar 2017) with default parameters was used to quality filter the sequences?"

# Evolutionary Genetics

LV 25600-01 | Lecture with exercises | 4KP

## Extension

## Digital Safety

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**GOOD**

**SAFETY**

**HABITS**

You are being tracked by:

Cookies

Evercookies

IP address

Flash cookies

HTML 5 storage

Fingerprinting

and more ...

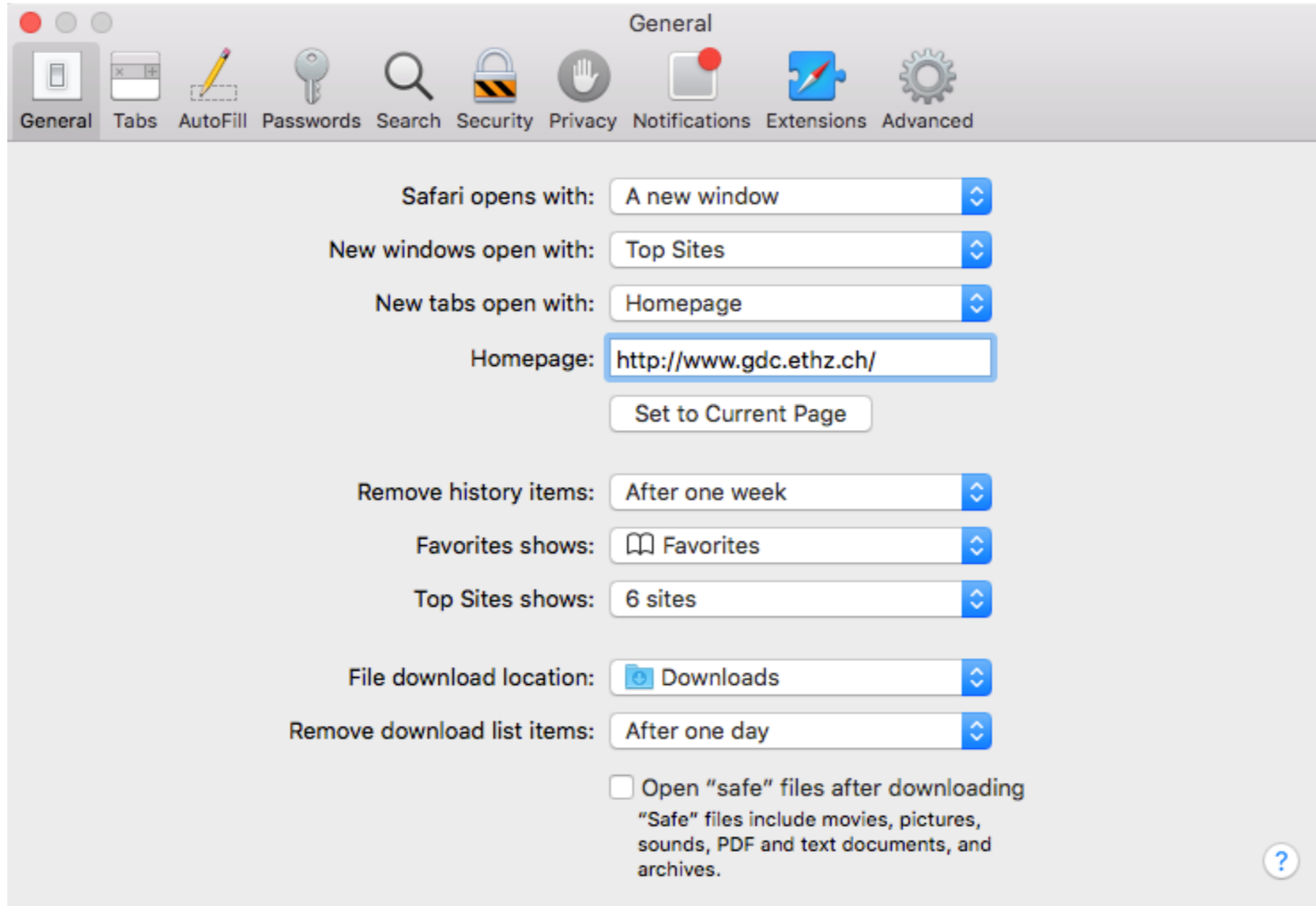
- ▶ Visiting the top 50 websites will install over 3,000 tracking files on your computer.
- ▶ Over 80% of websites use one or more tracking tools.
- ▶ The average number of tracking tools is six per website. Bigger websites can have up to 20 trackers.

## Webbrowser Safety



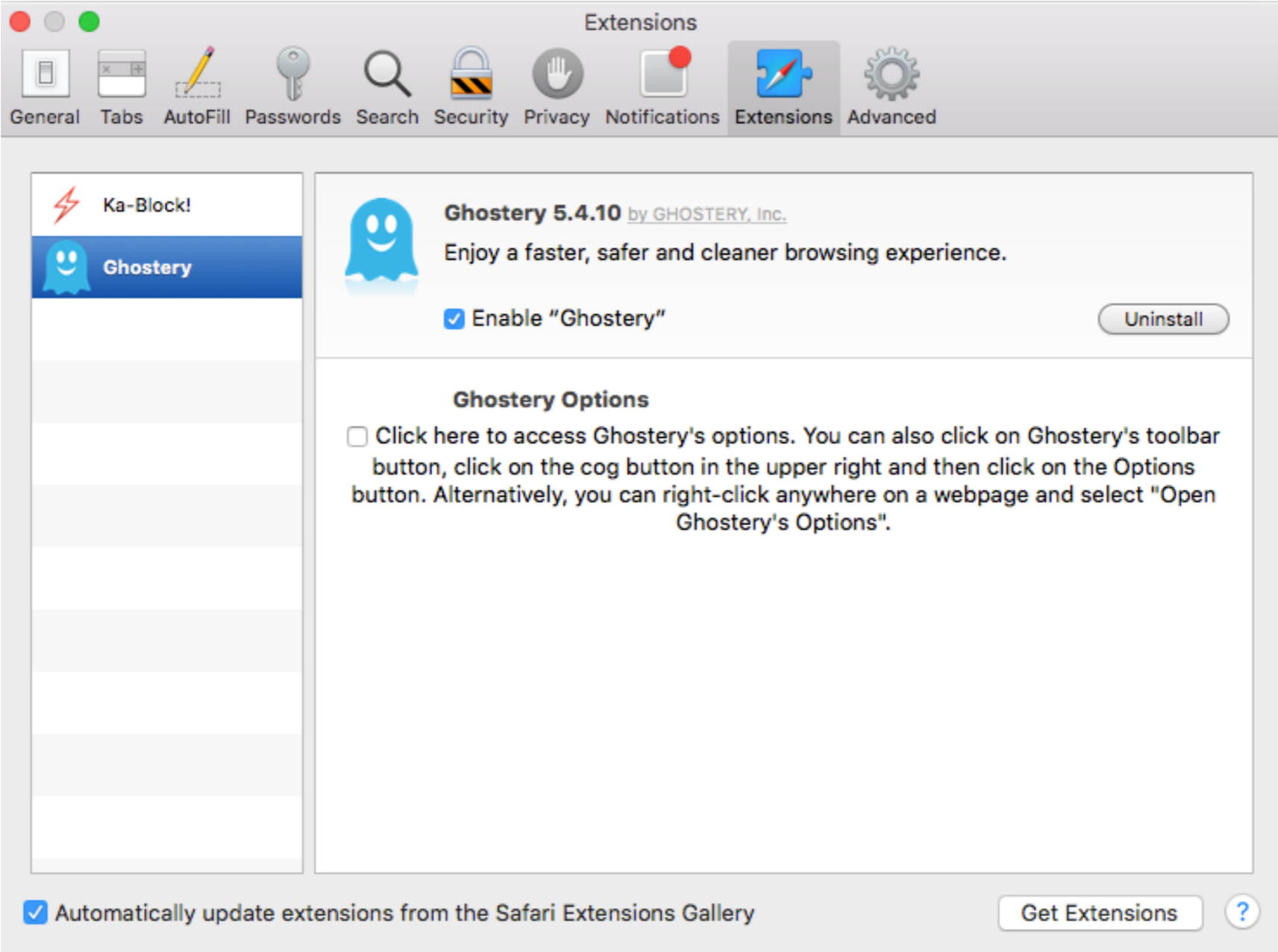
- Security settings
- Browser extensions
- VPN
- Incognito Mode

## Change / Check Settings





## Install Extensions



How to Enable **Private Browsing** on Any Web Browser



## Privacy Browser

- ▶ Epic Privacy Browser
- ▶ Comodo Dragon
- ▶ Brave
- ▶ Tor





VPN is a **virtual private network** that enables you to have a secure connection between your device and an Internet server that no one can monitor or access the data that you're exchanging. A VPN connection establishes a safe passageway through all the insecurities of public networks. You can benefit from a VPN connection for a number of reasons, including, security and privacy.

When you're connected to the Internet through a VPN connection, this private internet access ensures that you're not exposed to phishing, malware, viruses and other cyber threats. Your privacy is also guaranteed, as no one will be able to detect your online behaviour. Anyone can benefit from an added safety measure that the VPN connection feature provides.

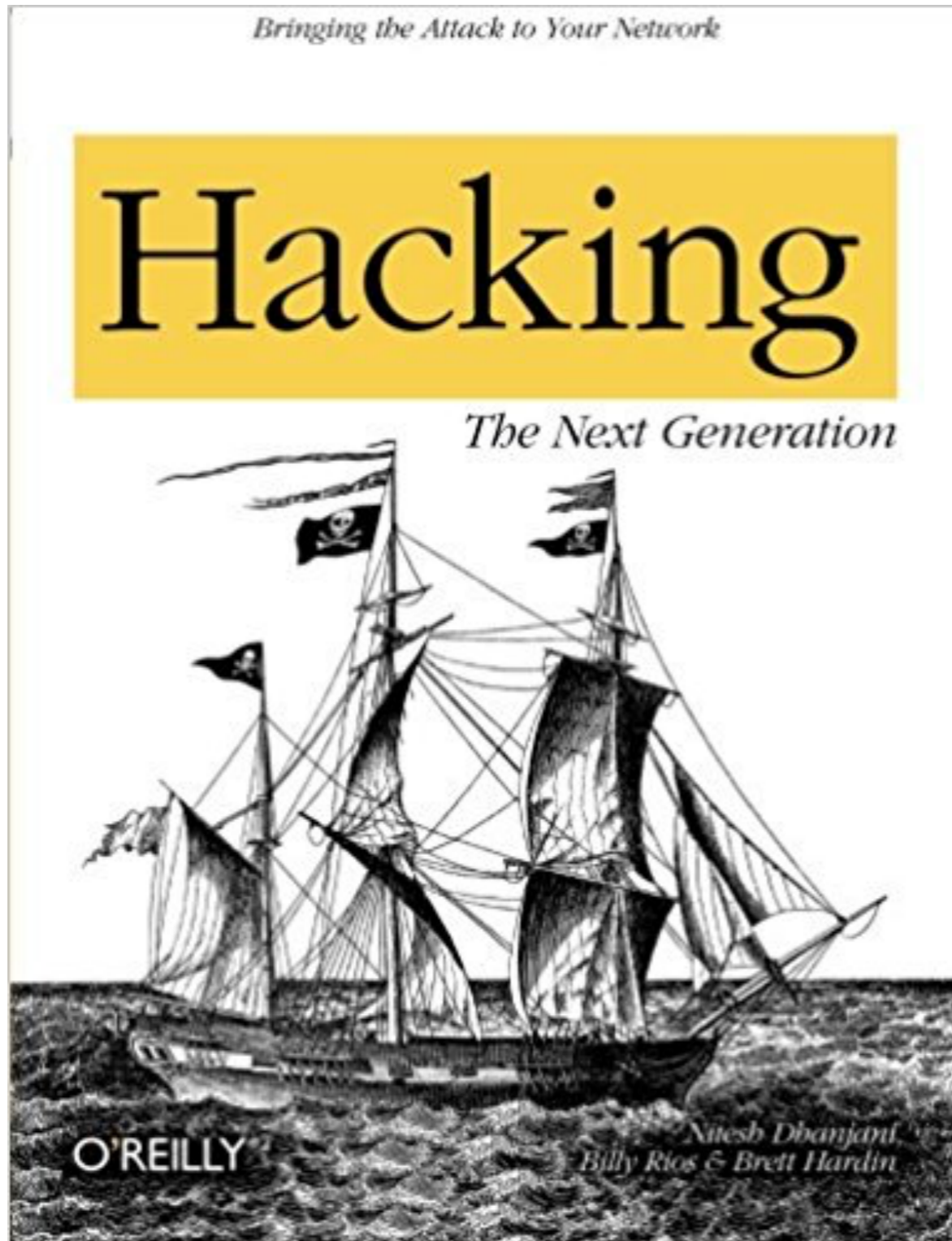
> <https://mobile.unibas.ch/home.html>







Metadata is data about data. It usually provides additional information. Metadata is essential for most data formats. You can't render a JPEG, send an email, or visit a web site without metadata. Metadata could be used for surveillance purposes, but it also has many common everyday purposes. **Metadata is not scary.**





## Outside In Clean Content

Outside In Clean Content addresses particularly challenging issues in native file processing. Focusing specifically on widely used formats (Microsoft Office and PDF), its extended extraction provides all text, properties, hidden information and system data emedded in native files. Its extended extraction includes the ability to analyze and process malformed documents, which is critical to accurate text extraction from PDFs. Clean Content can also programmatically modify native files enabling features such as scrubbing, property modification and document assembly. Outside In Clean Content is a pure Java technology that offers Java, C/C++ and .NET APIs.

- Extracts text, metadata and hidden information from Microsoft Office (Word, Excel and PowerPoint, versions 97-2010) and PDF documents
- Identifies, reports and optionally removes or modifies more than 40 metadata and hidden data elements
- Bursts and reassembles slides from multiple PowerPoint presentations
- Provides accurate text offset information to automate native search hit-highlighting of PDFs in Adobe Reader
- Architected for high document throughput required by the most performance sensitive environments
- Easy integration via a Java API for Java or any Java compatible environment like JSP and J2EE, or via a C/C++ or .NET APIs for integration with traditional languages
- No Microsoft Office dependency eliminating the reliability, scalability and platform dependency issues that arise when automating Office applications to process files in high volumes
- Available on Windows with Java and C/C++ and .NET interfaces, on Linux x86 with Java and C/C++ interfaces, and on Solaris SPARC with a Java interface. Supported on any Java 1.5 or above compliant JVM



Belkasoft Evidence Center makes it easy for an investigator to acquire, search, analyse, store and share digital evidence found inside computer and mobile devices. The toolkit will quickly extract digital evidence from multiple sources by analysing hard drives, drive images, cloud, memory dumps, iOS, Blackberry and Android backups, UFED, JTAG and chip-off dumps. Evidence Center will automatically analyse the data source and lay out the most forensically important artefacts for investigator to review, examine more closely or add to report.





## Pop Corn Concept



ICYWW\*, you totally can make popcorn with a hair straightener!

\*In case your were wondering.

## Dent Corn

*(Zea mays var. indentata)*

## Flint Corn

*(Zea mays var. indurata)*



## Popcorn

*(Zea mays var. everta)*



## Sweet Corn

*(Zea mays convar. saccharata var. rugosa)*

## Flour corn

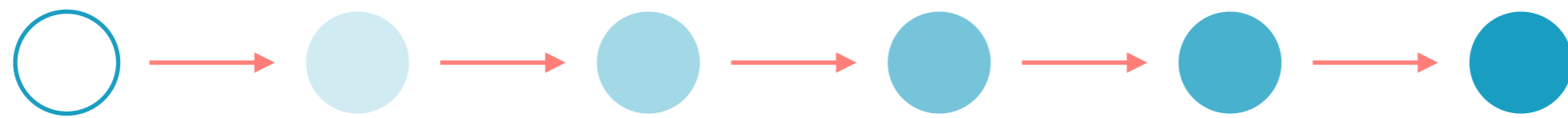
*(Zea mays var. amylacea)*



Each kernel of popcorn contains a certain amount of moisture and oil. Unlike most other grains, the outer hull of the popcorn kernel is both strong and impervious to moisture and the starch inside consists almost entirely of a hard type. As the oil and water within the kernel are heated, they turn the moisture in the kernel into pressurized steam. Under these conditions, the starch inside the kernel gelatinizes, softens, and becomes pliable. The internal pressure of the entrapped steam continues to increase until the breaking point of the hull is reached: a pressure of approximately **930 kPa** and a temperature of **180 °C**. The hull thereupon ruptures rapidly and explodes, causing a sudden drop in pressure inside the kernel and a corresponding rapid expansion of the steam, which expands the starch and proteins of the endosperm into airy foam. As the foam rapidly cools, the starch and protein polymers set into the familiar crispy puff. Special varieties are grown to give improved popping yield. Though the kernels of some wild types will pop, the cultivated strain is *Zea mays everta*, which is a special kind of flint corn.

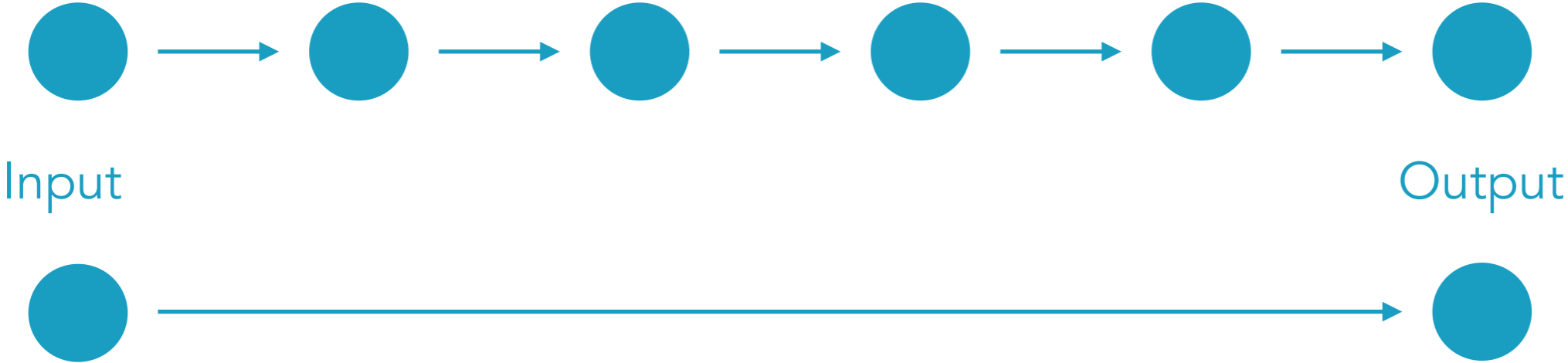


Source: Wikipedia



**Chain Concept**

# Bioinformatics - Concepts



# Bioinformatics - Concepts

