Humans and I the living world

Virginie COURTIER-ORGOGOZO





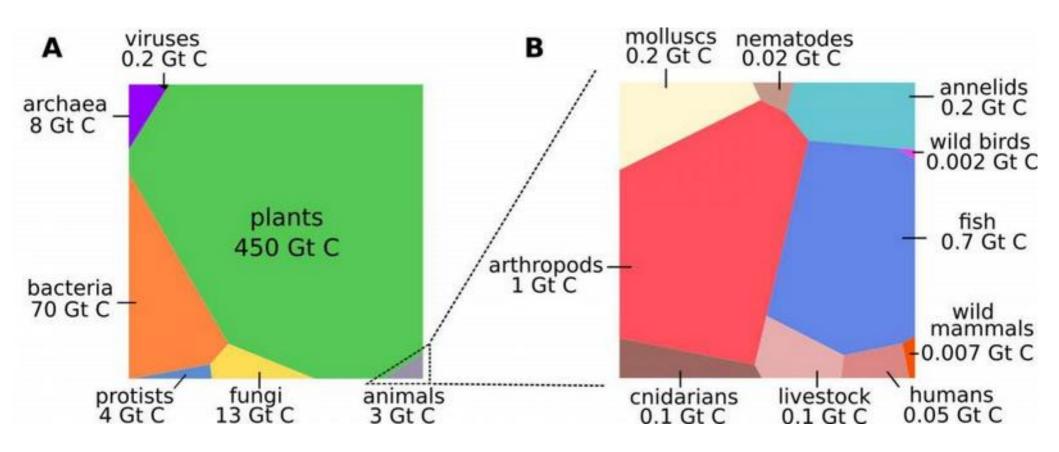




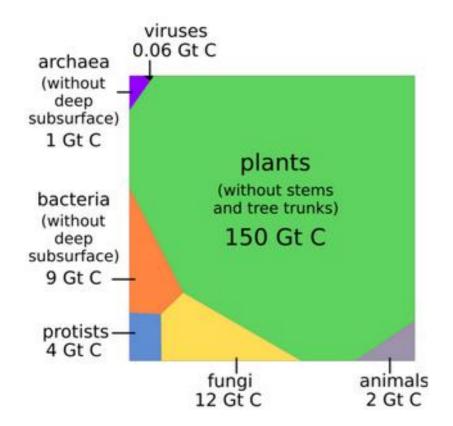
What is the largest biomass on earth ?

- □ Insects
- □ Viruses
- □ Bacteria
- □ Green plants

Biomass distribution on earth



Bar-On 2018 PNAS

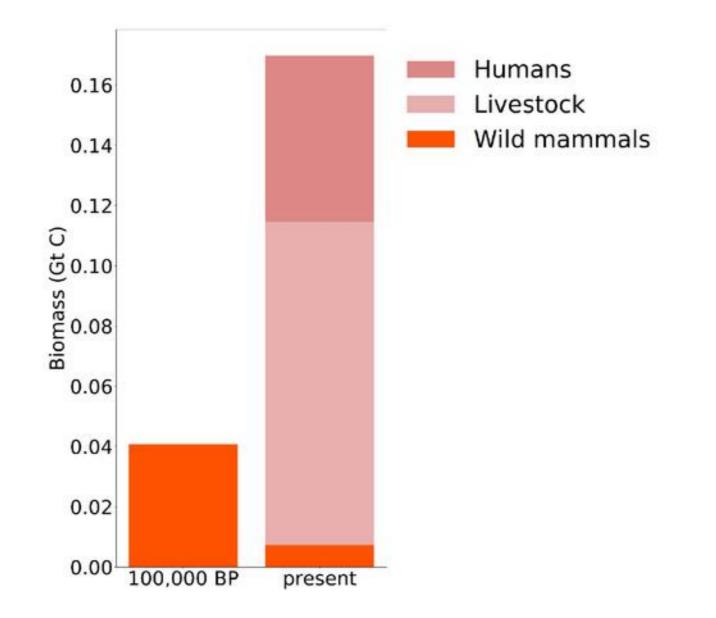


Bar-On 2018 PNAS

Which has the largest biomass ?

humanslivestock





Bar-On 2018 PNAS

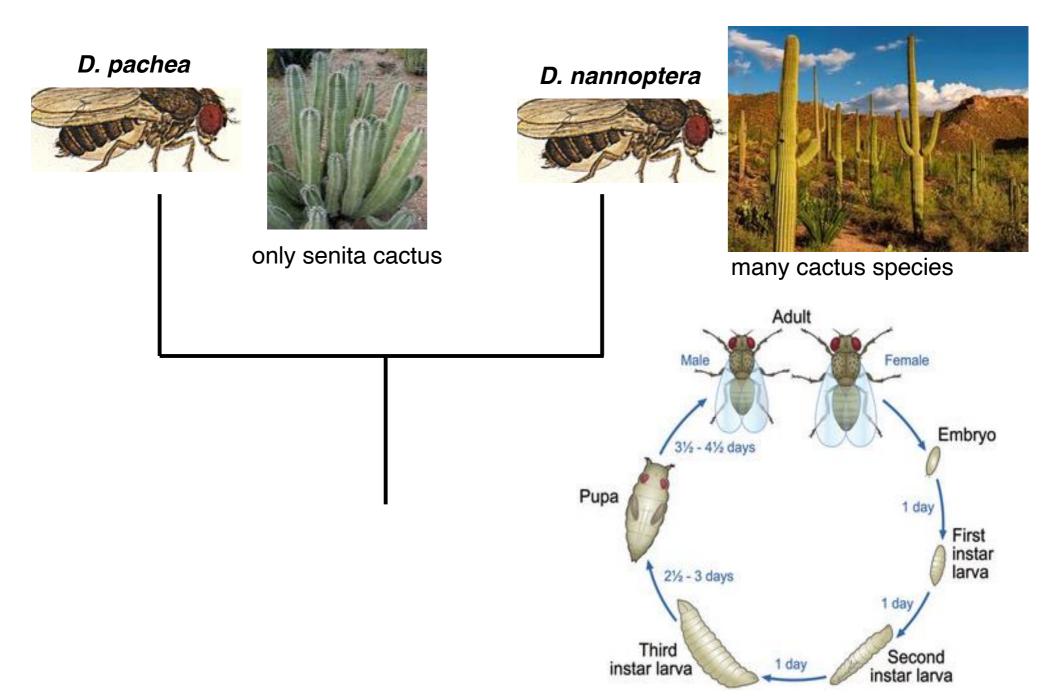
What is life ?

From molecules to ecosystems

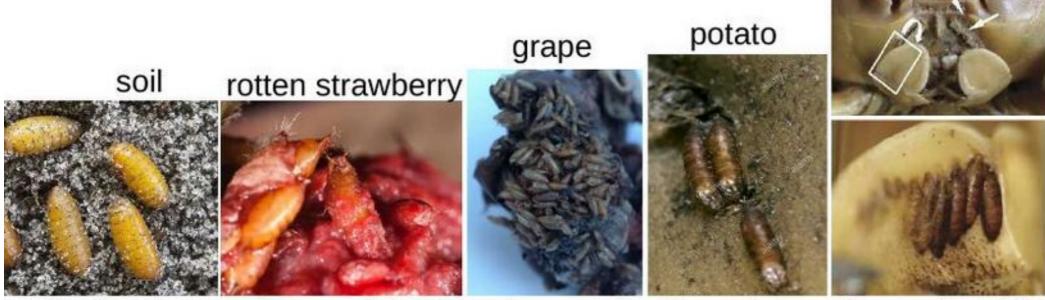
The tree of life Individuals and interconnections

Anthropocene

My lab topic: How do species diverge with time ?

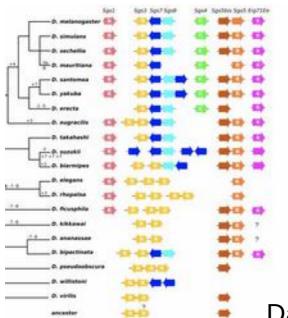


Evolution of Drosophila glue



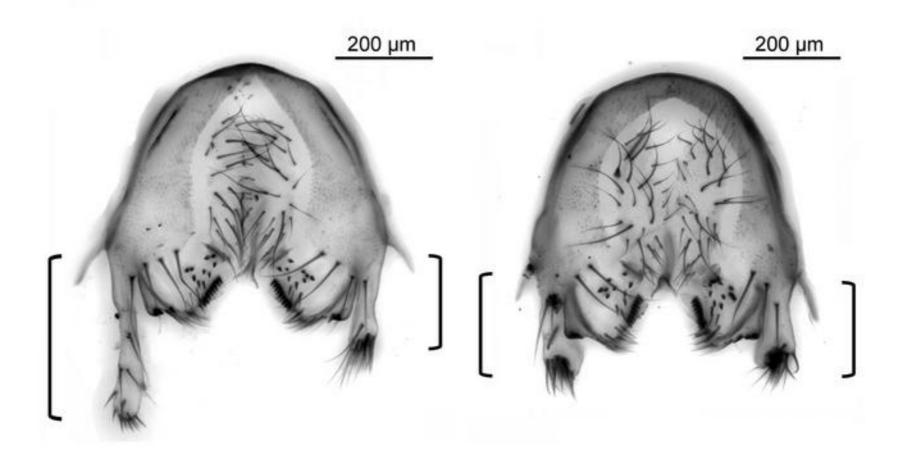
. grimshawi

D. suzukii D. melanogaster D. repleta D. carcinophila



Da Lage et al. 2019 BMC Evol Biol

Evolution of left-right asymmetry in *D. pachea*



What is life ?

From molecules to ecosystems

The tree of life Individuals and interconnections

Anthropocene

LIFE SCIENCES

LS1 Molecular Biology, Biochemistry, Structural Biology and Molecular Biophysics

LS2 Genetics, 'Omics', Bioinformatics and Systems Biology

LS3 Cellular and Developmental Biology

LS4 Physiology, Pathophysiology and Endocrinology

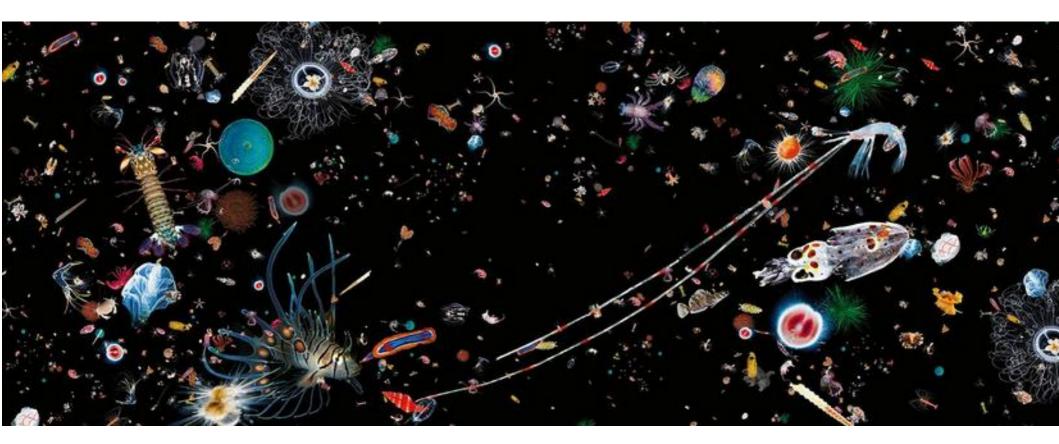
LS5 Neurosciences and Neural Disorders

LS6 Immunity and Infection

LS7 Applied Medical Technologies, Diagnostics, Therapies, Public Health

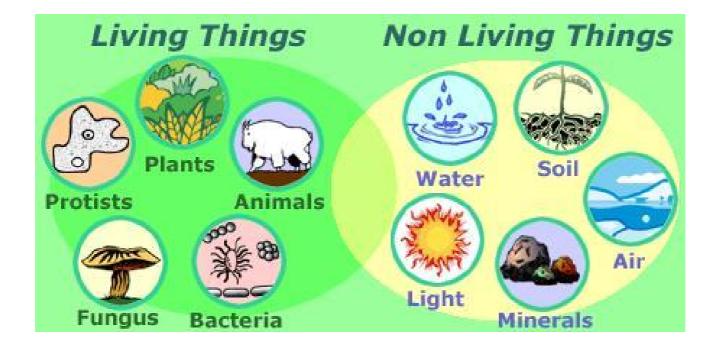
LS8 Ecology, Evolution and Environmental Biology

LS9 Applied Life Sciences, Biotechnology and Molecular and Biosystems Engineering « La nature n'est jamais aussi grande que dans ses créatures les plus petites » (Pline l'Ancien)



What is life ?

What are the properties of the living world ?



Fire The sun A virus A computer virus A mule (sterile hybrids) A foctus in the mother's womb A sperm cell A spore that will never germinate A sea urchin oocyte that will not be fertilized The earth

Properties of the living world

Metabolism Ability to reproduce Auto-organization properties

Delimited by a membrane Made of cells Contains nucleic acids Can mutate Derives from another living being

What is the difference between humans and other organisms ?

Humans versus other organisms

Several conceptions in different cultures

Our conception: **Naturalism** (same physicality between humans and other organisms, all can be explained by this physicality)

Other views: **Animism** (each living species has its own physicality and interiority and thus views the world in its own way)

Totemism (certain groups of humans have the same physicality and internality as certain groups of animals) Ex: catcher (white cockato *cacatoès*) / watcher (raven)

Analogism (all types of physicalities and interiorities can be combined)

Descola 2005

Theodore Schwann, 1839

"All living things are composed of cells and cell products"

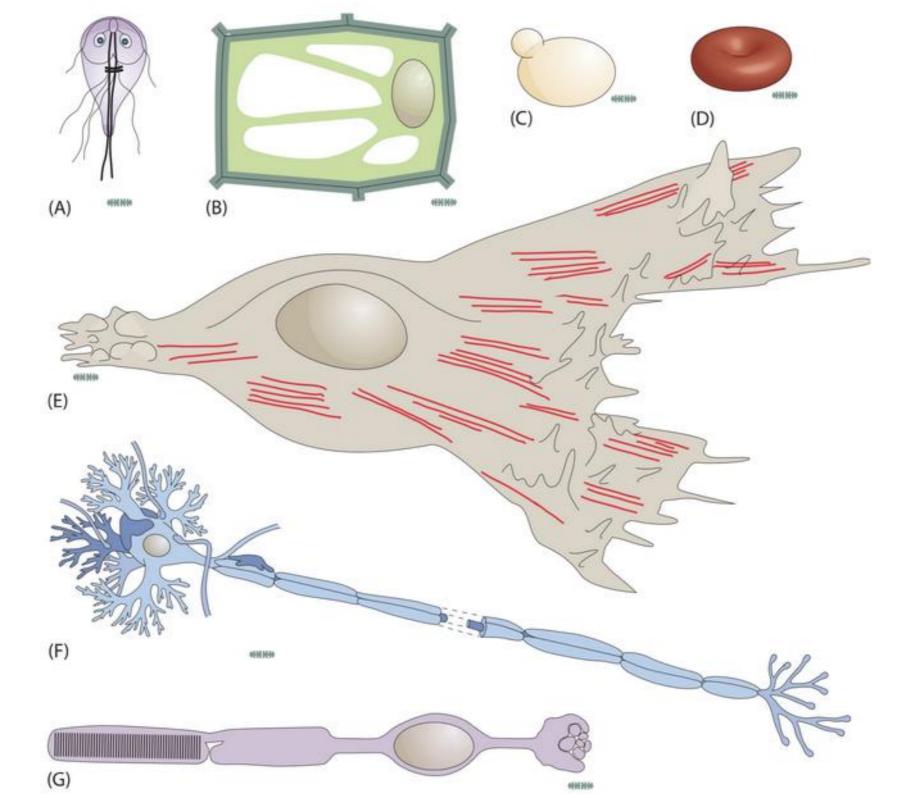
1) The cell is the unit of structure, physiology, and organization in living things.

2) The cell retains a dual existence as a distinct entity and a building block in the construction of organisms.

Rudolf Virchow, 1857

Every cell arises from another cell





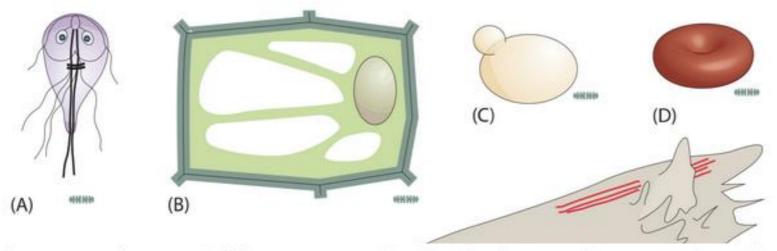
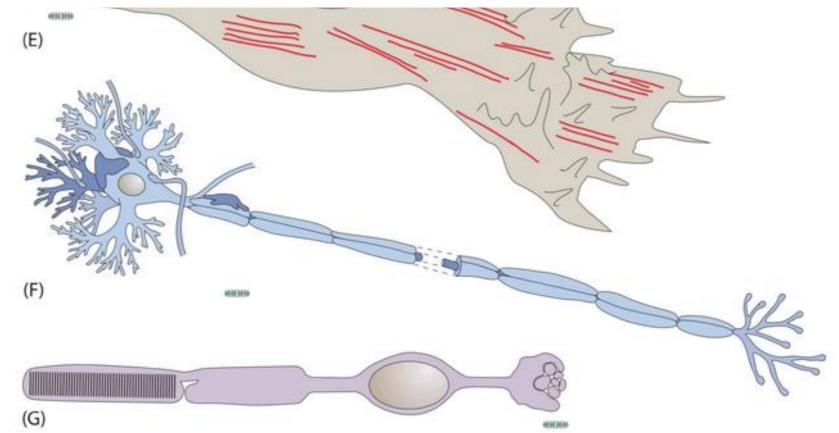
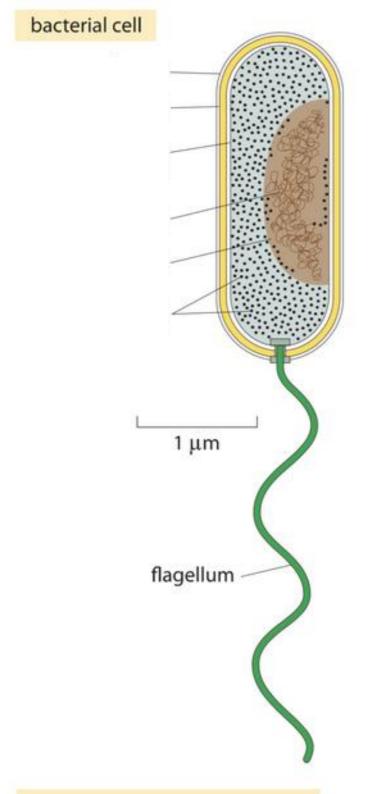
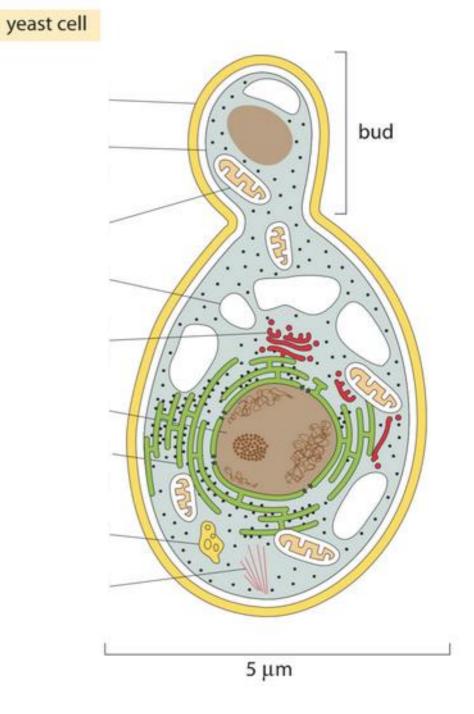


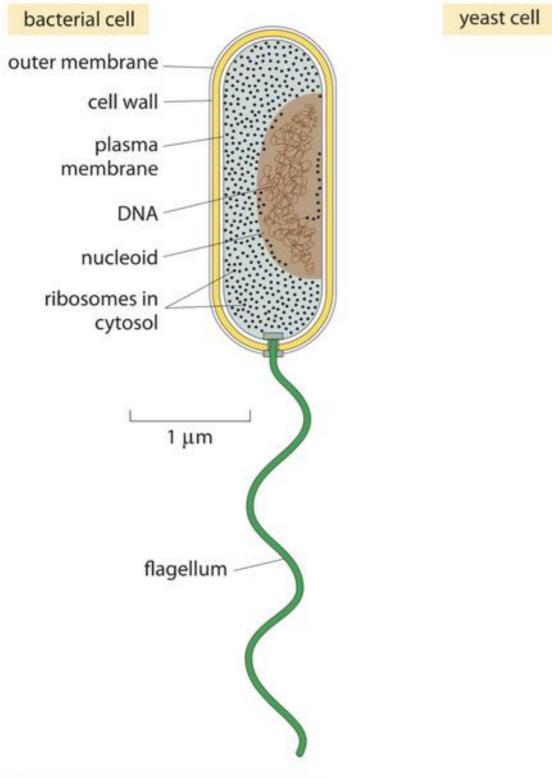
Figure 4: Cartoons of several different types of cells all referenced to a standard *E. coli* ruler of 1 micron width drawn in grey. (A) The protist Giardia lamblia, (B) a plant cell, (C) a budding yeast cell, (D) a red blood cell, (E) a fibroblast cell, (F) a eukaryotic nerve cell, and (G) a rod cell from the retina.

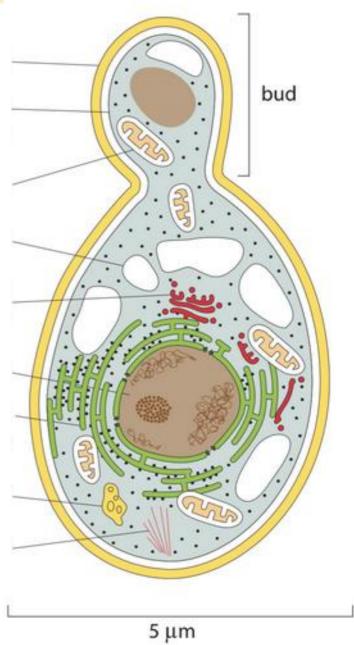


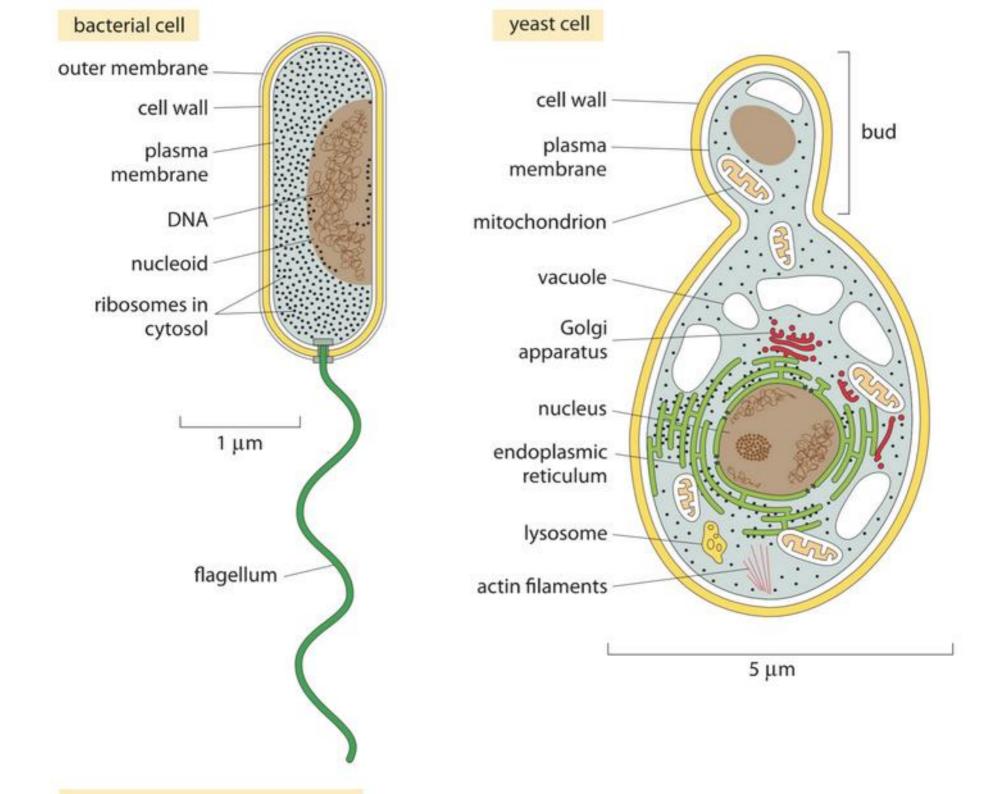
From cells to molecules



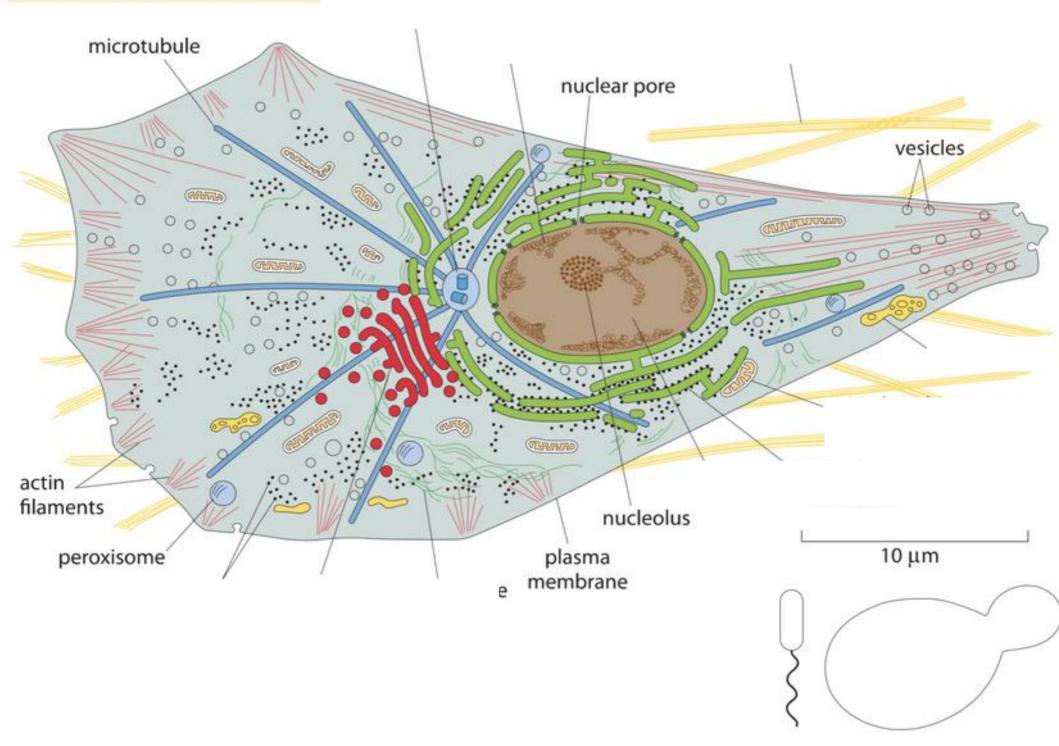


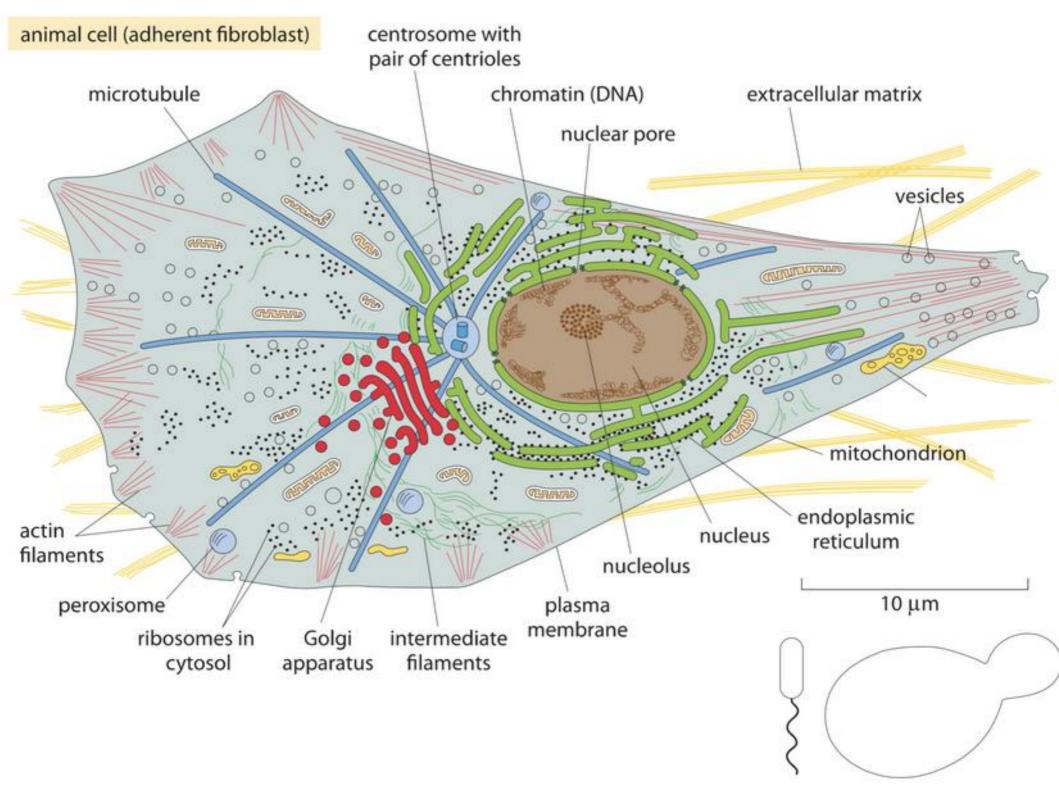


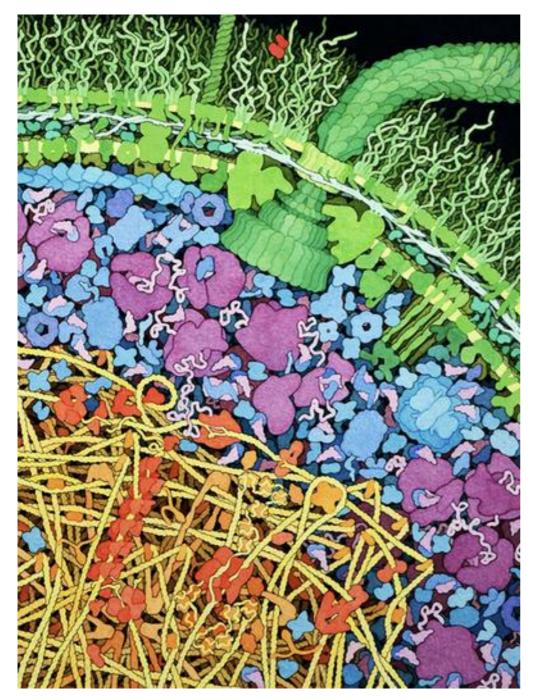




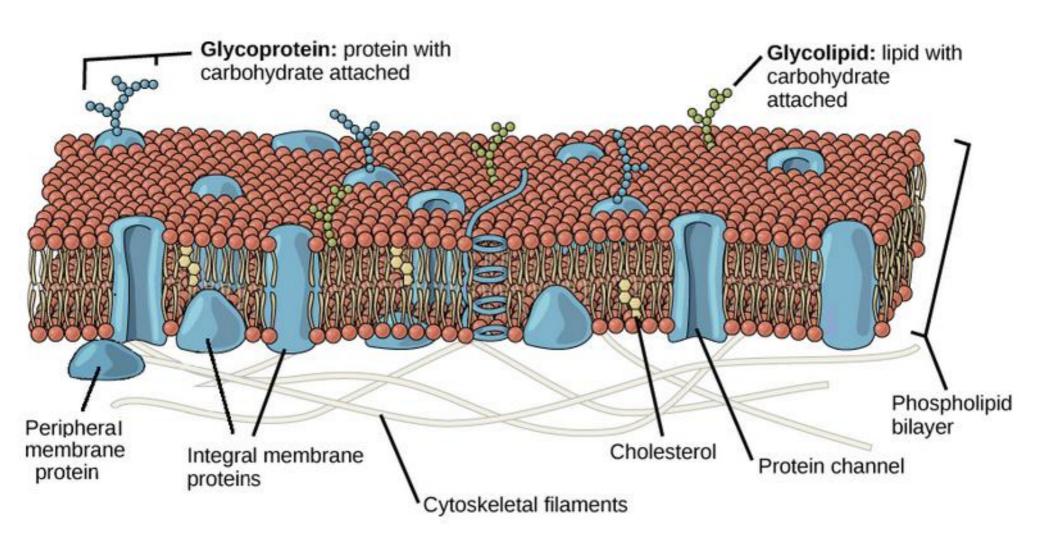
animal cell (adherent fibroblast)







David Goodsell



nucleic acid: biopolymer composed of nucleotides, which are the monomers made of three components: a 5-carbon sugar, a phosphate group and a nitrogenous base. If the sugar is a compound ribose, the polymer is RNA (ribonucleic acid); if the sugar is deoxyribose, the polymer is DNA (deoxyribonucleic acid).

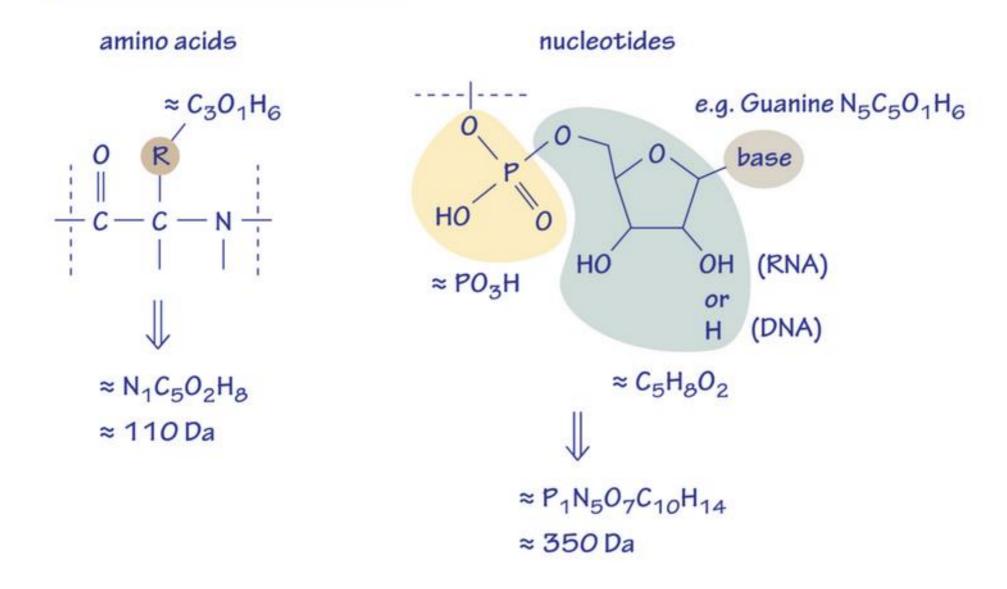
protein: large biomolecule consisting of one or more long chains of amino acid residues.

amino acid: organic molecule that contains amine (-NH2) and carboxyl (-COOH) functional groups, along with a side chain (R group) specific to each amino acid.

lipid: biomolecule soluble in nonpolar solvents. Functions of lipids include storing energy, signaling, and acting as structural components of cell membranes.

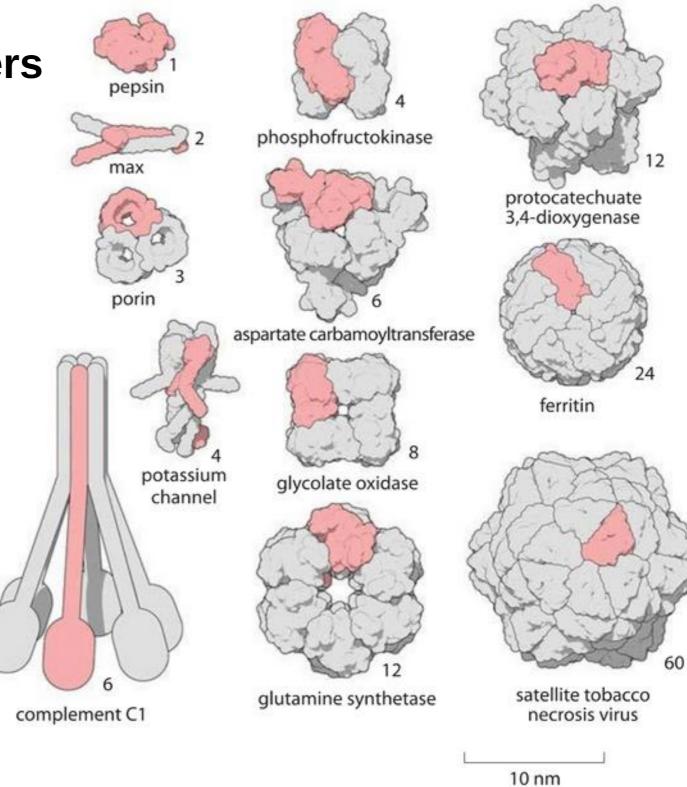
carbohydrate: biomolecule consisting of carbon (C), hydrogen (H) and oxygen (O) atoms, usually with the empirical formula Cm(H2O)n. Functions include storage of energy (e.g. starch and glycogen) and acting as structural components (e.g. cellulose in plants and chitin in arthropods).

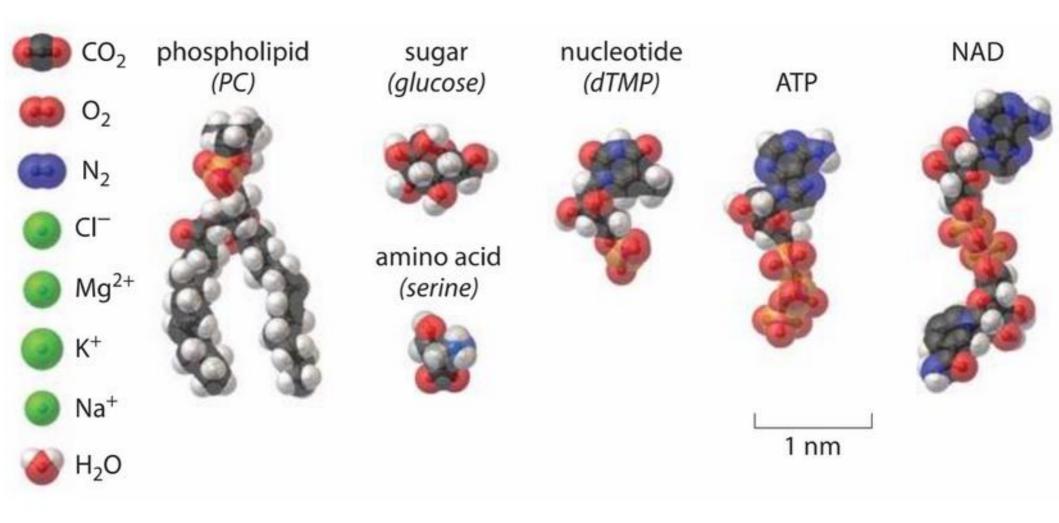
main constituents of a cell



total protein mass in cell ≈ 3 times larger than RNA + DNA

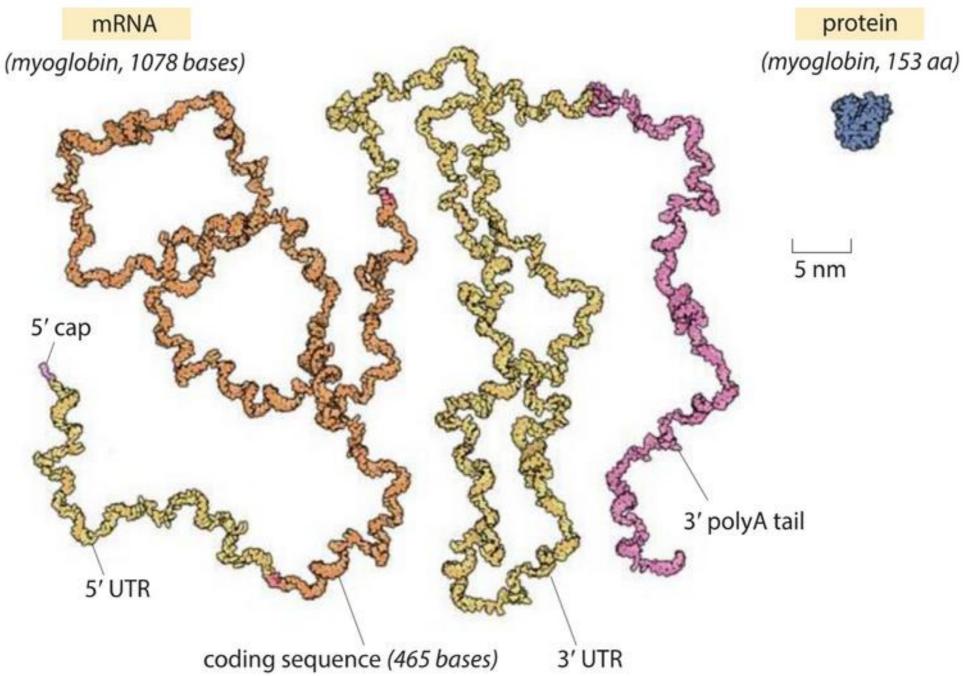
Homooligomers





Which is larger: mRNA or the protein it codes for?

Which is larger: mRNA or the protein it codes for?



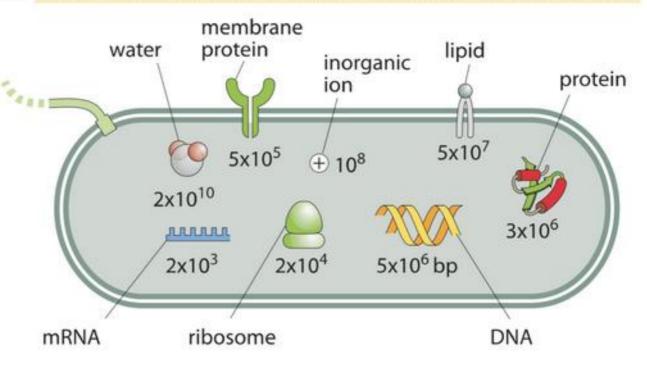
Median protein length

organism	median protein length (amino acids)
H. sapiens	375
D. melanogaster	373
C. elegans	344
S. cerevisiae	379
A. thaliana	356
5 eukaryotes (above)	361
67 bacteria	267
15 archaea	247

property	E. coli	budding yeast	mammalian (HeLa line)
cell volume	0.3–3 μm ³	30–100 μm ³	1,000–10,000 μm ³
proteins per µm ³ cell volume		2-4×10 ⁶	
mRNA per cell	10 ³ -10 ⁴	10 ⁴ -10 ⁵	10 ⁵ -10 ⁶
proteins per cell	~10 ⁶	~10 ⁸	~10 ¹⁰
mean diameter of protein	n	4–5 nm —	
genome size	4.6 Mbp	12 Mbp	3.2 Gbp
number protein coding genes	4300	6600	21,000
regulator binding site length	2	10-20 bp	
promoter length	~100 bp	~1000 bp	~10 ⁴ -10 ⁵ bp
gene length	~1000 bp	~1000 bp	~10 ⁴ -10 ⁶ bp (with introns)
concentration of one protein per cell	~1 nM	~10 pM	~0.1–1 pM
diffusion time of protein across cell ($D \approx 10 \ \mu m^2/s$)	~0.01 s	~0.2 s	~1-10 s
diffusion time of small molecule across cell (D \approx 100 μ m ² /s)	~0.001 s	~0.03 s	~0.1-1 s
time to transcribe a gene	<1 min (80 nts/s)	~1 min	~30 min (incl. mRNA processing)
time to translate a protein	<1 min (20 aa/s)	~1 min	~30 min (incl. mRNA export)
typical mRNA lifetime	2–5 min	~10 min to over 1 h	5-100 min to over 10 h
typical protein lifetime	1 h	0.3–3 h	10–100 h
minimal doubling time	20 min	1 h	20 h
ribosomes/cell	~10 ⁴	~10 ⁵	~10 ⁶
transitions between protein states (active/inactive)		1–100 μs	
timescale for equilibrium binding of small molecule to protein (diffusion limited)	1–1000 ms (1 μM–1 nM affinity)		
timescale of transcription factor binding to DNA site	~1 s		
mutation rate			

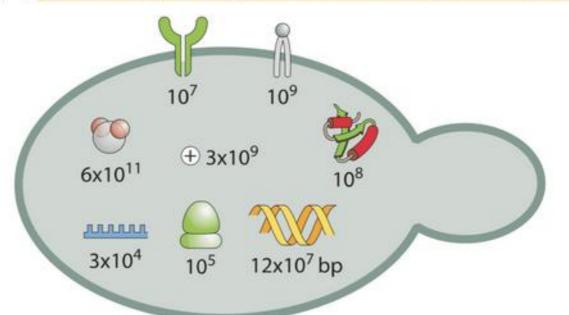
(A) bacterial cell (specifically, *E. coli*: $V \approx 1 \ \mu m^3$; $L \approx 1 \ \mu m$; $\tau \approx 1 \ hour$)



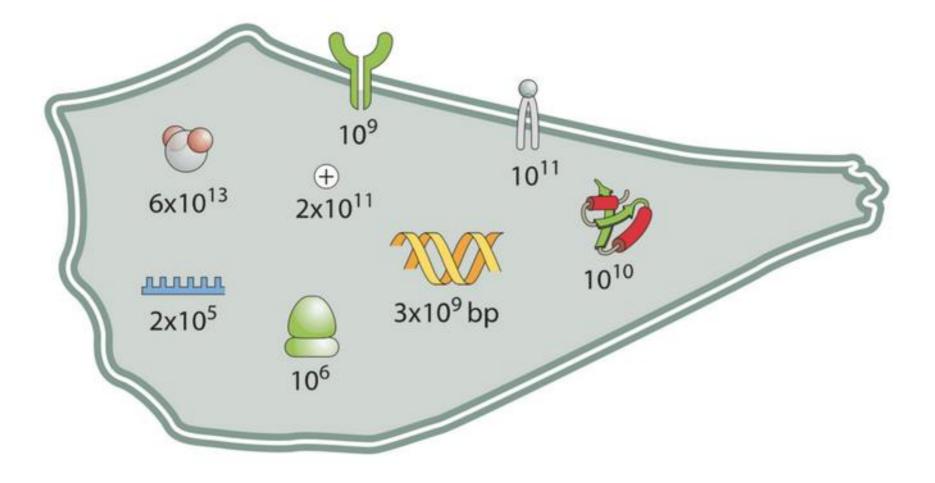


(B) yeast cell (specifically, S. cerevisiae: $V \approx 30 \ \mu m^3$; $L \approx 5 \ \mu m$; $\tau \approx 3 \ hours$)

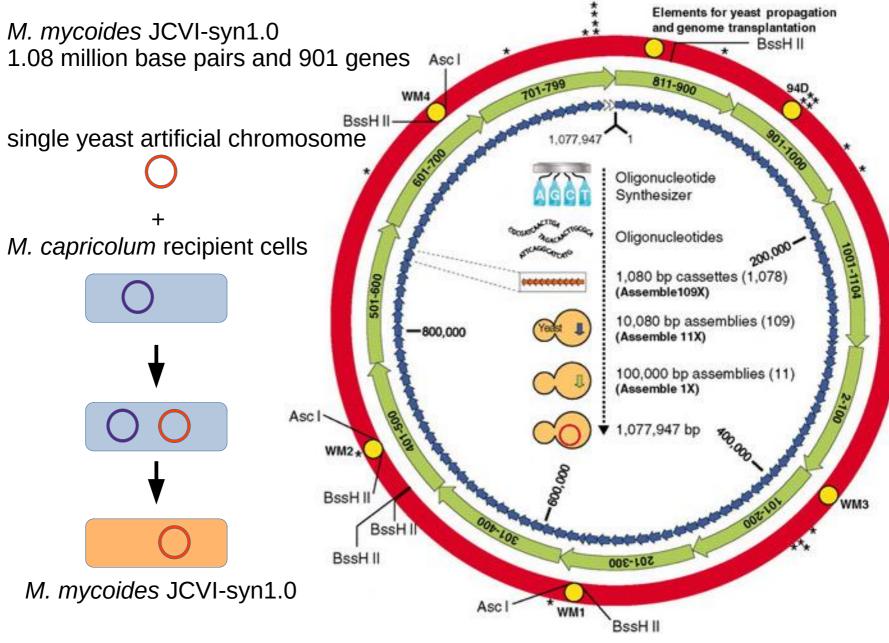




(C) mammalian cell (specifically, HeLa: $V \approx 3000 \ \mu m^3$; $L \approx 20 \ \mu m$; $\tau \approx 1 \ day$)



First "synthetic" cell developed by scientists



Gibson et al. 2016

First "synthetic" cell developed by scientists

M. mycoides JCVI-syn1.0 contains strings of bases that, in code, spell out:

a web address to send emails to if you can successfully crack the new code,

the names of 46 authors and other key contributors,

three famous quotations. One of which by James Joyce, perfectly encapsulates the ups and downs of a the 15 year project—"To live, to err, to fall, to triumph, to recreate life out of life."

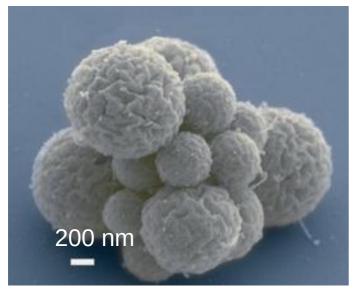
Smallest "synthetic" cell

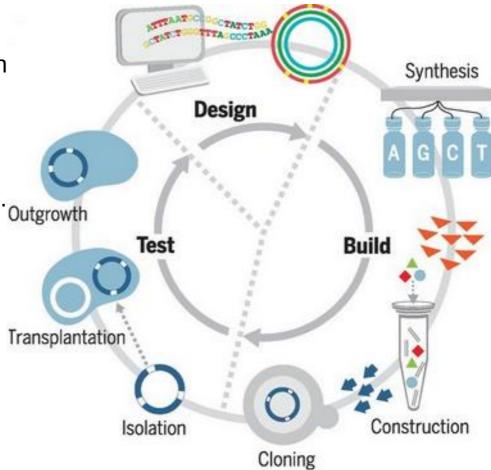
M. mycoides JCVI-syn3.0 531,560 base pairs and 473 genes

retains genes involved in key processes such as transcription and translation, but also contains 149 genes of unknown function

genome smaller than that of any autonomously replicating cell found in nature.

doubling time of ~180 min colonies morphologically similar to those of JCVI-syn1.0

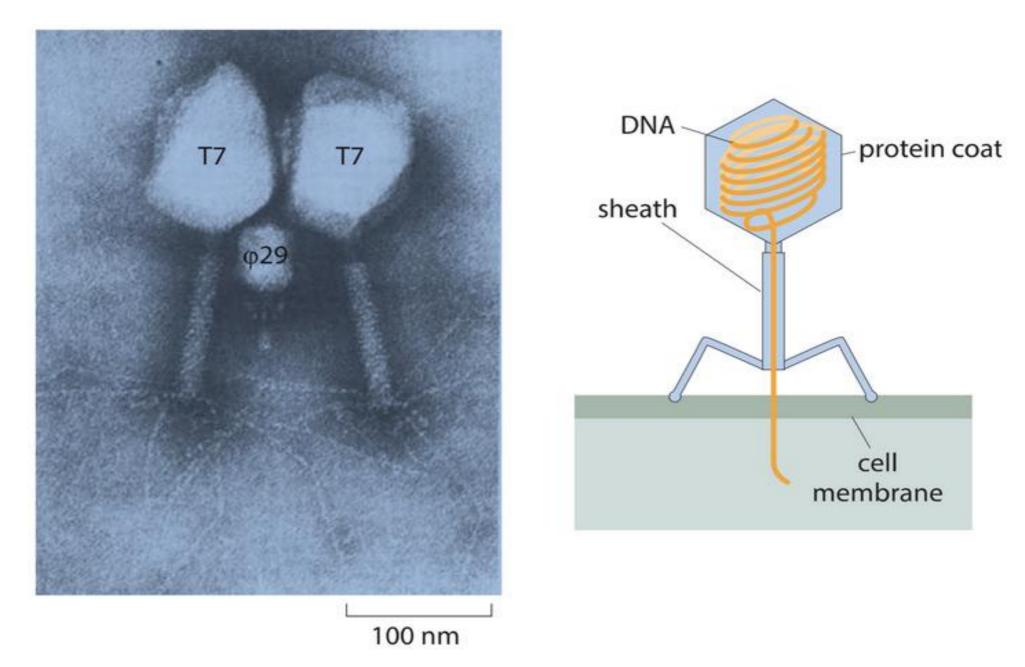


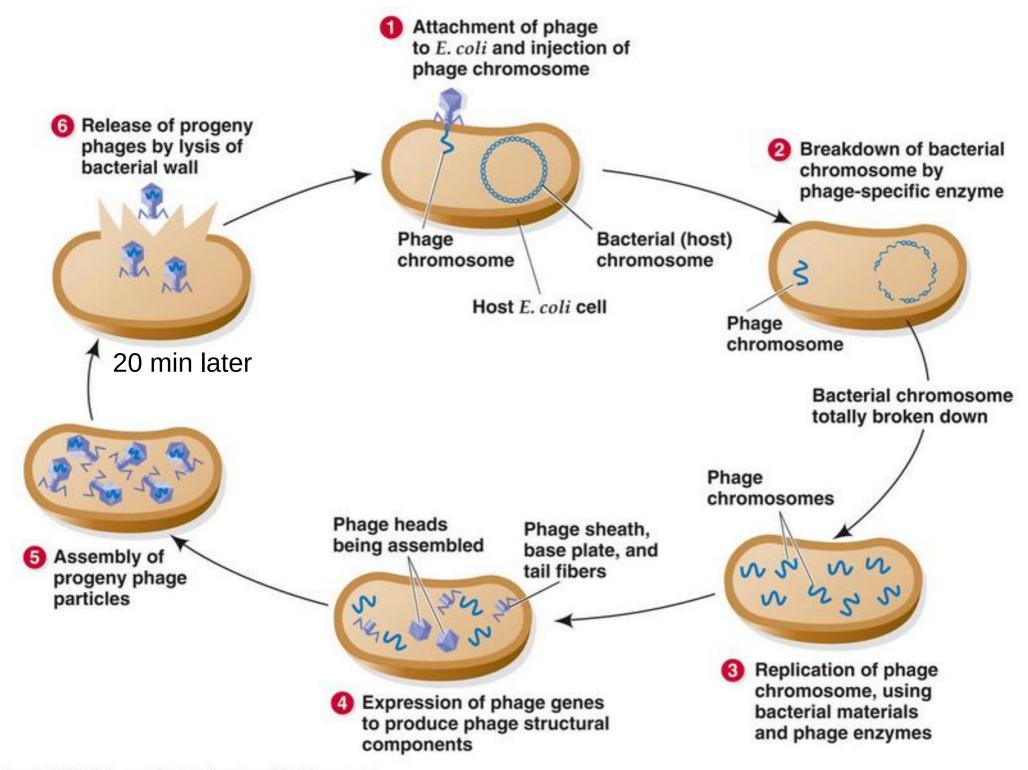


Hutchison III et al. 2016

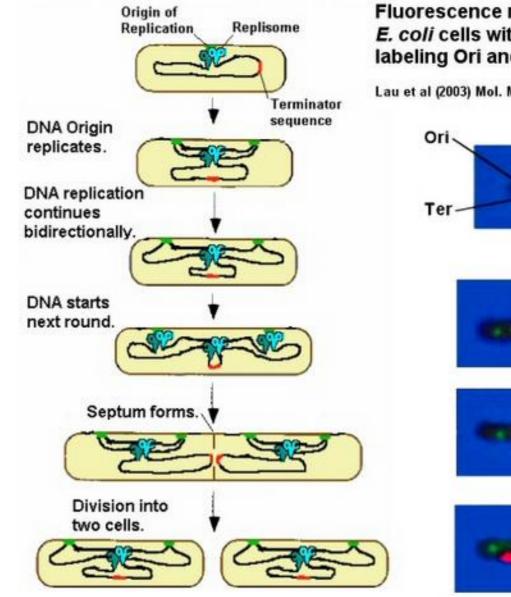
Viruses, cells, mitochondria, reproduction

Virus infecting bacteria = phages



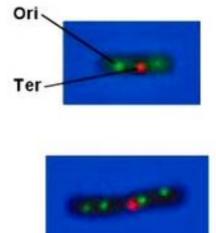


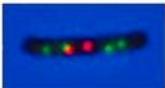
Copyright @ 2006 Pearson Benjamin Cummings. All rights reserved.

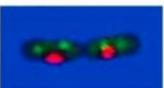


Fluorescence microscopy: E. coli cells with fluorophores labeling Ori and Ter

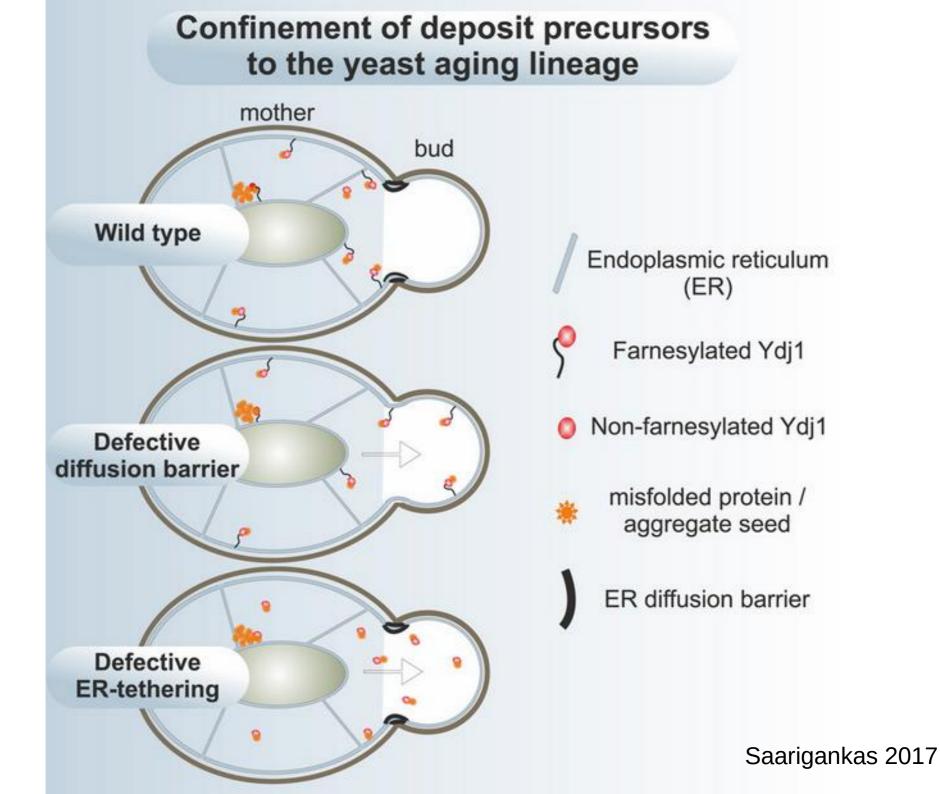
Lau et al (2003) Mol. Micro. 49:731



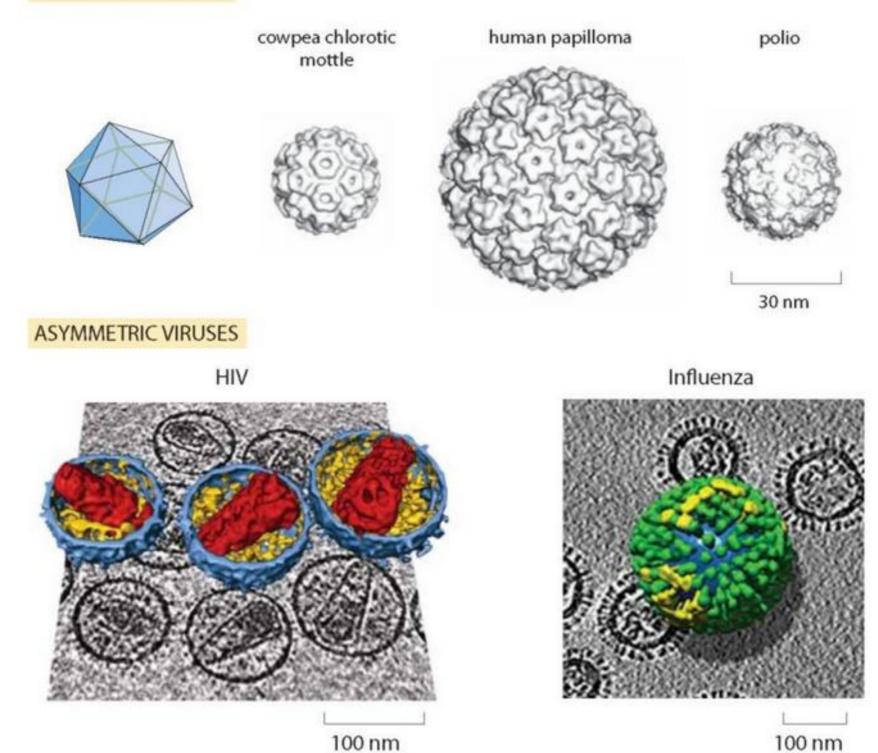




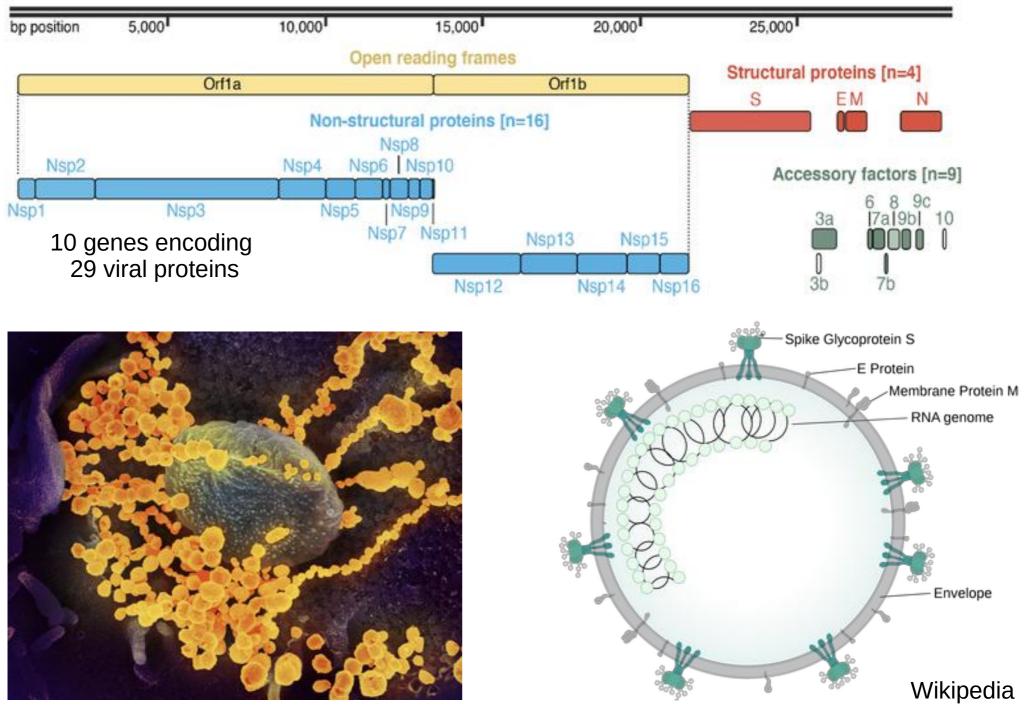
http://biology.kenyon.edu/courses/biol114/Chap01/cell.html



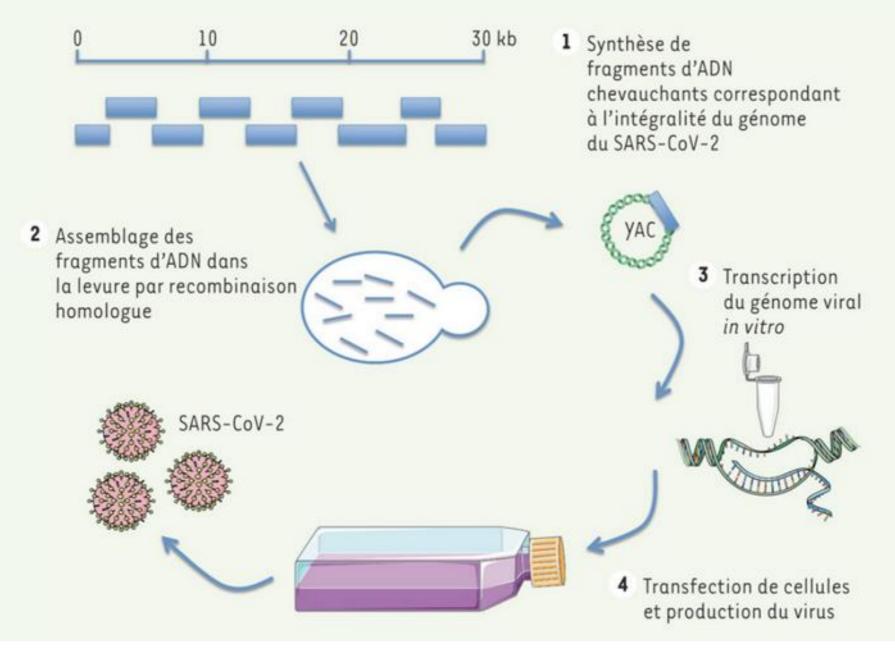
SYMMETRIC VIRUSES



SARS-CoV-2 genome



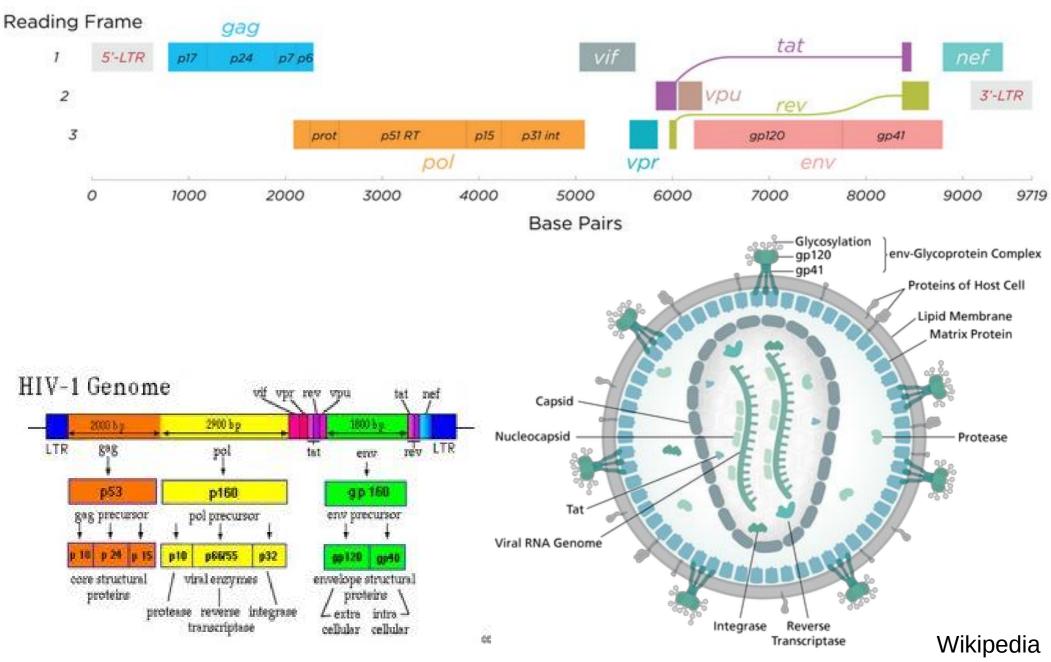
Synthesis of SARS-CoV-2 in a few weeks



Iseni, Frédéric, and Jean-Nicolas Tournier. 2020. 'Une Course Contre La Montre-Création Du SARS-CoV-2 En Laboratoire, Un Mois Après Son Émergence !' Médecine/Sciences 36 (8–9): 797–802

HIV genome

9 genes encoding 15 viral proteins



virus	size (nm)	genome size (base pairs)
porcine circovirus (PCV)	17	1,760
cowpea mosaic virus (CPMV)	28	9,400
cowpea chlorotic mottle virus (CCMV)	28	7,900
φX174 (<i>E. coli</i> bacteriophage)	32	5,400
tobacco mosaic virus (TMV)	40×300	6,400
polio virus	30	7,500
φ29 (<i>Bacillus</i> phage)	45x54	19,000
lambda phage	58	49,000
T7 bacteriophage	58	40,000
adenovirus (linear DNA)	88-110	36,000
influenza A	80-120	14,000
HIV-1	120-150	9,700
herpes simplex virus 1	125	153,000
Epstein-Barr virus (EBV)	140	170,000
mimivirus	500	1,200,000
pandora virus	500x1000	2,800,000

Criteria to define viruses

Established by André Lwolff in 1957

1) does not contain ribosomes (needs the host machinery for replication)

2) cannot divide to produce 2 entities

3) reproduce via only part of their constituents

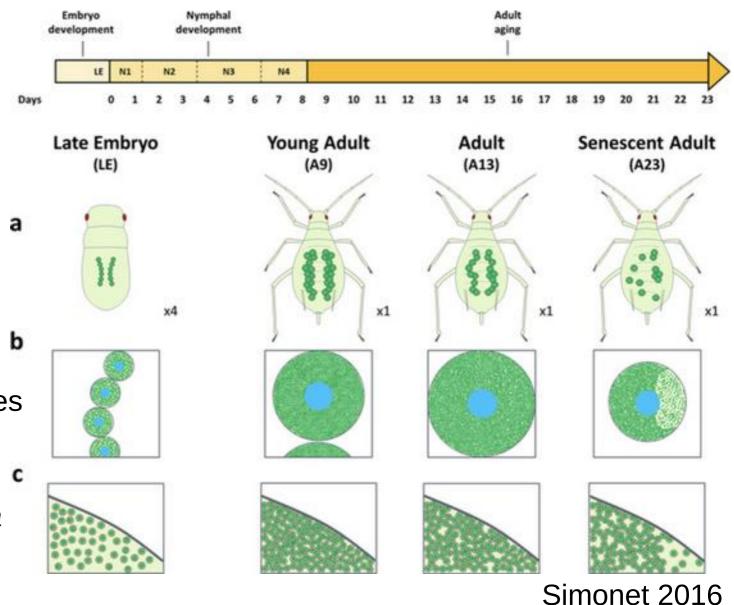
4) does not harbour a minimal metabolism to reproduce

Points 3 and 4 were later refuted.

Buchnera : an obligate symbiont

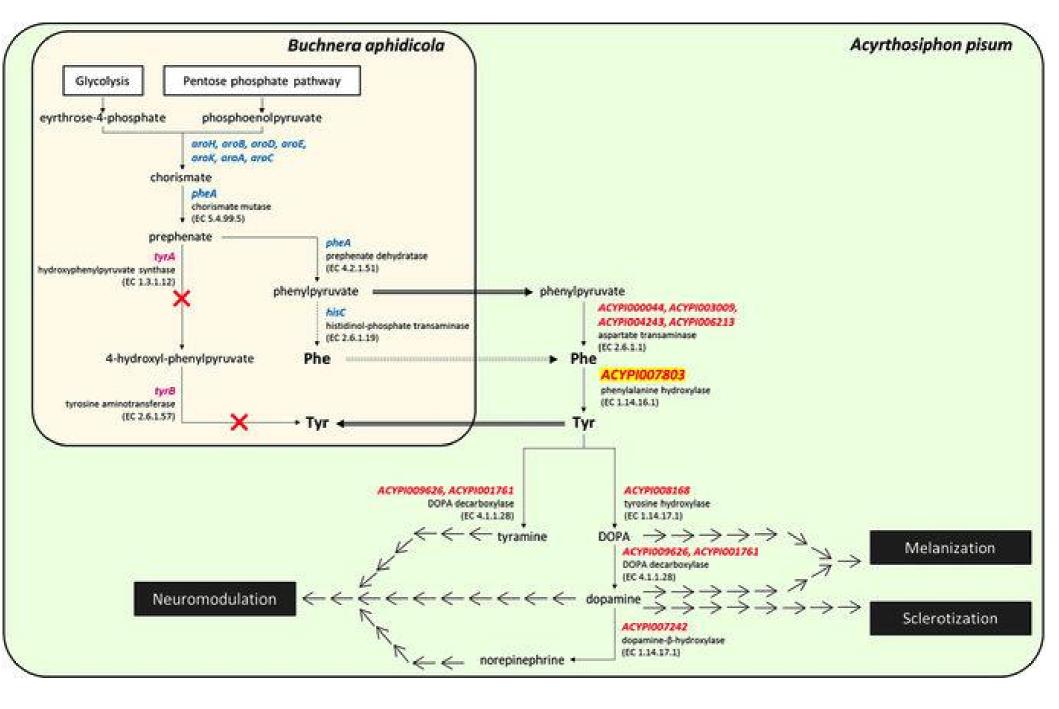


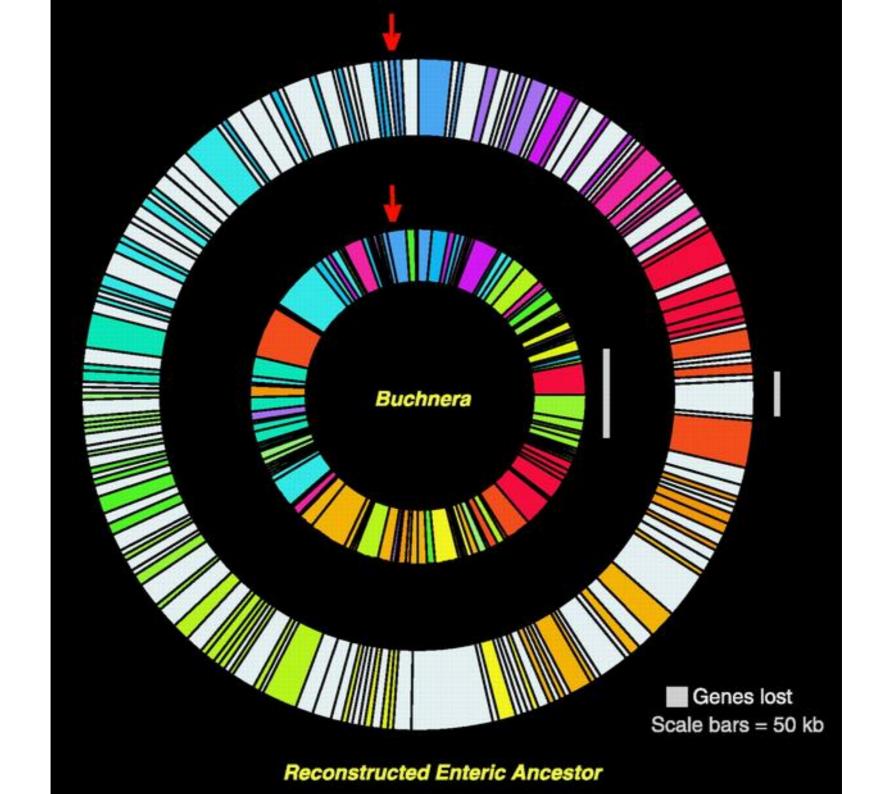
Phloem sap rich in carbohydrates but devoid of essential amino acids

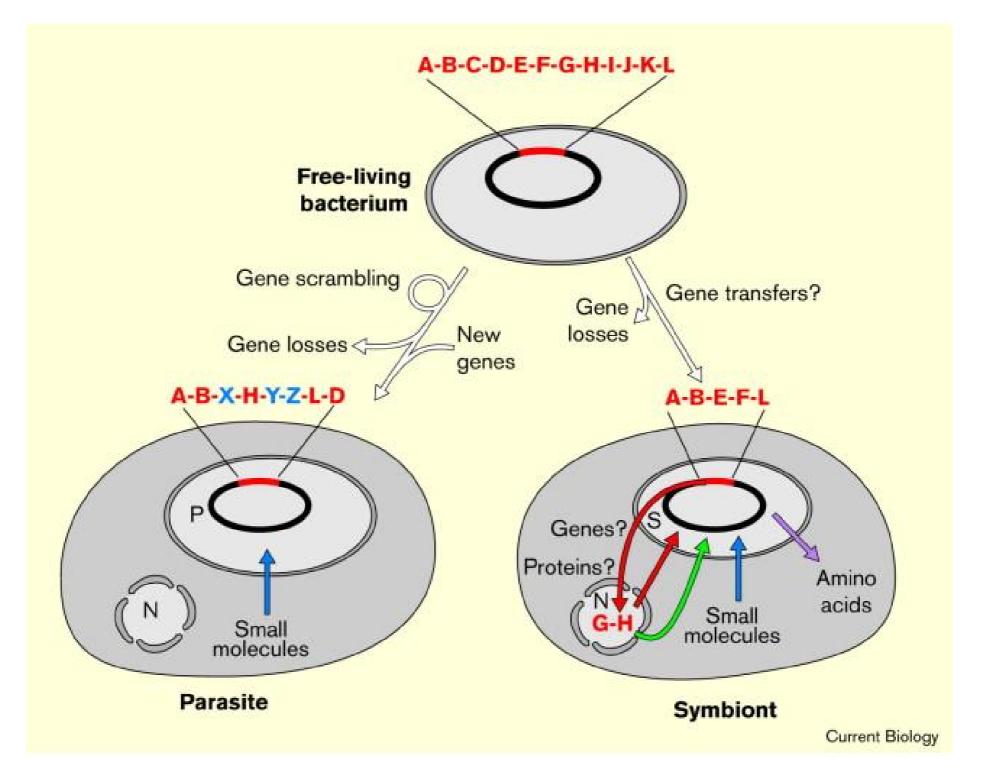


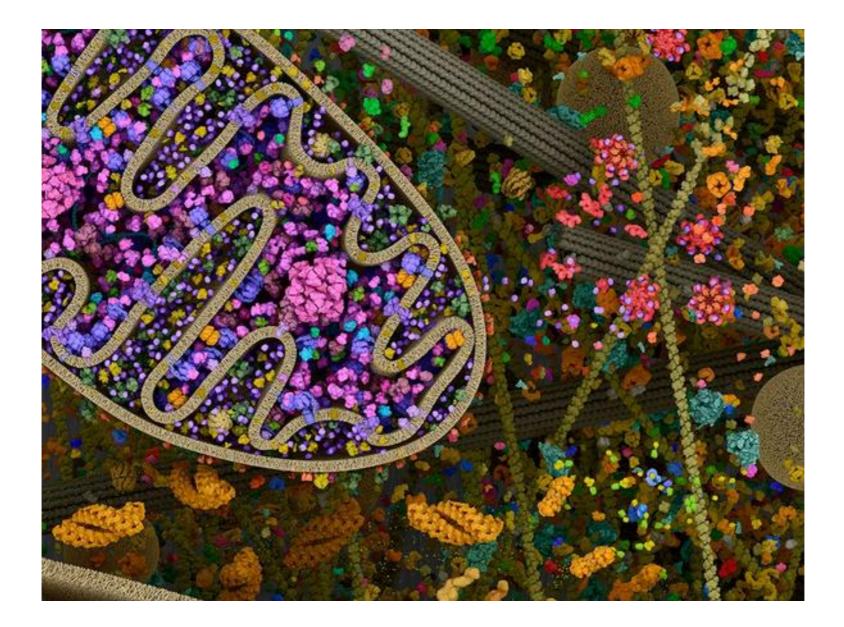
bacteriocytes

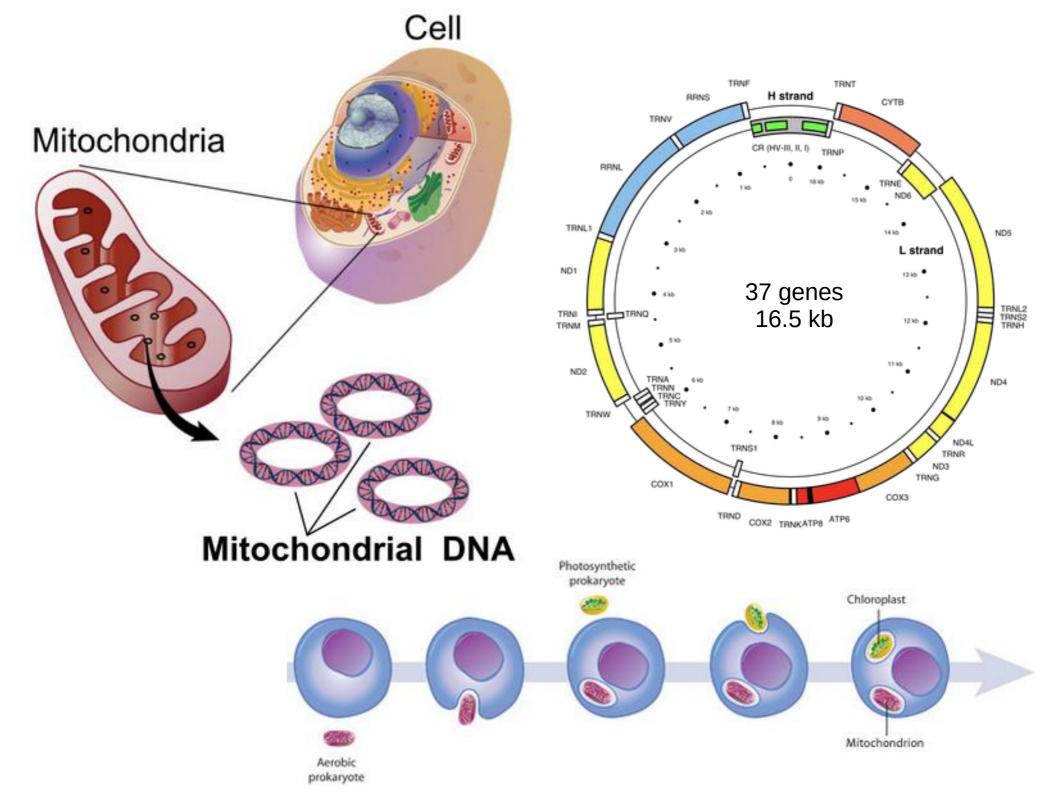
Buchnera bacteria



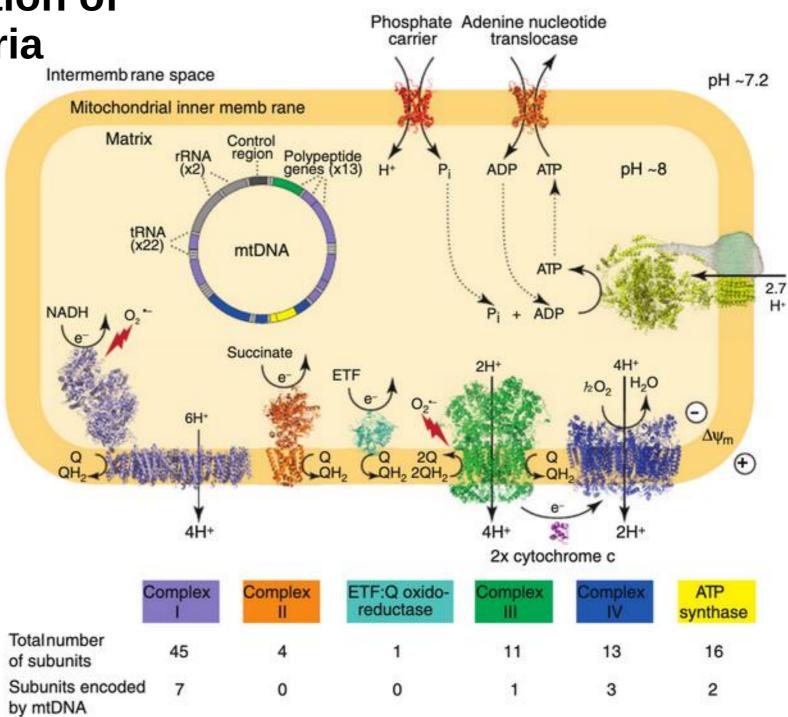








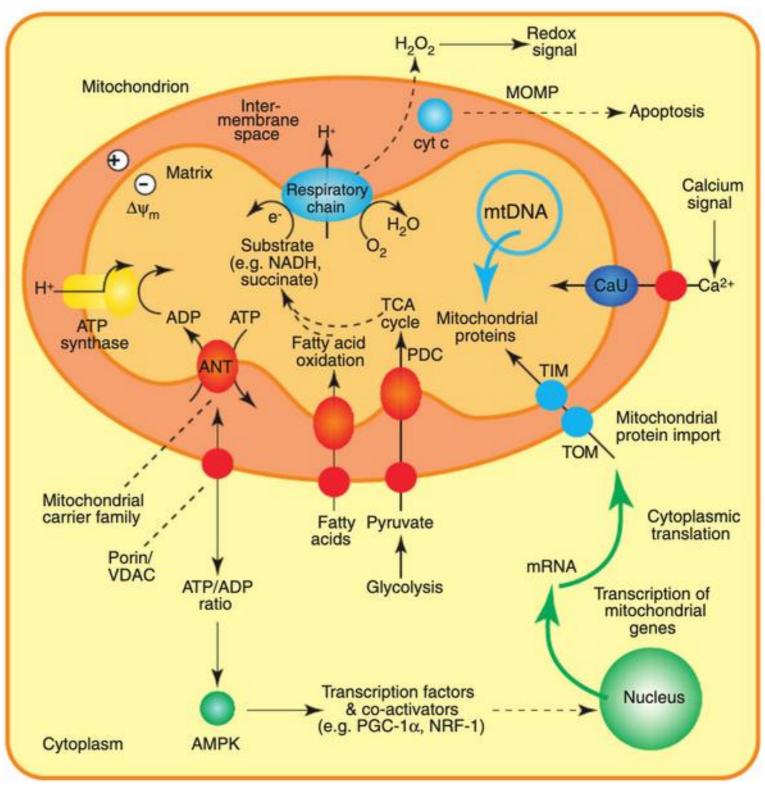
Major function of mitochondria is to produce ATP



Smith 2012

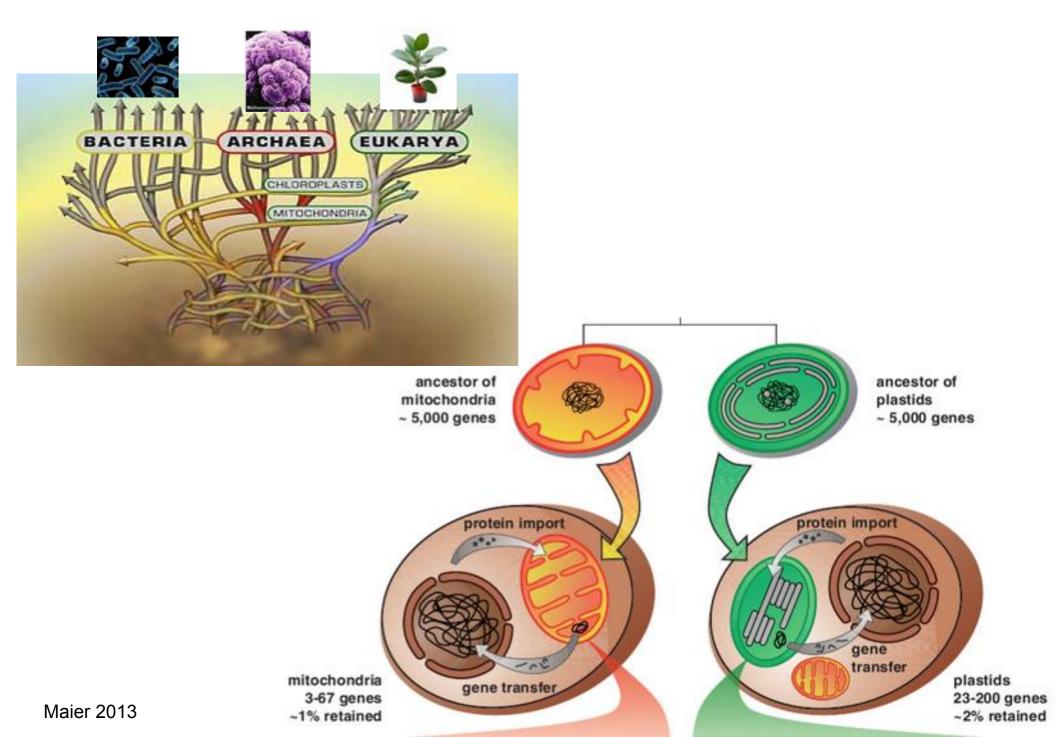
TRENDS in Pharmacological Sciences

Functions And Biogenesis Of Mitochondria



Smith 2012

Mitochondria and chloroplasts

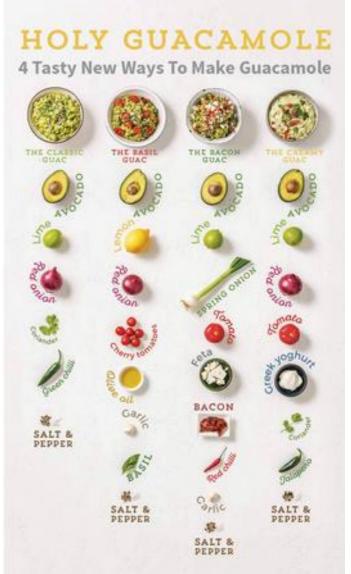


A living organism is not made by assembling pieces together



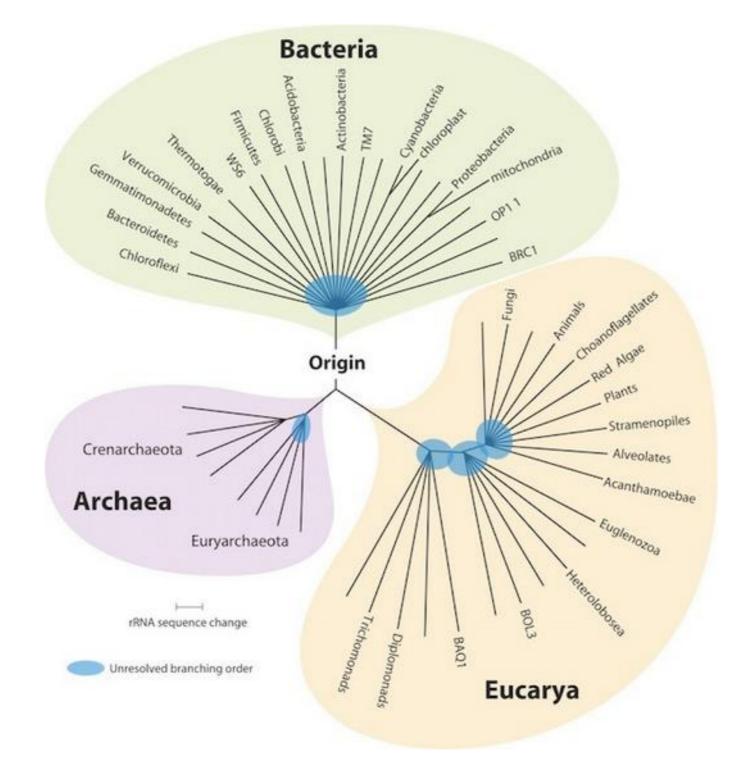
..but results from changes that occurred successively across evolutionary time



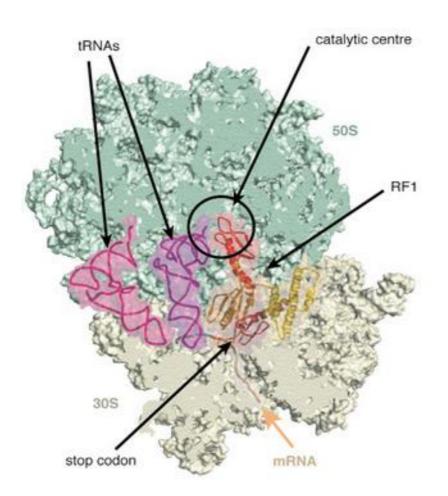


Orgogozo et al 2015 Frontiers Genetics

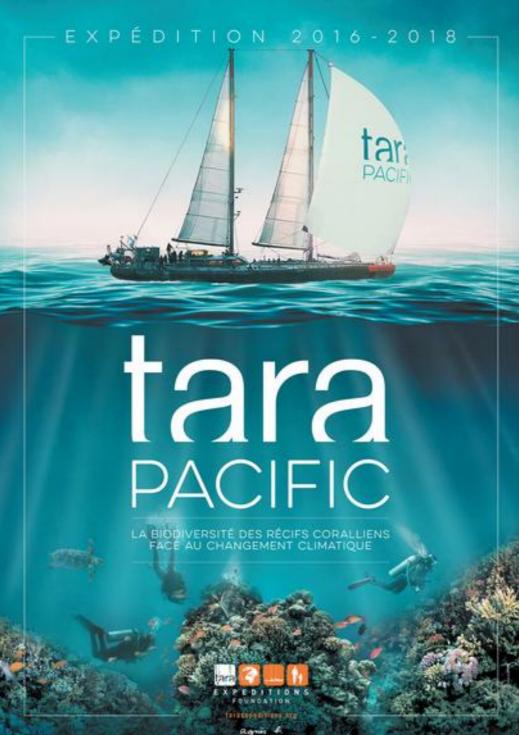
The tree of life, sampling the living world



Ribosome on mRNA



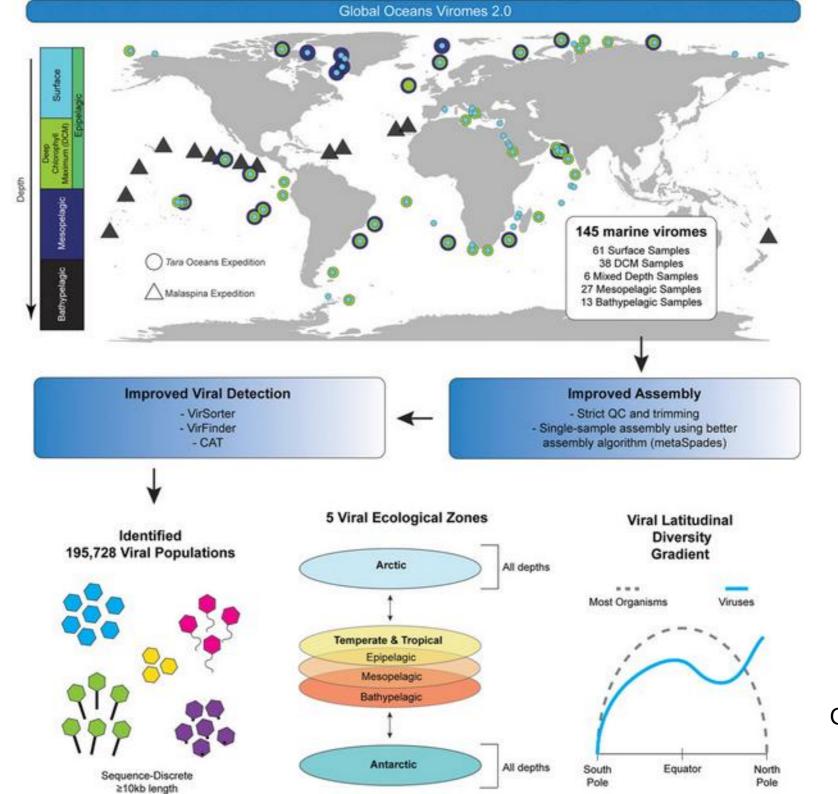
Eukaryote: 18S RNA, part of the 40S subunit Bacteria, Mitochondria : 16S RNA, part of the 30S subunit



QVIOUS Septemen sussesses Loger 🐨 🔄

EMBL

22a



Gregory et al 2019

Living organisms in extreme conditions

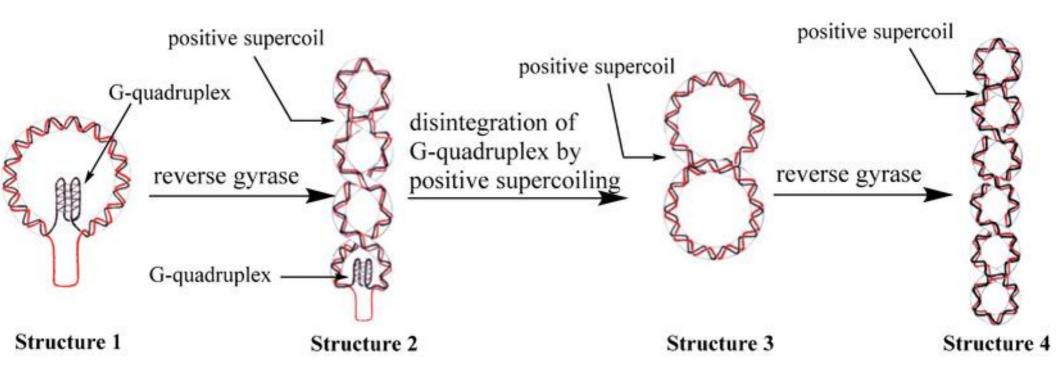
Tardigrad: mountaintops, deep sea, mud volcanoes, tropical rain forests, Antarctic

Temperatures between -272 and +150 °C, pressure up to 6 000 atm, air deprivation, radiation, dehydration, starvation, outer space



Hyperthermophile bacteria

All bacteria living at >80°C have a reverse gyrase enzyme Maintains DNA stability



Ecology

Tracking king penguins

1993

1994

1992

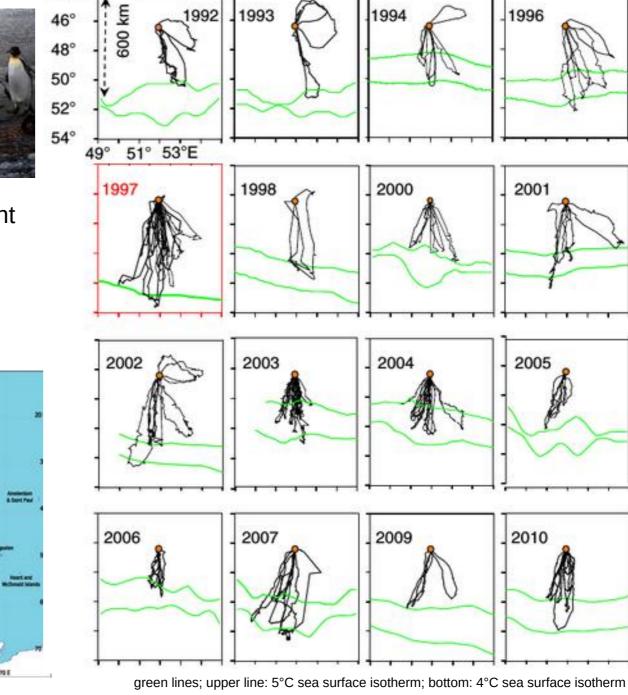
44°S

46°

48°



Must go to the polar front to capture fishes

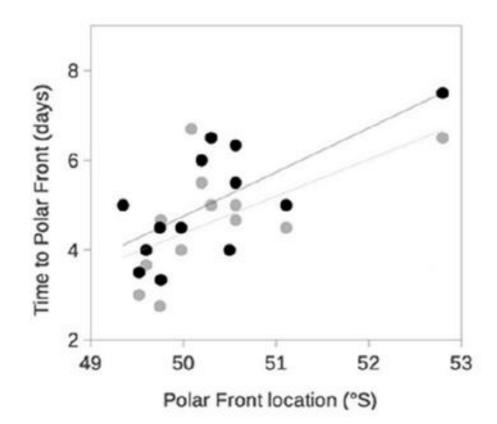


1996



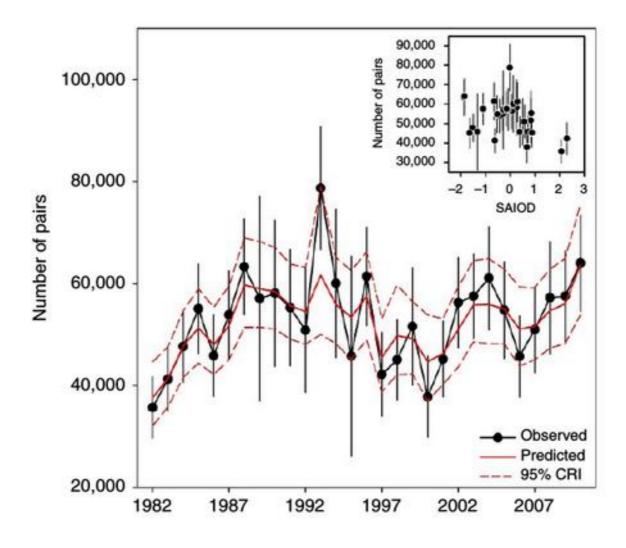
Bost et al 2015

The further, the longer it takes



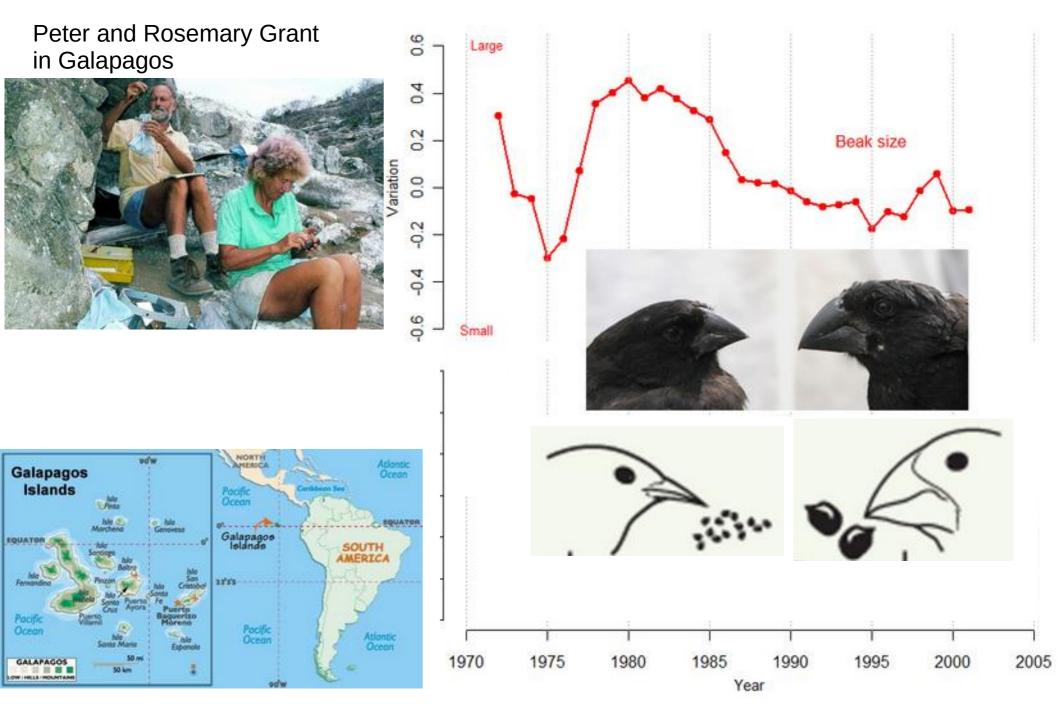
Bost et al 2015

Decrease in population numbers in 1997

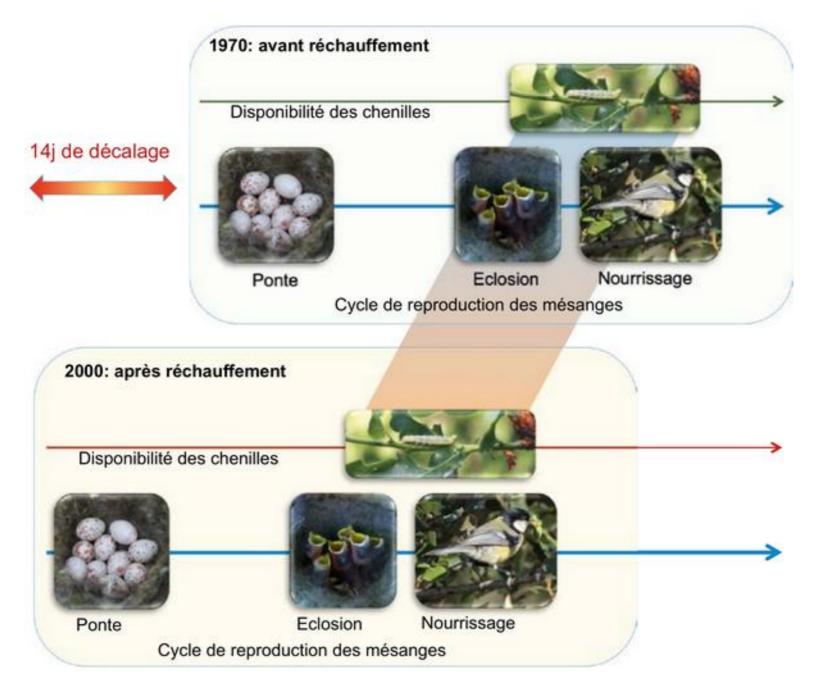


Bost et al 2015

Monitoring birds



Rapid evolution of great tits



Etonnant vivant : découvertes et promesses du XXIe siècle (2017)

proposed geologic timescale dating from the commencement of significant human impact on Earth's geology and ecosystems, including anthropogenic climate change

proposed geologic timescale dating from the commencement of significant human impact on Earth's geology and ecosystems, including anthropogenic climate change

Plastic

Pollution

Transports

Humans move sediments at rates higher than all rivers together (mines, constructions)

Extensive Agriculture (deforestation)

Climate change

Biodiversity crisis

proposed geologic timescale dating from the commencement of significant human impact on Earth's geology and ecosystems, including anthropogenic climate change

Plastic

Pollution

Transports

Humans move sediments at rates higher than all rivers together (mines, constructions)

Extensive Agriculture (deforestation)

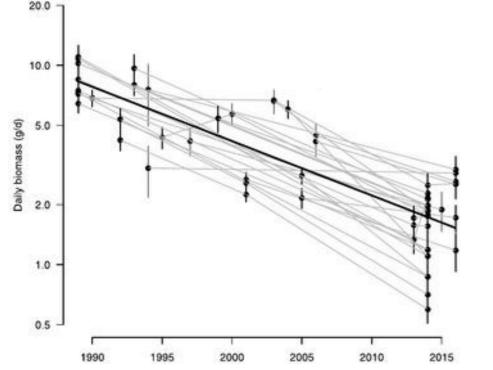
Climate change

Biodiversity crisis

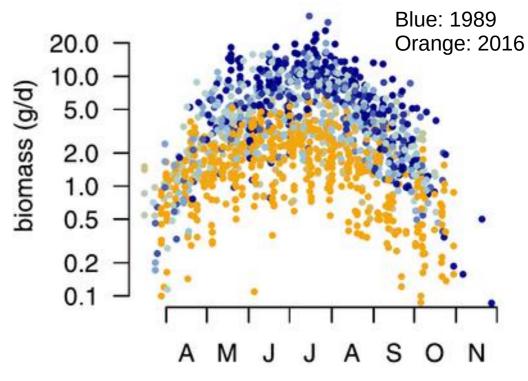
Not all human, certain human activities - Capitalocene (D. Haraway, J. Moore)

75% decline of flying insect biomass in 27 years





Malaise trap 96 unique location-year samplings in Germany from 1989 to 2016 (27 years) ~50 kg of insects collected



Hallmann 2017

Finding causes for the decline

Check various parameters: climate (T, precipitations, frost days, light...), habitat (plants, pH...), land use

Decline = evident throughout the growing season irrespective of habitat type or landscape configuration

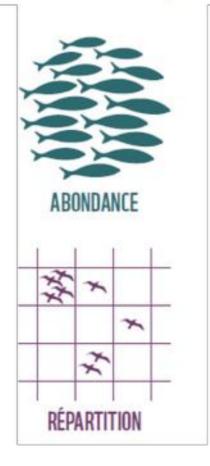
Temperature: increase in 0.5°C, should have increased insect biomass

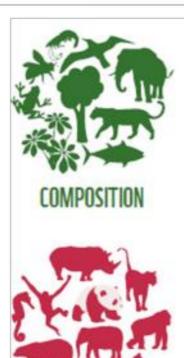
Pesticide usage not tested

Hallmann 2017

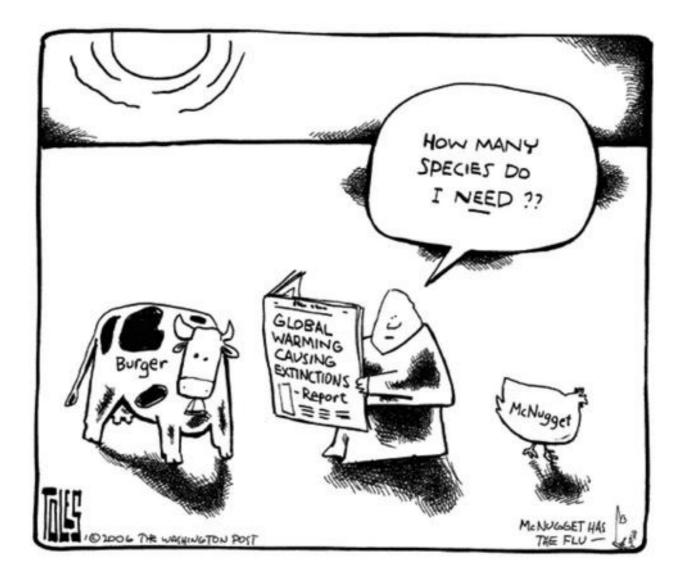
DIFFÉRENTS INDICATEURS, MÊME CONCLUSION

La biodiversité : un concept à multiples facettes qui nécessite de multiples indicateurs





RISQUE D'EXTINCTION



Services ecosystémiques (valeurs extrinsèques)

Support

Production primaire Recyclage des éléments Fertilité des sols Pollinisation Habitat pour les espèces

Approvisionnement

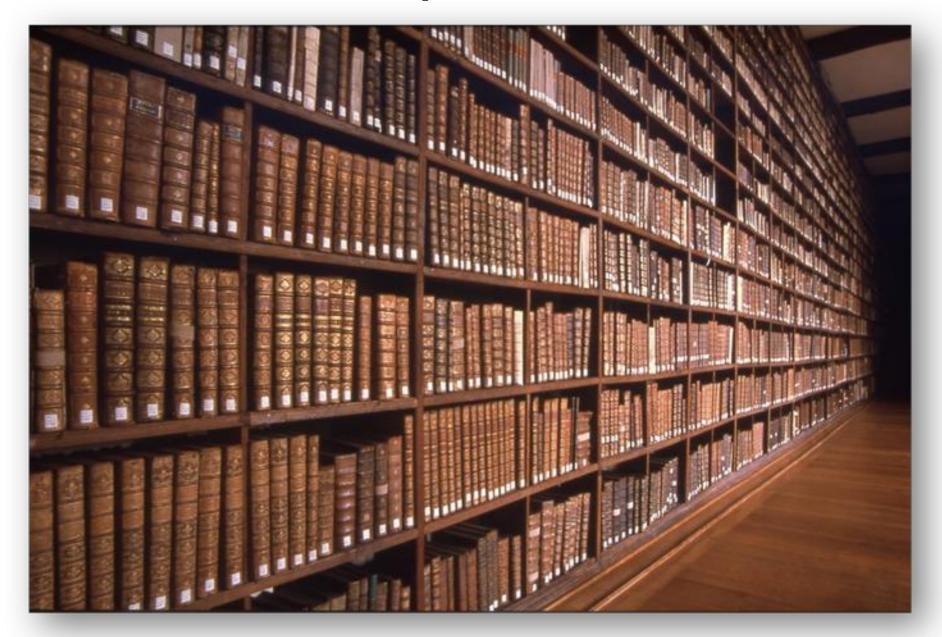
Cultures Elevage Pêche Aquaculture Gibier, Cueillette Bois Ressources génétiques Eau potable

Culture *Ecotourisme Loisirs Ethique Esthétique Education*

Régulation Environnementale

Régulation du climat Purification des eaux et de l'air Régulation des flux hydriques Atténuation des perturbations Contrôle de l'érosion Contrôle biologique Prévention des épidémies

La perte d'une espèce n'est pas comme la perte d'un livre



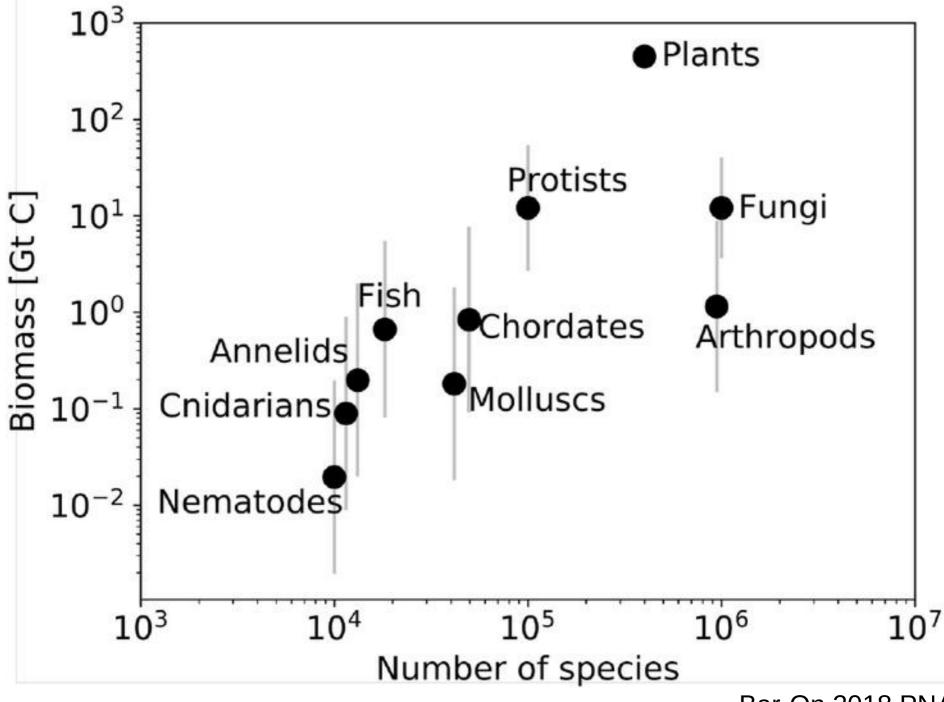
La perte des rivets d'un avion



Le tissu vivant de la planète



http://sogeco31.blogspot.com/2019/08/non-lamazonie-nest-pas-le-poumon-de-la.html



Bar-On 2018 PNAS

From molecules to ecosystems

Characteristics of living organisms

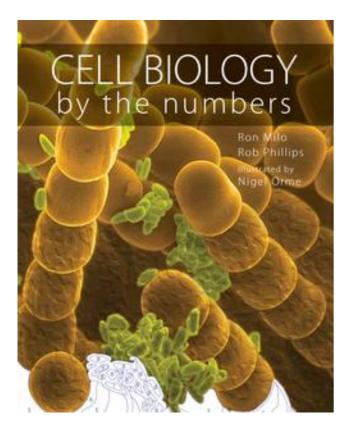
Main molecules in living organisms

Virus, Bacteria, Eucaryote, Archaea

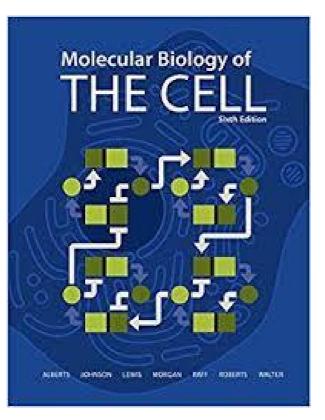
Cell division, Virus amplification

Ecological observations

Anthropocene



Bionumbers.org





inserm

CNRS EDITIONS

CATHERINE JESSUS