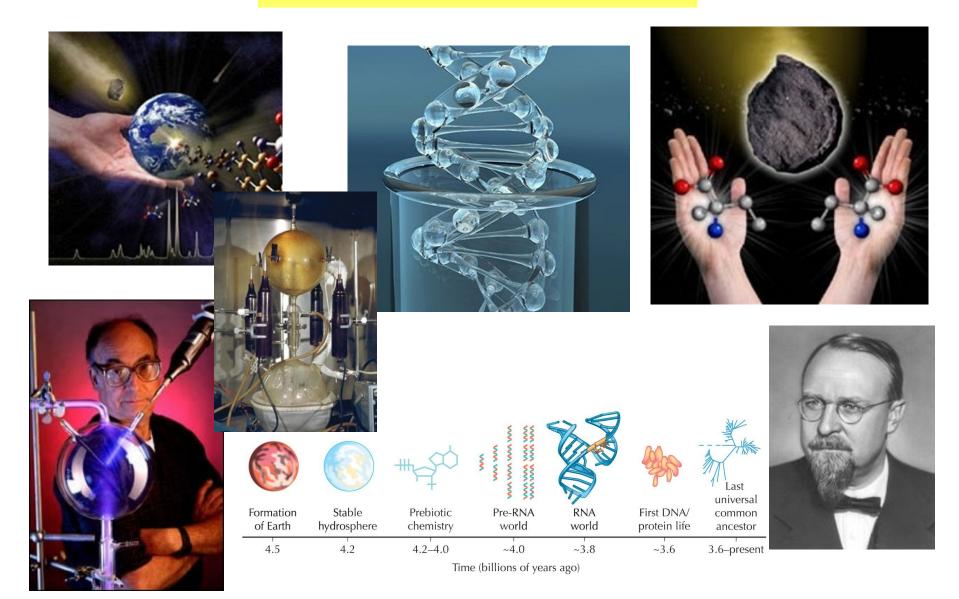
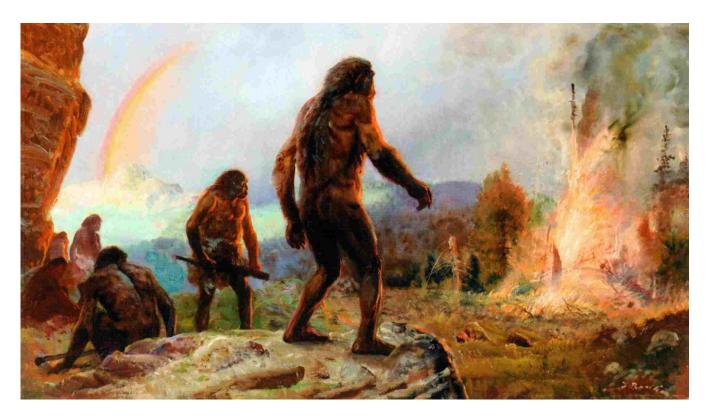
ORIGIN OF LIFE

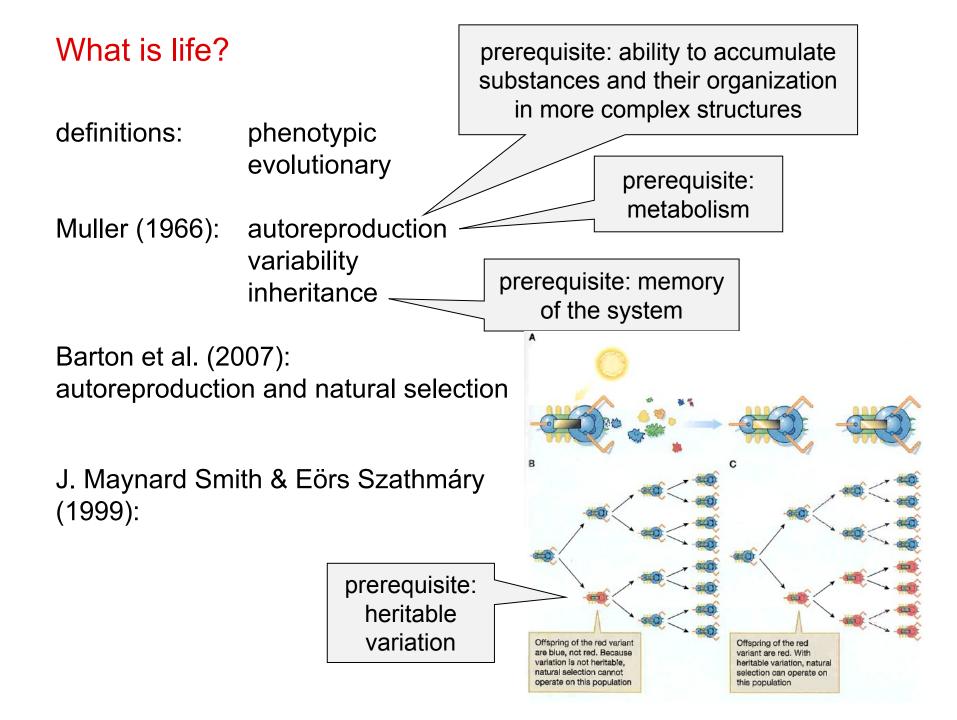


Origin of life is to large extent outside evolutionary biology

→ in essence, it is interdisciplinary study: chemistry (nature of substances composing organisms), geology, study of atmosphere (nature of environment in which life has emerged) etc.

Actually, what is life?





Problem of study of the origin of life:

- J. Monod: evolutionary tinkering, always short-term advantage or coincidence, never long-term perspective × assessment of evolution from backward view, with respect to long-term consequences
- \Rightarrow present-day life cannot help with solving

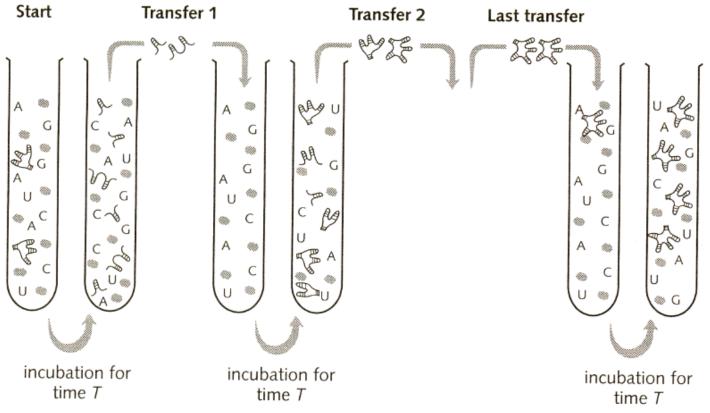
criticisms from creationists: nobody has succeeded to create life in the tube



Evolution in the tube:

Sol Spiegelman et al. (1970):

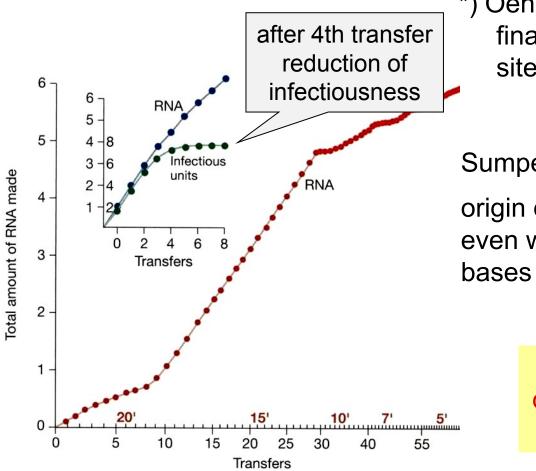
RNA (template ~ 4500 bp) and bacteriophage Q β replicase, + nucleotides





Sol Spiegelman

- \Rightarrow evolution \rightarrow "Spiegelman's monster":
- → reduction of size after 74th transfer to 218 bp \approx 5% size of the original RNA*) \Rightarrow increase of replication rate
- \rightarrow decrease of the ability to infect *E. coli*



 *) Oehlenschläger a Eigen (1997): finally only 48-54 bp (~ binding site for RNA replicase)

Sumper a Luce (1975):

origin of the "Spiegelman's monster" even without template (only RNA bases and $Q\beta$ replicase)

But these experiments don't explain <u>origin</u> of life (enzyme supplied)

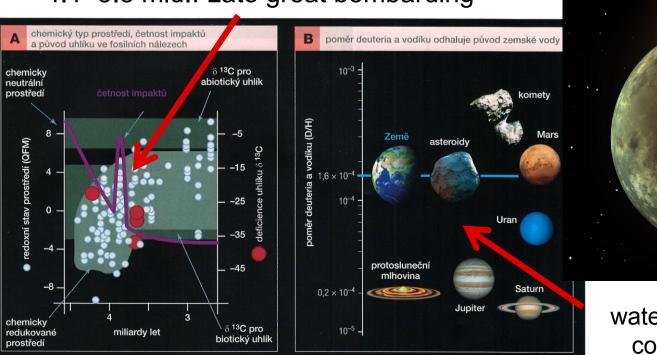
ORIGIN OF LIFE

Age of Solar system: 4.5682×10^9 years

age of primordial Earth: $4.54 \pm 0.04 \times 10^9$ years

Moon: different estimates, oldest 4.5 mld. \rightarrow great oimpact of Earth and Theia^{*)}

^{*)} size of Mars, 10% of Earth mass



4.1–3.8 mld.: Late great bombarding



water on Earth has the same composition as asteroids!

Kejnovský et al., Vesmír 97, 2018

ORIGIN OF LIFE

According to radiometric measurements age of Earth \sim 4,54 \pm 0,04 GYA

(but according to some theories Earth has been created secondarilly and so it is younger)

lower limit: oldest rocks gneiss in Great Slave Lake (Canada) – 4 GY zircon crystals (Australia) – 4,3 GY some meteorites – 4,5 GY end of bombarding of Earth – ~4 GY

upper limit: microfossils, chemical fossils

chert in Warrawoona Group (Z Australia) 3,45 GY: resemblance to present stromatolites ... now questioned



Precambrian stromatolites Siyeh Formation, Glacier Natl. Park

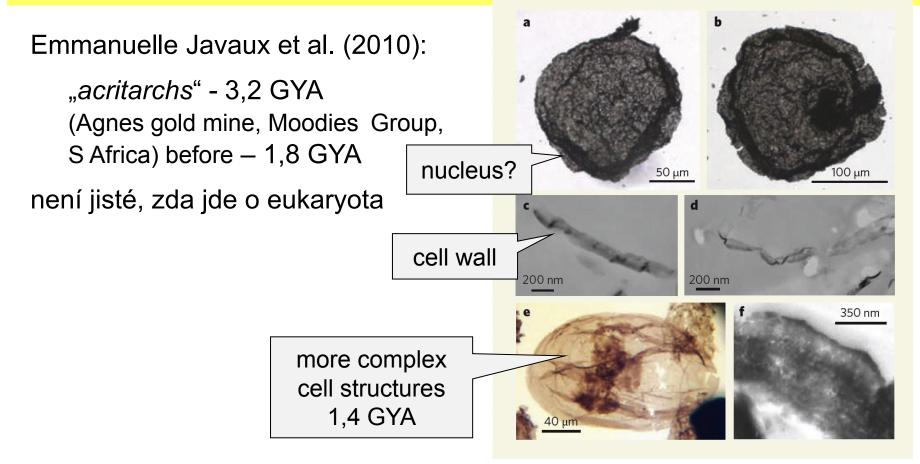


present-day stromatolites Shark Bay, W Australia

chemical fossils – kerogen = organic matter created by decay and transformation of living organisms

Greenland glacier: 3,85 GY, confirmed by C¹²/C¹³ ratio

Conclusion: life has probably emerged during 200 MY between 4 and 3,8 GY



How has life arisen?

origin of simple organic molecules \rightarrow chemical evolution, primitive metabolism

origin of autoreplication, compartmentation and origin of cells, ...

First chemical experiments:

1828: ammonium chloride + silver cyanate + heat → urine
 (= Wöhler synthesis)

1850s: formamide + H_2O + UV, electricity \rightarrow alanine

formaldehyde + NaOH \rightarrow saccharides

 \Rightarrow evidence against vitalism (claims that chemistry in living systems is fundamentally different from non-living, ie. organic \neq inorganic)

Chemical evolution

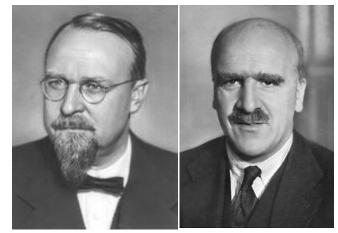
Alexandr Ivanovich Oparin (1924) J. B. S. Haldane (1928)

reducing atmosphere: hydrogen, water, methane, ammonia

Stanley L. Miller, Harold C. Urey (1953):

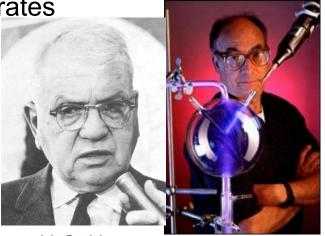
methane + ammonia + H_2 + H_2O \rightarrow 10-15 % carbon in organic compounds 2 % carbon \rightarrow amino acids, lipids, carbohydrates

building components of nucleic acids



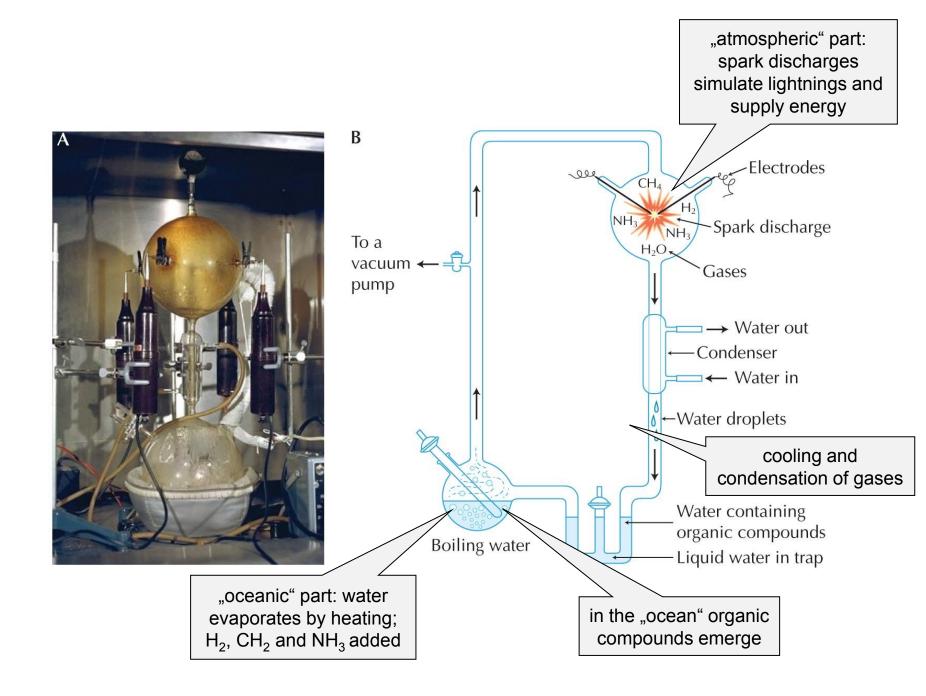
A.I. Oparin

J.B.S. Haldane



H.C. Urey

S.L. Miller



Problems:

according to current knowledge the atmosphere then less reducing: CO_2 , N_2 , H_2O and others \Rightarrow consequently much less molecules arising

not all nucleotides synthesized

phosphorus in nature rare

some compounds in minimal amounts

some products highly unstable (eg. along with ribose also other carbohydrates inhibiting ribose synthesis are produced)

limited production of long polymers

origin of both D and L AA and NA stereoisomers

H₃ +N H₃ +N H H H H R L- amino acid D- amino acid

000

spontaneous origin of ramose, not linear, lipids

Where has life originated?

Darwin: "hot little pond", prebiotic soup

alternatives:

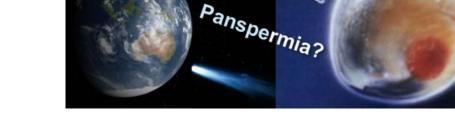
extraterrestric origin:

panspermia: Svante August Arrhenius

existence of organic compounds in universe (comets, meteorites): eg. Murchison meteorite (1969, Australia): 4,6 GYA; many compounds as in the Miller-Urey experiment

bubbles: clouds, sea spume

Thomas Gold (1970): life deep under beneath the surface – existence of extremophilic archaebacteria up to 5 km beneath surface





S. A. Arrhenius

hydrotermal vents = "black smokers"

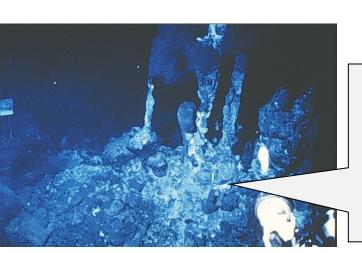
Günter Wächtershäuser

thermal energy rather than Sun chemical synthesis: carbon fixation by chemical energy protection against UV radiation and meteorite impacts fixation of unstable molecules by cold water around the vents



G. Wächtershäuser





1977: thermophilic bacteria and archaebacteria, threemeter tube worms, bivalves, starfish, barnacles, limpets, crabs, annelids, shrimps G. Wächtershäuser:

life on the pyrite surface = the Fe-S world hypothesis

"prebiotic pizza"

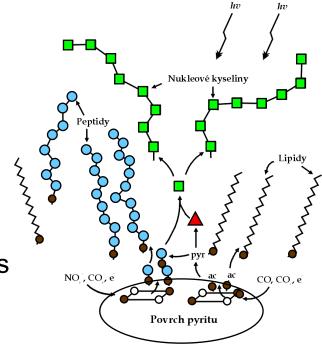
on the pyrite surface molecule clusters [2Fe-2S] or [4Fe-4S] → potential precursors of ferredoxins, pyridoxalphosphates, folates, and cofactors (NAD)

central role of acetyl-CoA

chemoautotrophy

Advantages of flat surface:

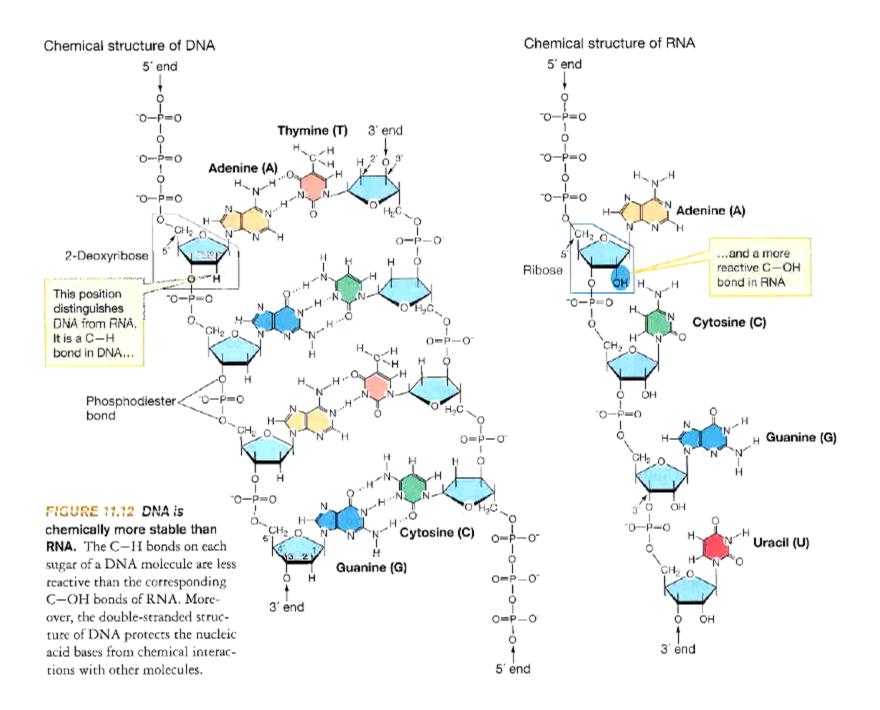
<u>thermodynamics</u>: lower entropy <u>kinetics</u>: higher probability of molecule colisions supply of <u>ions</u> to reactions (not clay!) production of <u>linear</u> lipids easier removing of water molecules



Origin of replicators – RNA world

Experiments of Spiegelman, Sumper and others have shown that on the replicator level there is not only heredity and mutation but also <u>selection</u> but WHAT was replicated?





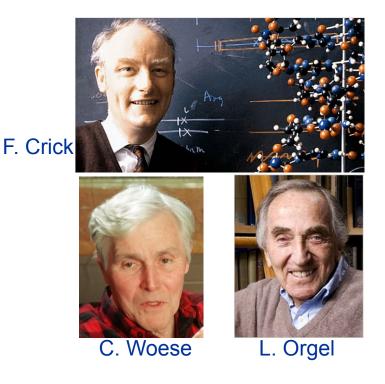
Origin of replicators – RNA world

Experiments of Spiegelman, Sumper and others have shown that on the replicator level there is not only heredity and mutation but also <u>selection</u> but WHAT was replicated?

proteins DNA <u>RNA</u> something else

Francis Crick, Carl Woese, Leslie Orgel (1967):

dual role of RNA: heredity + enzyme = ribozyme



RNA characteristics:

simpler than DNA

Evolution © 2007 Cold Spring Harbor Laboratory Press

absence of complex repair mechanisms

ability to build multiple 3D conformations

more reactive than DNA (OH-group on 2' carbon)



HC

Base

OH

Function	Type of RNA	Role of RNA
Translation	mRNA	Product of DNA transcription
	tRNA	Involved in translation of the genetic cod
	rRNA	Serves as part of a ribosomal subunit
DNA replication	RNA primers	Replication of the lagging DNA strand initiates with an RNA primer
	Telomerase RNA	Needed at the ends of linear chromosomes
Splicing and RNA	Small nuclear RNA (snRNA)	Involved in splicing
processing	Small nucleolar RNA (snoRNA)	Required for posttranscriptional processing of rRNA
	RNase P	Essential for tRNA processing
Translation quality control	tmRNA	Targeting aberrant protein products for degradation in bacteria
Protein translocation	Signal recognition particle (srpRNA)	A component of the signal recogni- tion particle (SRP)
RNA interference (RNAi)	Many types	Involved in regulating RNA stability and translation in euykaryotes
Transcription regulation	6S	Regulates the function of bacterial RNA polymerase

many functions have arisen long ago RNA as "molecular fossils"

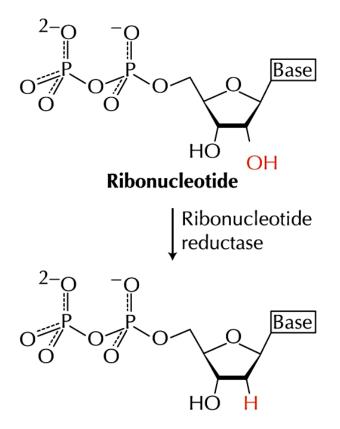
many essential coenzymes, eg. NAD⁺, flavin adenin dinucleotide (FAD)

= ribonucleotide derivates

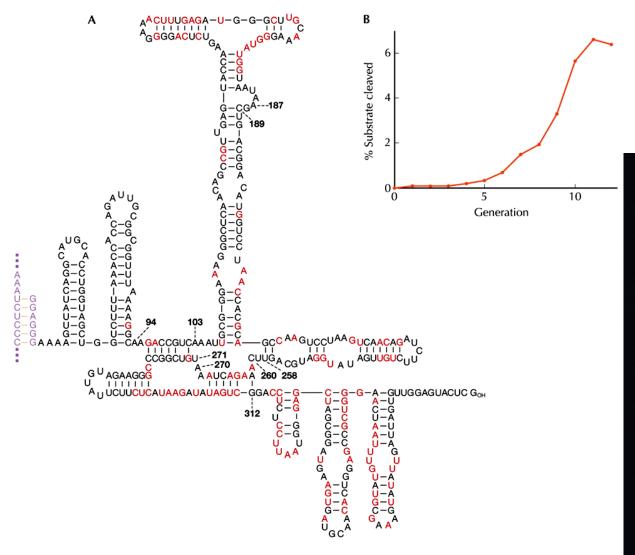
deoxyribonucleotides arise from ribonucleotides

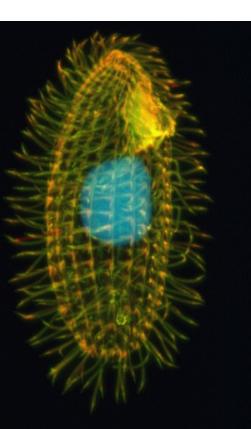
RNA primer is used during DNA replication

ATP = ribonucleotide

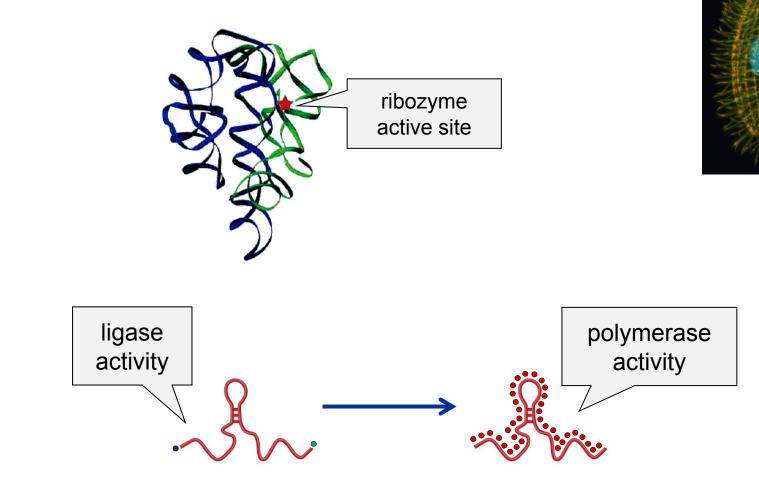


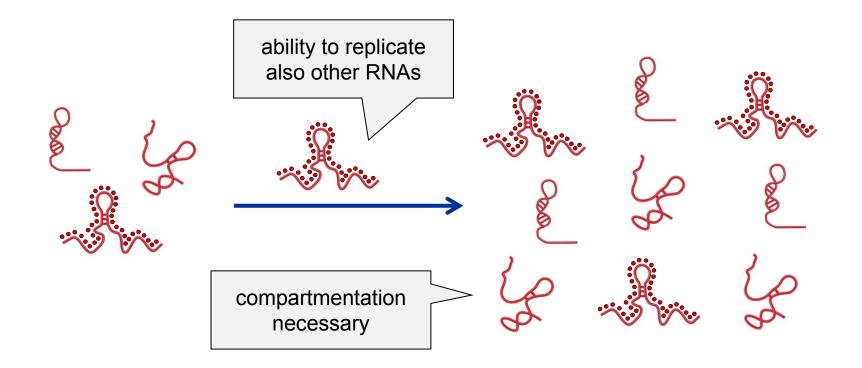
Kruger et al. (1982): self-cleaving of intron in pre-mRNA of Tetrahymena





Kruger et al. (1982): self-cleaving of intron in pre-mRNA of *Tetrahymena* Zaug a Cech (1986): IVS (intervening sequence) \rightarrow ribozyme





Doudna a Szostak (1989): modification of IVS → catalysis of synthesis of complementary strand according to external template – max. 40 nucleotides, only 1% complete

Doudna (1991):

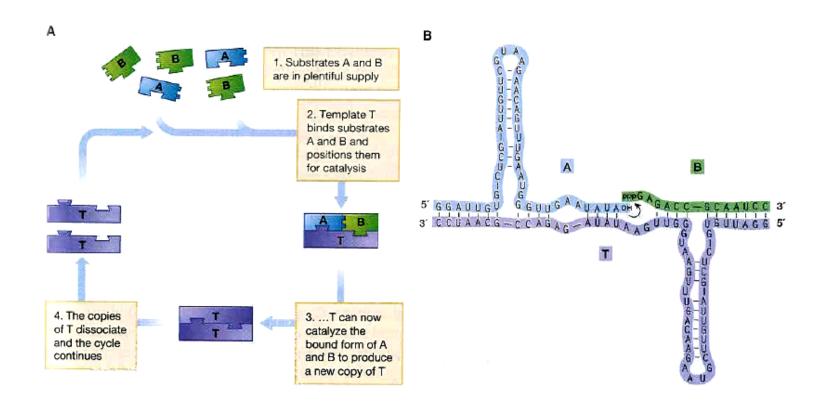
three-subunit ribozyme from sequence of *sunY* of T4 bacteriophage

Paul & Joyce (2002):

R3C ribozyme – ligation of two RNA molecules

R3C modified so that the ligation product is identical to R3C \rightarrow catalysis of own replication

× only two rounds of replication and absence of selection (no variation) \rightarrow these problems later solved (Lincoln & Jozce 2009)



Some known natural ribozymes:

peptidyl transferase 23S rRNARNase P introns of groups I and II hairpin ribozyme GIR branching ribozyme leadzyme hammerhead ribozyme HDV ribozyme mammal CPEB3 ribozyme VS ribozyme glmS ribozyme CoTC ribozyme

TABLE 4.4. Ribozymes		
Ribozyme	Description	
Self-splicing introns	Some introns splice themselves by an autocatalytic process. There is also growing evidence that the splicing pathway of GU-AG introns includes at least some steps that are catalyzed by snRNAs.	
Ribonuclease P	This enzyme creates the 5' ends of bacterial tRNAs. It consists of an RNA subunit and a protein subunit, with the catalytic activity residing in the RNA.	
Ribosomal RNA	The peptidyl transferase activity required for peptide bond formation during protein synthesis is associated with the 23S rRNA of the large subunit of the ribosome.	
Virus genomes	Replication of the RNA genomes of some viruses involves self-catalyzed cleavage of chains of newly synthesized genomes linked head to tail. Examples are the plant viroids and virusoids and the animal hepatitis delta virus. These viruses form a diverse group with the self-cleaving activity specified by a variety of different base-paired structures, including a well-studied one that resembles a hammerhead.	
Telomeres	In some species, replication of DNA ends is catalyzed by an RNA subunit of its telomerases.	

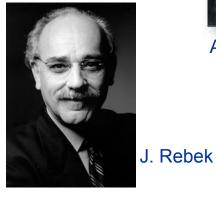
From Brown T.A. 2002. *Genomes*, 2nd ed., Table 10.4, BIOS Scientific Publishers Ltd., Oxford. snRNA, small nuclear RNA; tRNA, transfer RNA.

Alternatives to nucleic acids:

Alexander Graham Cairns-Smith: crystalic clay as <u>"urgene</u>" – initially anorganic replication, a kind of "scaffold"

Julius Rebek: autoreplication using AATE (amino adenosin triacid esther)

Ronald Breaker (2004): DNA can bahave as ribozymes





A.G. Cairns-Smith

Problem of ribozyme-aided replication:

Manfred Eigen (1971):

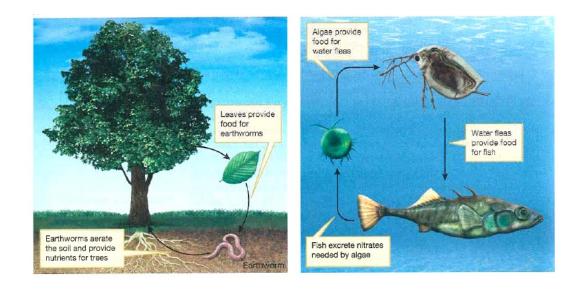
individual genes will compete

without repair mechanisms maximum size of replicating molecules \approx 100 bp

length of DNA segment encoding functional enzyme much exceeds 100 bp

= Eigen paradox

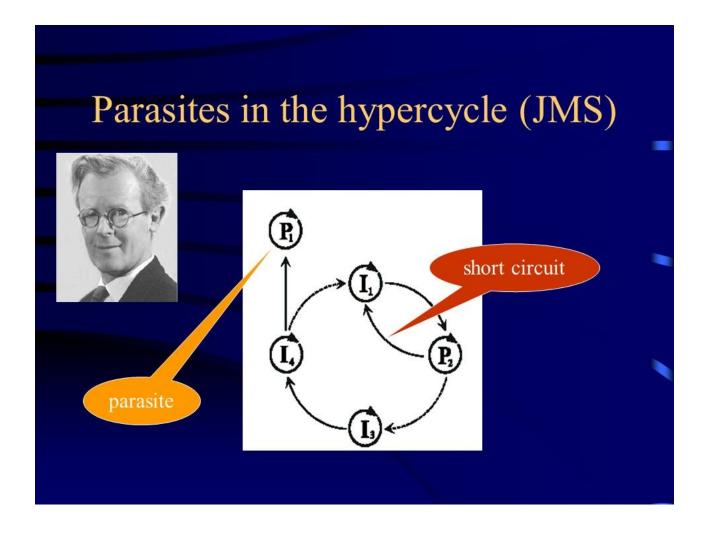
hypercycles = stable coexistence of two or more cooperating replicators



hypercycles:

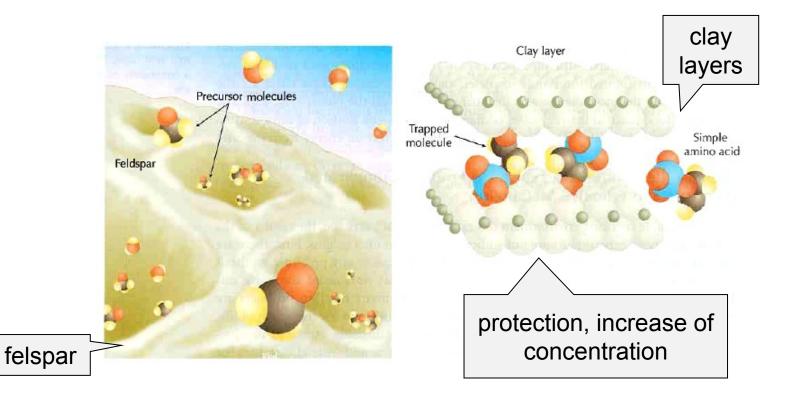
molecular mutualism: reciprocal altruism (win-win relationship)

competition of the whole systém with other cycles risk of systém "parasitation" ⇒ need for compartmentation



Compartmentation and origin of cells

role of tiny crevices and unevenness of the mineral surface



Compartmentation and origin of cells

role of tiny crevices and unevenness of the mineral surface

```
proteins: microspheres (Sidney W. Fox)
```

lipids: spontaneous production of liposomes

spontaneous production of lipidic membranes: "oil on water" \rightarrow "water on oil"

 $\mathsf{semi-cell} \to \mathsf{proto-cell} \to \mathsf{cell}$

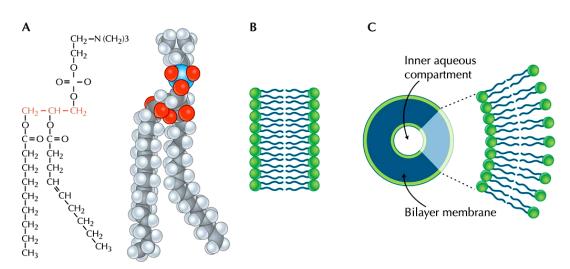


FIGURE 4.14. Lipids. (*A*) General structure of phospholipids. Phospholipids are made up of fatty acids, glycerol, and a phosphate group. They are amphipathic, with one hydrophobic end and one hydrophilic end. (*B*) Bilayers form when phospholipids spontaneously aggregate in water. The hydrophobic ends attract each other in the center of the layer and the hydrophilic ends are surrounded by water. (*C*) Liposomes are formed when a lipid bilayer folds over itself.

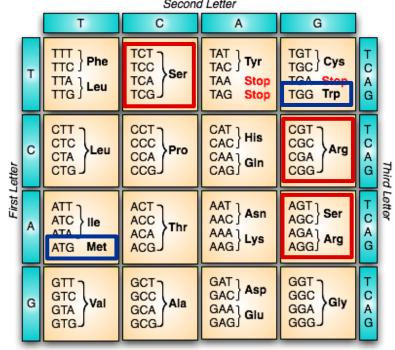


Origin of chromosomes and genetic code

fusion of replicators \Rightarrow longer replication \Rightarrow selective disadvantage possible benefits:

- 1. reduction of competition between functionally connected replicators
- 2. products of functionally connected replicators at the same place

genetic code: redundant, redundancy random (Ser, Arg, Leu: 6 codons × Met, Trp: 1 codon)

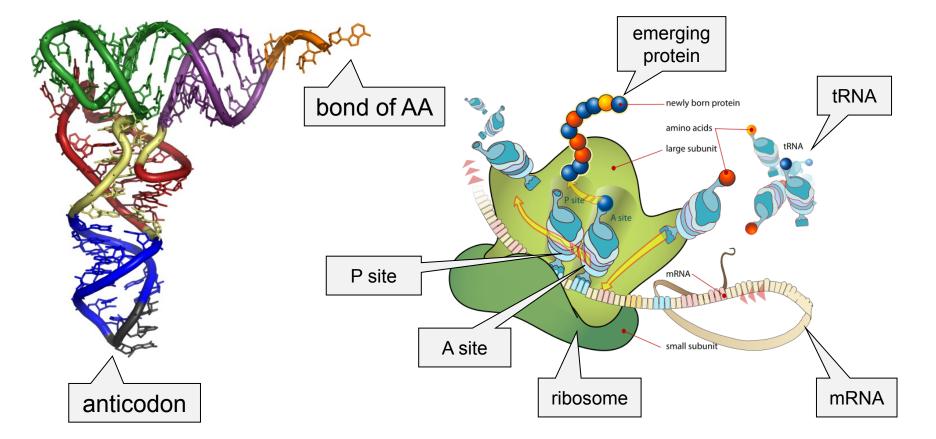


chemically related AAs \rightarrow similar code

genetic code is not by far "universal" – exceptions in some organisms (eg. *Mycoplasma*) or organelles (mitochondria)

AAs perhaps initially helped to stabilize RNAs or

as enzymatic co-factors enhancing RNA activity \rightarrow step by step emergence of function in translation system



Association AAs and RNAs:

synthesis of proteins governed by RNA mapping of RNA sequence onto AA origin of tRNA

"frozen accident" – F. Crick (1968)

some RNA molecules evolved ability to transfer AA to other RNA selection gradually favours one or more RNAs for each AA association between AA and RNA <u>random</u>

stereochemical theory: Carl Woese

different RNAs tend to preferentially bind particular AAs \rightarrow some experiments show that RNA molecules may be selected according to their preferential bond to particular AAs

Transition RNA \rightarrow DNA

RNA world: RNA = both genotype and phenotype

with translation proteins adopt most catalytic RNA functions (they can create broader range of polymers) ⇒ much more diverse catalytic activities → eg. no RNA molecule can catalyze redox reactions or break C–C bonds

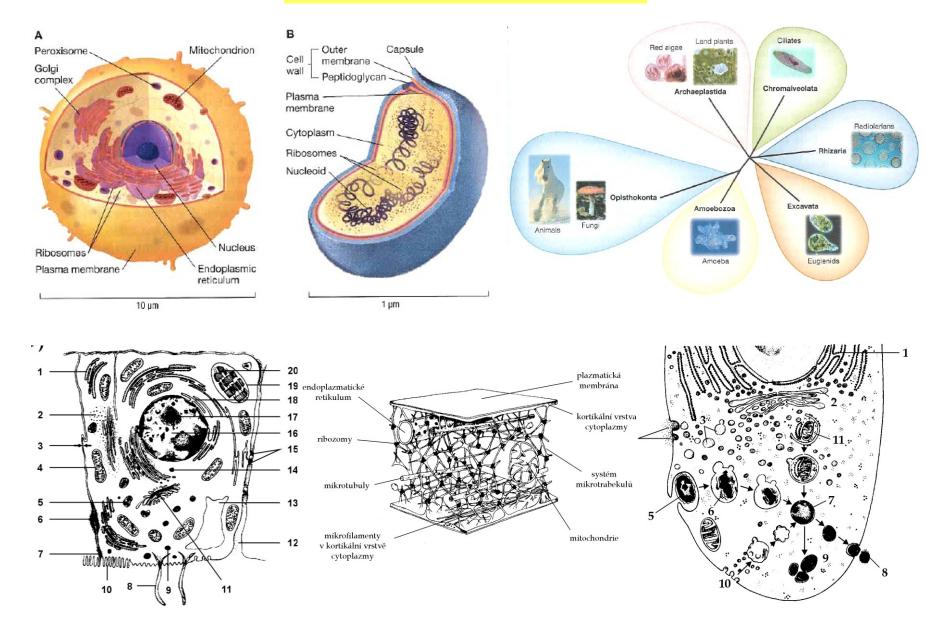
DNA advantages:

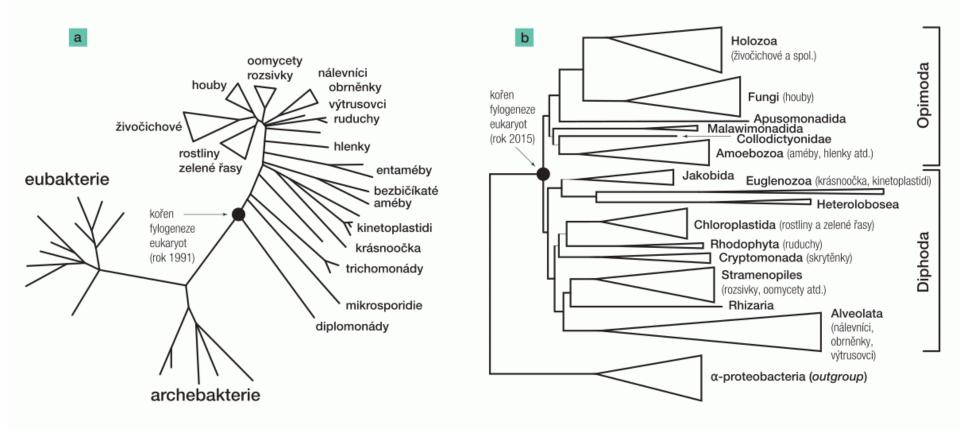
lower reactivity \Rightarrow higher stability \Rightarrow longer genes

division of labour between RNA and DNA

with loss of genetic function RNA could have carry out catalytic and structural functions with smaller restrictions

Origin of eukaryotic cell





Thomas Cavalier-Smith:

loss of cell wall \Rightarrow necessity to create endoskeleton \Rightarrow flexibility, movement, fagocytosis

invagination of membrane $\rightarrow \text{ER}$

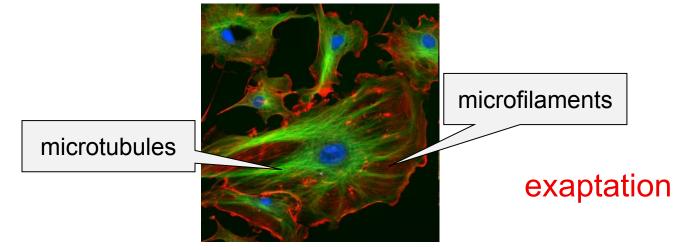
Prokaryotic cytoskeleton:

FtsZ: tubuline analogue, function in cell division

MreB: actin analogue, rod-shaped cell shape

Crescentin: analogue of intermediary microfilaments, helix creation

MinD, ParA: no analogue, cell division, separation of plasmids





Origin of cell organelles:

Konstantin Sergeyevich Merezhkovsky (1905, 1909):

term symbiogenesis

chloroplasts = originally alien organisms

(Andreas Schimper, 1883: similarity between chloroplasts and cyanobacteria; Richard Altmann, 1890: mitochondria [bioblasts] = originally bacteria)

first animal cell: anucleate amoeba + bacterium (nucleus)

Lynn Margulis (1966, 1970): endosymbiosis

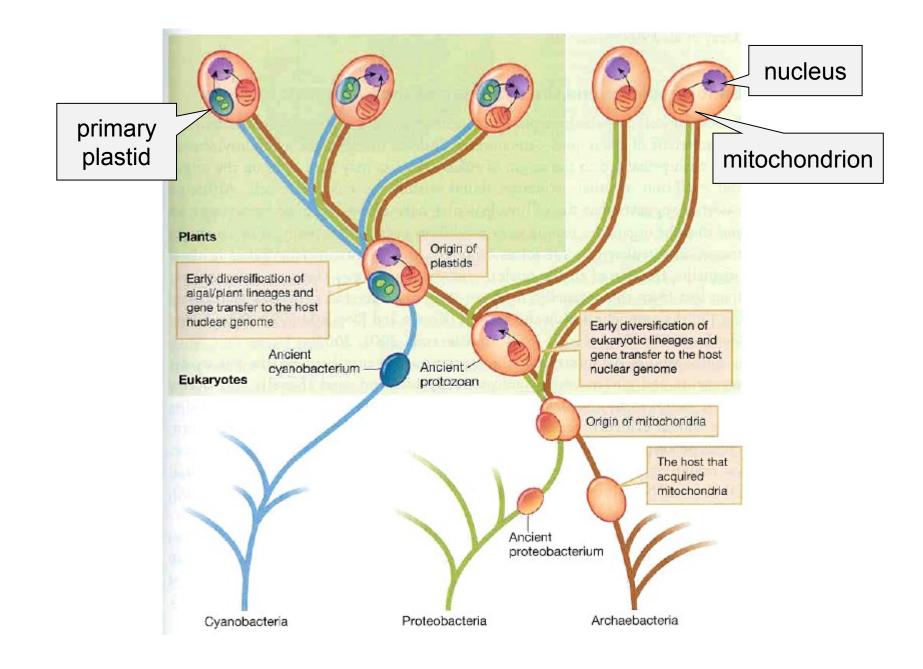
mitochondria: bacteria related to rickettsias or other α -proteobacteria (eg. *Rhodospirillum*), gradually loss of photosynthesis

chloroplasts: cyanobacteria, loss of respiration



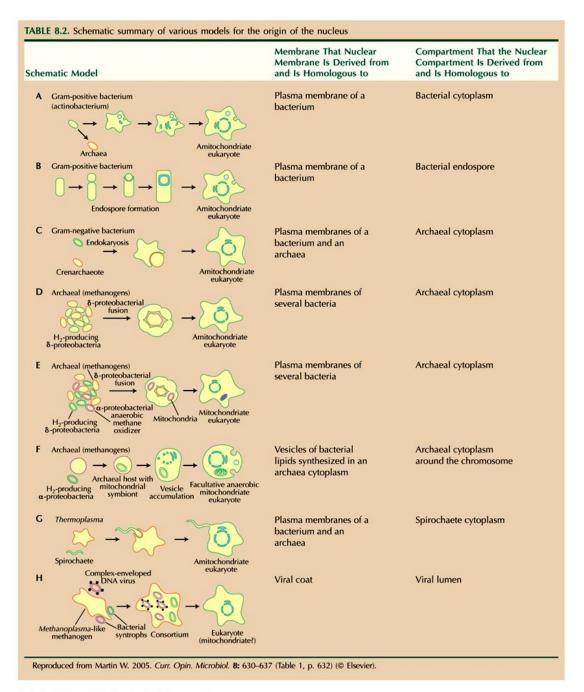


Lynn Margulisová



Theories of the origin of nuclear membrane:

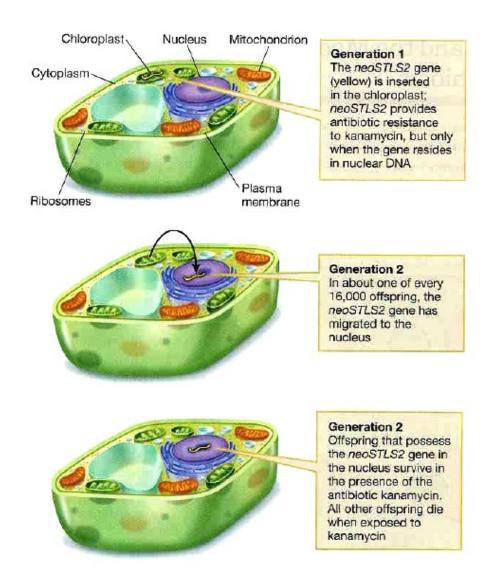
- 1. fusion of vesicles of cytoplasmatic membrane
- 2. fusion of eubacterium and archaebacterium (archaebacterial membrane = nuclear, bacterial membrane = cellular)
- 3. viral origin (several alternatives) ... controversial
- 4. first origin of the 2nd cytoplasmatic membrane, from the inner eventually nuclear membrane



Evolution © 2007 Cold Spring Harbor Laboratory Press

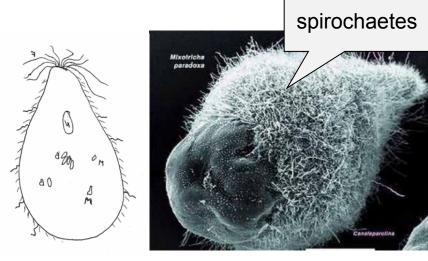
Transfer of genes to nucleus:

eg. *neoSTLS2* gene, tobacco chloroplast \rightarrow in 16 of 250 000 (\approx 1/16 000) daughter cells transfer of the gene to nucleus \Rightarrow kanamycin resistence



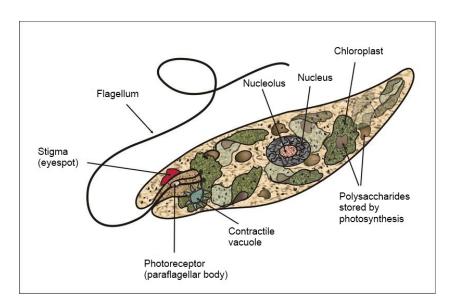
peroxisomes: G+ bacteria

microtubules: spirochaetes × současné poznatky nepotvrzují



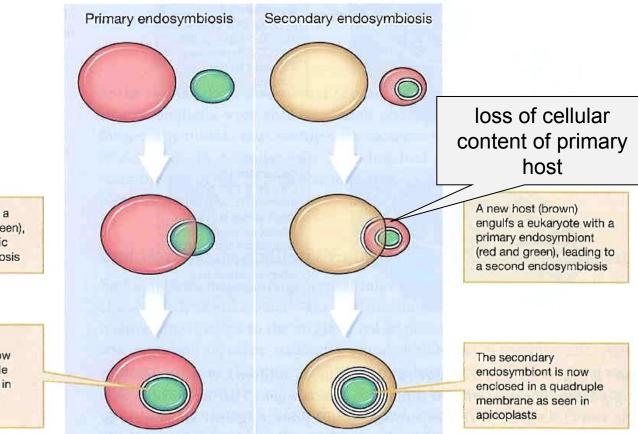
Mixotricha paradoxa

secondary and tertiary endosymbiosis \rightarrow complex plastids: eg. euglena + green alga





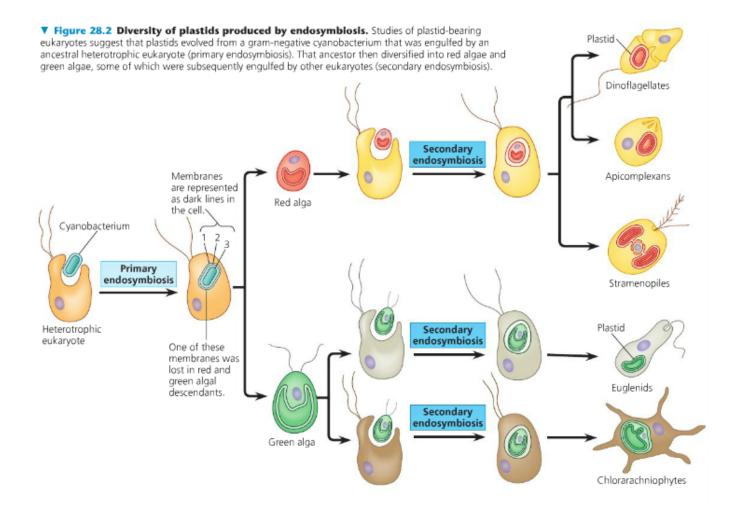
Secondary endosymbiosis:

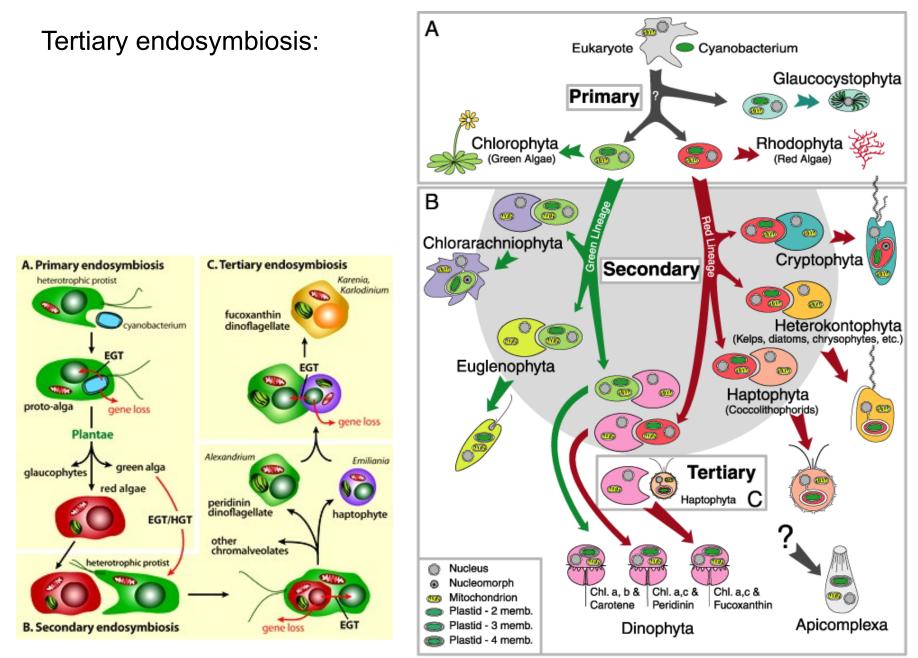


A host (red) engulfs a cyanobacterium (green), creating a eukaryotic primary endosymbiosis

The primary endosymbiont is now enclosed in a double membrane as seen in mitochondria and chloroplasts

Secondary endosymbiosis:

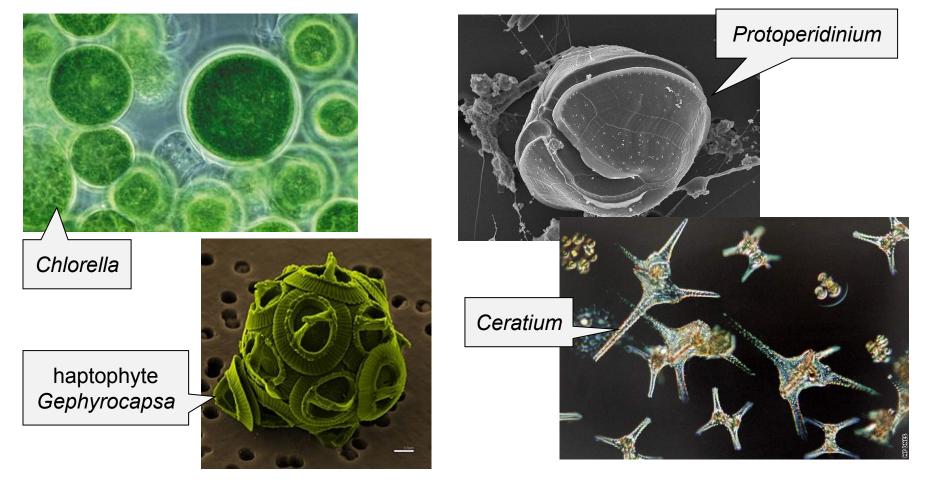




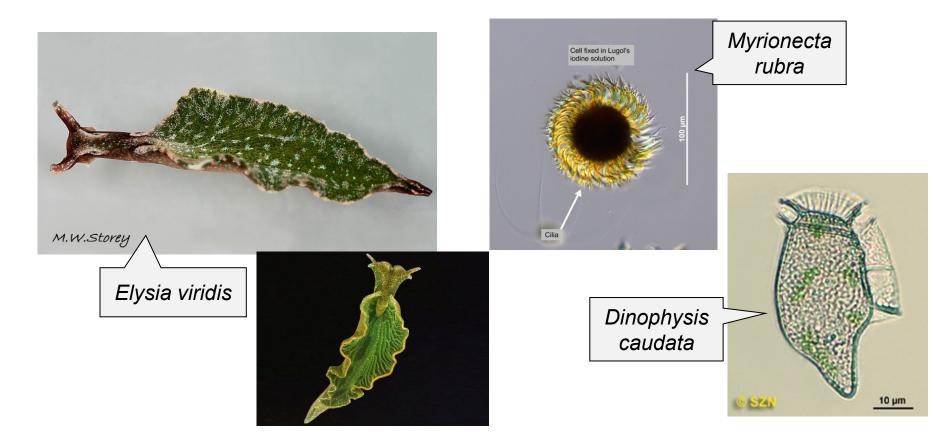
Modified from Delwiche, C.F. 1999. Tracing the thread of plastid diversity through the tapestry of life. Am. Nat. 154:S164-S177.

sometimes the existence of a secondary endosymbiont can be revealed only according to presence of its DNA (eg. chlamydia genes in plant plastids and primary algae)

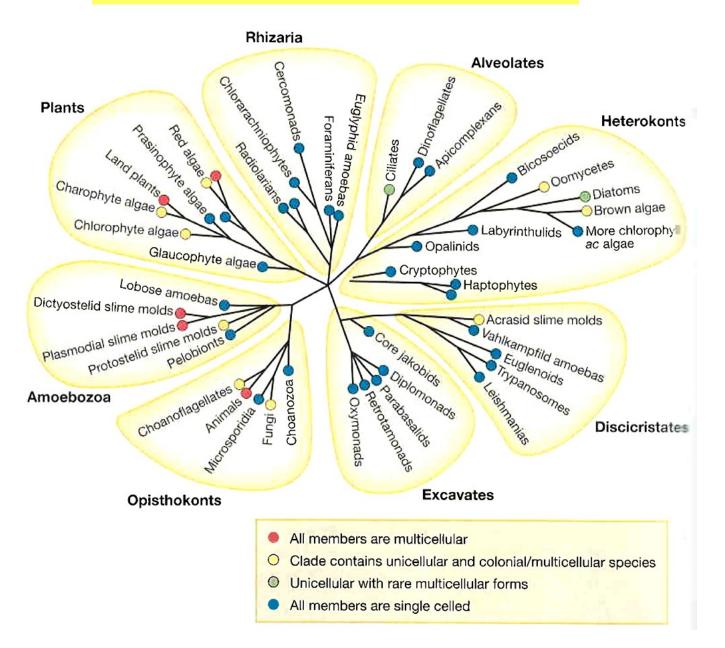
in other cases endosymbionts still able of independent life, eg. photosynthetic algae (chlorellas, dinoflagellates, haptophytes) in cells of corals, foraminiferans, radiolarians, and some ciliates

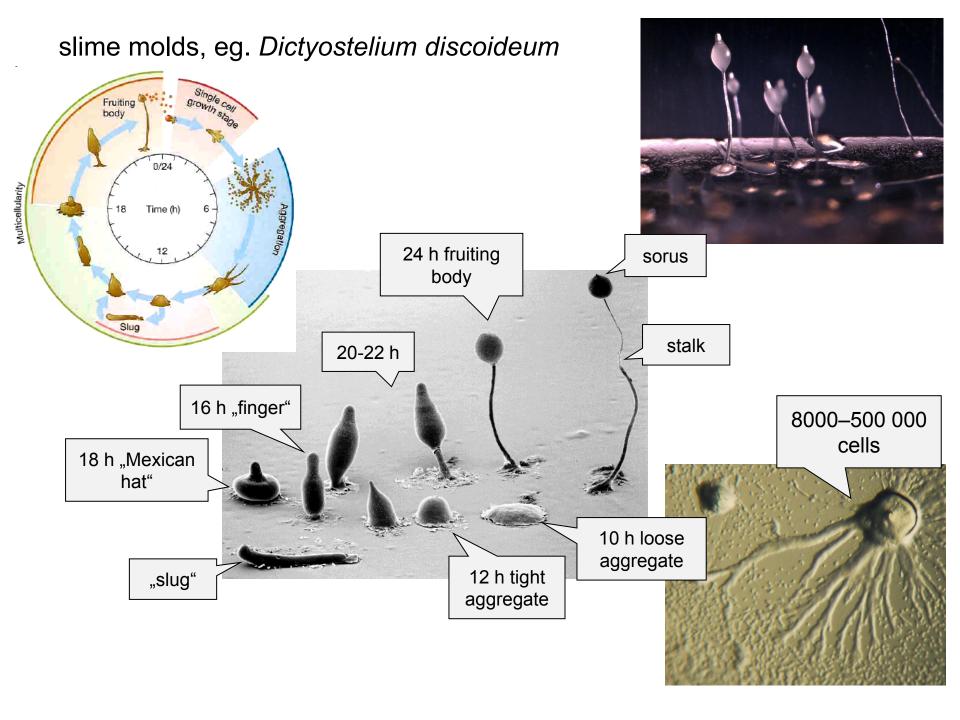


"kleptoplastids" (eg. ciliate *Myrionecta rubra*, dinoflagellate of the genus *Dinophysis*, marine gasteropod *Elysia viridis*)



Origin of multicellular organisms





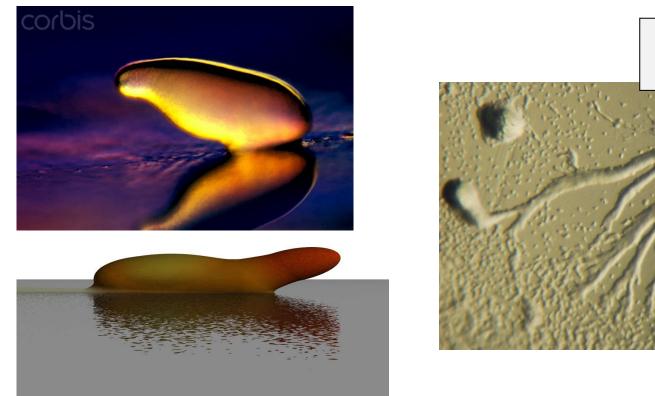
How can "slug", composed of independent amoebas, orient itself in its environment?

cAMP (cyclic adenosine monophosphate): emission in area of the densest aggregation \rightarrow signal for "downstream" cells \rightarrow gradual aggregation

cAMP emission

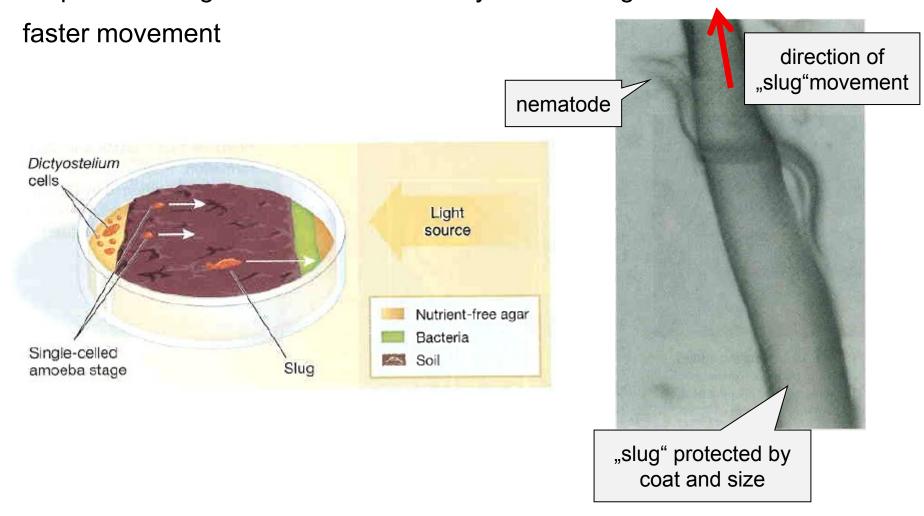
production of protein enabling mutual attachment of amoebas

reaction to external stimuli: light, teperature, oxygen and ammonium in the soil



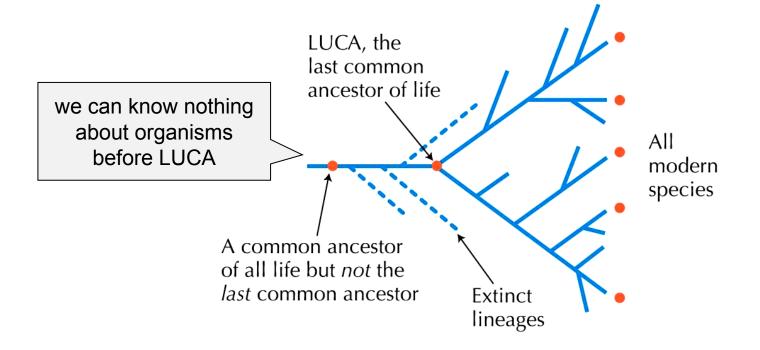
Advantages of *D. discoideum* aggregation?

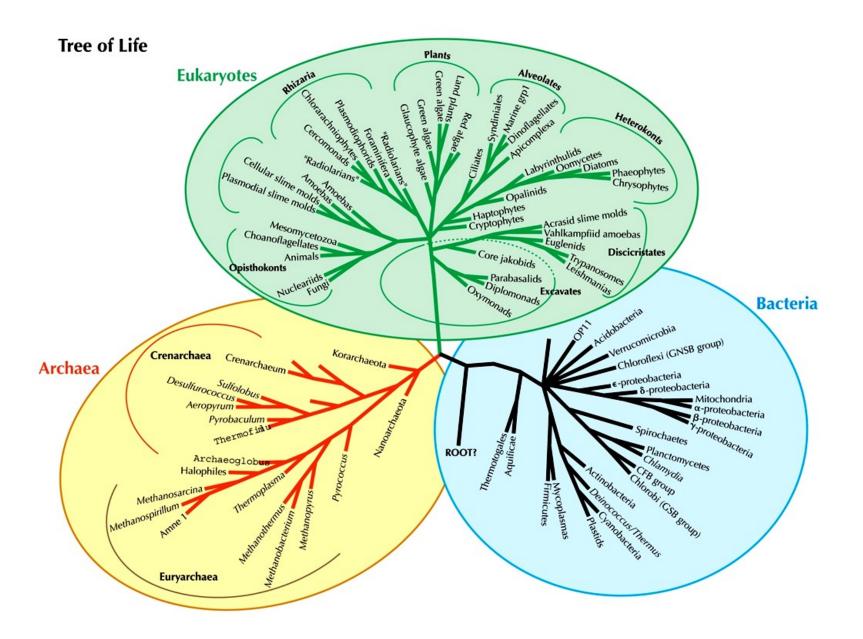
production of coat made of cellulose and substances rich of proteins \rightarrow protection against nematodes – only on the "slug" surface

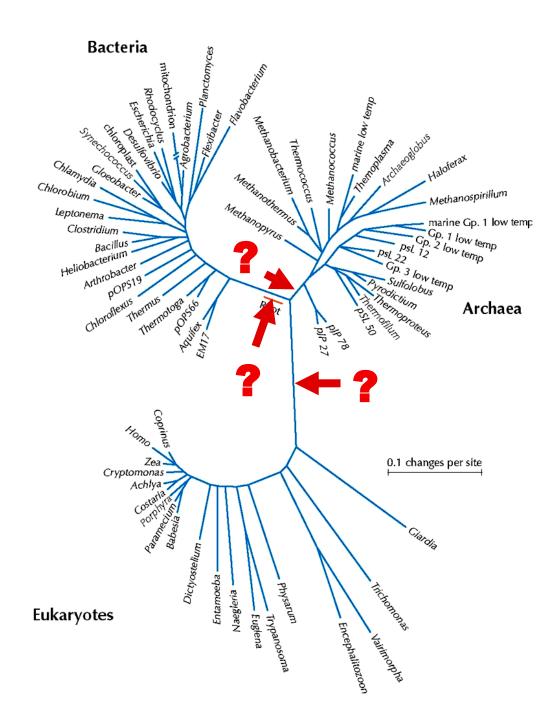


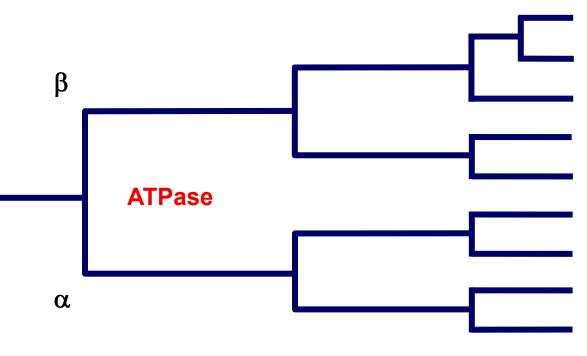
Tree of life:

Last Universal Common Ancestor (LUCA)

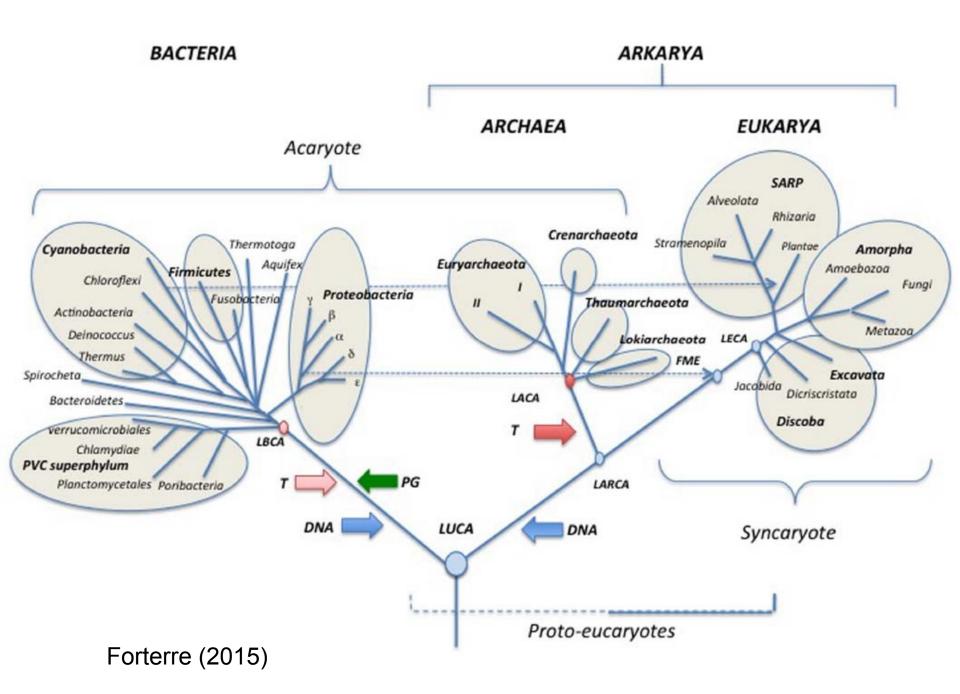




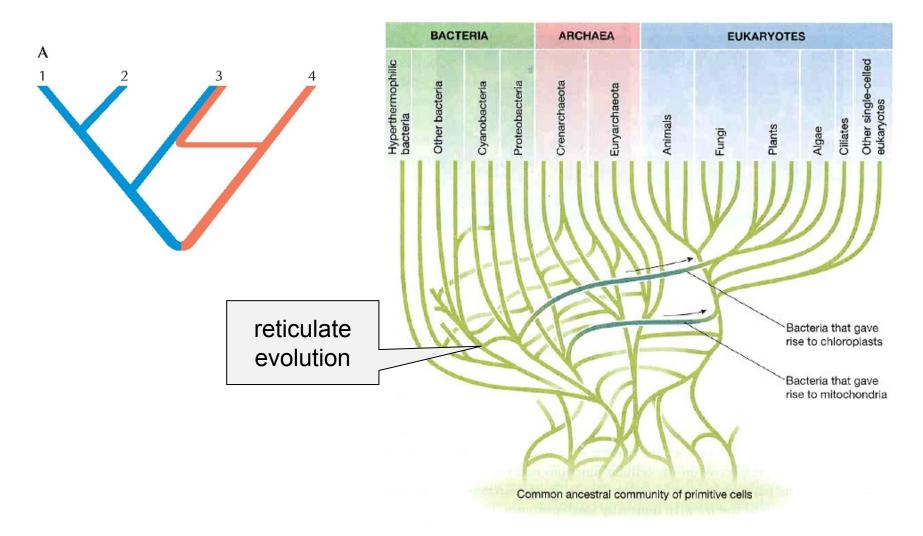




Eubacteria (*E. coli*)
mitochondria (cows)
chloroplasts (tobacco)
Archaea (*Sulfolobus*)
Eukaryotes (plants, fungi)
Eubacteria (*E. coli*)
chloroplasts (tobacco)
Archaea (*Sulfolobus*)
Eukaryotes (plants, fungi)



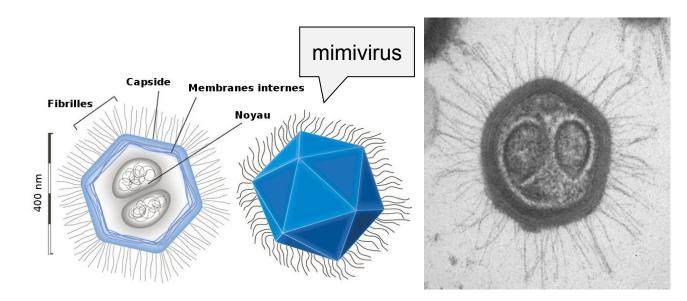
Horizontal transfer of genes



 \Rightarrow no LUCA of recent organisms \times trees for individual genes <u>can</u> have LUCA

Where to place viruses on the Tree of Life?

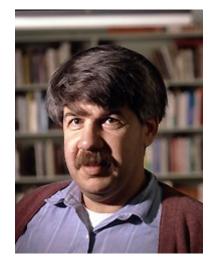
- relics of pre-cellular world: some processes and genes ancient many genes only very distantly related to their cellular counterparts × how could they independently exist in pre-cellular world?
- 2. similarity with transposons \rightarrow "escaped" parts of cellular organisms (eg. RNA or DNA elements, plasmids)
- 3. originally free-living organisms eg. mimivirus: genome size = 1,2 Mb, > 900 proteins, ie. more than some bacteria and archaebacteria!



Increase of complexity:

Stephen Jay Gould:

evolution moves as a drunk person which cannot return to the starting point however he wants even at present most organisms prokaryotic secondary simplification (eg. parasites)



X

John Maynard Smith a Eörs Szathmáry:

the contingent irreversibility theory): steady tendency to increase

of complexity major transitions complexity emrges withou selection

ratchet





Major transitions in evolution:

origin of replicators compartmentation, origin of cell origin of chromosomes origin of genetic code, DNA origin of Eukaryotes origin of sex multicellularity societies origin of language

JOHN MAYNARD SMITH & EÖRS SZATHMÁRY THE MAJOR TRANSITIONS IN EVOLUTION



Individuals cease to reproduce independently
Bigger size → bigger pray, specialization, division of labour
Origin of more effective ways of acquiring, processing, transmission, and saving of informations

But advantages of a transition to a "higher level" do not imply group selection!

conflict of selections at different levels:

replication control \times B chromosomes, transposition fair meiosis \times meiotic drive differentiation of somatic cells \times oncogenic growth non-reproducing castes \times egg-laying workers