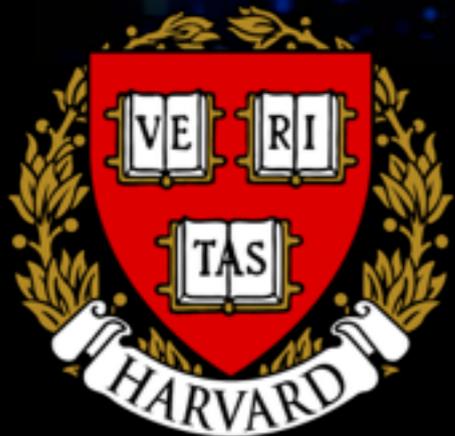


Life Resurrected

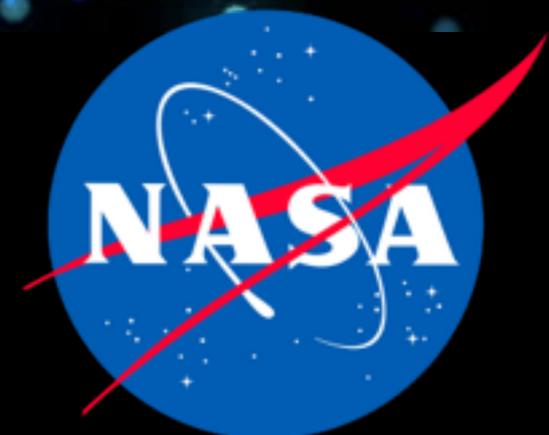
Traveling backwards in evolutionary time

Betul Kacar (Harvard)

Host: NASA



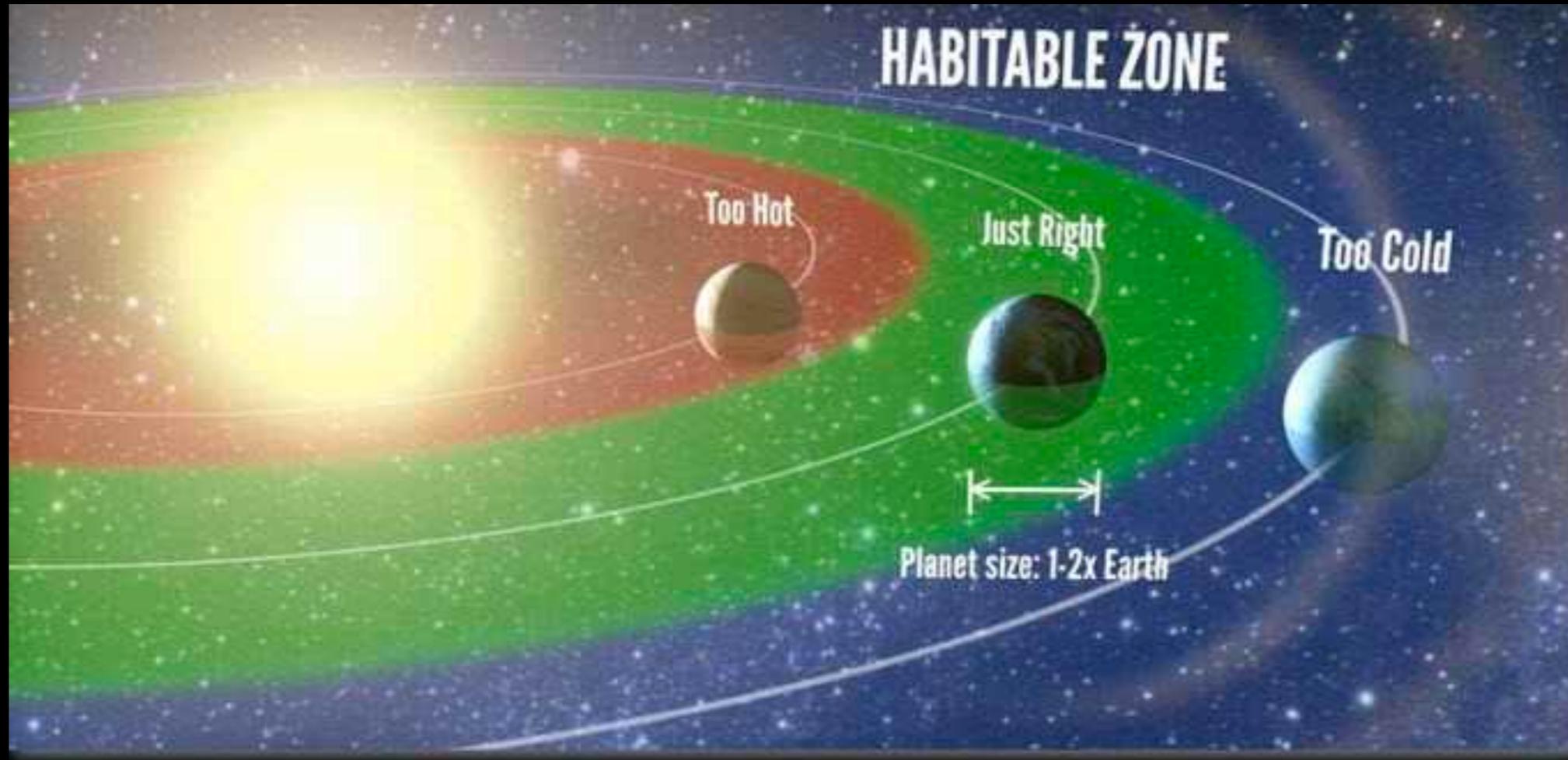
John
Templeton
Foundation



- **Are we alone?**
- **Is there life elsewhere?**
- **How did life on Earth begin?**

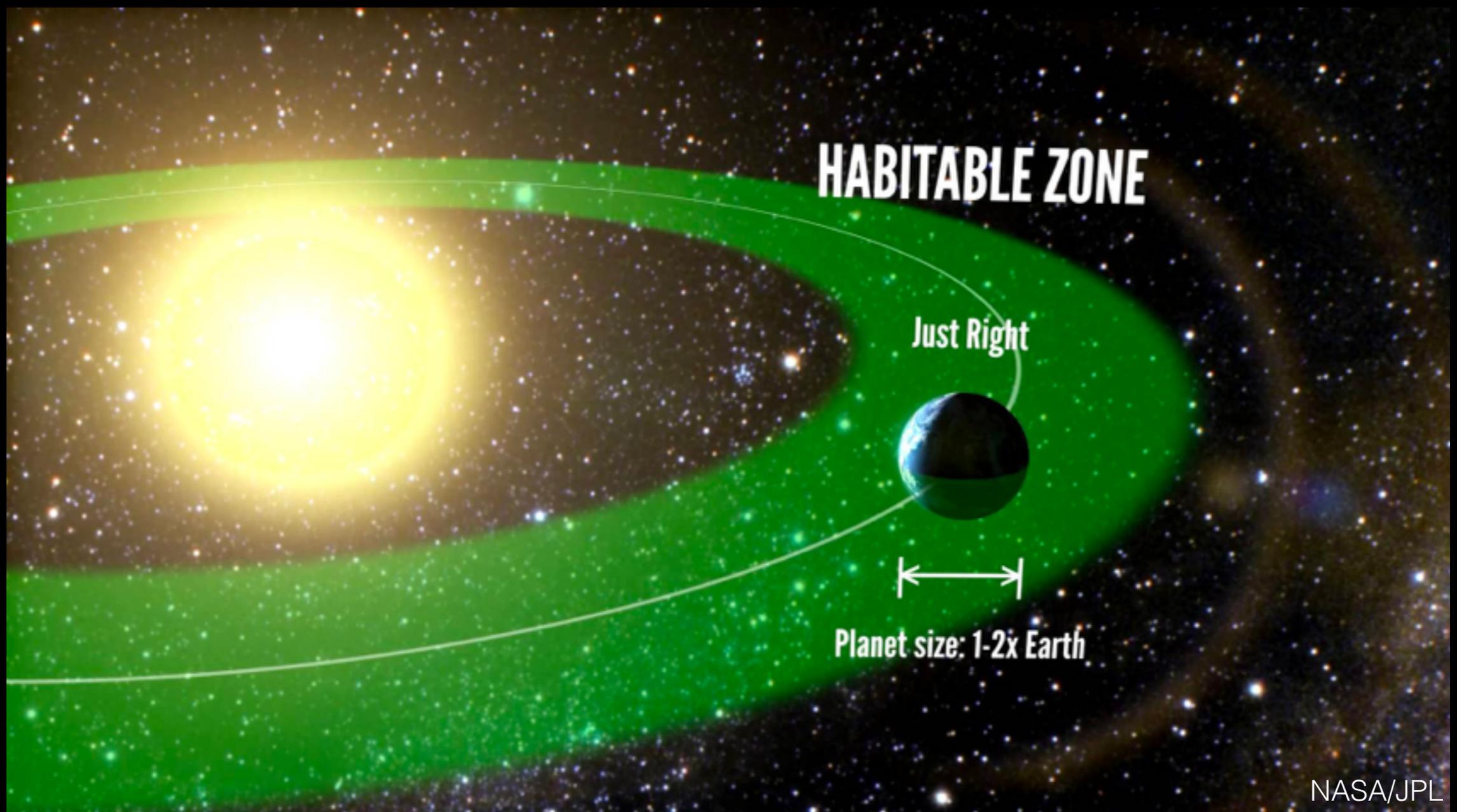


There could be as many as 40 billion Earth-sized planets orbiting in the habitable zones of Sun-like stars



The habitable zone corresponds to the range of orbital distances where liquid water can exist on a planet's surface.

Hubble reveals an estimated 100 billion galaxies in the universe



NASA/JPL

Understand (and protect) life here on Earth
(One of the main goals of NASA)

Astrobiology is the study of the origin, evolution, distribution, and future of life in the **universe**:
extraterrestrial life and life on **Earth**.

Curiosity Rover is the first applied astrobiologist



Curiosity Rover is the first applied astrobiologist

Followed by Matt Damon...



Who am I and what do I do?

(My grandmother raised me, despite never having went to school)

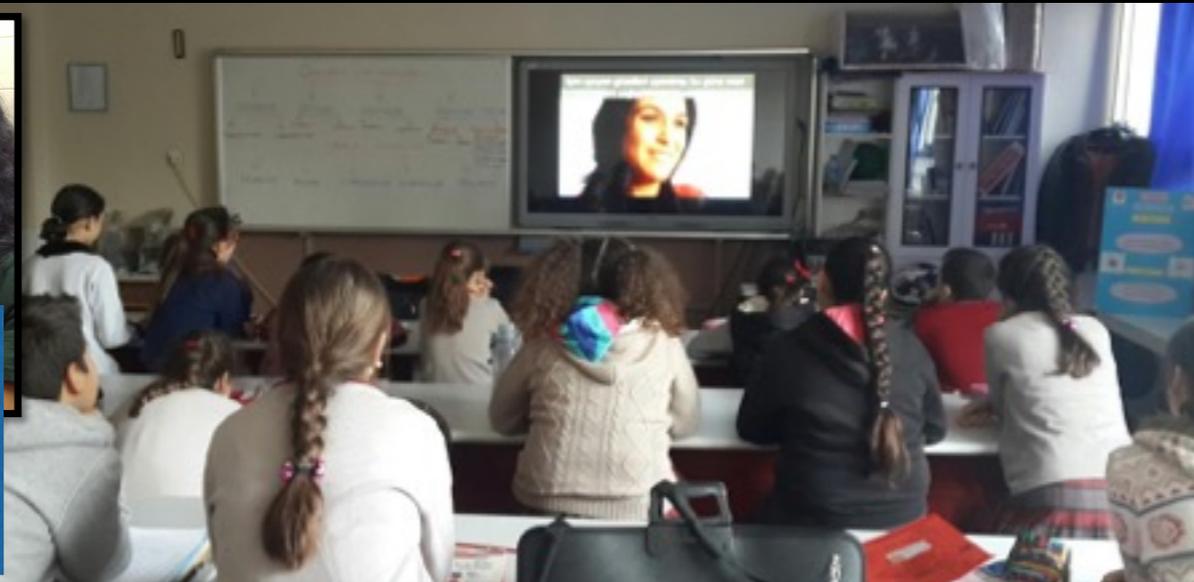


(I immigrated to the USA from Turkey, in 2003)
to study Biochemistry (undergraduate)

(My first day of school, 1990)



NASA
Astrobiology
Institute



Turkey



Georgia
Tech



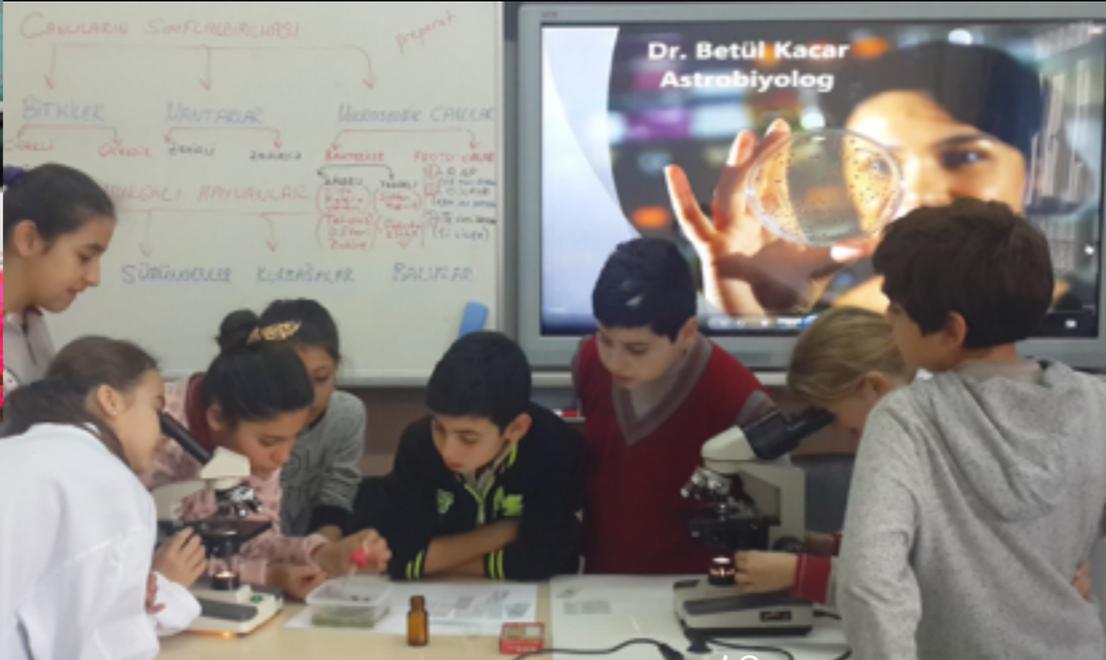
Uppsala,
Sweden



the scientist is



ELSI Origins of
Life Institute,
Tokyo



The Scientist Is In
Dr. Betül Kacar is a postdoctoral fellow from Georgia Institute of Technology and studies Experimental Paleogenetics.
Dr. Kacar is interested in understanding evolution at the molecular level. She uses computational tools to travel back in time and resurrect ancient genes. This method (paleogenetics) allows her to create a molecular time machine in the laboratory. She follows the evolutionary history of our DNA to unravel the molecular events that gave rise to extinction and speciation. She uses this information to understand how climate change directly affects the quality of our lives.

Smithsonian
Institute



Explore Life on Mars?



National Standards

Benchmarks for Science Literacy

5F/H4a (Grades: 9-12): Heritable characteristics can be observed at molecular and whole-organism levels—in structure, chemistry, or behavior.

5F/H5 (Grades: 9-12): New heritable characteristics can result from new combinations of existing genes or from mutations of genes in reproductive cells. Changes in other cells of an organism cannot be passed on to the next generation.

5F/H6b (Grades: 9-12): Chance alone can result in the persistence of some heritable characteristics having no survival or reproductive advantage or disadvantage for the organism.

5F/H8 (Grades: 9-12): Life on earth is thought to have begun as simple, one-celled organisms about four billion years ago. Once cells with nuclei developed about a billion years ago, increasingly complex multi-cellular organisms evolved.

5F/M3a (Grades: 6-8): Many thousands of layers of sedimentary rock provide evidence for the long history of the earth and for the long history of changing life forms whose remains are found in the rocks.

5F/M3b (Grades: 6-8): More recently deposited rock layers are more likely to contain fossils resembling existing species.

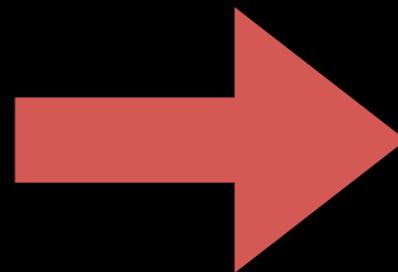
LPI Girls! Mars Now! PBS Learning Media Origins of Life

<http://www.lpi.usra.edu/education>

<http://www.pbslearningmedia.org/collection/origins-of-life/>



I am a project leader at Harvard University



Moving to University of Arizona as an Astrobiology faculty!

Goal of my research:

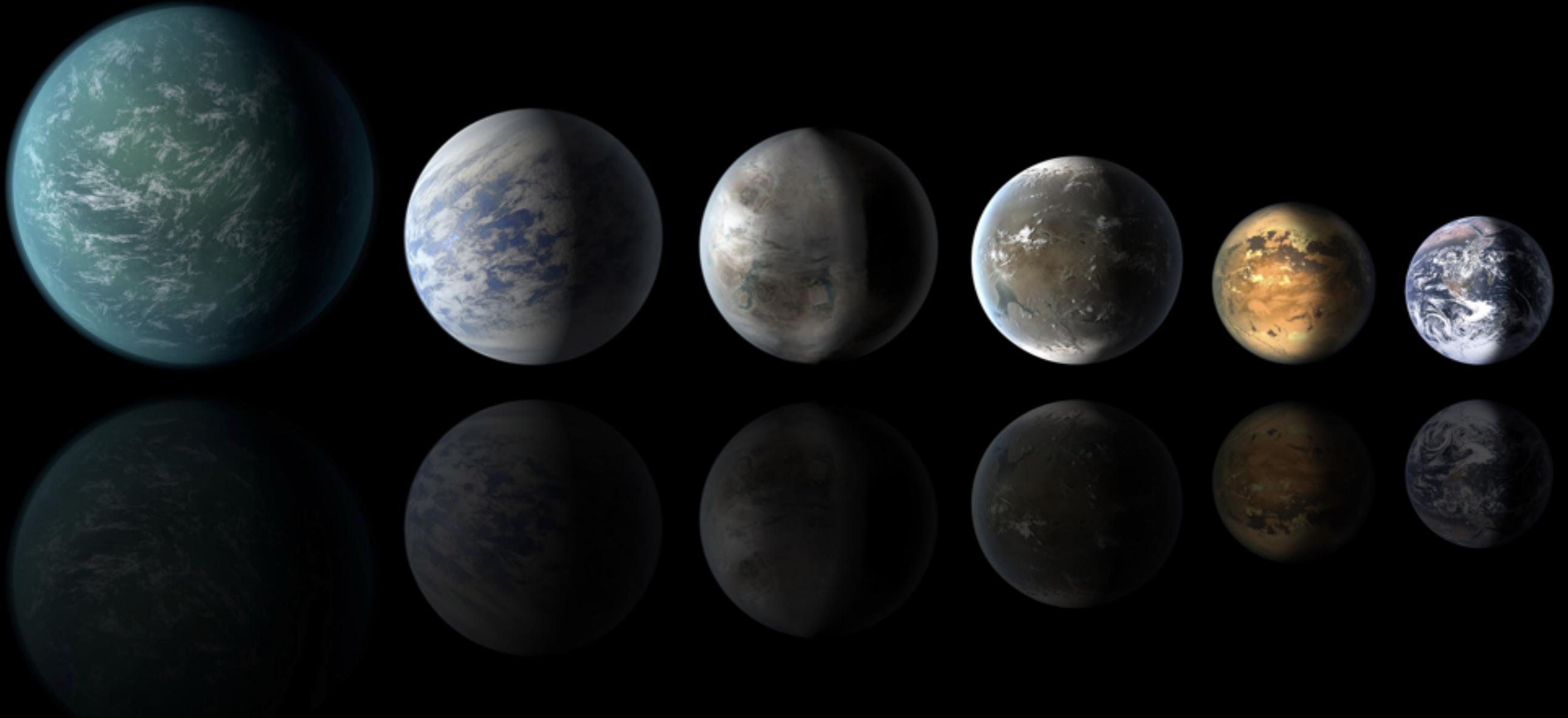
I aim to understand ancient life:

How did life evolve?

How did life, at the cellular level, behave in the past?



Earth's past is a strange planet (different atmosphere, temperature, etc.)



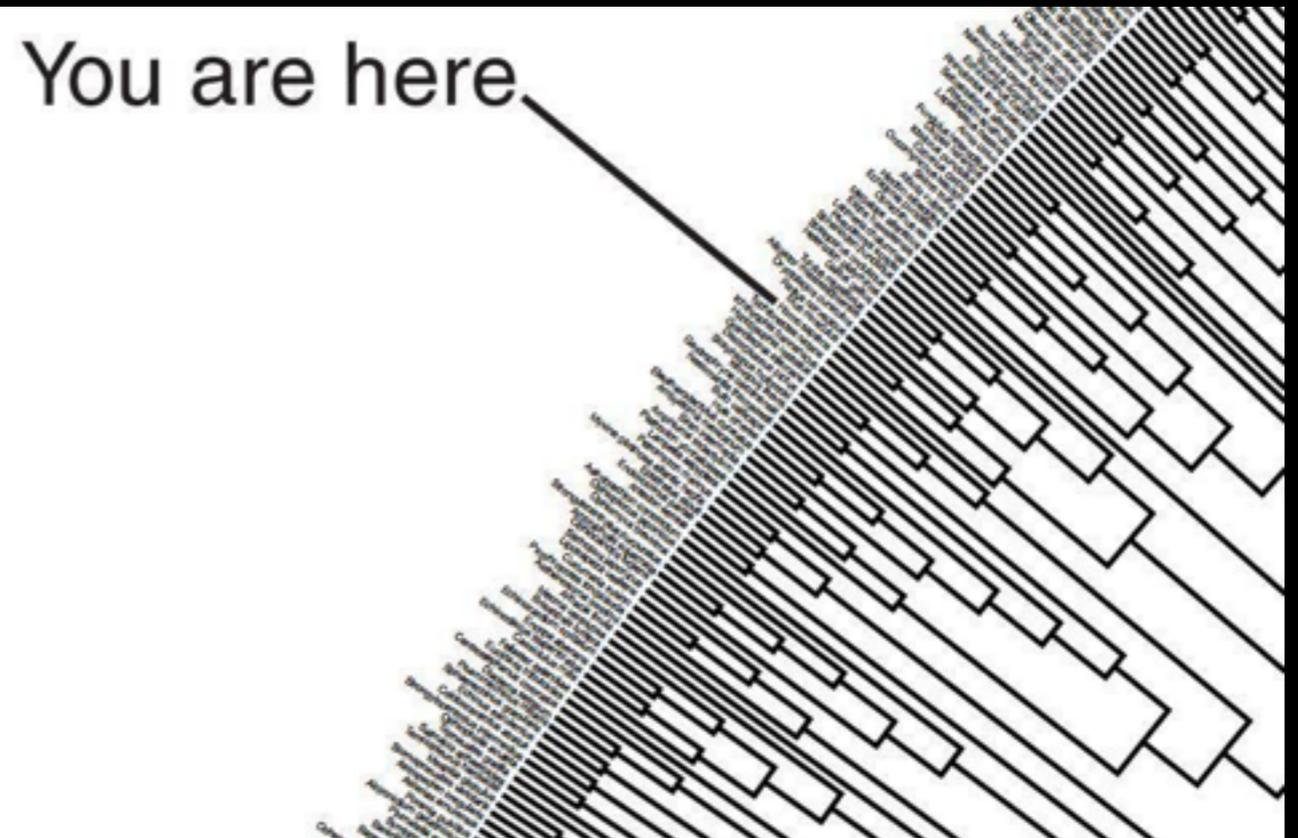
How did microorganisms behave in the past?

Rocks provide the global chemical picture

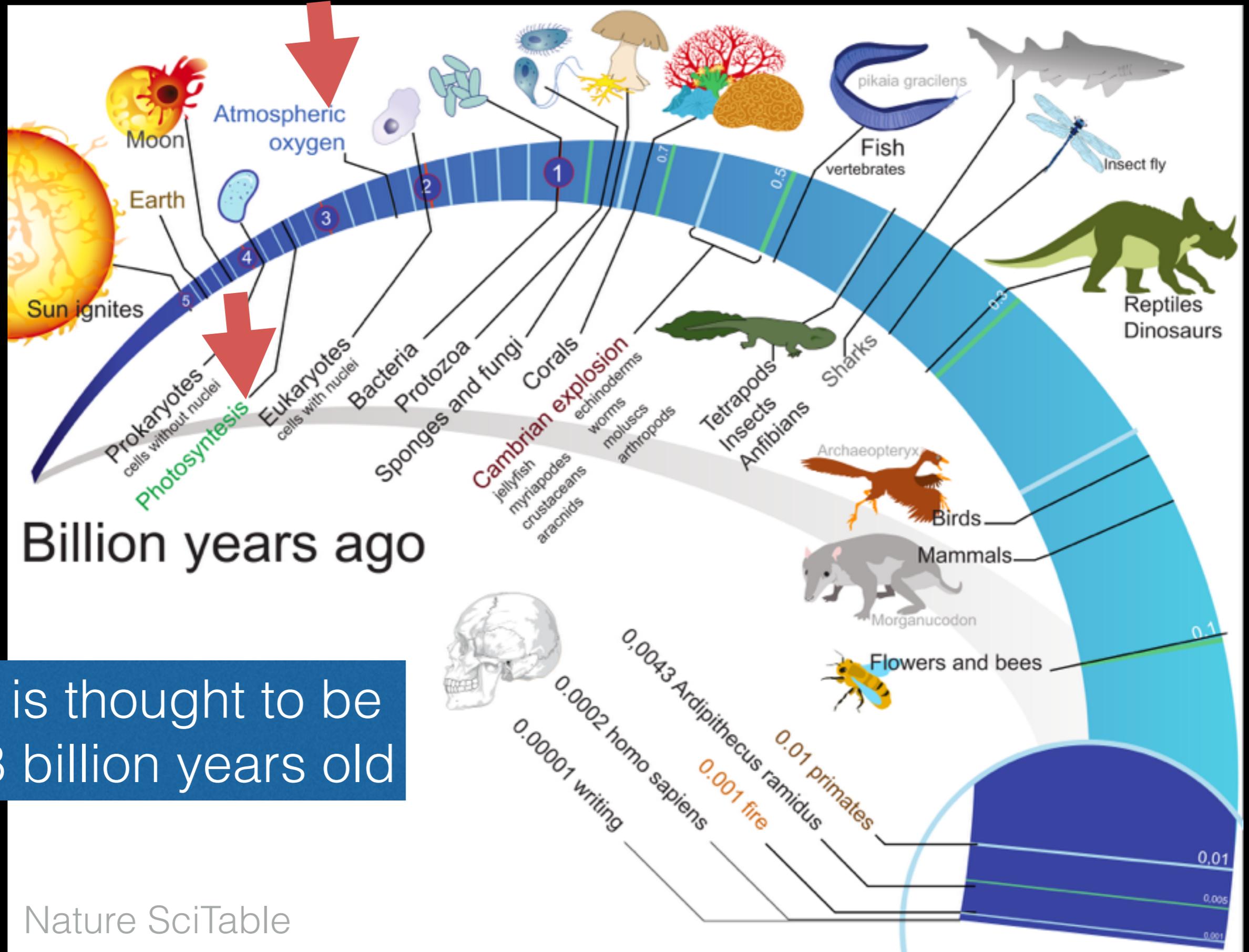
Two main approaches to understand the past:



Life provides a mix of specific molecular patterns



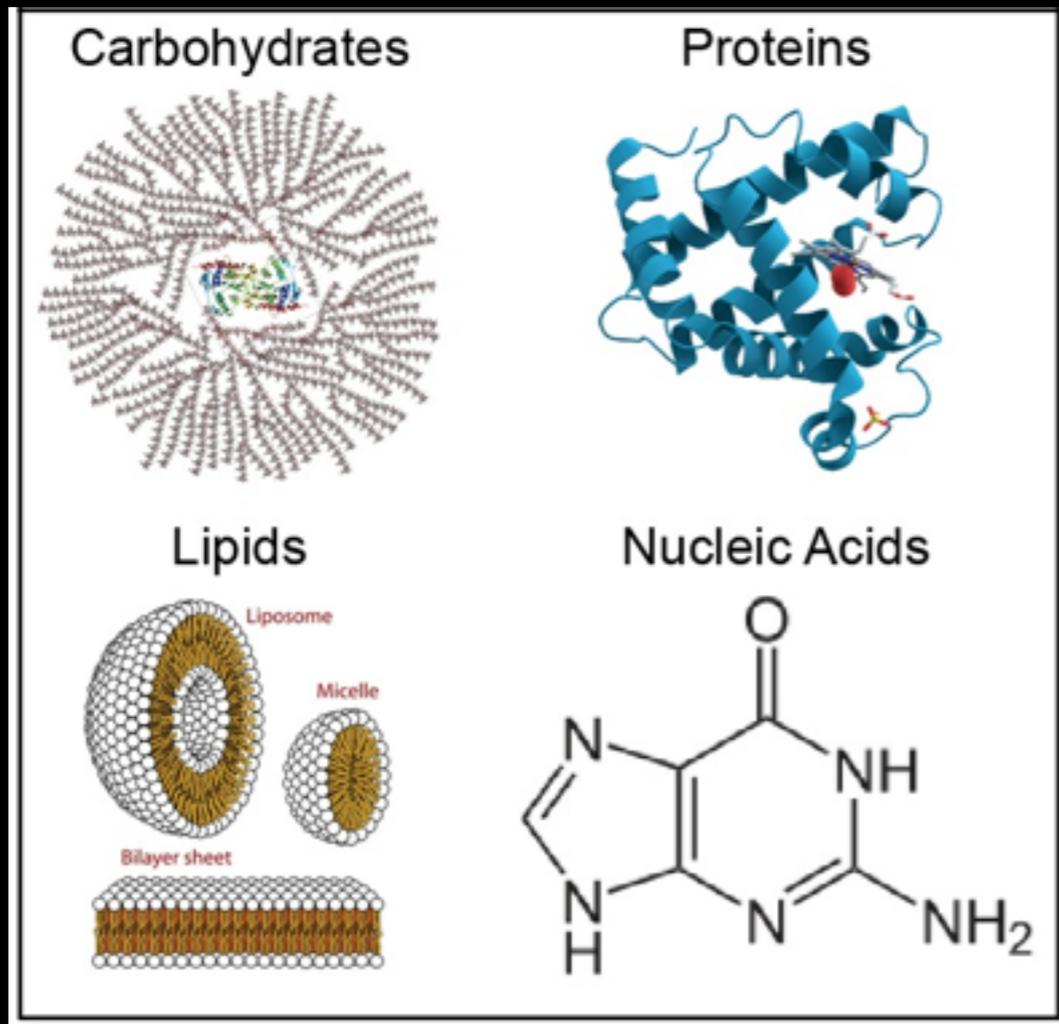
Evolution of Life Timeline



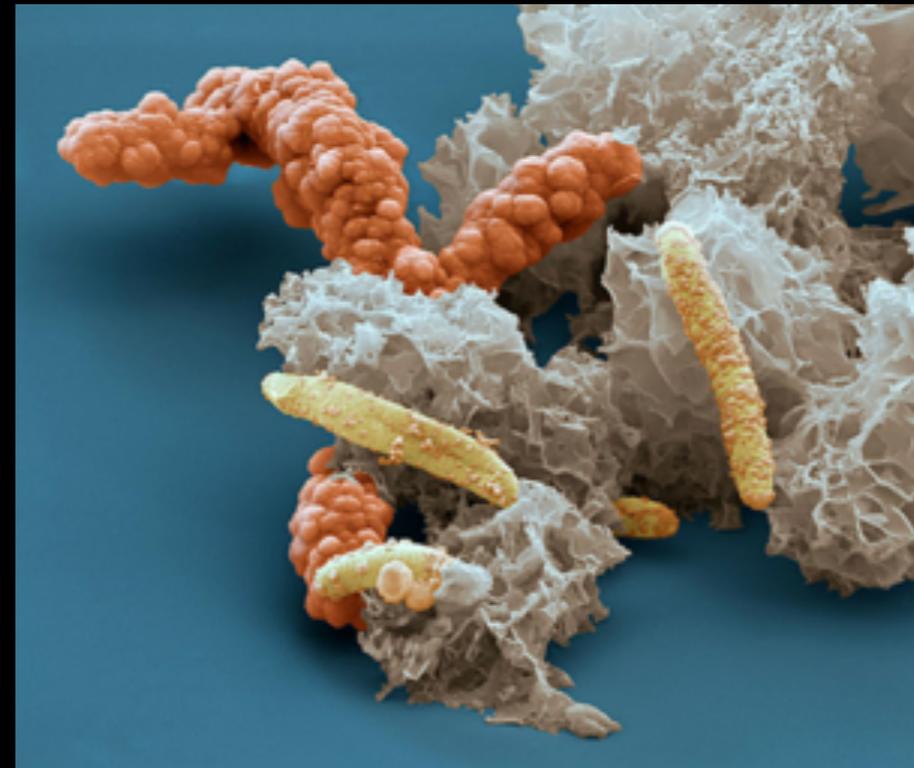
Life is thought to be ~3.8 billion years old

Biosignatures

Any substance – such as an element, isotope, molecule, or phenomenon – that provides scientific evidence of past or present life.



Biomolecules

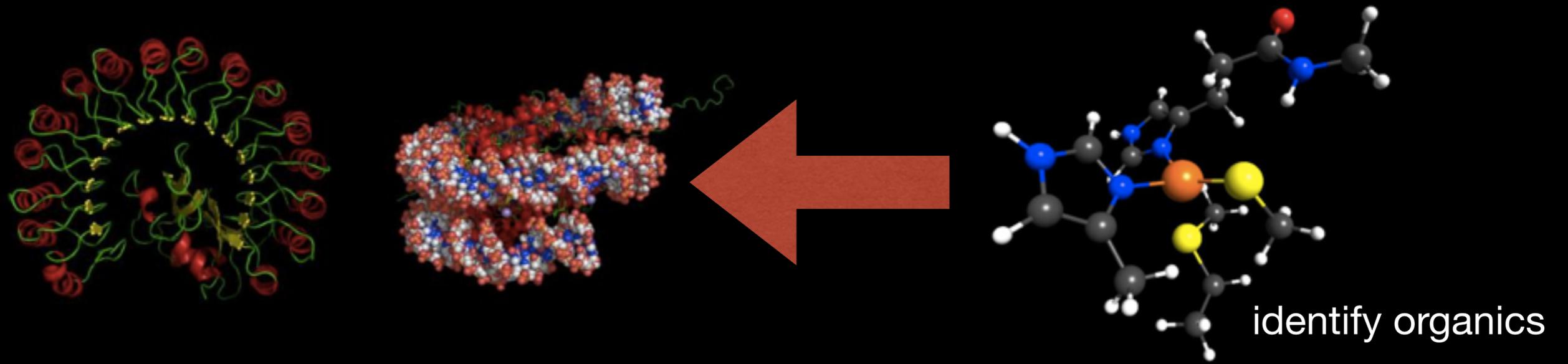


Microbes, Minerals



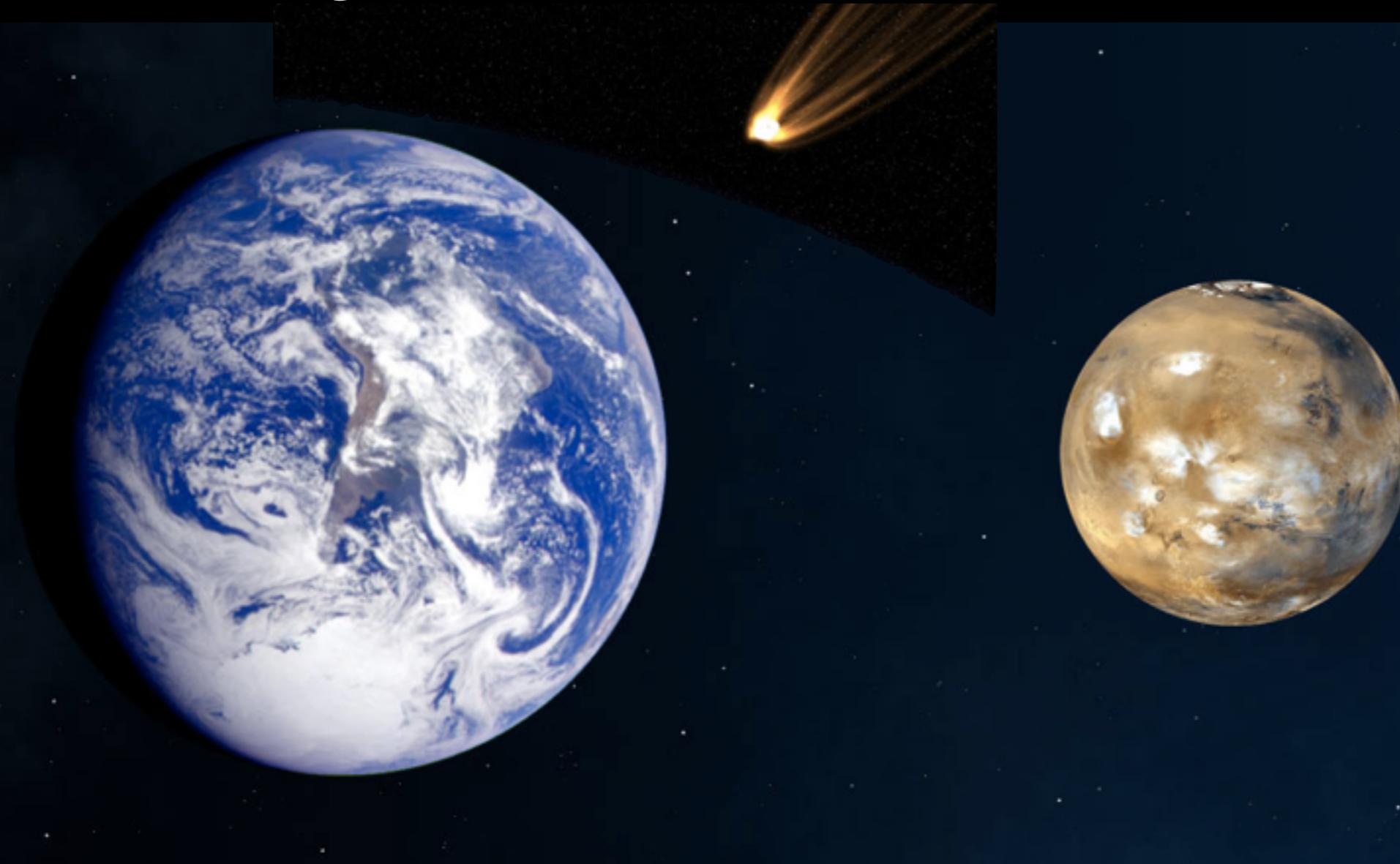
17 Genetics

Search for Biosignatures

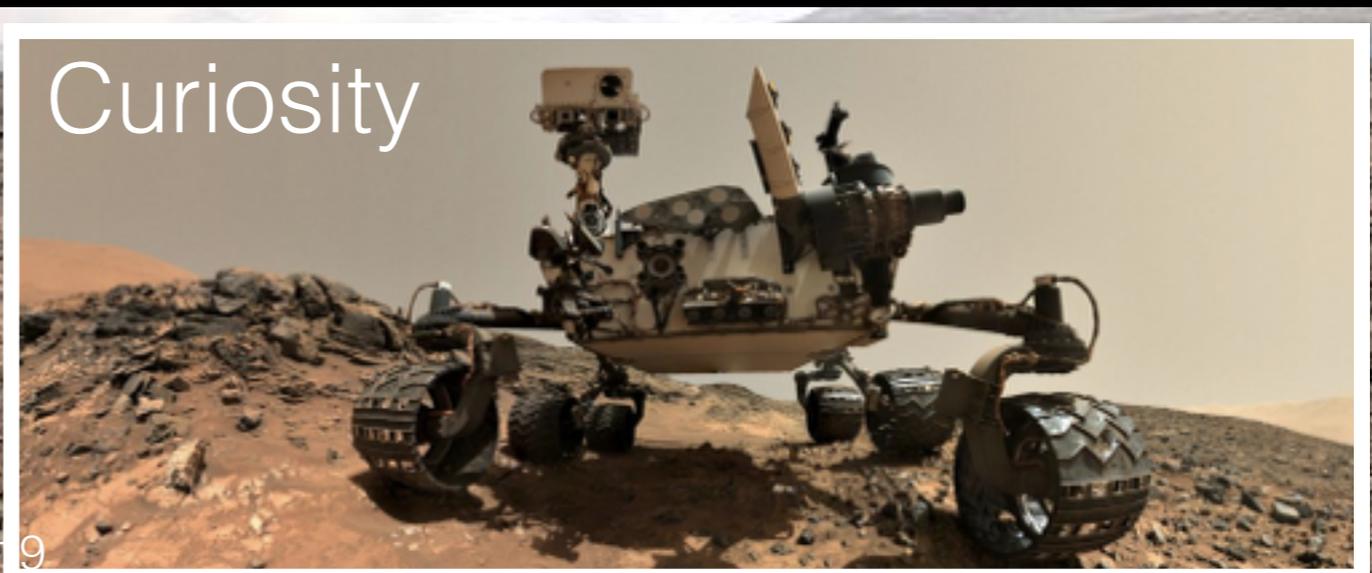


Asteroid landing missions

Search for Biosignatures



Earth's past and Mars' present (any resemblance?)

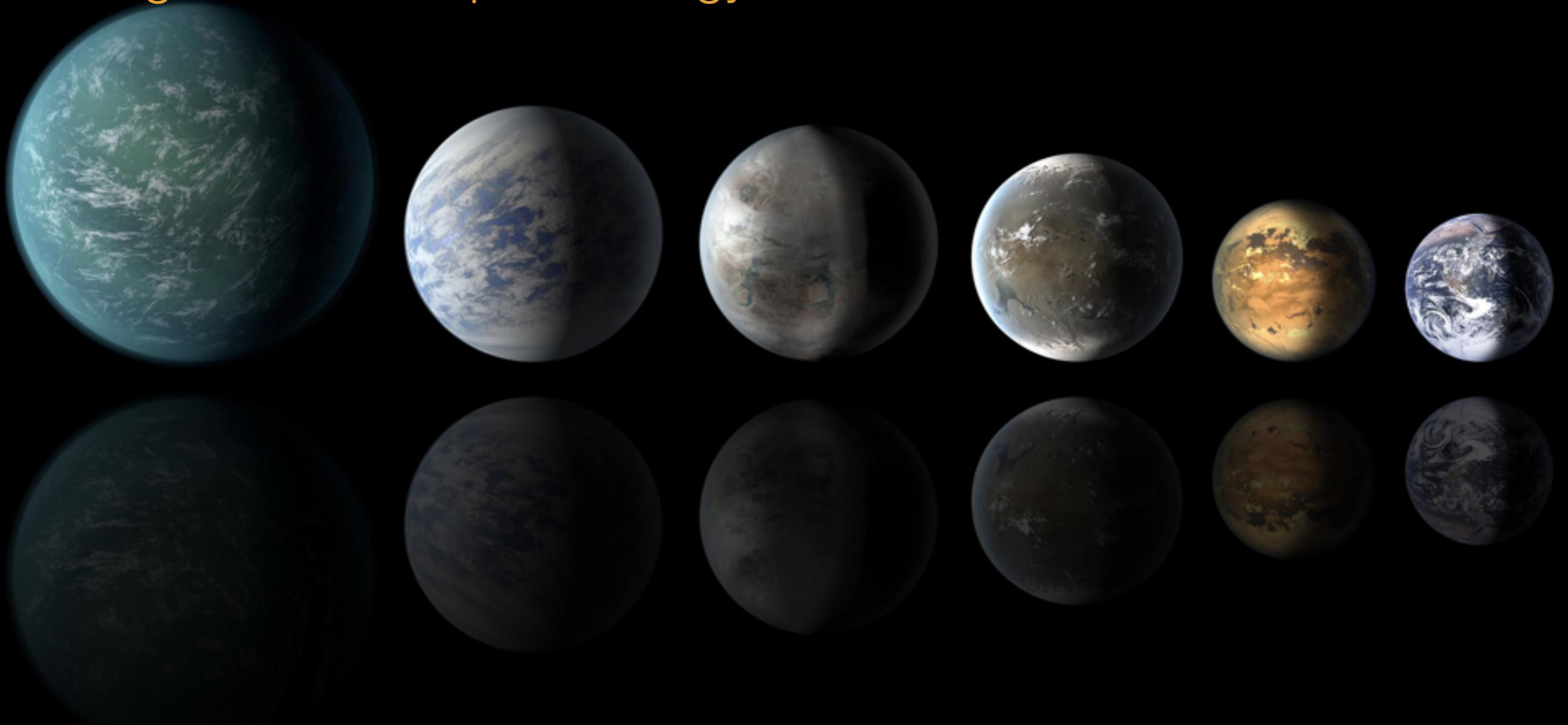


Earth's past is an alien planet

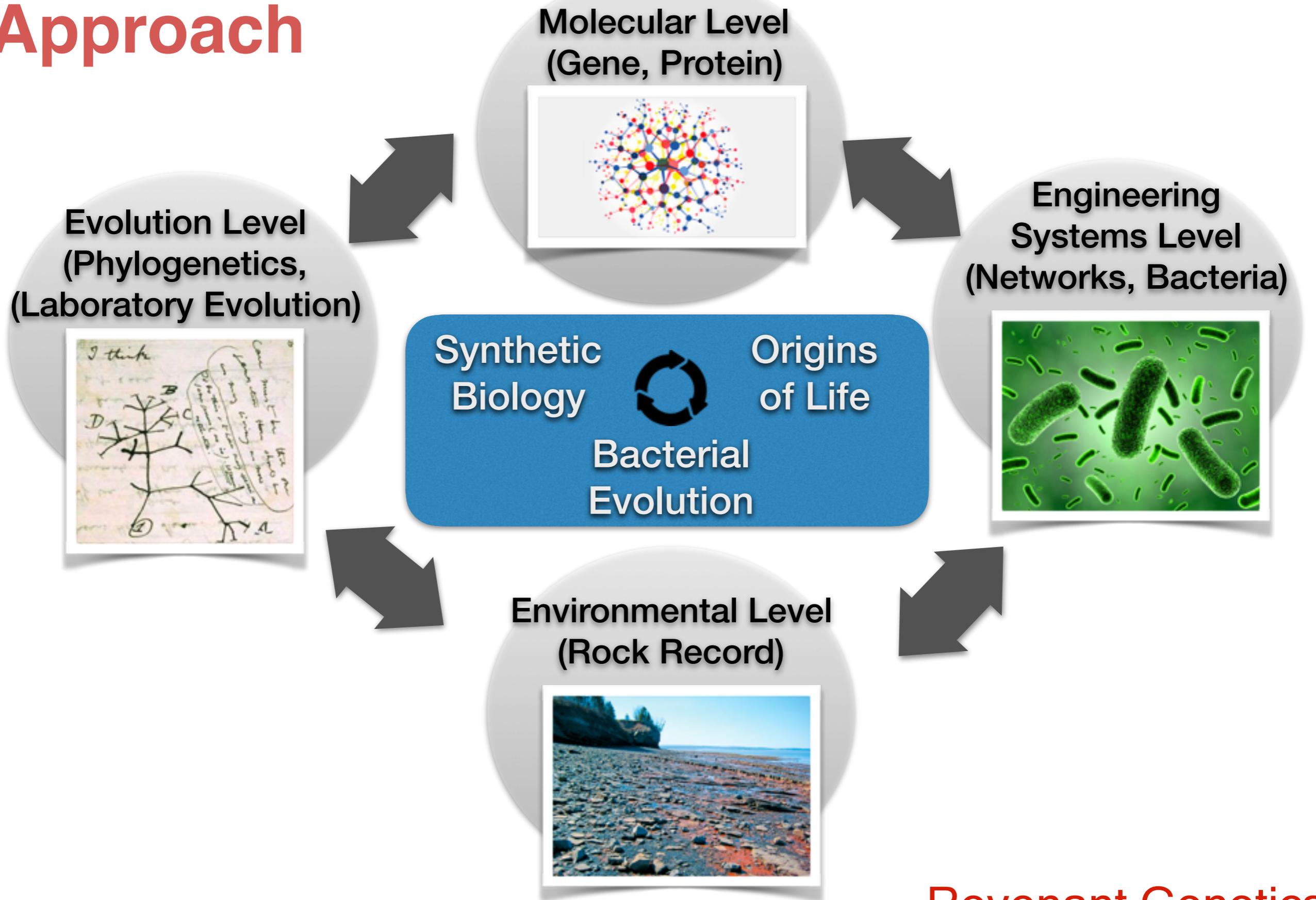
Can we reconstruct its past (alien) biology?

My lab:

Learning more about past biology could aid search for life elsewhere



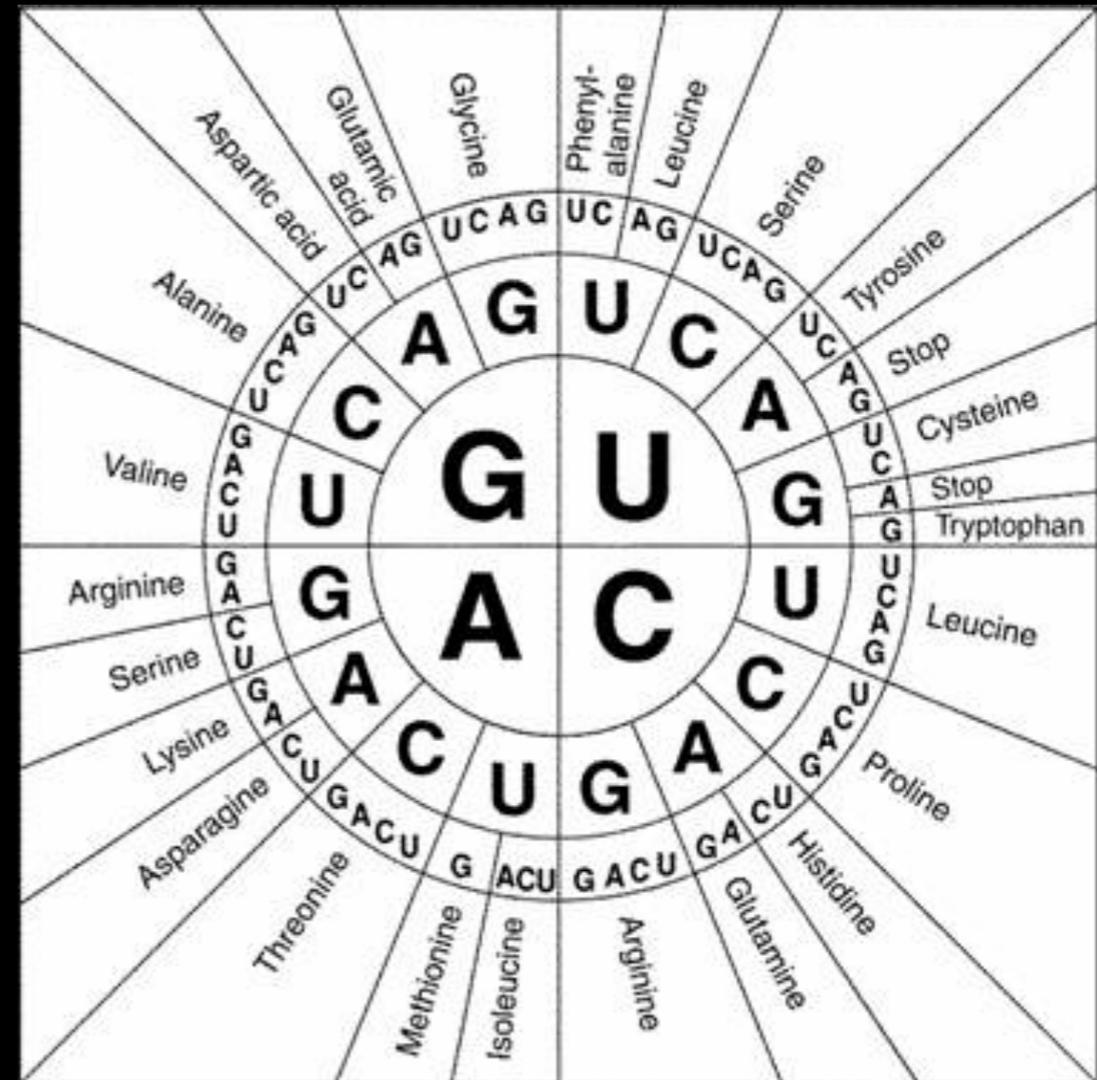
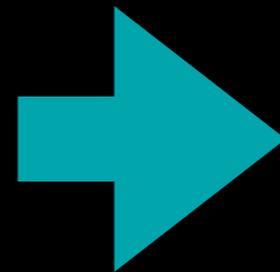
Approach



Revenant Genetics
kacarlab.org

- Linguists make arrangements in life's alphabet to make inferences about the past
- Can we use 'Nature-made' words (amino acids) to make inferences about the past?

A **B** **C** **Ç** **D** **E** **F**
G **Ğ** **H** **I** **J** **K** **L**
M **N** **O** **Ö** **P** **R**
S **Ş** **T** **U** **Ü** **V** **Y**
Z abcçdefgğhijklmnoöprsstuüvyz



How do we travel back in time?



Hollywood's version

How do we travel back in time?

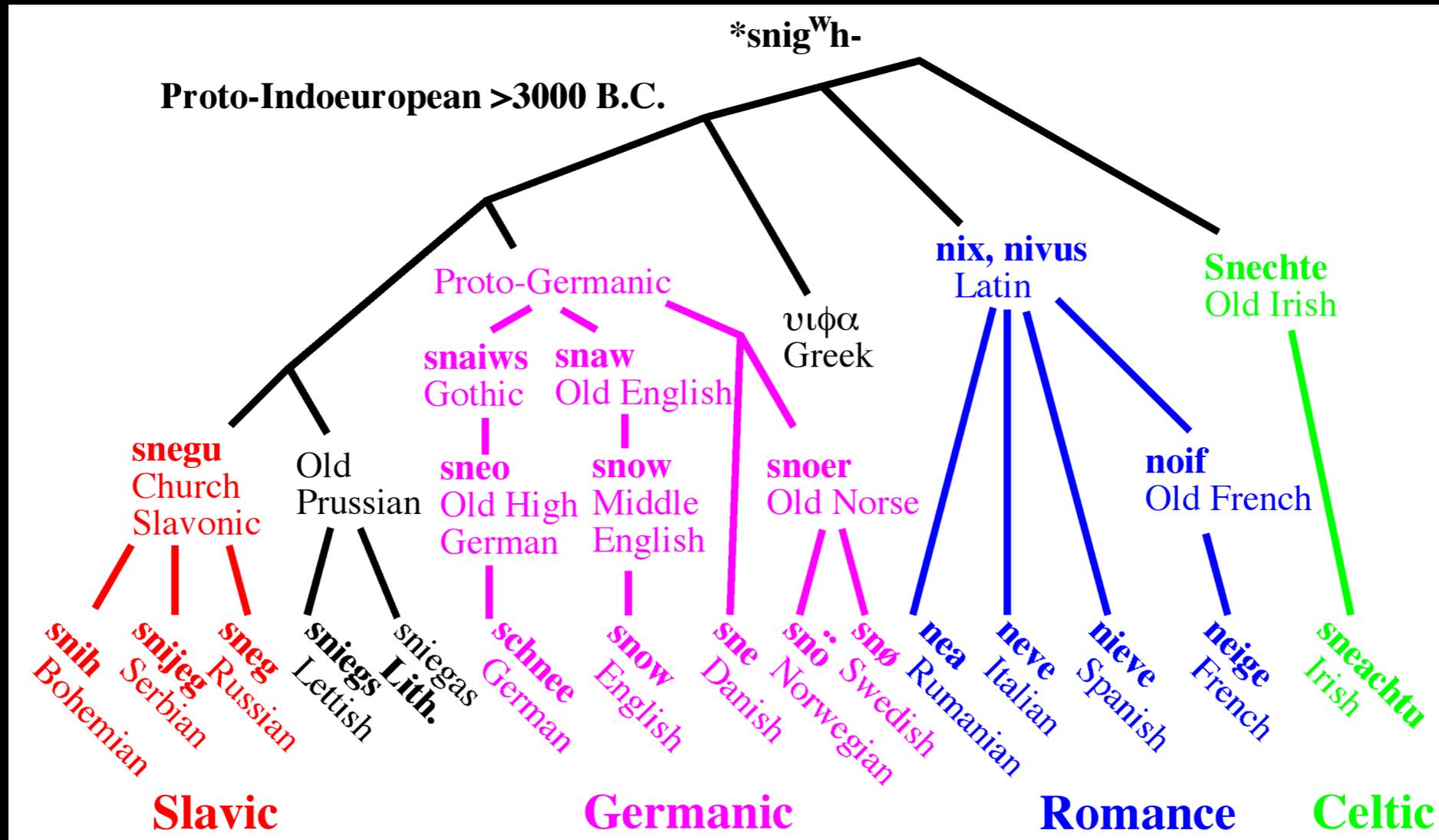


Hollywood's version

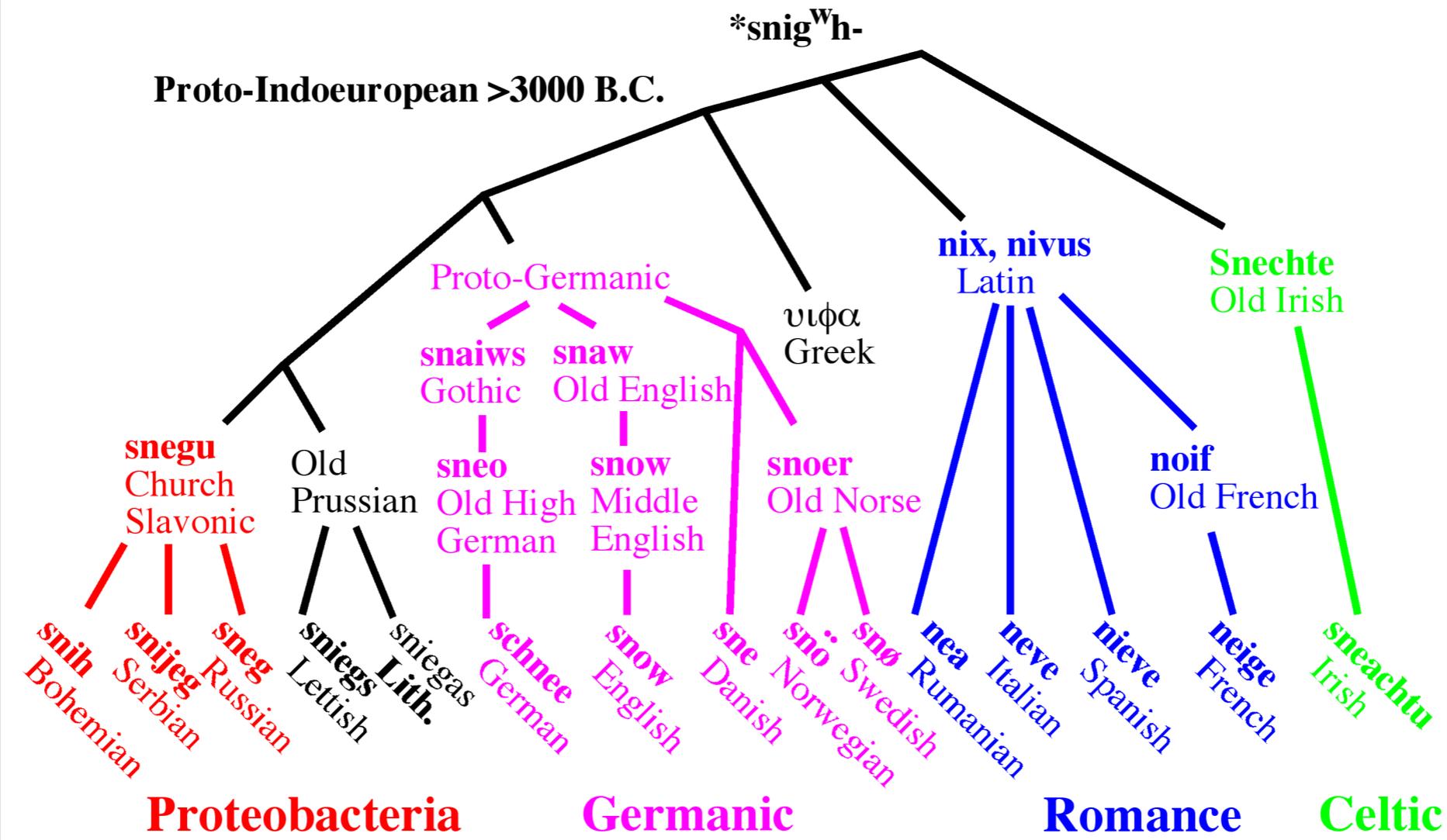


Science's version:

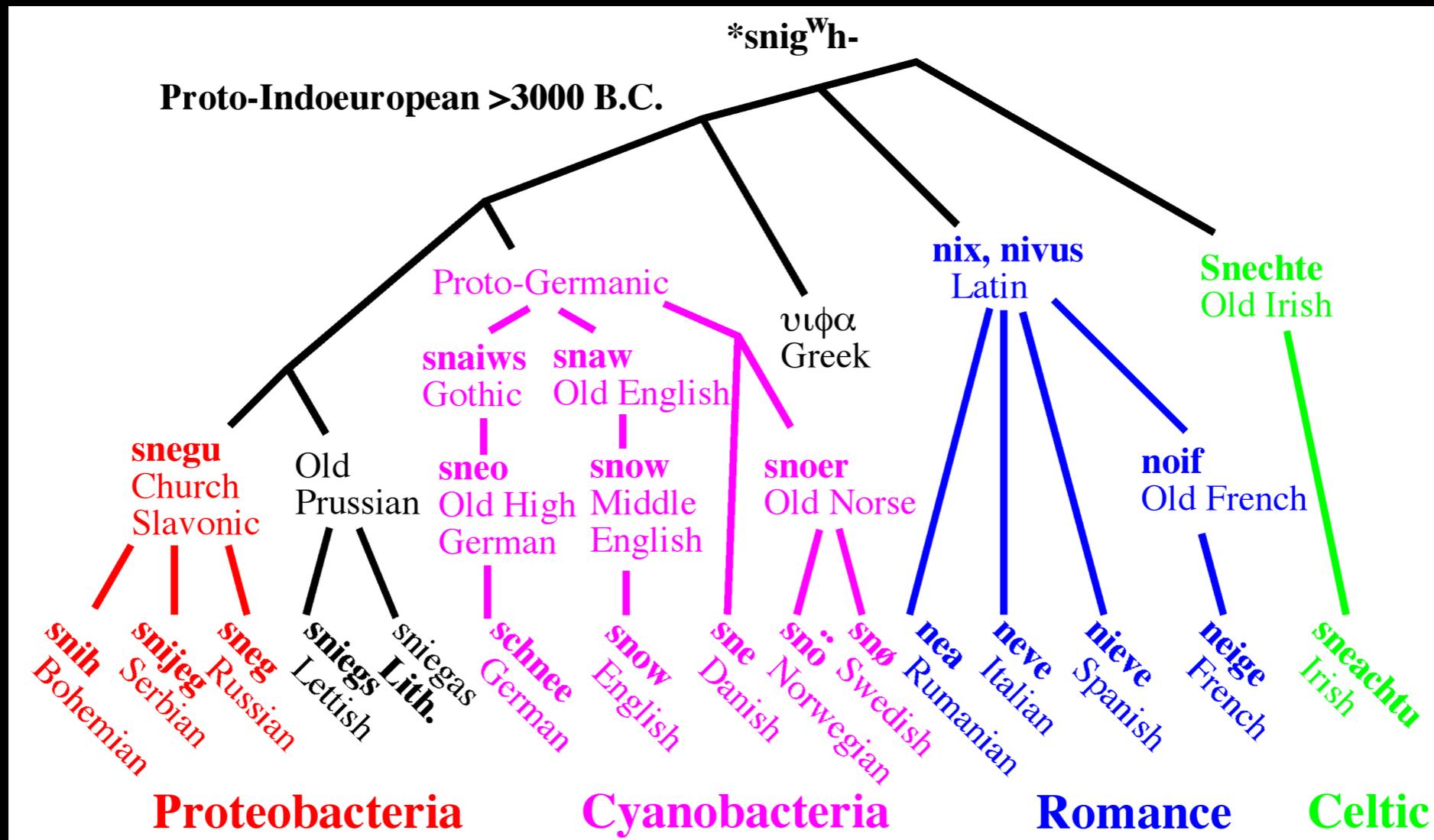
Rewinding the tape: Phylogenetics



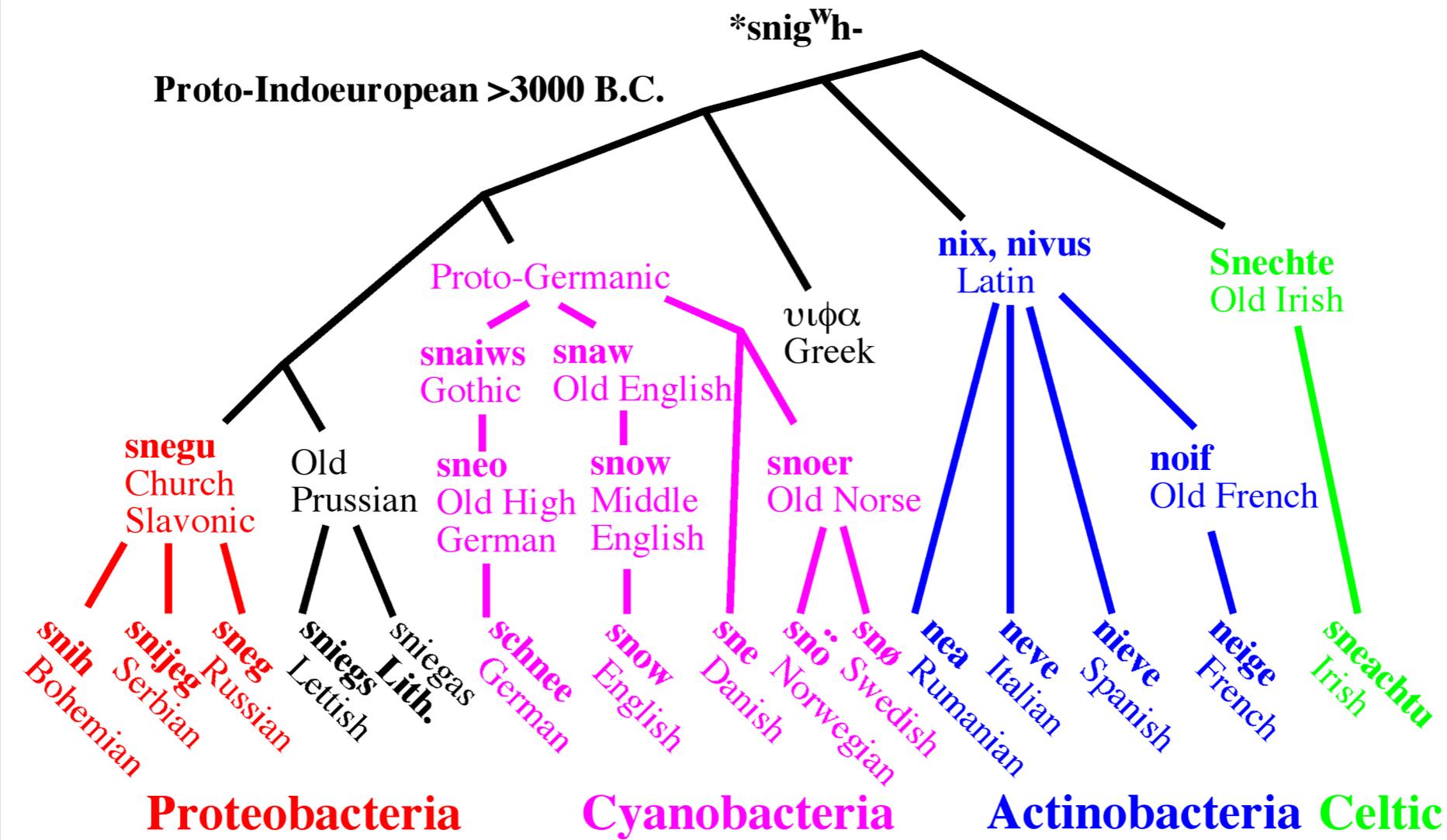
Rewinding the tape: Phylogenetics



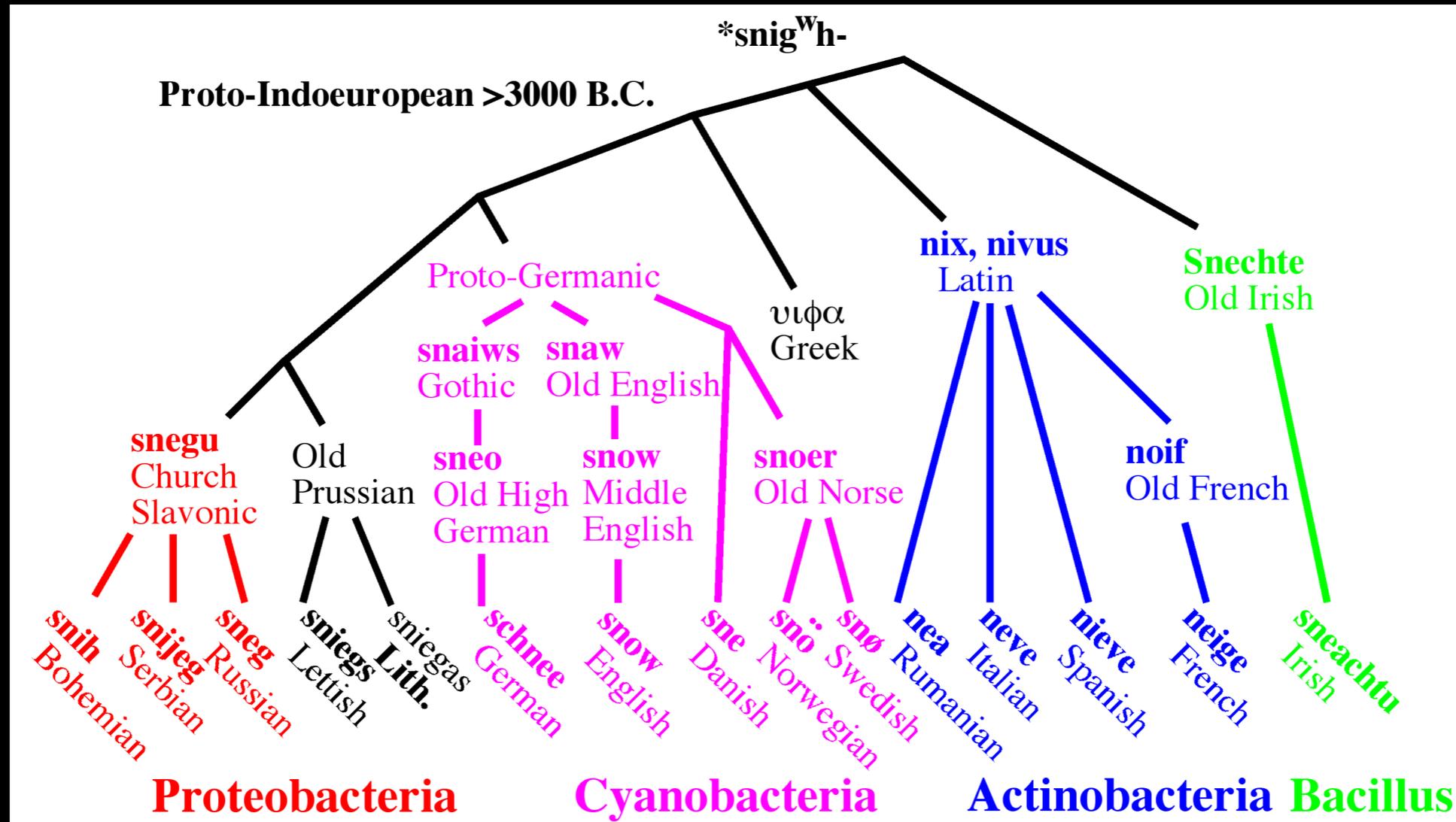
Rewinding the tape: Phylogenetics



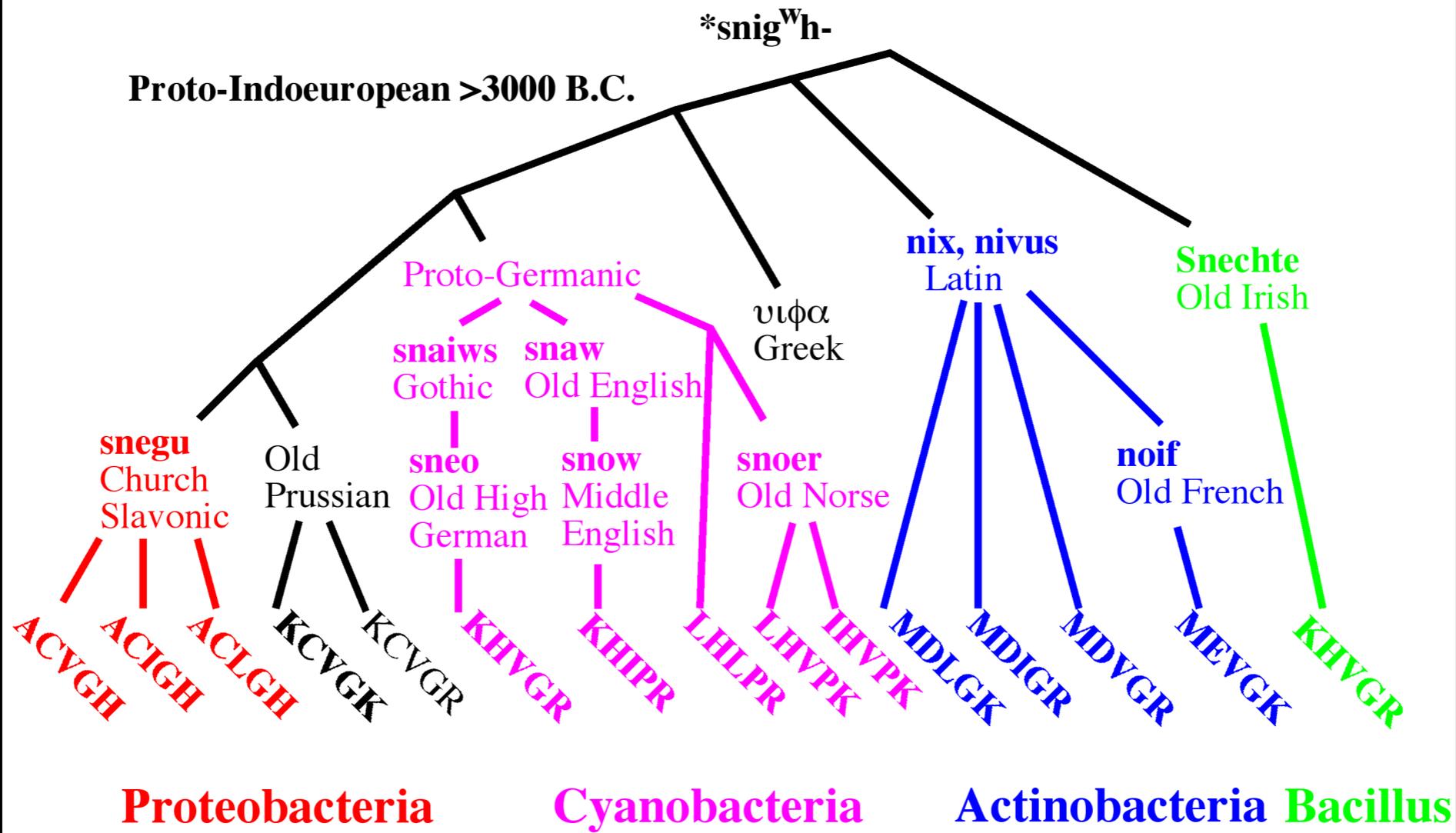
Rewinding the tape: Phylogenetics



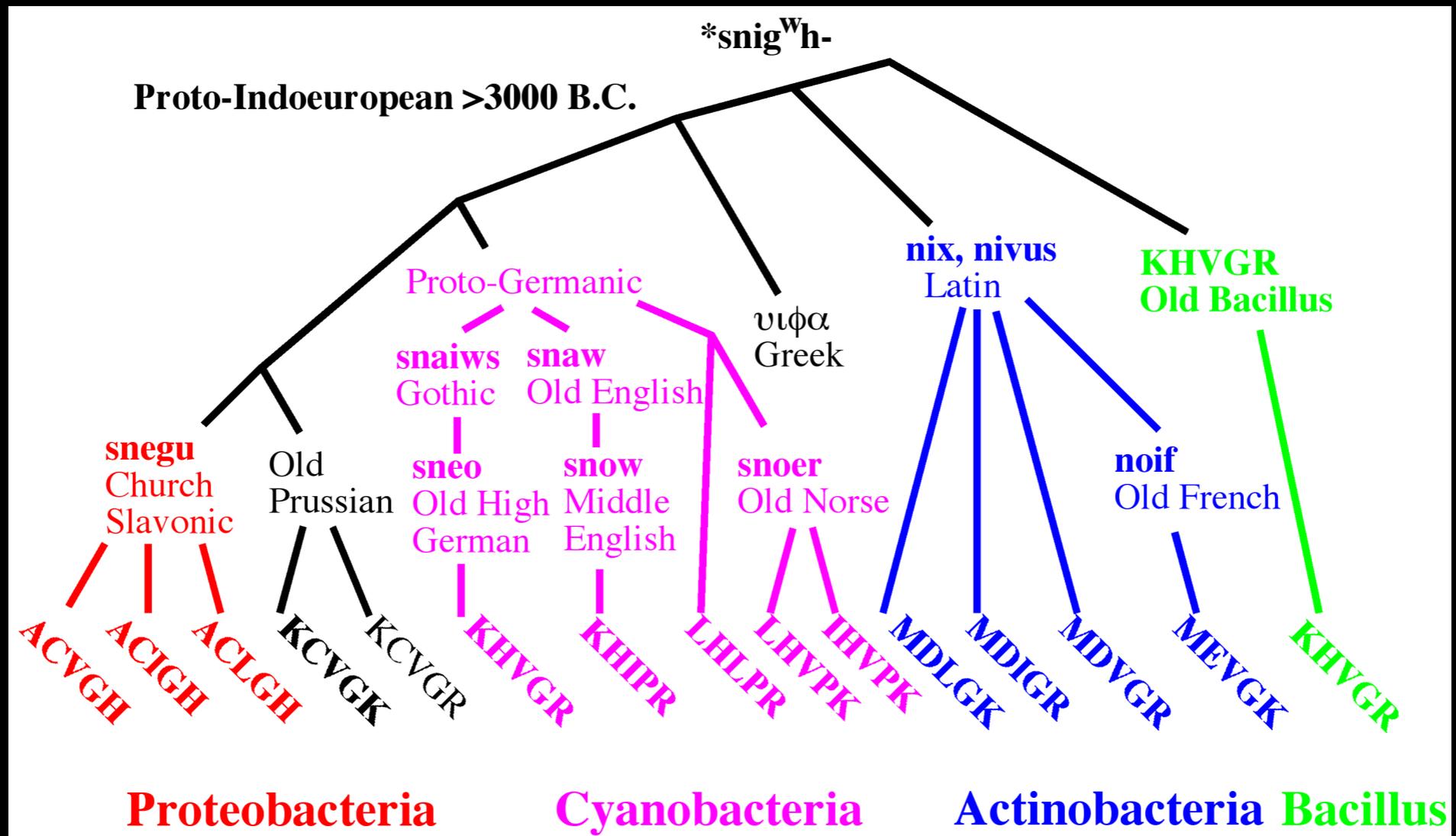
Rewinding the tape: Phylogenetics



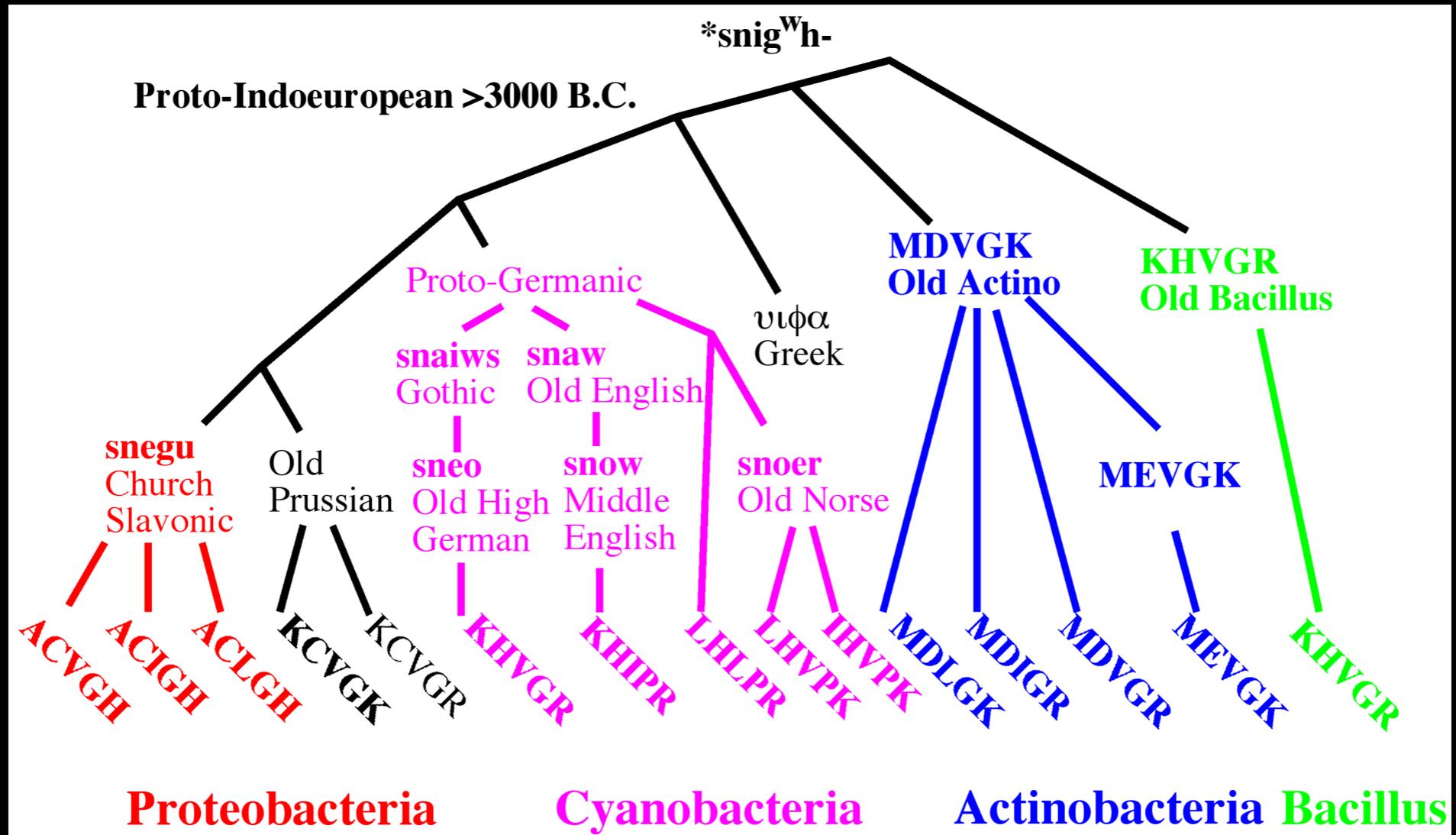
Rewinding the tape: Phylogenetics



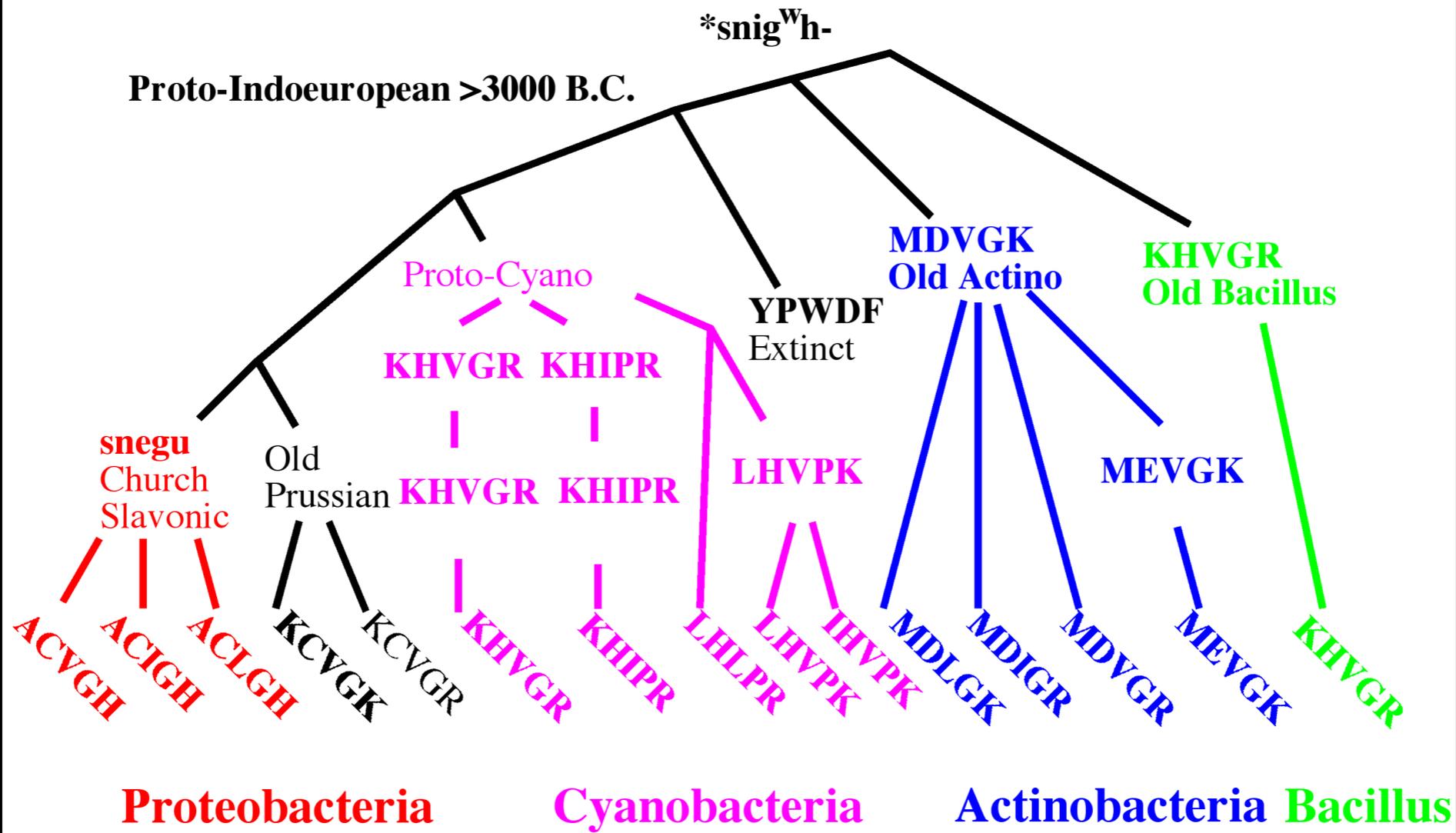
Rewinding the tape: Phylogenetics



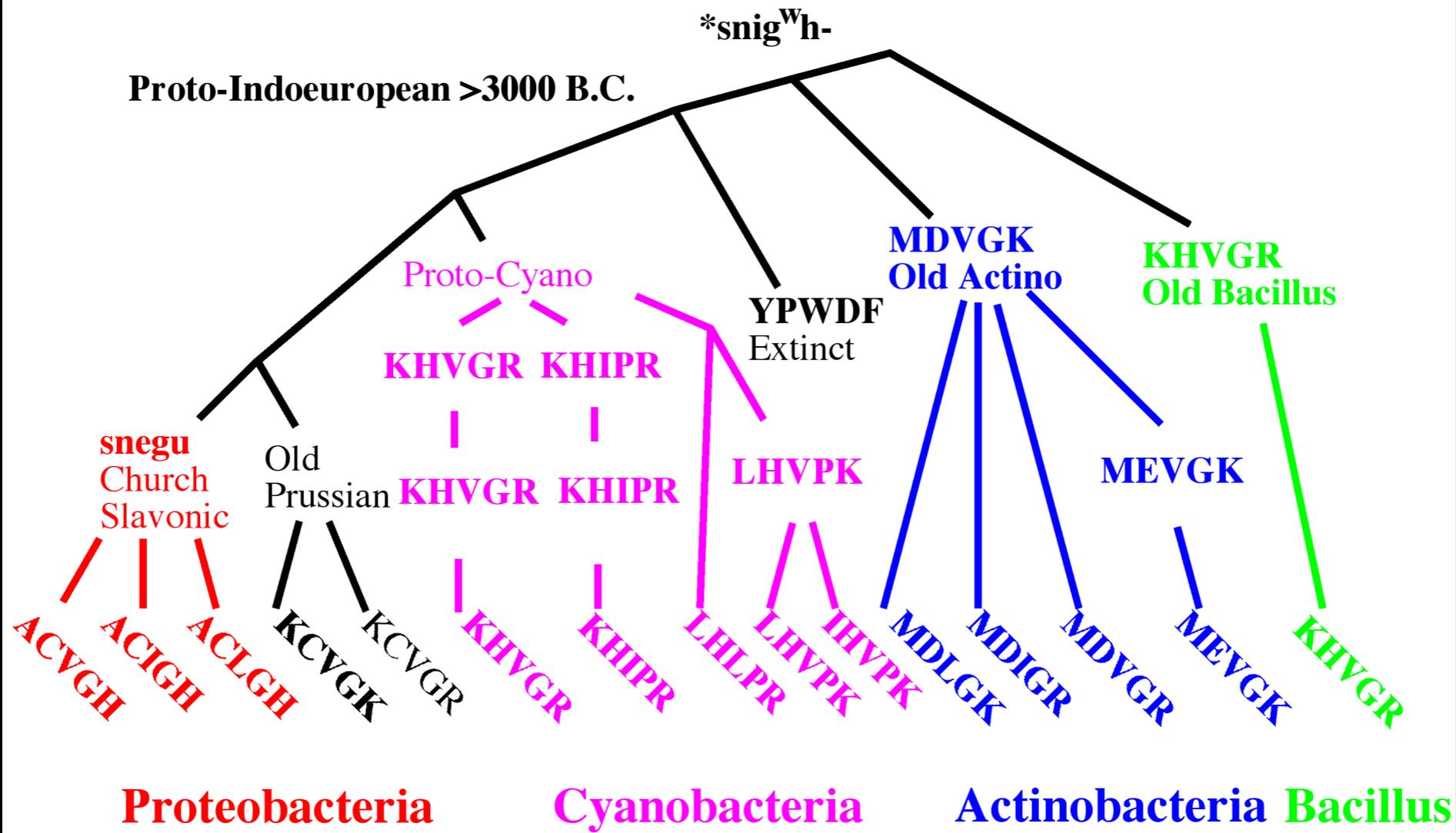
Rewinding the tape: Phylogenetics



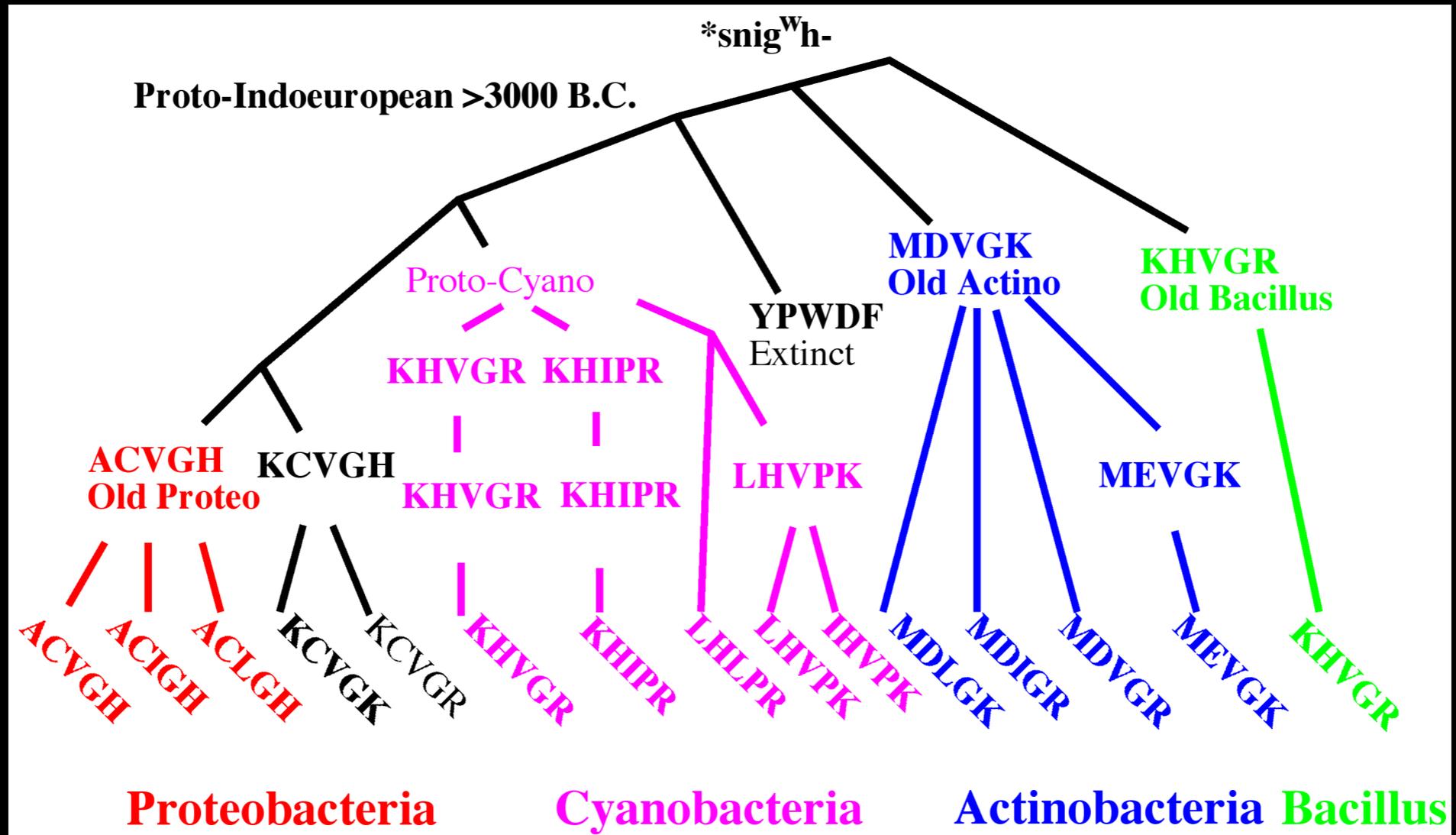
Rewinding the tape: Phylogenetics



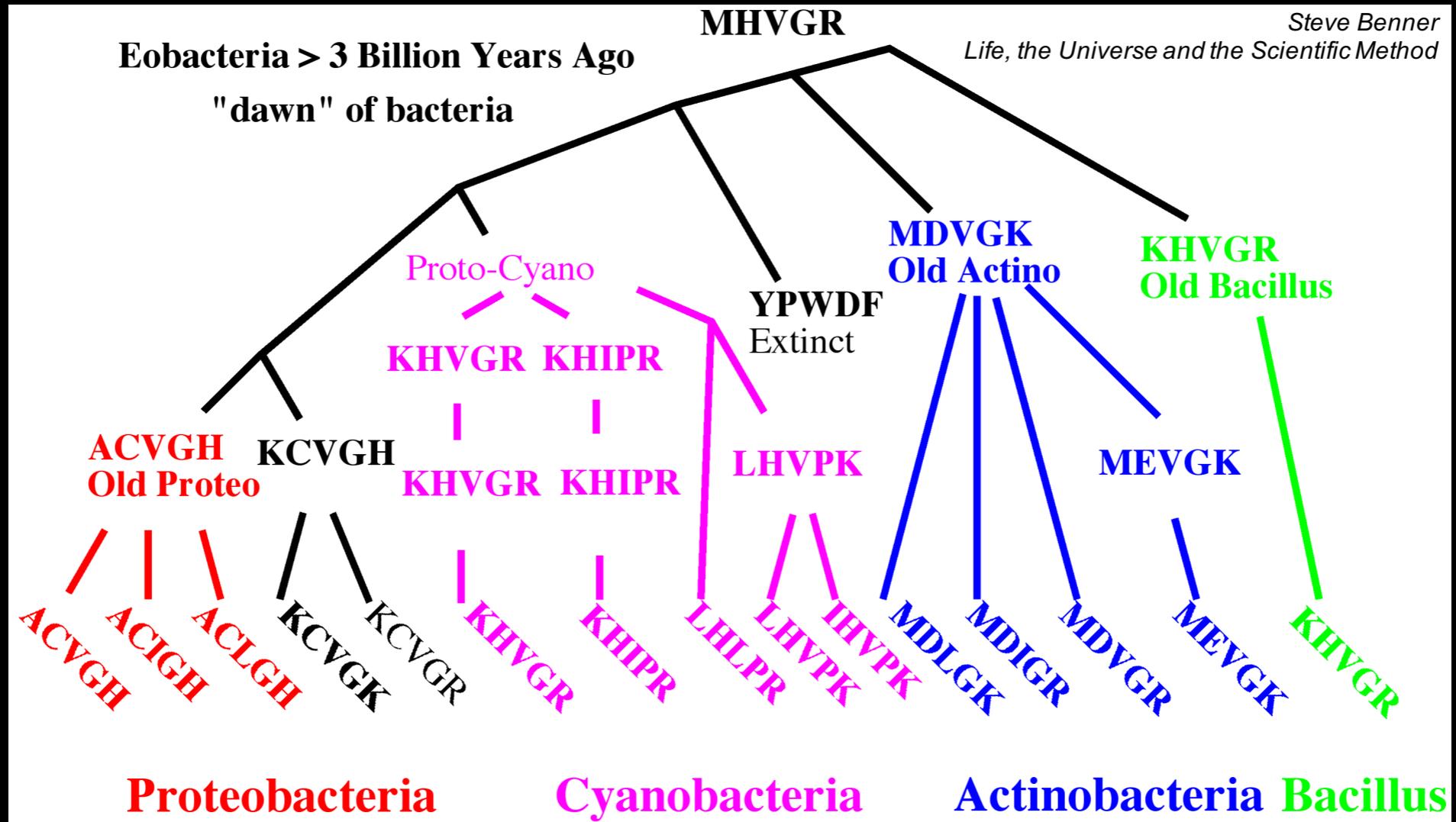
Rewinding the tape: Phylogenetics

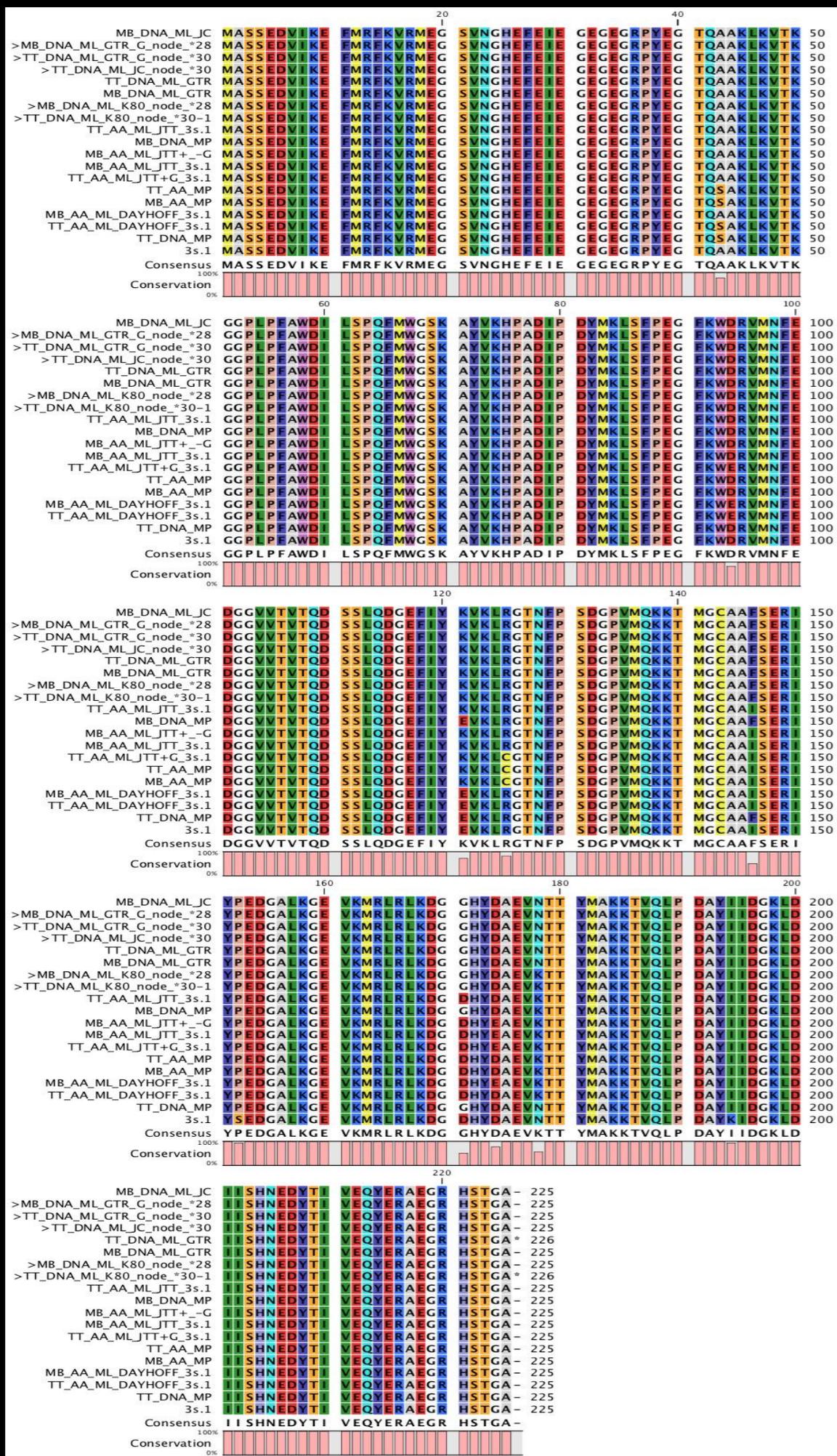


Rewinding the tape: Phylogenetics



Rewinding the tape: Phylogenetics



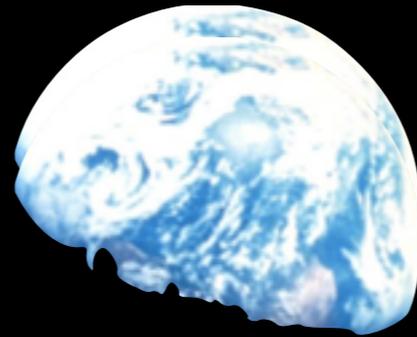


Ancestral Sequence Reconstruction

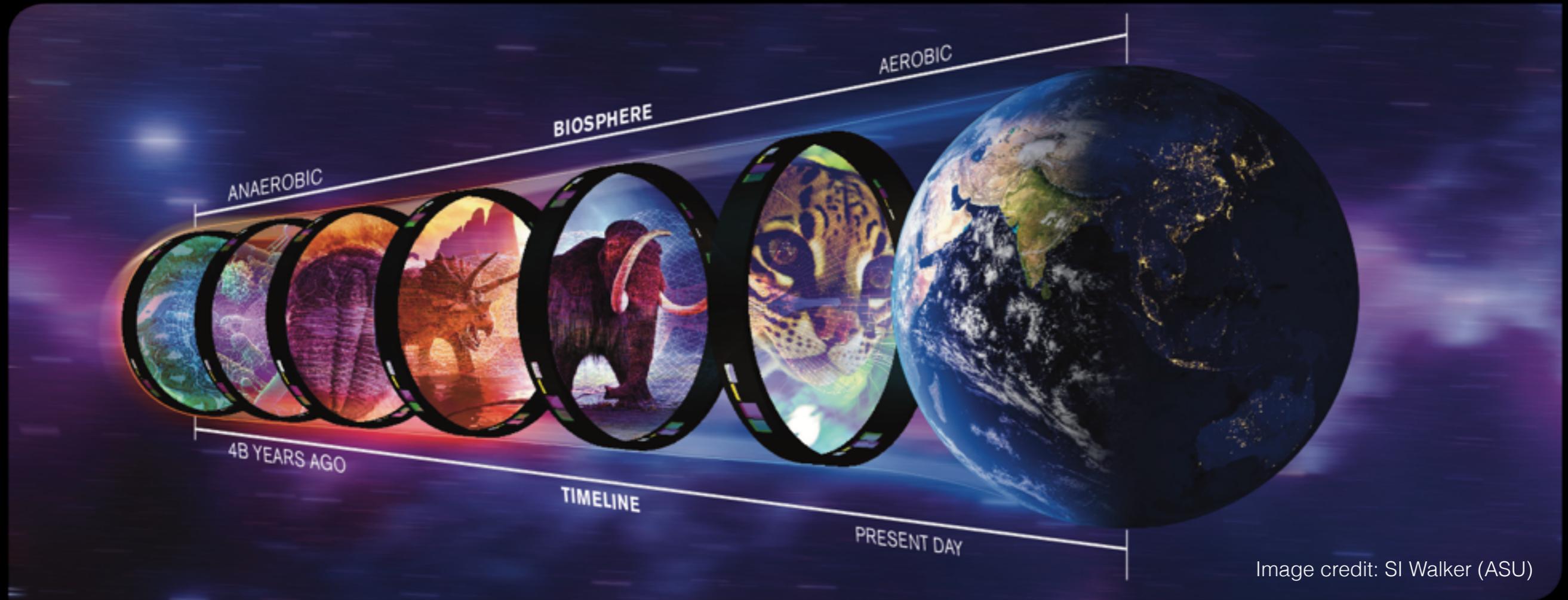
Reconstruct ancestral (ancient) proteins using modern biology

Motivating Questions

- How can we travel back in evolutionary time?
- How repeatable is life's evolution?
- Can we resurrect early life biosignatures?



Big astrobiology question: Is life repeatabl?



“I call this experiment **"replaying life's tape"** .

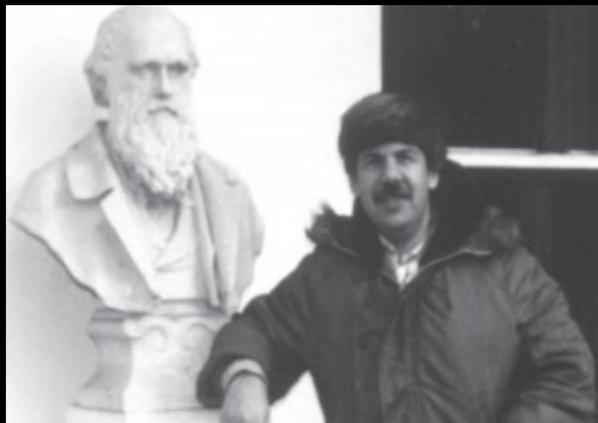
You press the rewind button and, making sure you thoroughly erase everything that actually happens, go back to any time and place in the past. Then let the tape run again and see if the repetition looks at all like the original” .



Stephen Jay Gould
Wonderful Life, 1989

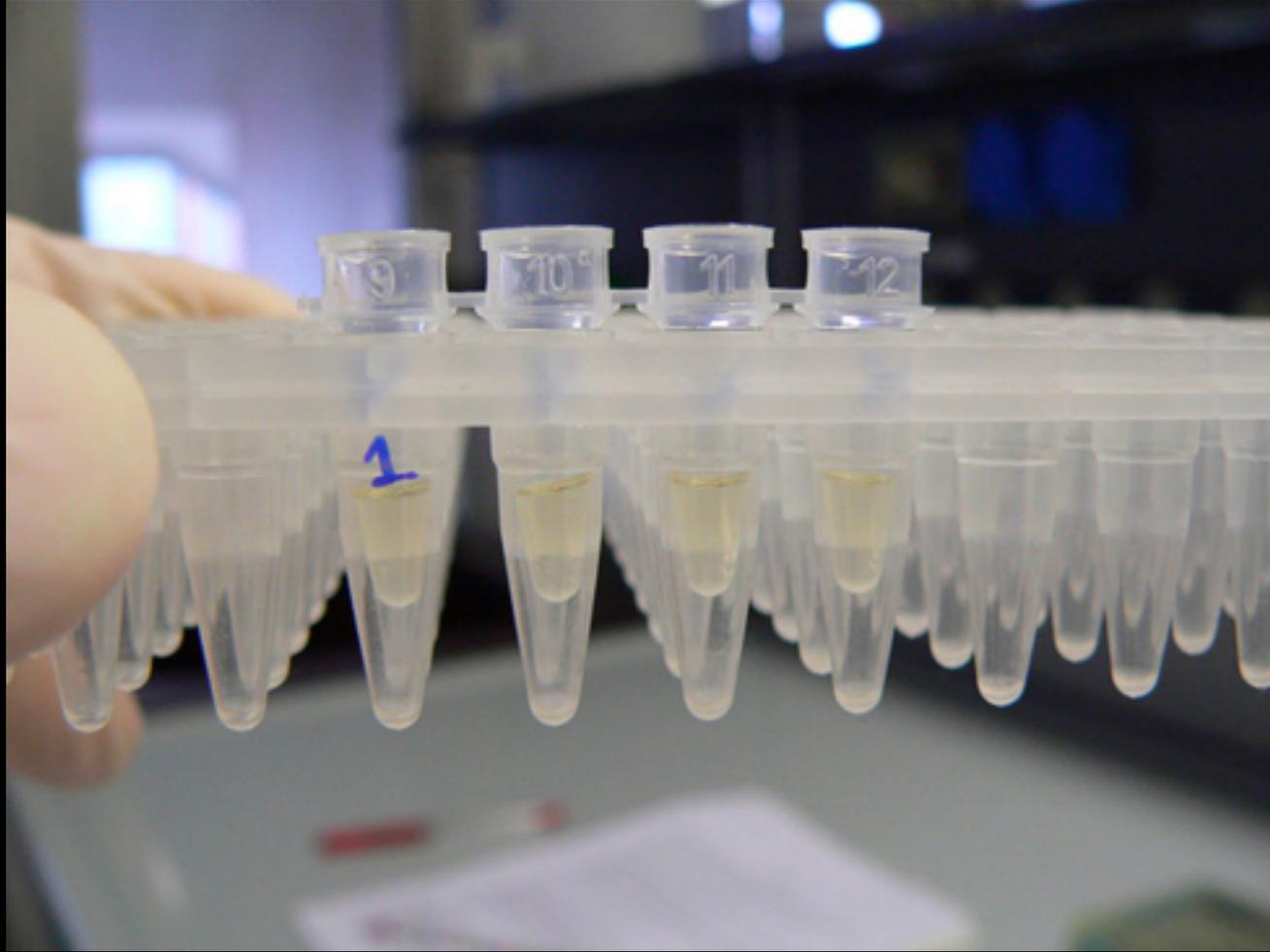
“I call this experiment "replaying life's tape".

You **press the rewind button** and, making sure you thoroughly **erase everything** that actually happens, **go back to any time** and place in the past. Then let the tape run again and see if the repetition looks at all like the original”.



Stephen Jay Gould
Wonderful Life, 1989

Experimental evolution of ancient genes in the laboratory using engineered microbes

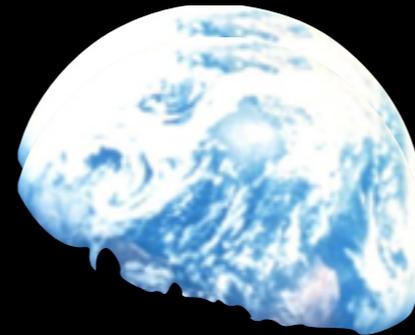


Rewind and replay the tape of life (one molecule at a time)

Kacar et al. J Mol Evol 2017

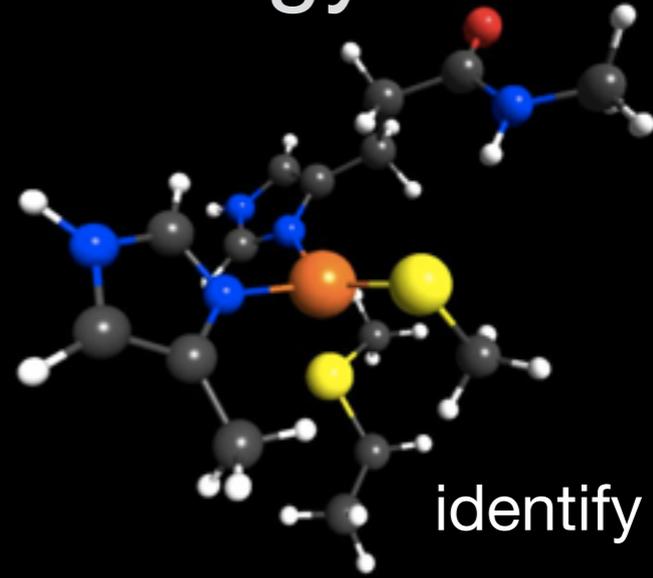
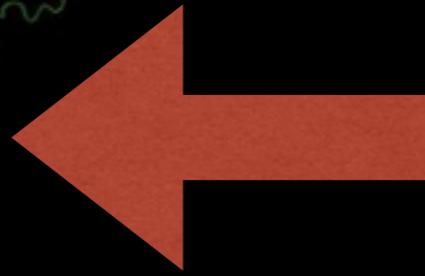
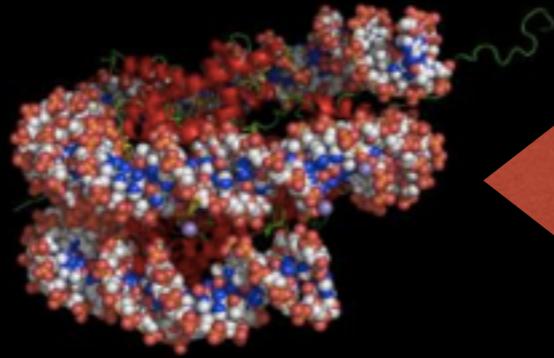
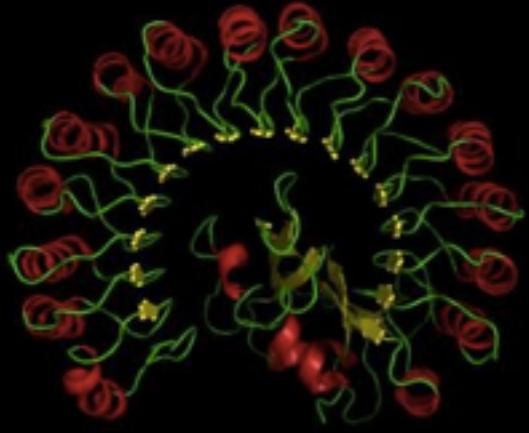
Motivating Questions

- How can we travel back in evolutionary time?
- How repeatable is life's evolution?
- Can we resurrect early life biosignatures?



Modern life preserves signs of ancient protobiology

How are the chemicals used in early biology?



identify organics

reconstruct protobiology that used these types of primitive molecules



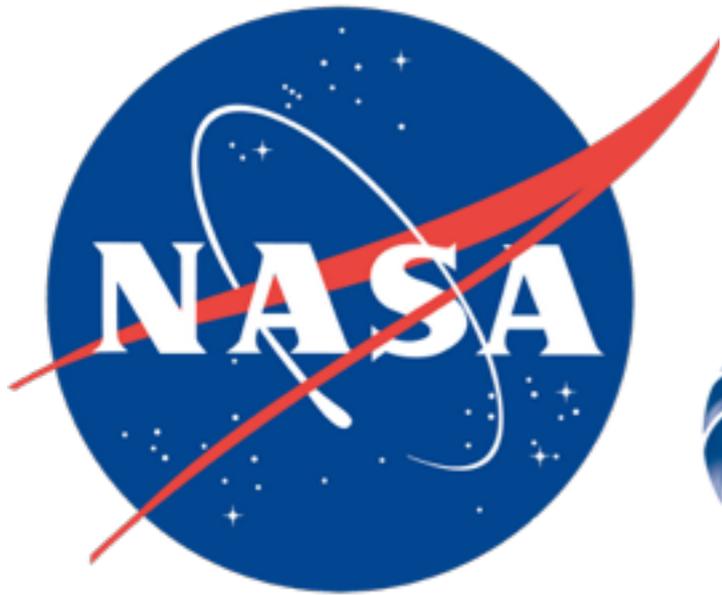
CHNOPS

Current / Future



Asteroid landing missions

Funding



John Templeton Foundation



- NASA Postdoctoral Program
- NASA Exobiology and Evolutionary Biology Program
- NASA Early Career Science Award
- NASA Astrobiology “Ribosome and Evolution” CAN5
- NASA Astrobiology “Reliving the Past” CAN7

Current Collaborators

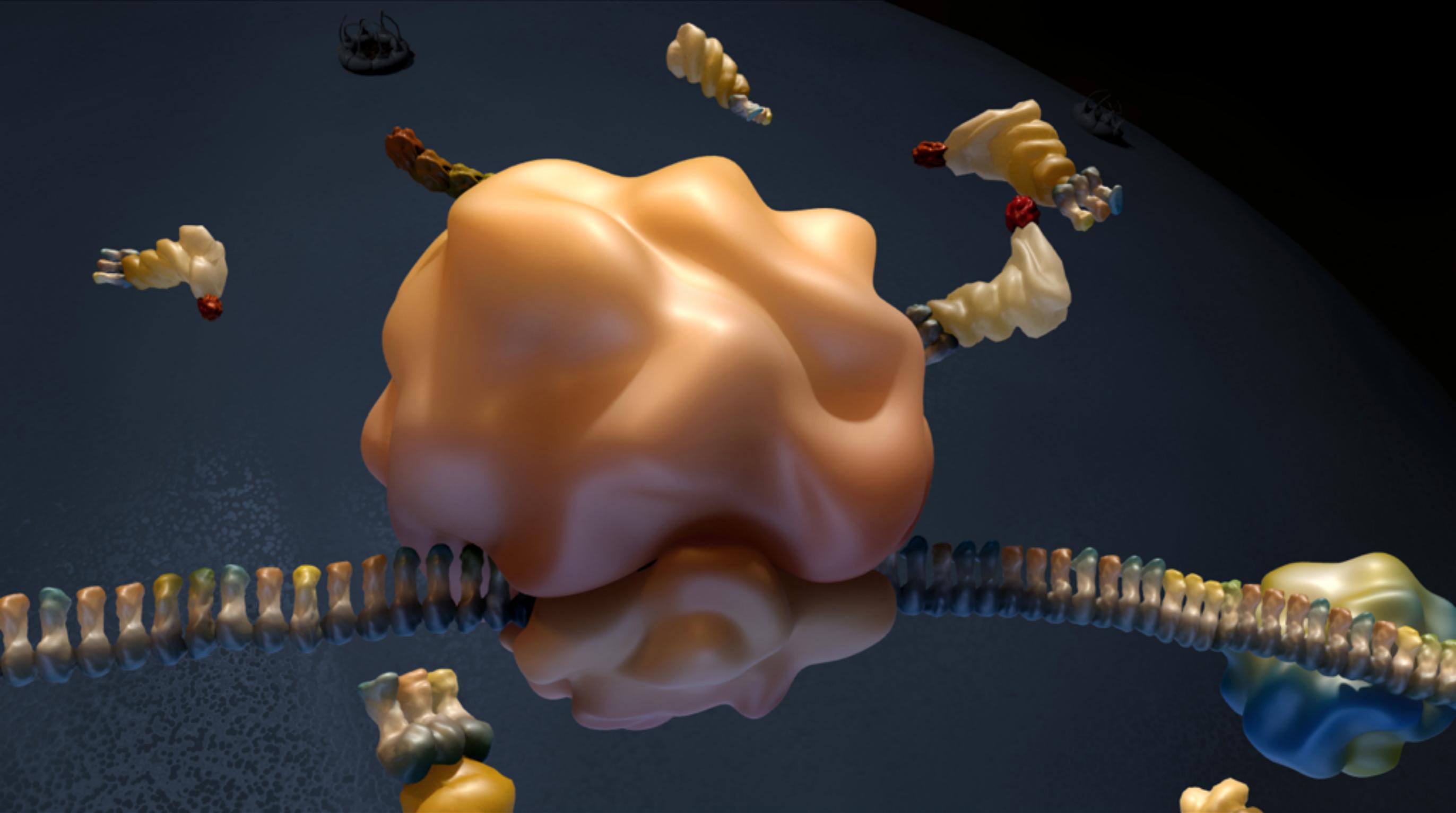
- Kate Adamala (U Minnesota)
- Dan Andersson (Uppsala)
- John Baross (U Wash)
- Michael Brenner (Harvard)
- Church Lab (HMS)
- Greg Fournier (MIT)
- Lionel Guy (Uppsala)
- Dave Johnston (Harvard)
- O’Shea Lab (Harvard)
- Margie Kinnerseely (U Montana)
- Sergey Kryazhimskiy (UCSD)
- Ann Pearson (Harvard)
- Desai Lab (Harvard)
- Daniel Segre (Boston U)
- Eric Smith (ELSI, Santa Fe)



Thank you NASA!

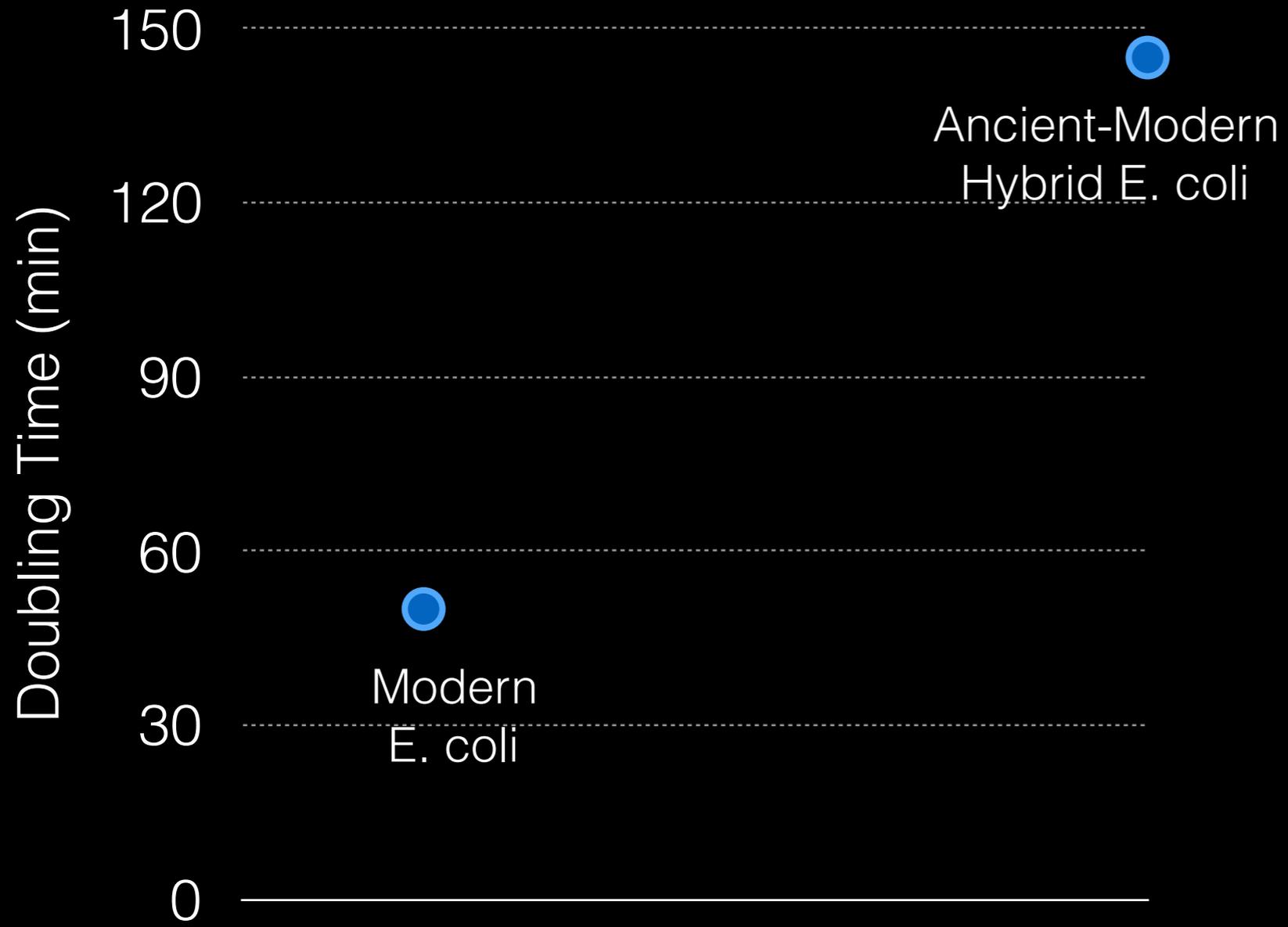
contact: kacar@g.harvard.edu

Ribosome is the protein making machinery of every terran cell



Replace ribosome components with ancient counterparts

Impact of ancient gene on bacterial growth



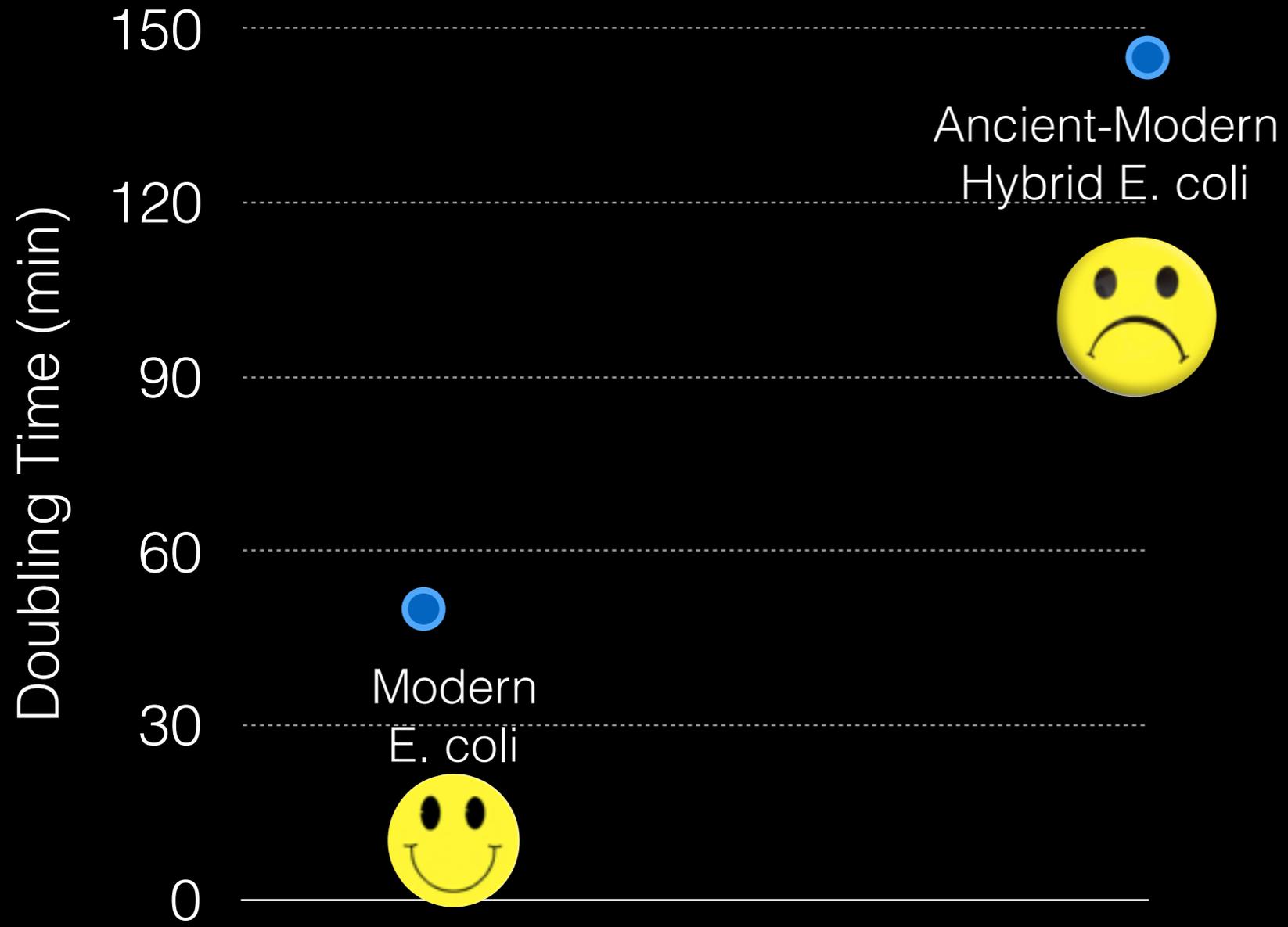
[Kacar and Gaucher, Artificial Life, 2013]

[Kacar and Gaucher, Biochem J, 2014]

[Kacar, Chance in Evolution, U Chicago Press, 2016]

[Kacar, Ge, Sanyal and Gaucher, J Mol Evol, in press]

Impact of ancient gene on bacterial growth



It's alive! (but sick)

[Kacar et al, Artificial Life, 2013]
[Kacar et al, Biochem J, 2014]
[Kacar, Chance in Evolution, U Chicago Press, 2016]
[Kacar et al, J Mol Evol, 2017]