

Range Expansions & Spatial Population Genetics

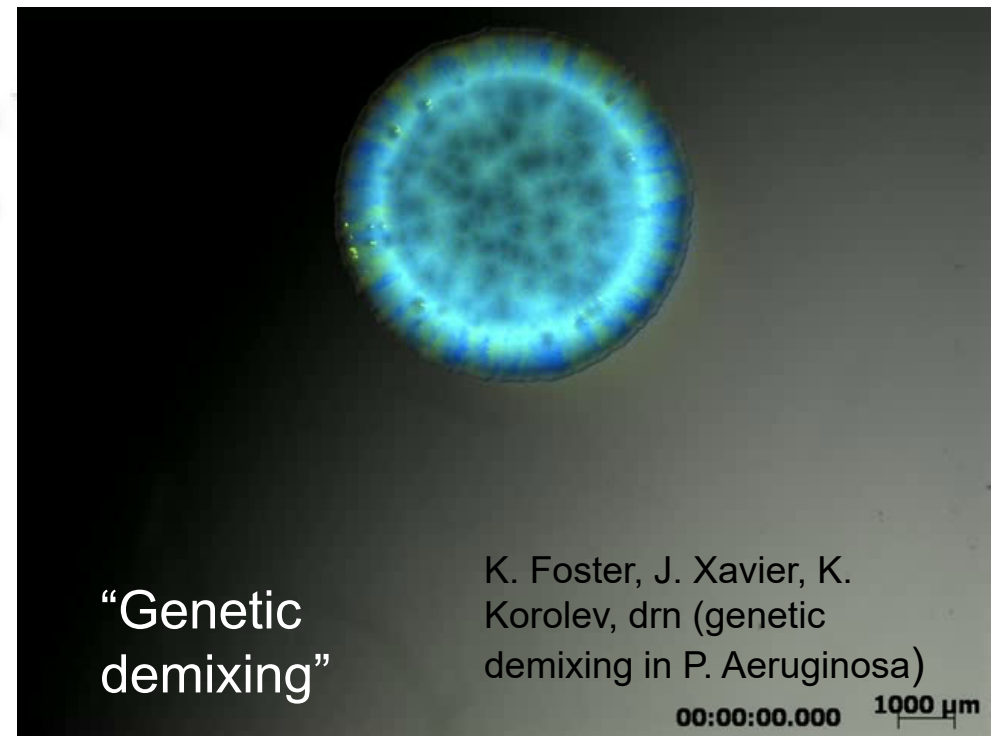
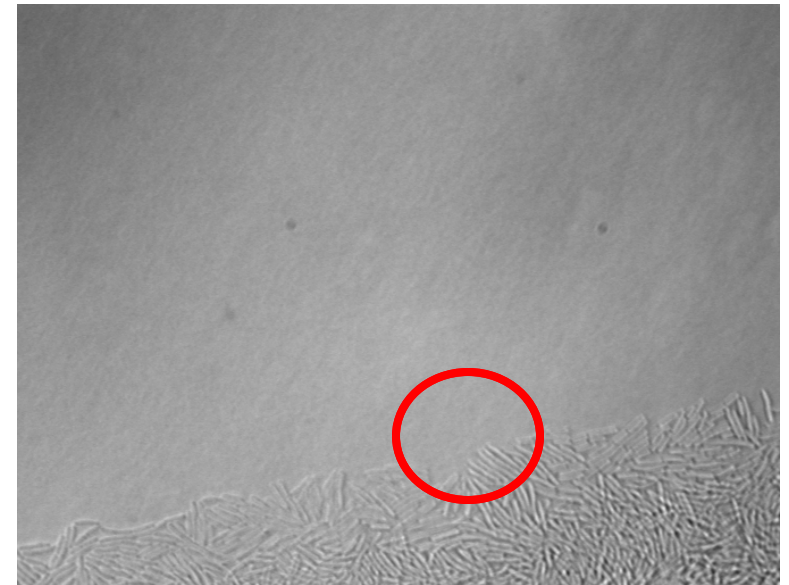


In 500 generations....

Large mammals expand over $\sim 10^4$ km

But bacteria (in a Petri dish) only expand ~ 1 cm

Oskar Hallatschek & drn, (*E. coli*)



“Genetic demixing”

K. Foster, J. Xavier, K. Korolev, drn (genetic demixing in *P. Aeruginosa*)

00:00:00.000 1000 μ m

Gene Surfing and Survival of the Luckiest

Mutations and competition in a spreading population

J. Xavier
K. Foster
W. Moebius
M. Mueller
A. Murray



Oskar Hallatschek



Kirill Korolev



vs.

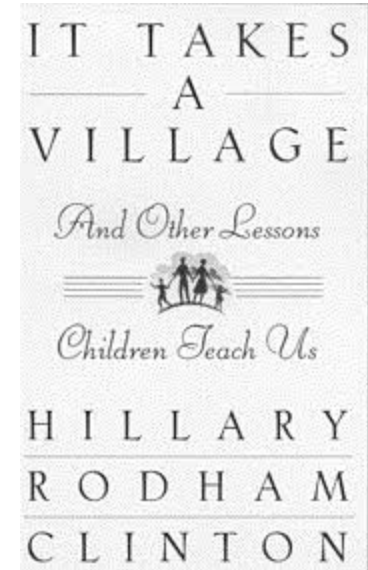
*Is this
Biology,
Physics,
or....?*



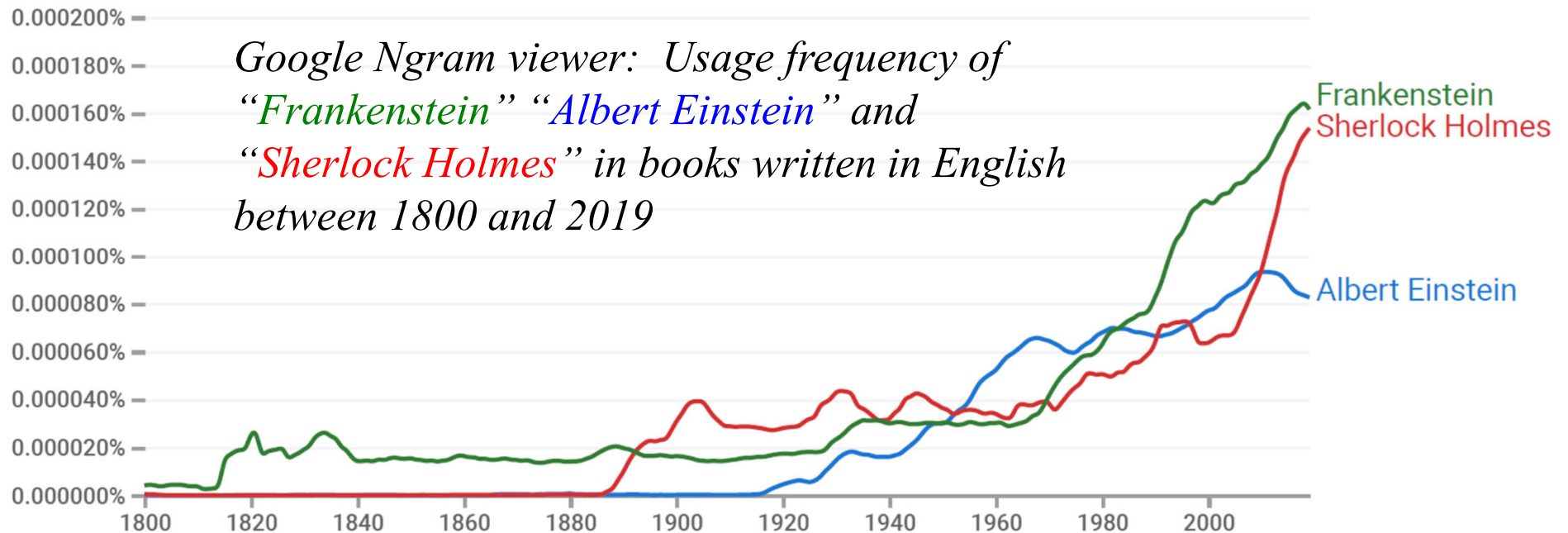
Will the 21st century be the “Century of Biology”?

•Bill Clinton’s exchange with his science advisor upon completion of the first draft of the human genome.....

“If, in scientific terms, the twentieth century has been the century of physics, then the twenty-first will surely be the century of biology.”



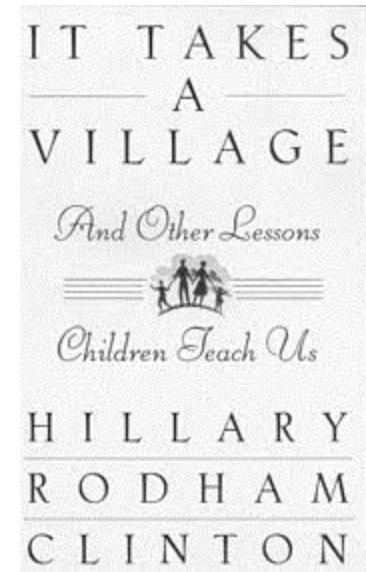
Is this really true?



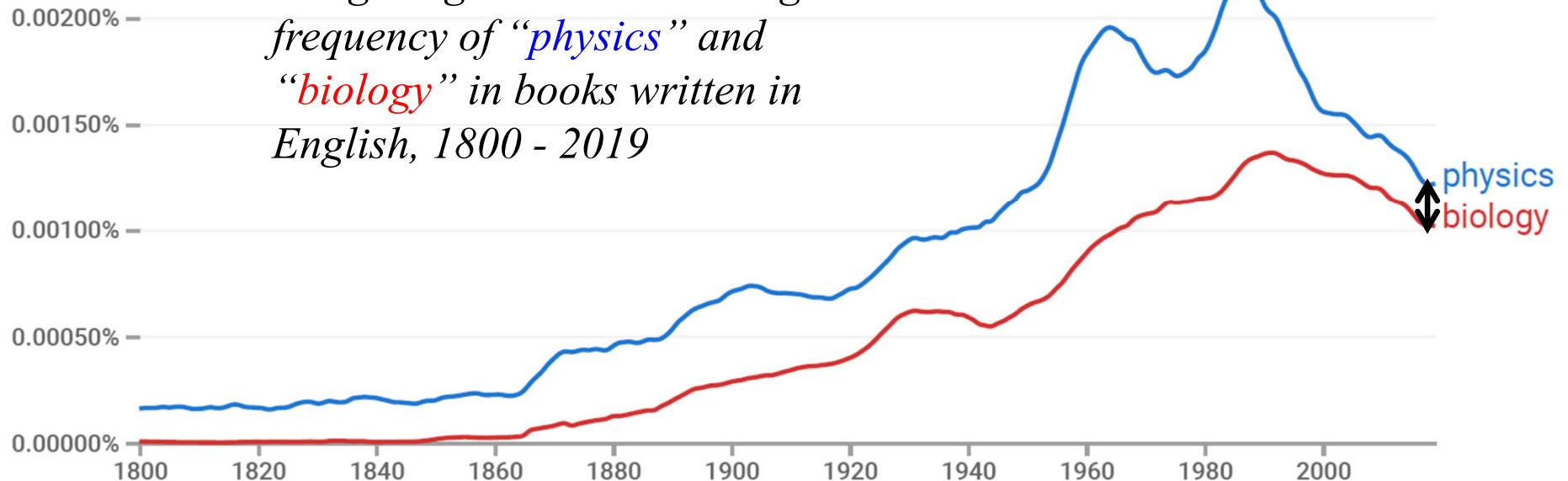
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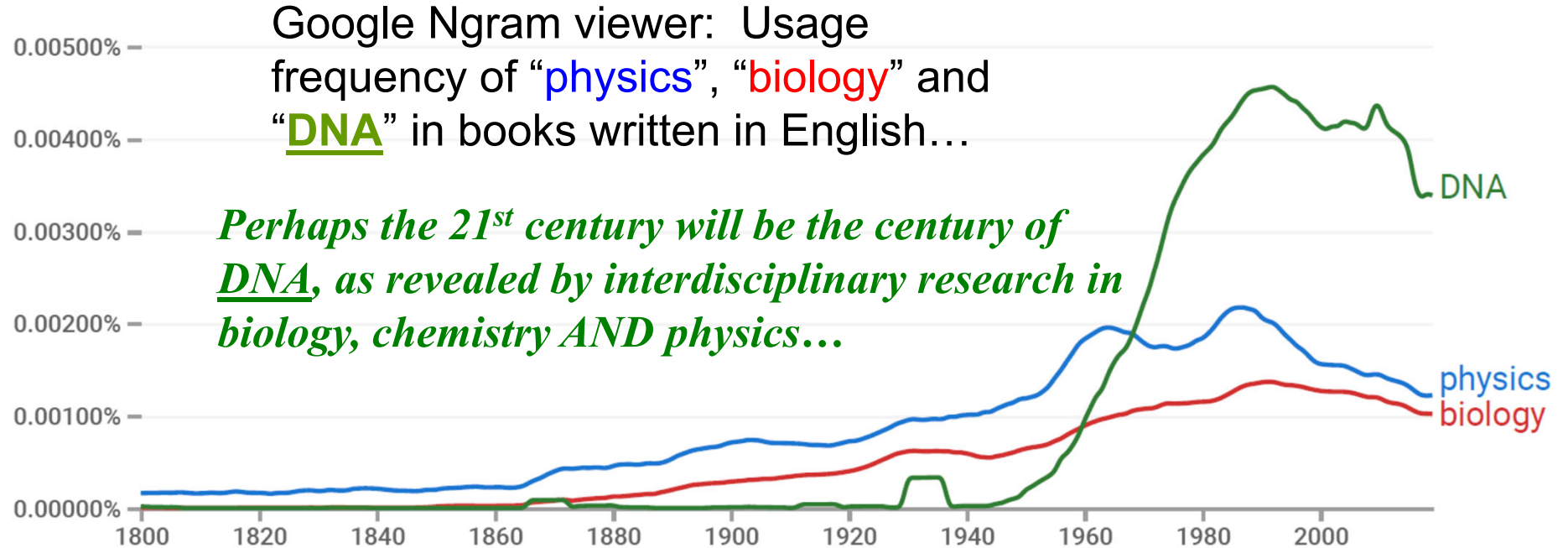
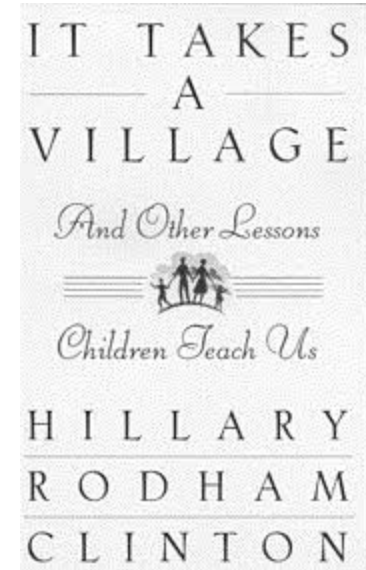
Google Ngram viewer: Usage frequency of “*physics*” and “*biology*” in books written in English, 1800 - 2019



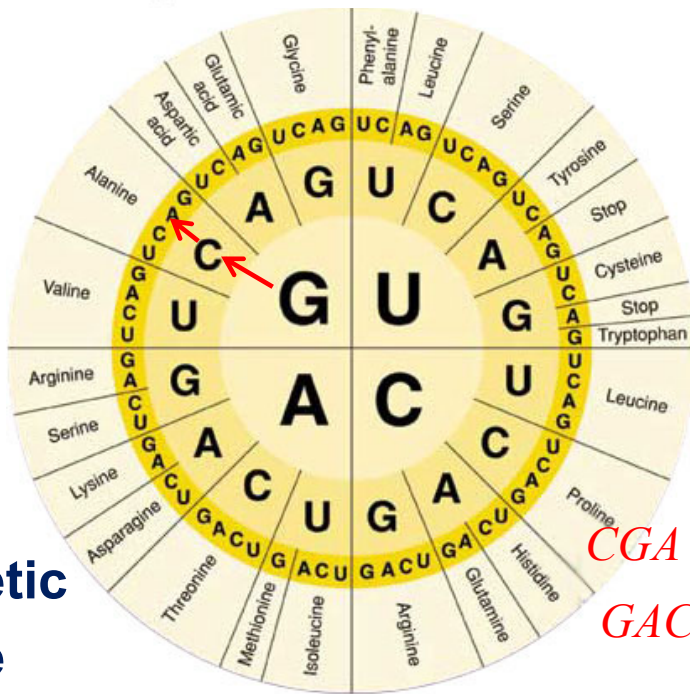
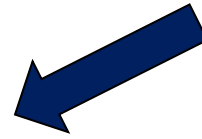
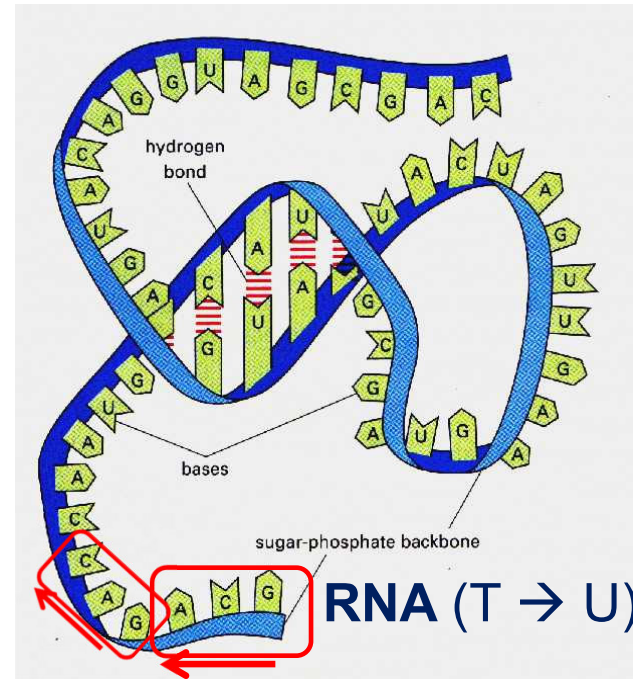
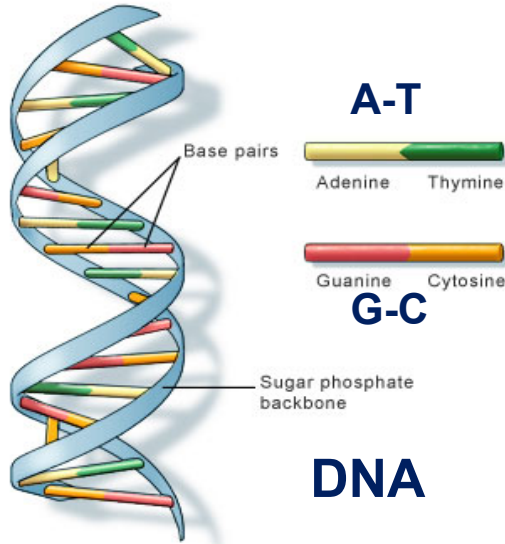
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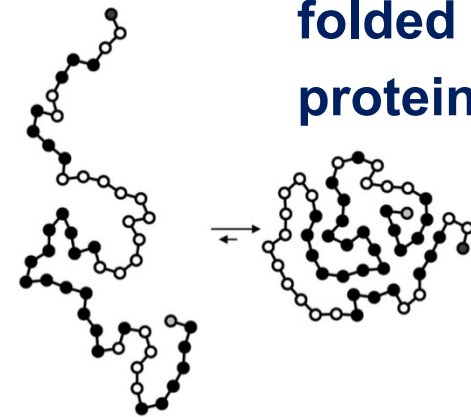


“Central Dogma” of Biology



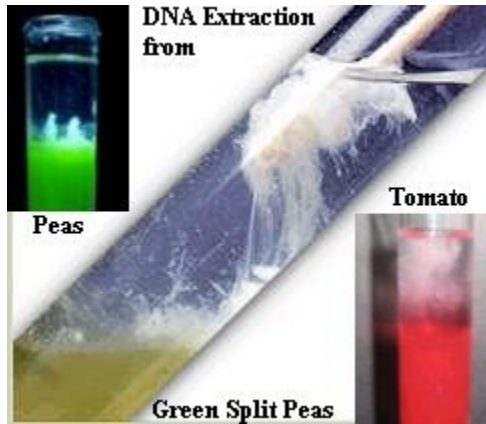
CGA = alanine

GAC = aspartic acid

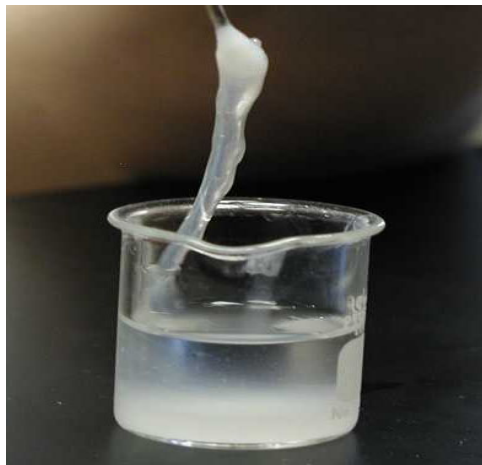


- = water-loving amino acid
- = water-hating amino acid

Diffraction from oriented fibers of DNA: “Photo 51”



•<http://universe-review.ca/R11-16-DNAsequencing.htm>



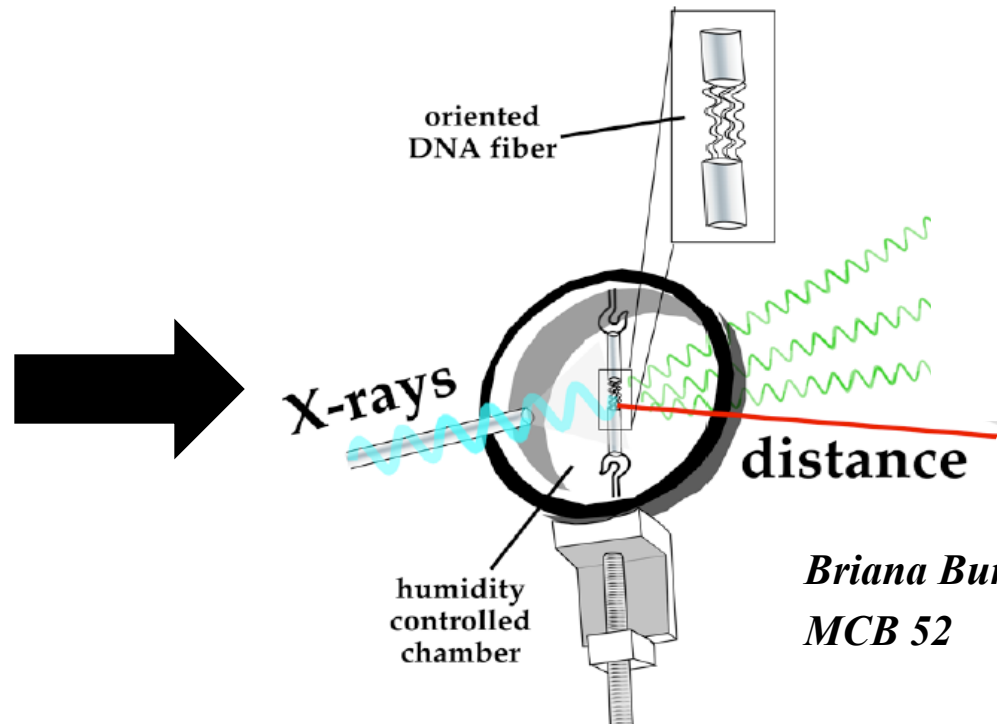
•http://biology.clc.uc.edu/fankhauser/labs/genetics/dna_isolation/thymus_dna.htm



Rosalind Franklin

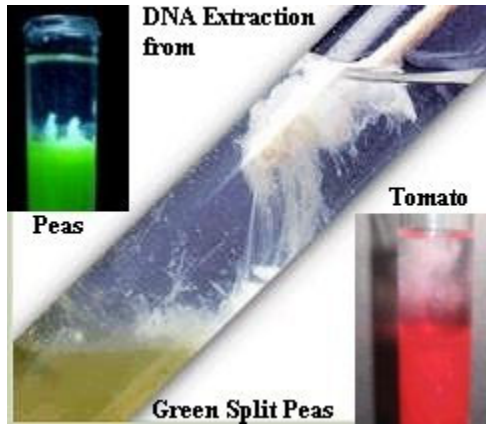


Maurice Wilkins

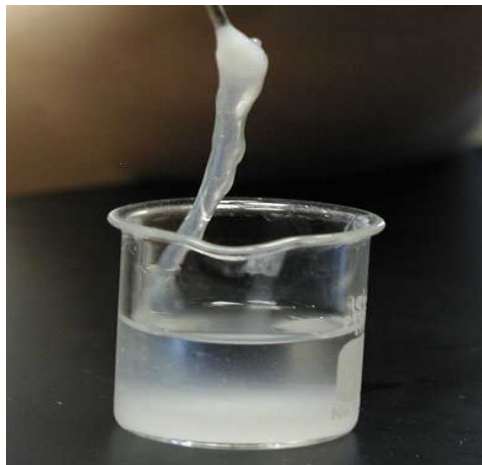


*Briana Burton
MCB 52*

Diffraction from oriented fibers of DNA: “Photo 51”



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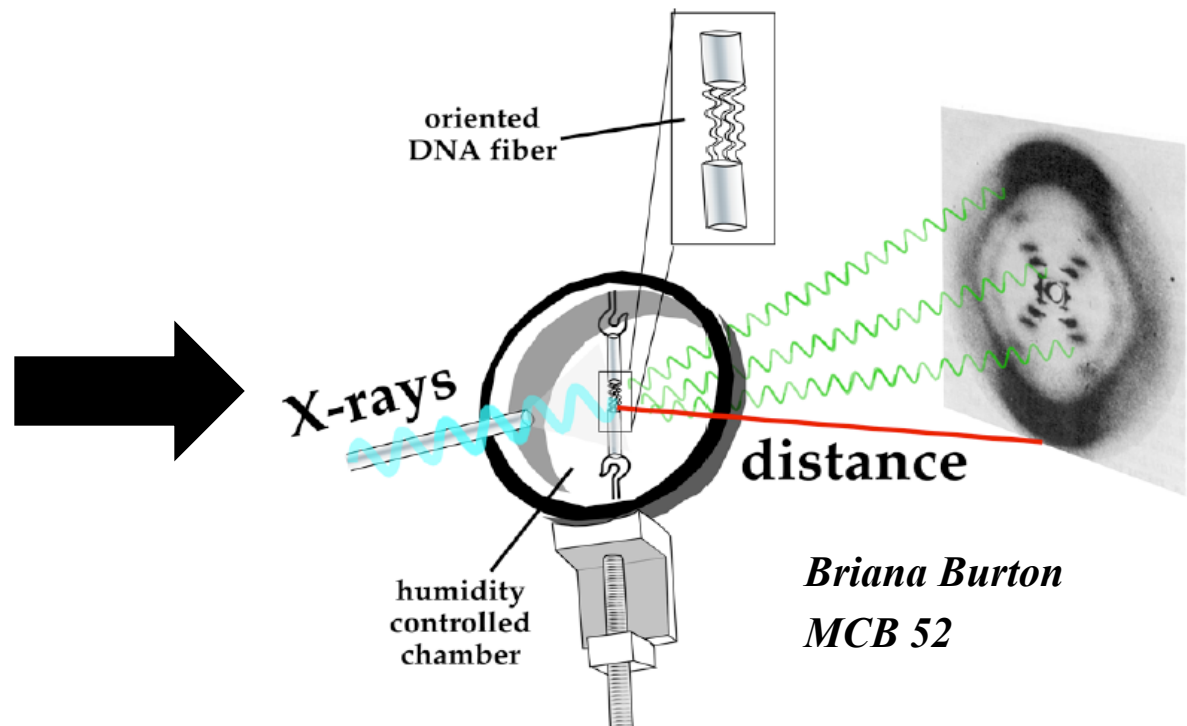
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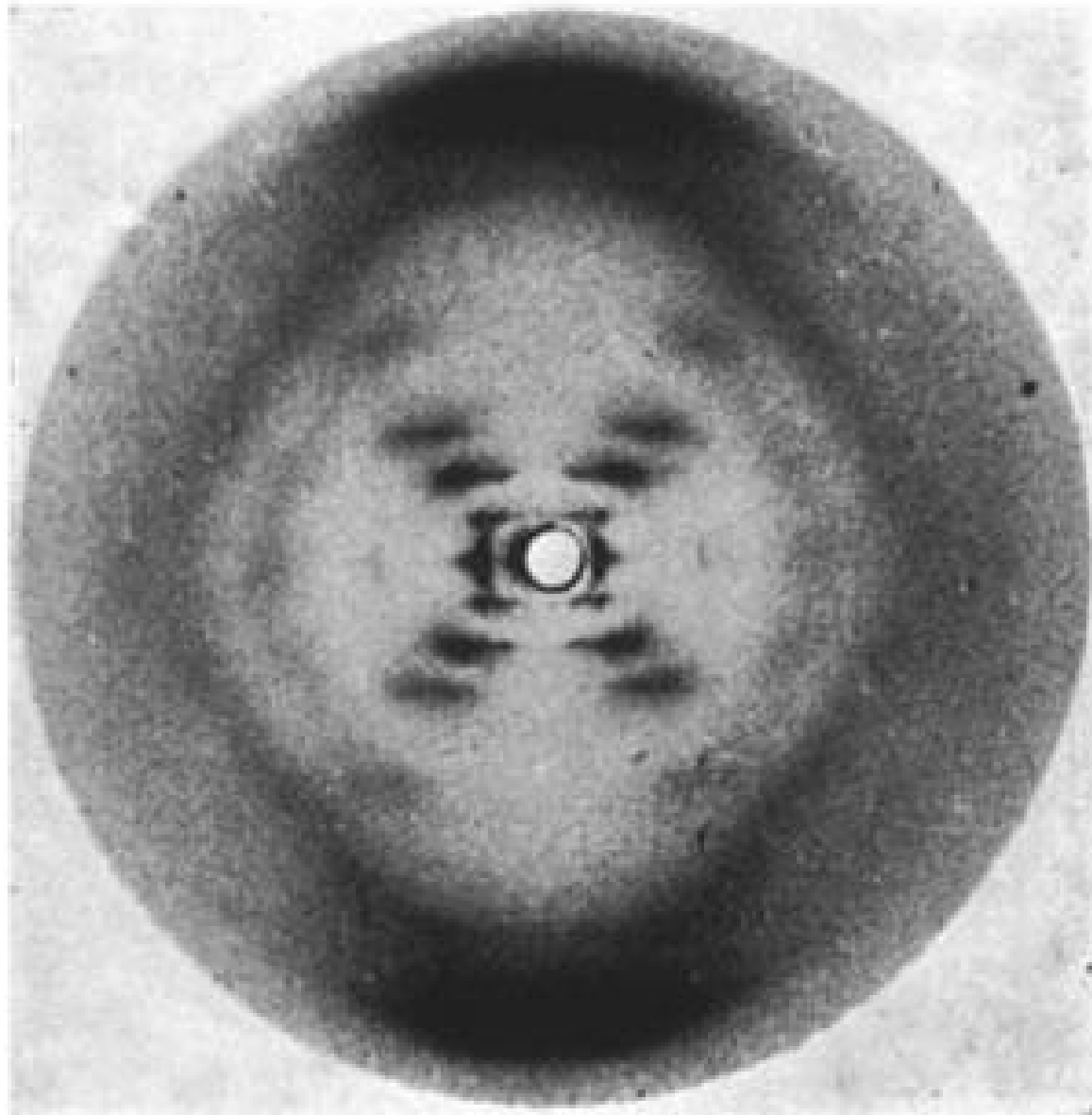
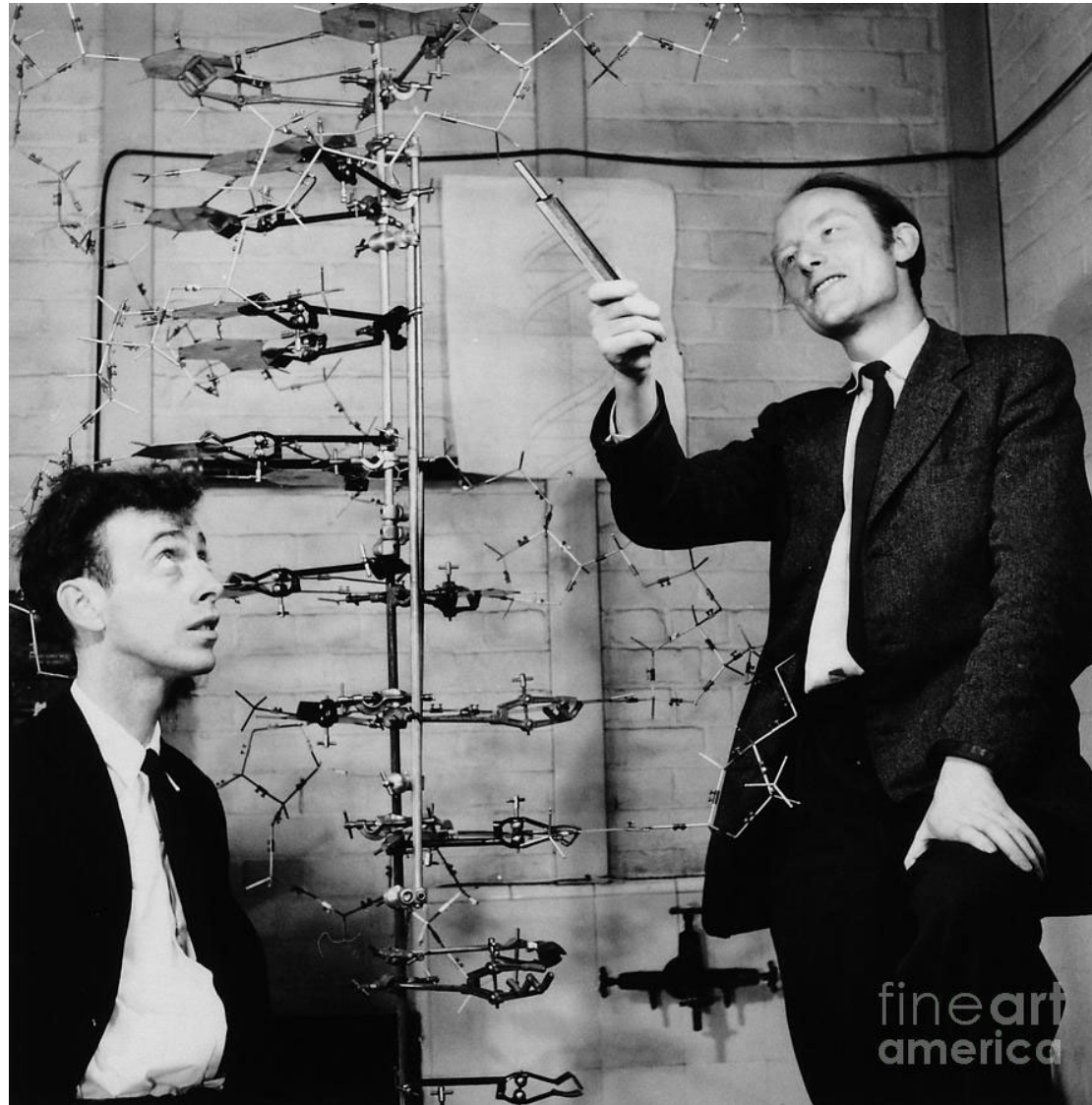


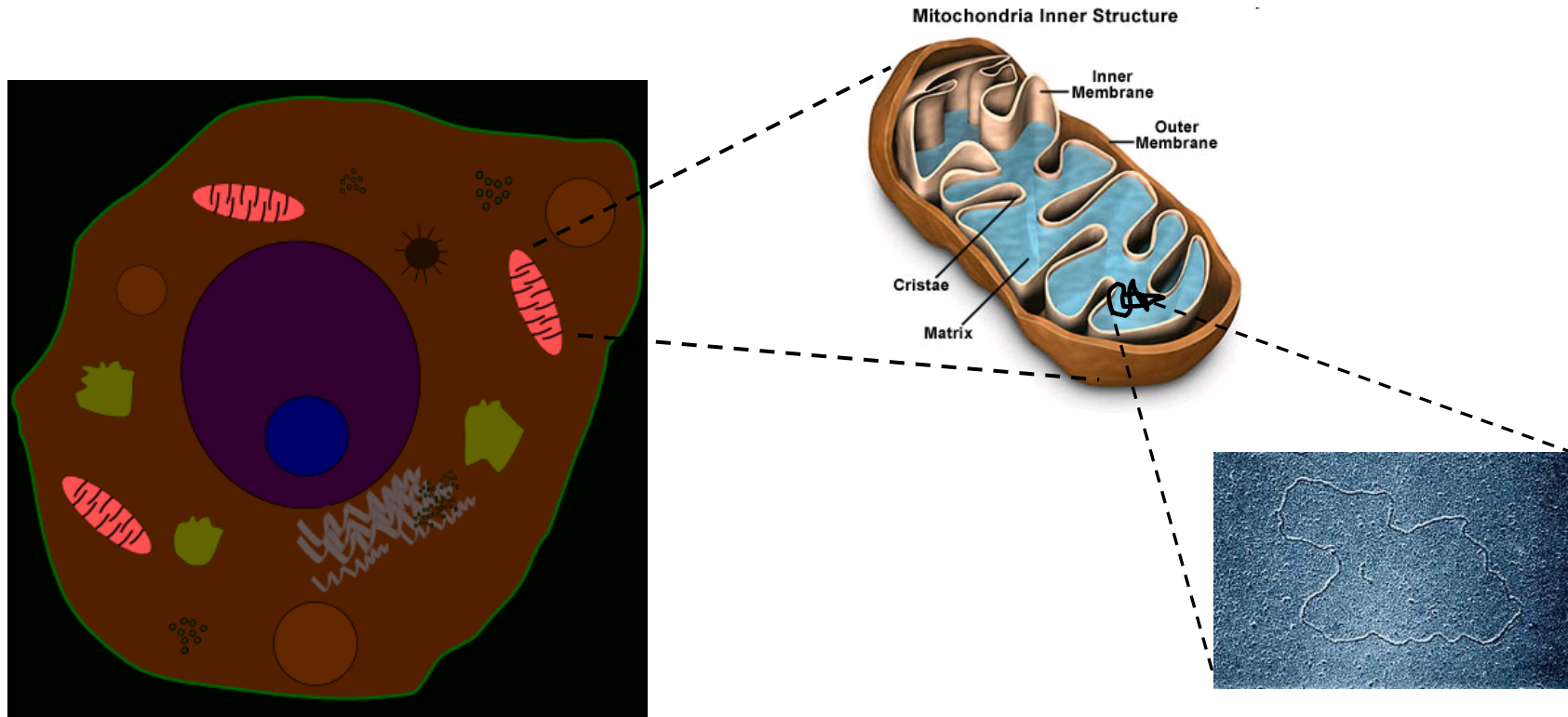
Photo 51: May 2, 1951

Double helix structure of DNA



James Watson and Francis Crick

Mitochondria supply energy to plant and animal cells

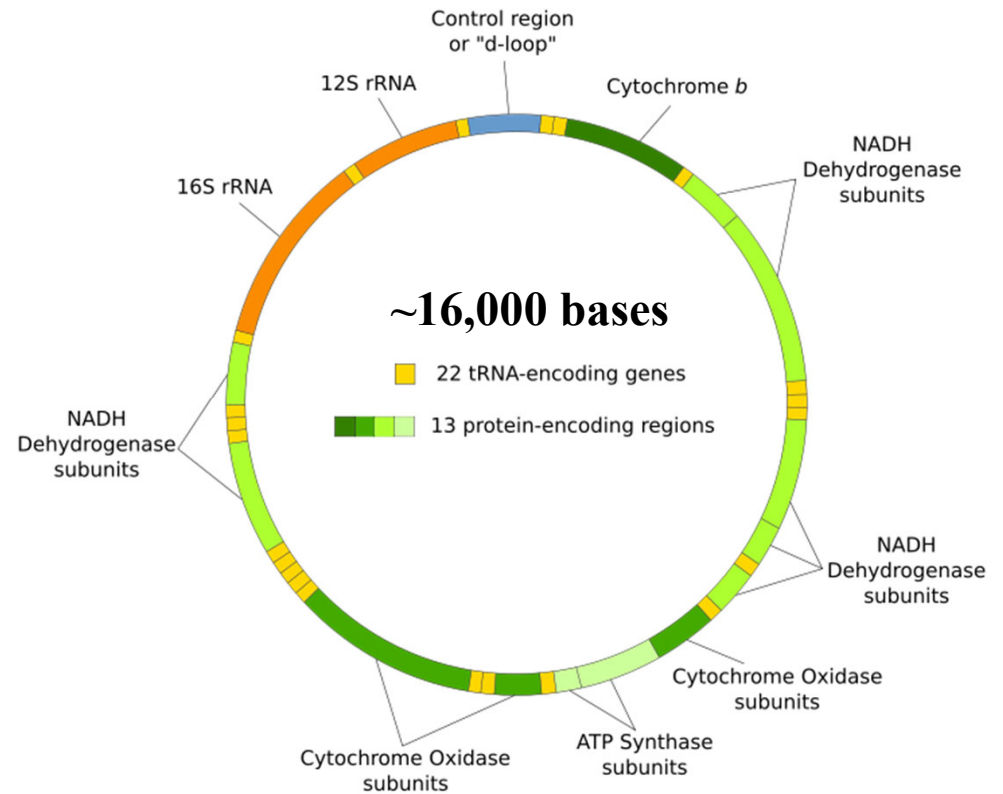


- *Each mitochondrion contains a small circular chromosome, with some “junk DNA”, as well as genes that allow it to supply energy to the cell.*
- *100's of mitochondria in each human cell, each with 2-10 copies of a circular genome ($\sim 10^3$ gene replicas per cell)*
- *Mitochondria likely have a very ancient origin, and even possess a slightly different genetic code! (different start and stop codons, tryptophan)*

Divergence times from mitochondrial DNA



Lynn Margulis

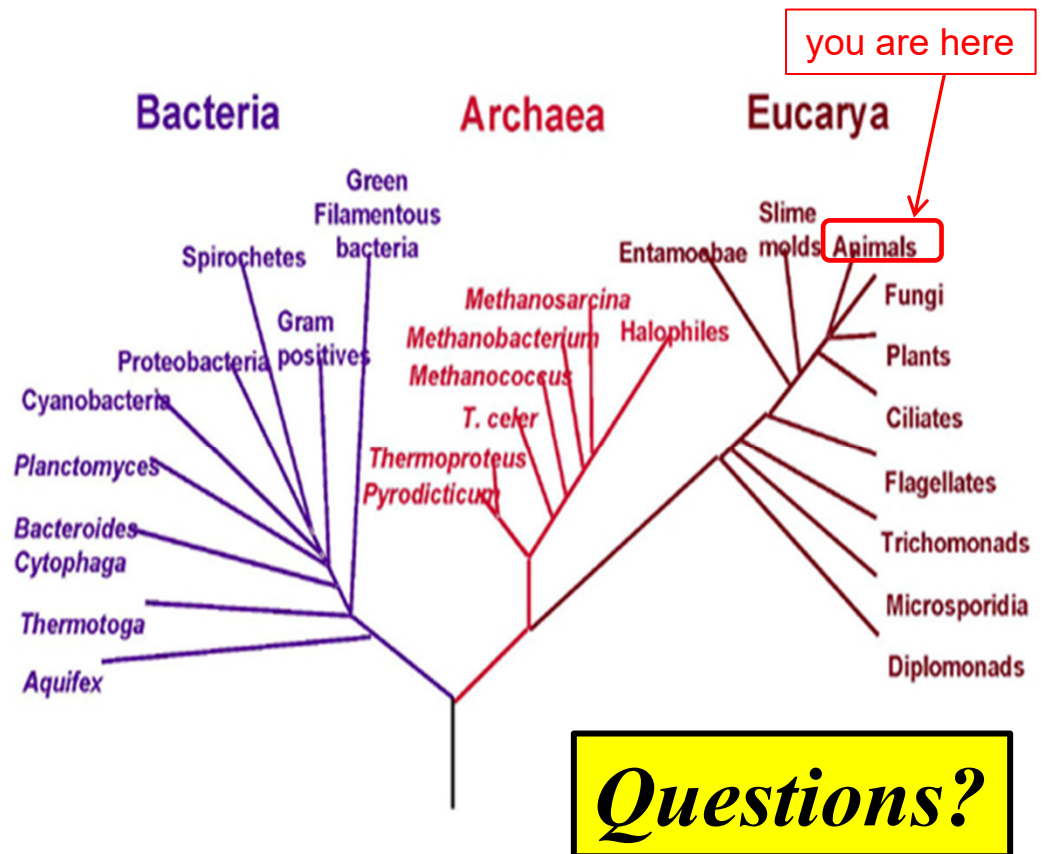
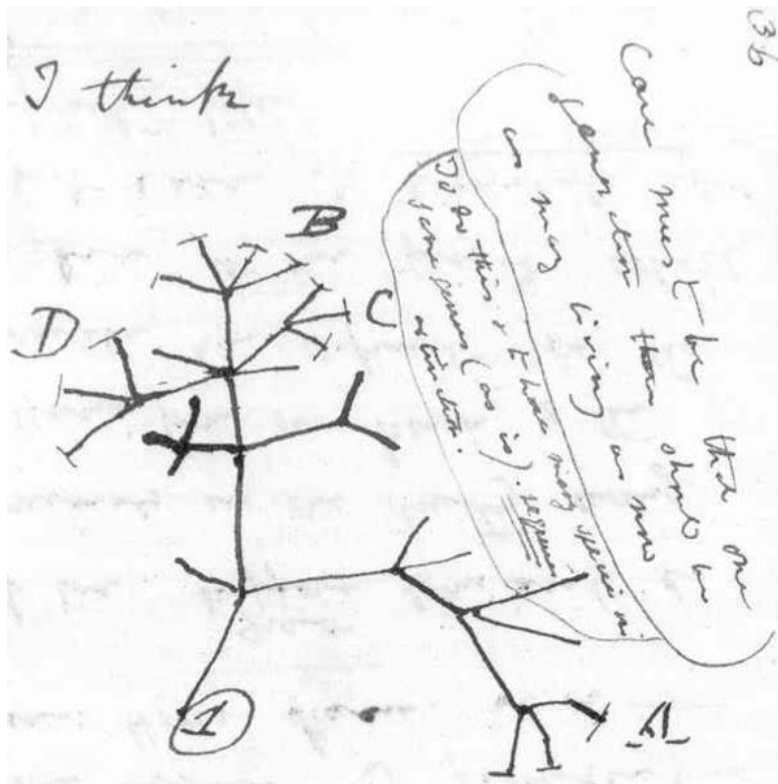


- *Mitochondria descended from bacteria engulfed by ancestral animal cells*
- *Inherited along matrilineal lines*
- *Substitution rate 5-10 faster than nuclear DNA*
- *Conclude, e.g., that chimps and humans diverged about 6-7 M years ago...*

Genes and genealogies

*Charles Darwin's sketch
of an evolutionary tree
of life (circa 1837),
Museum of Natural
History, NYC*

*Modern tree of life obtained by
sequencing DNA encoding for
ribosomal RNA*



Spatial population genetics

What's in a name?



*Hyskenstræde 9, 1207
København K. Denmark*

*I am not related to the
Lord Nelson who fought
the Danes at the battle of
Copenhagen in 1801....*



What is the distribution of genes in space?

Haplogroup I1 is a Y chromosome haplogroup occurring at greatest frequency in Scandinavia, associated with SNP (“single nucleotide polymorphism”) mutations such as M253.

Frequency is about 35% in southern Norway, southeastern Sweden, especially on the island of Gotland, and Denmark



http://en.wikipedia.org/wiki/Haplogroup_I_%28Y-DNA%29

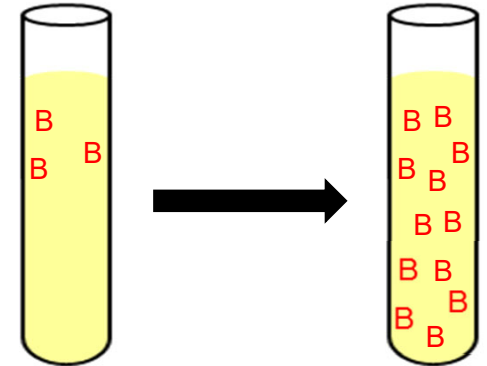
Malthus-Verhulst Theory of Population Dynamics

change in population size
per unit time = births - deaths + "saturation"

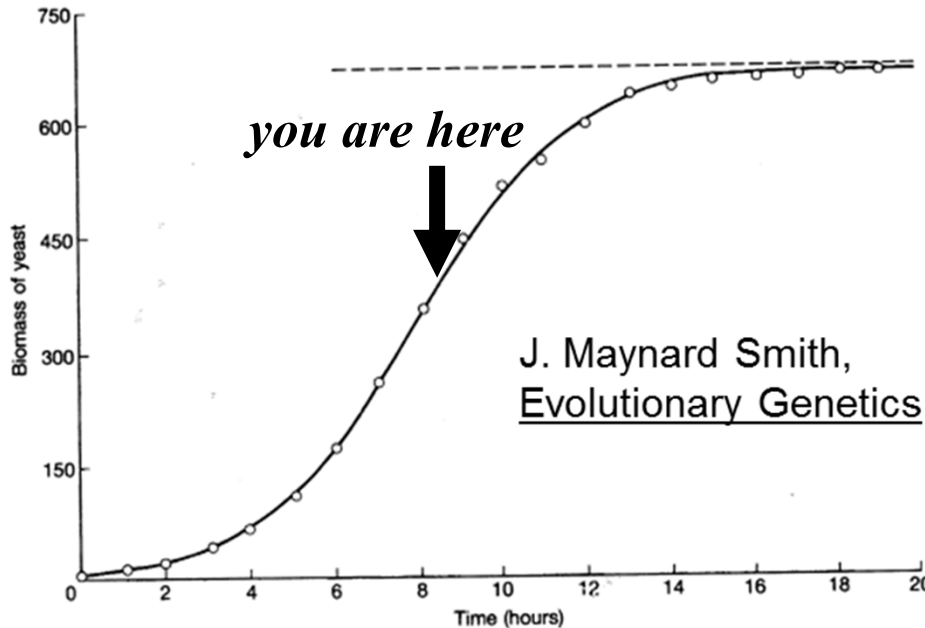
c_n = population size at generation n

$$c_{n+1} = 2c_n = 2^2 c_{n-1} = \dots = 2^{n+1} c_0$$

more generally, $c_{n+1} = 2 \cdot (1 - c_n / 2K) \cdot c_n$
in a doubling time, K = carrying capacity



1798 T.R. Malthus



1836 P.F. Verhulst



Fisher Population Waves In One Dimension

change in population size
per unit time = births - deaths + saturation + spatial diffusion

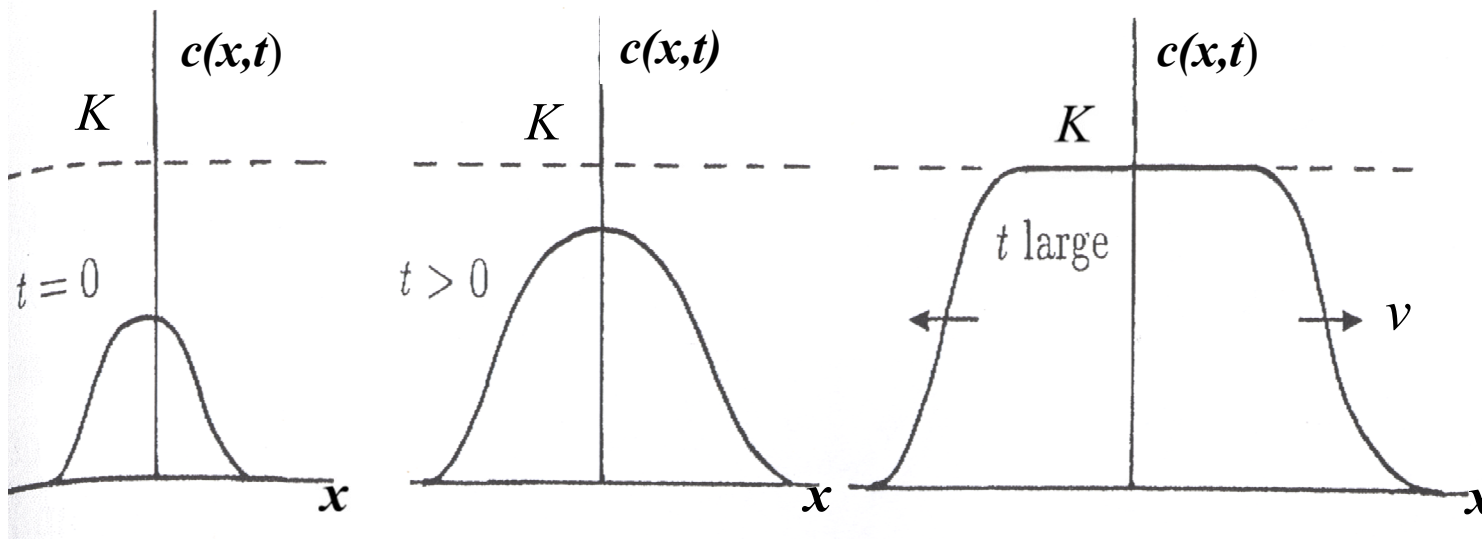
Let $c(x,t)$ be the density of organisms at position x at time t ...

$$\frac{\partial}{\partial t} c(x,t) = D \frac{\partial^2}{\partial x^2} c(x,t) + ac(x,t) - bc^2(x,t); \text{ let } c(x,t) = f(x-vt)$$



1932

R. A. Fisher

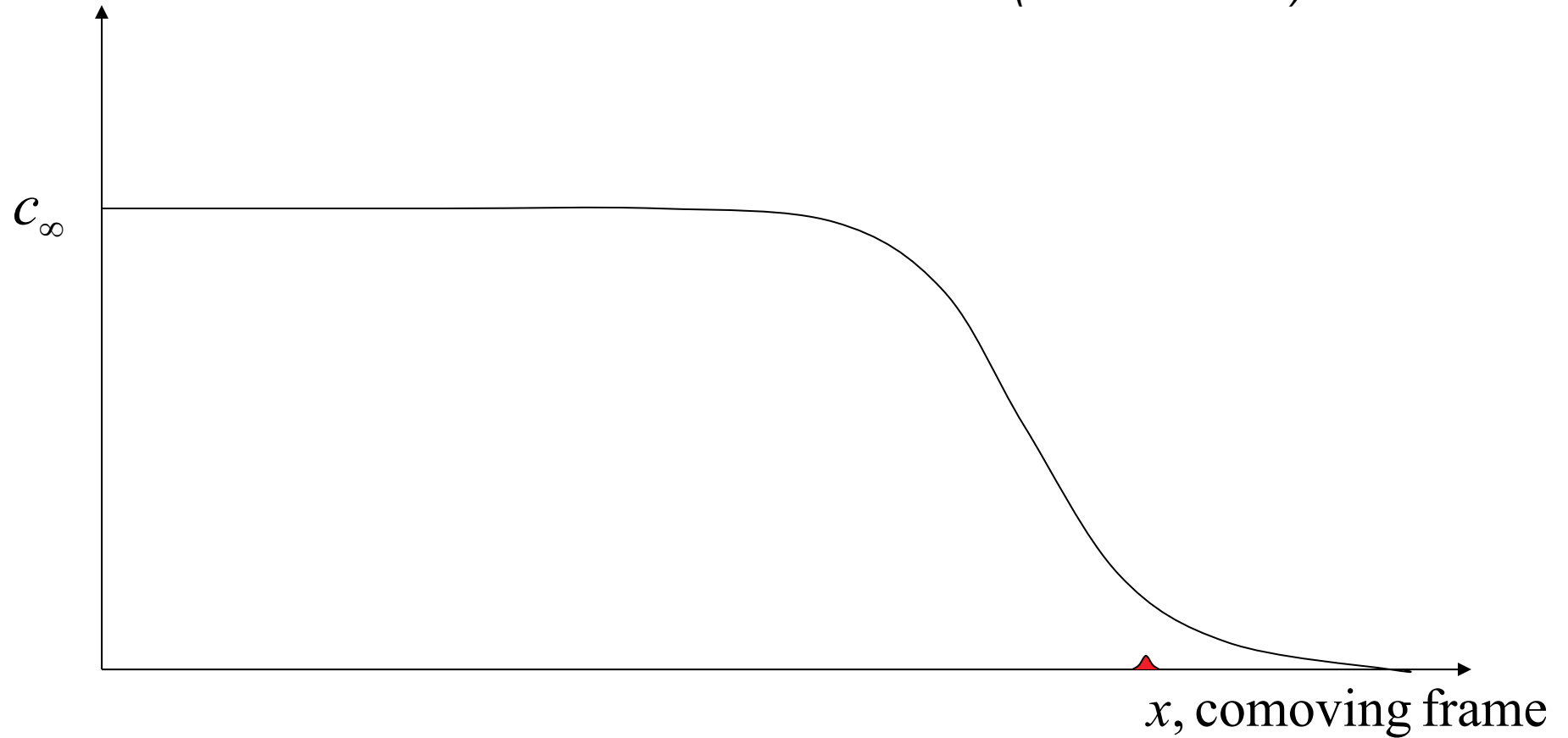


Schematic time development of a wavefront solution of Fisher's equation on the infinite line. (J.D. Murray, Mathematical Biology)

Successful Surfing (1d)

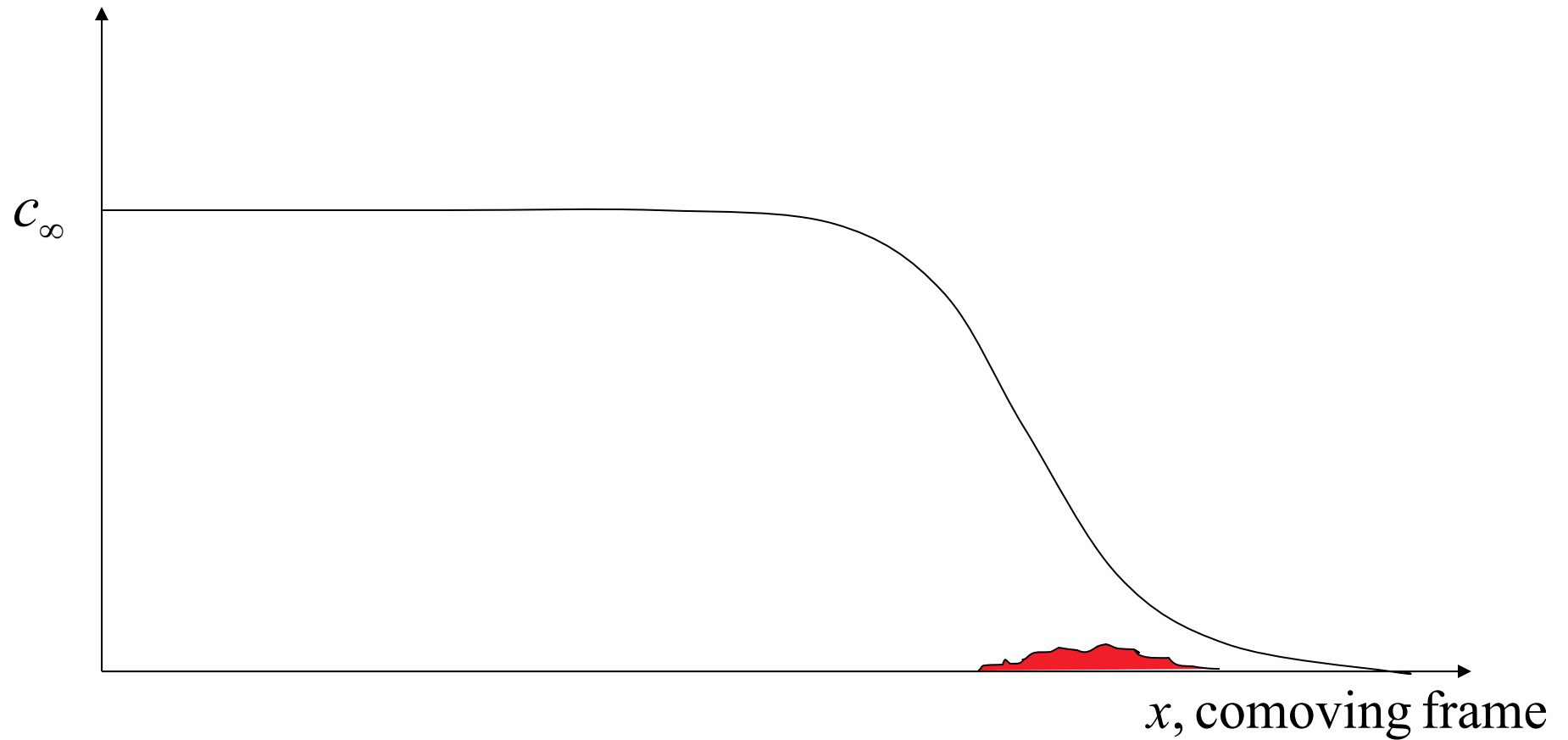
$c_s(x)$, steady state population density

(O. Hallatschek)



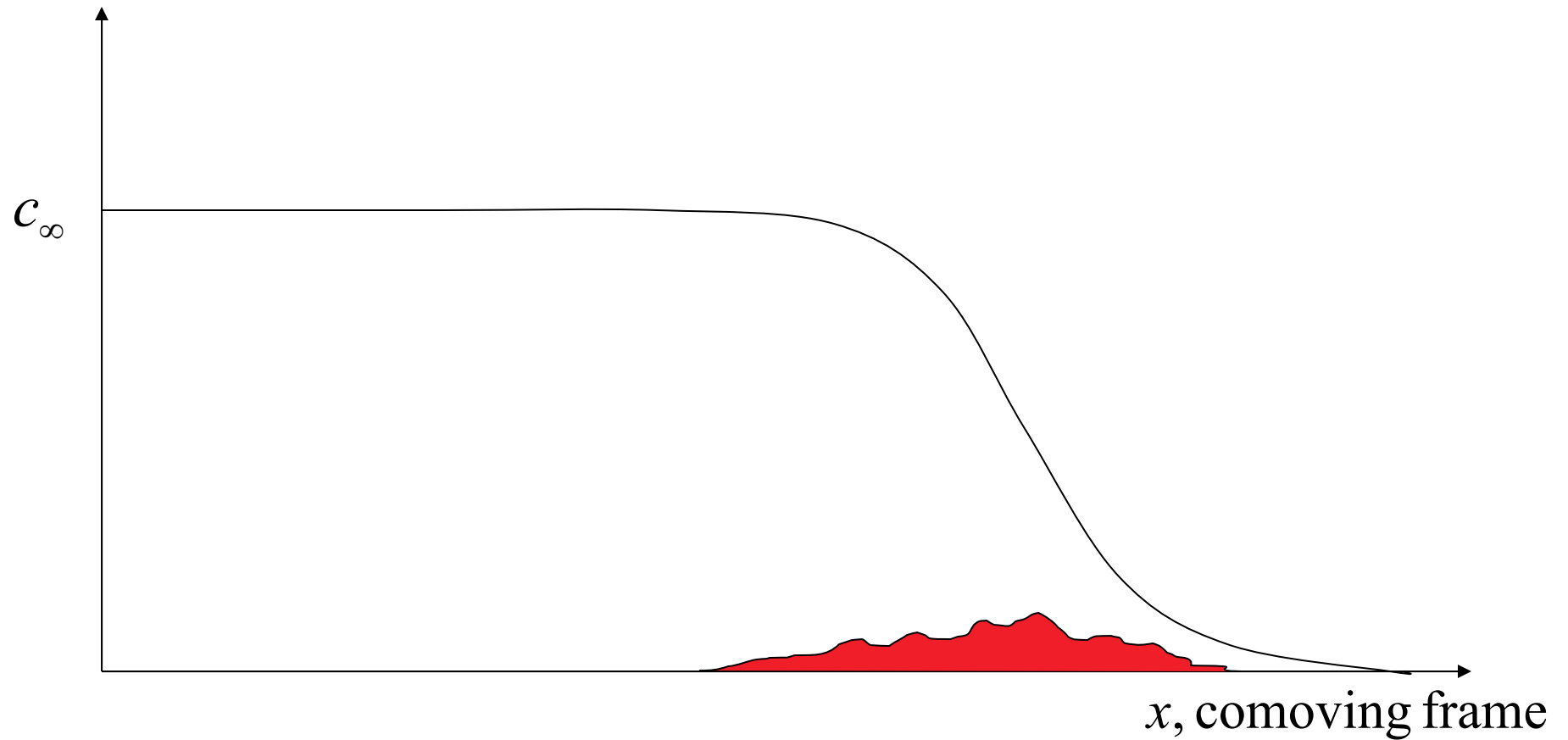
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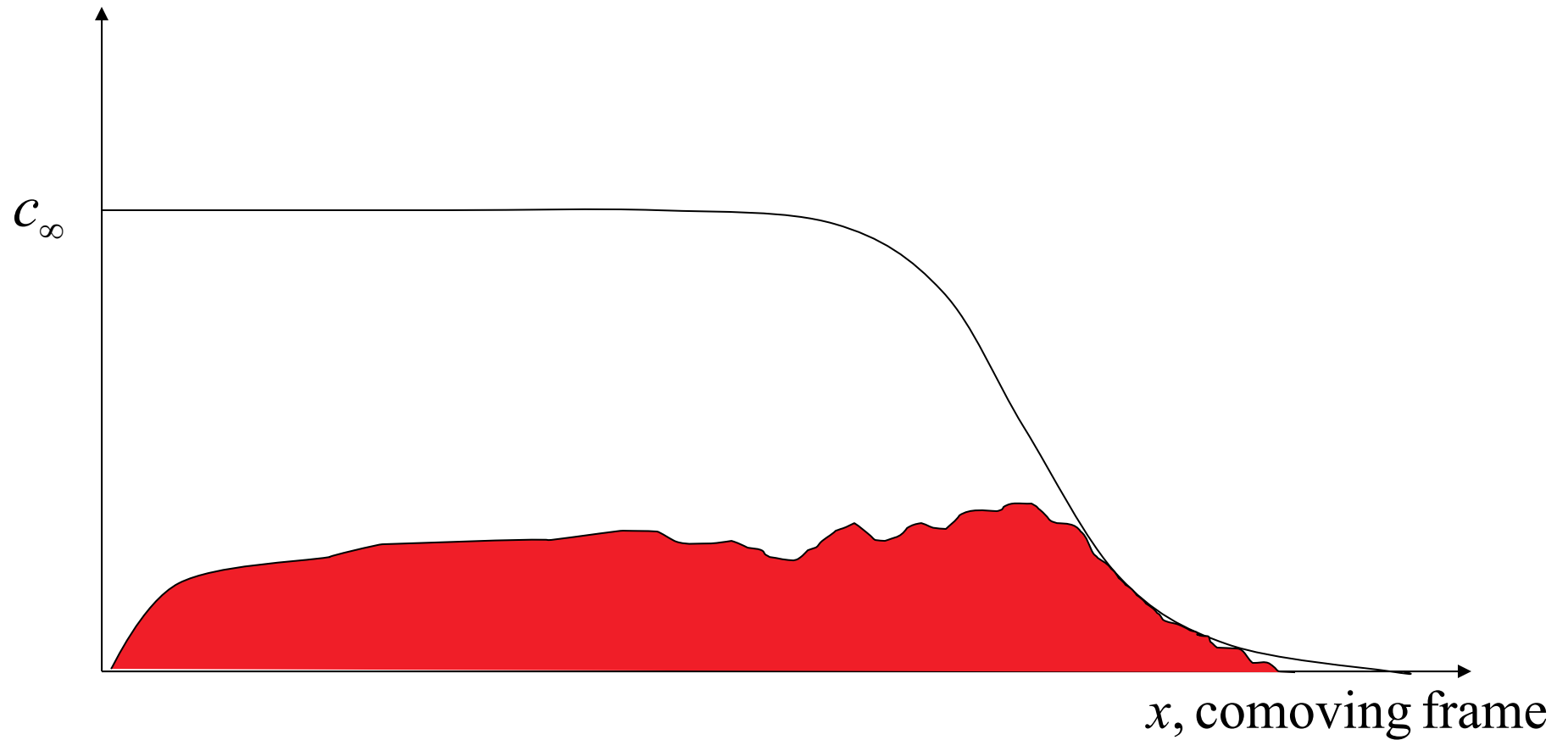
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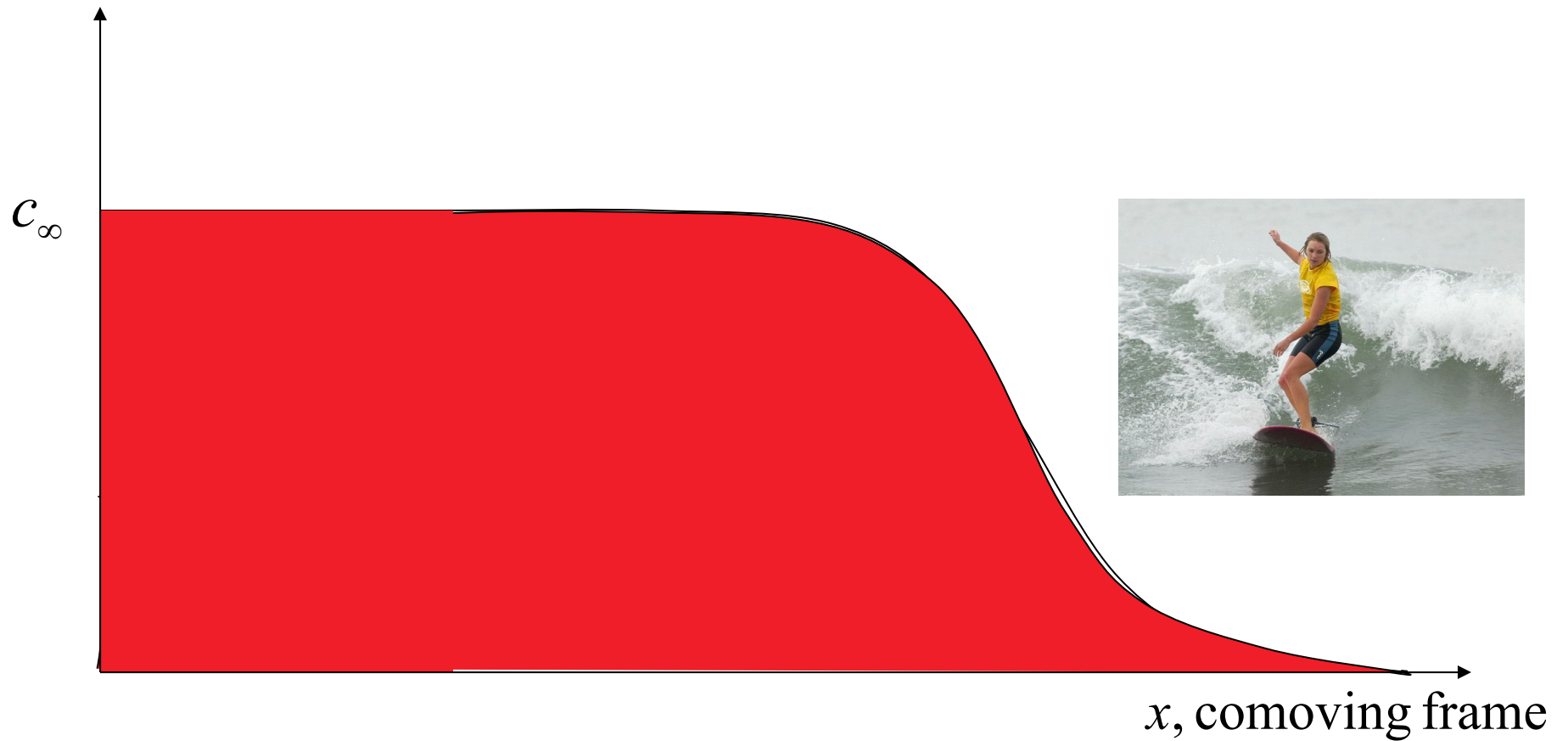
Successful Surfing (1d)

$c_s(x)$, steady state population density



Successful Surfing (1d)

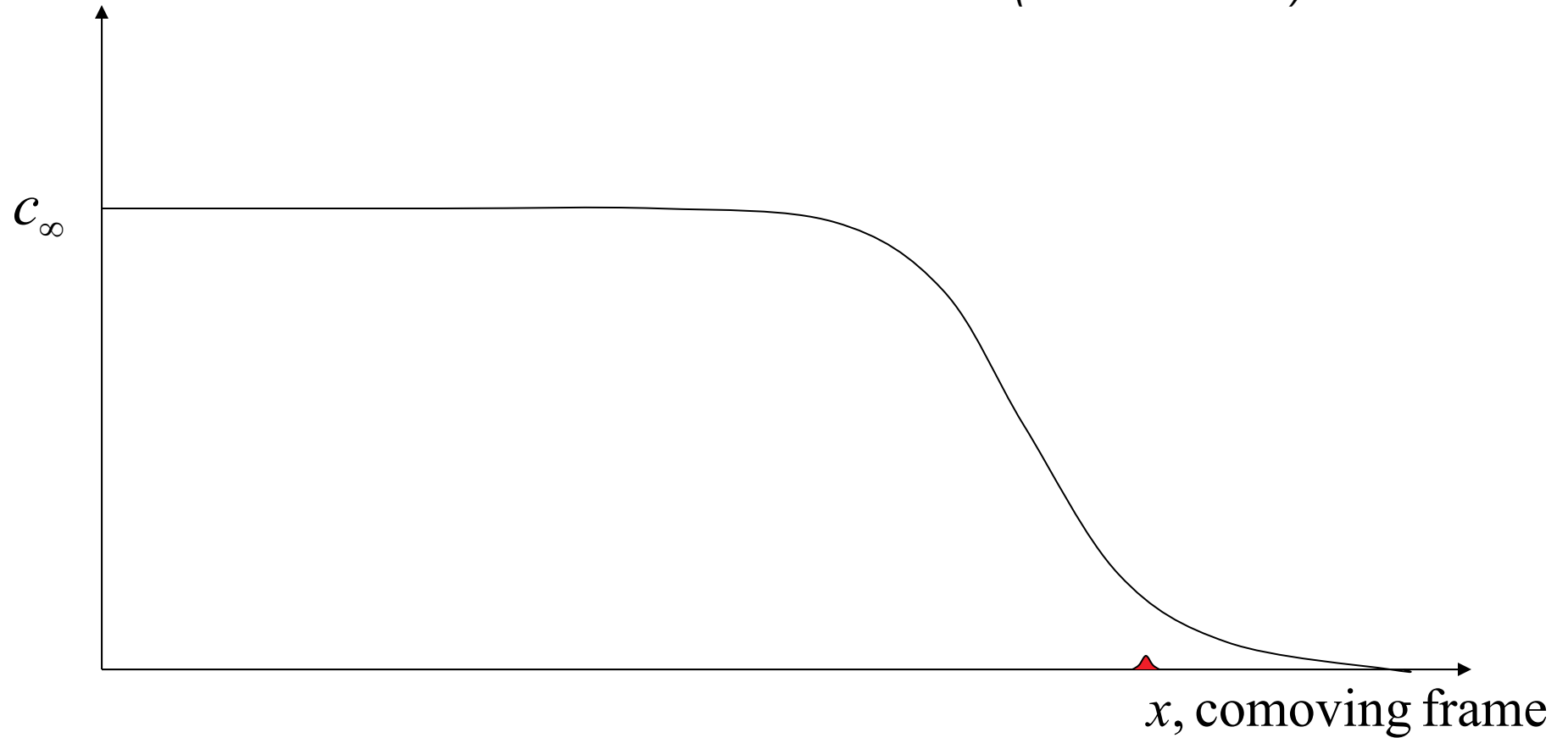
$c_s(x)$, steady state population density



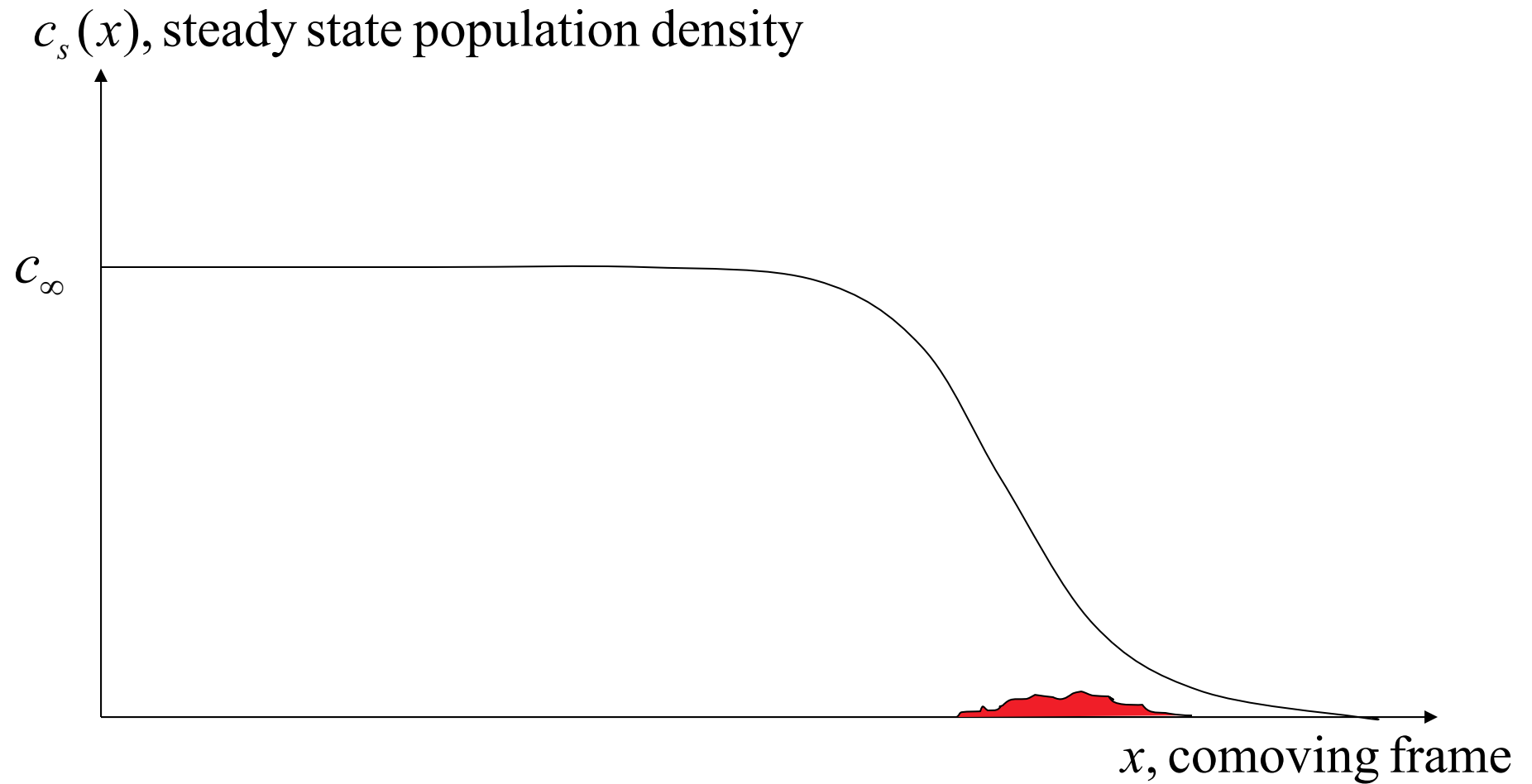
Often however ...

$c_s(x)$, steady state population density

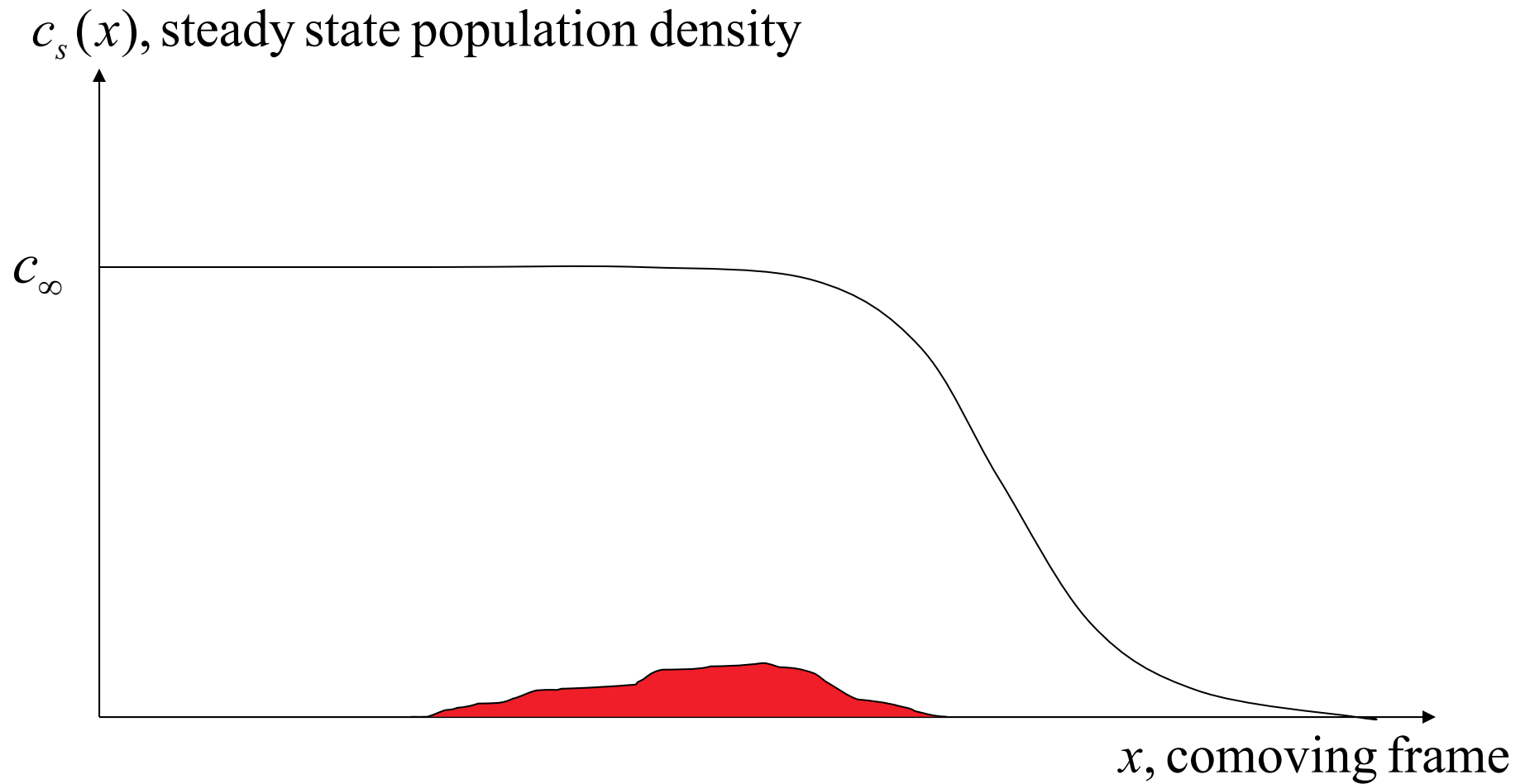
(O. Hallatschek)



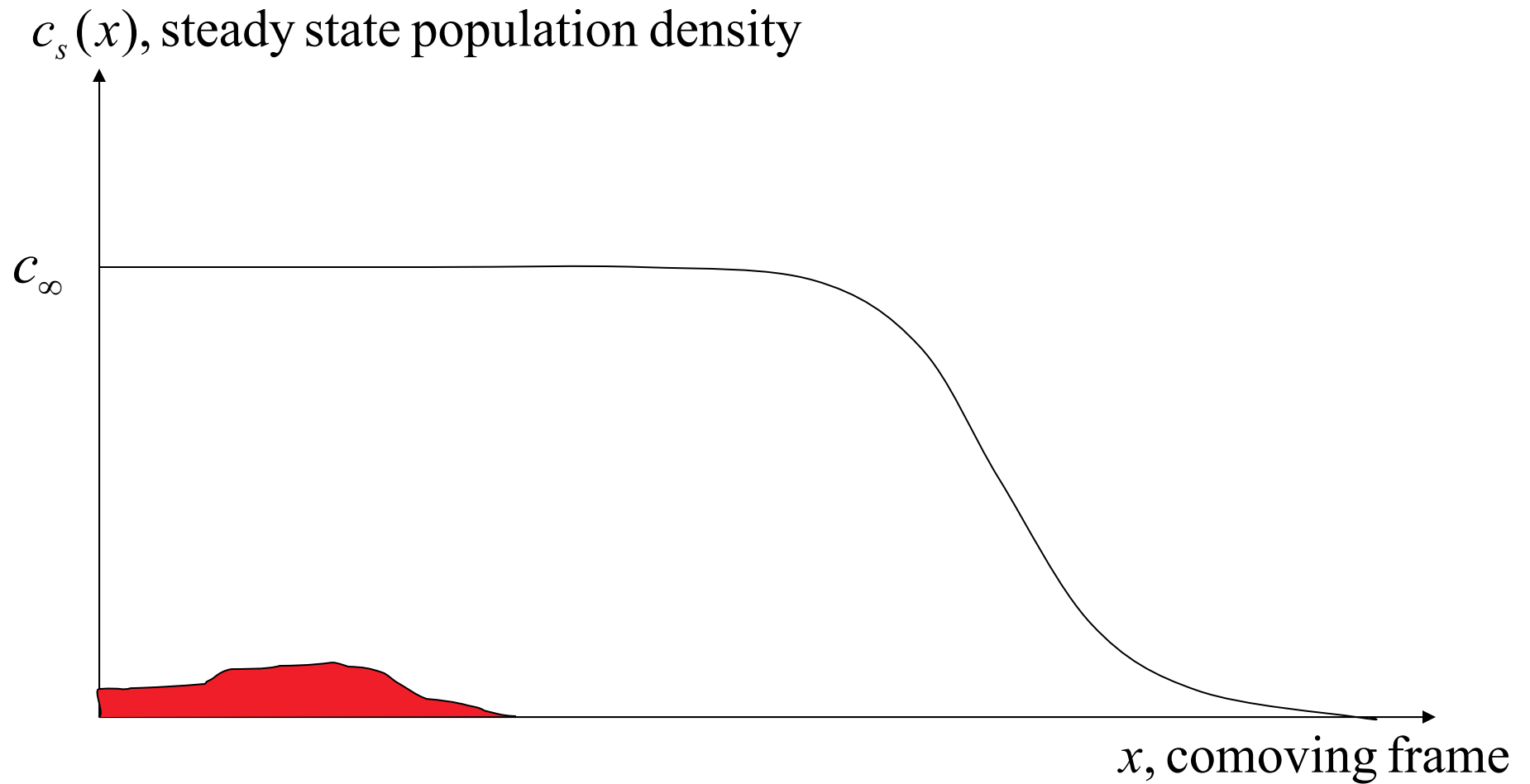
Often however ...



Often however ...



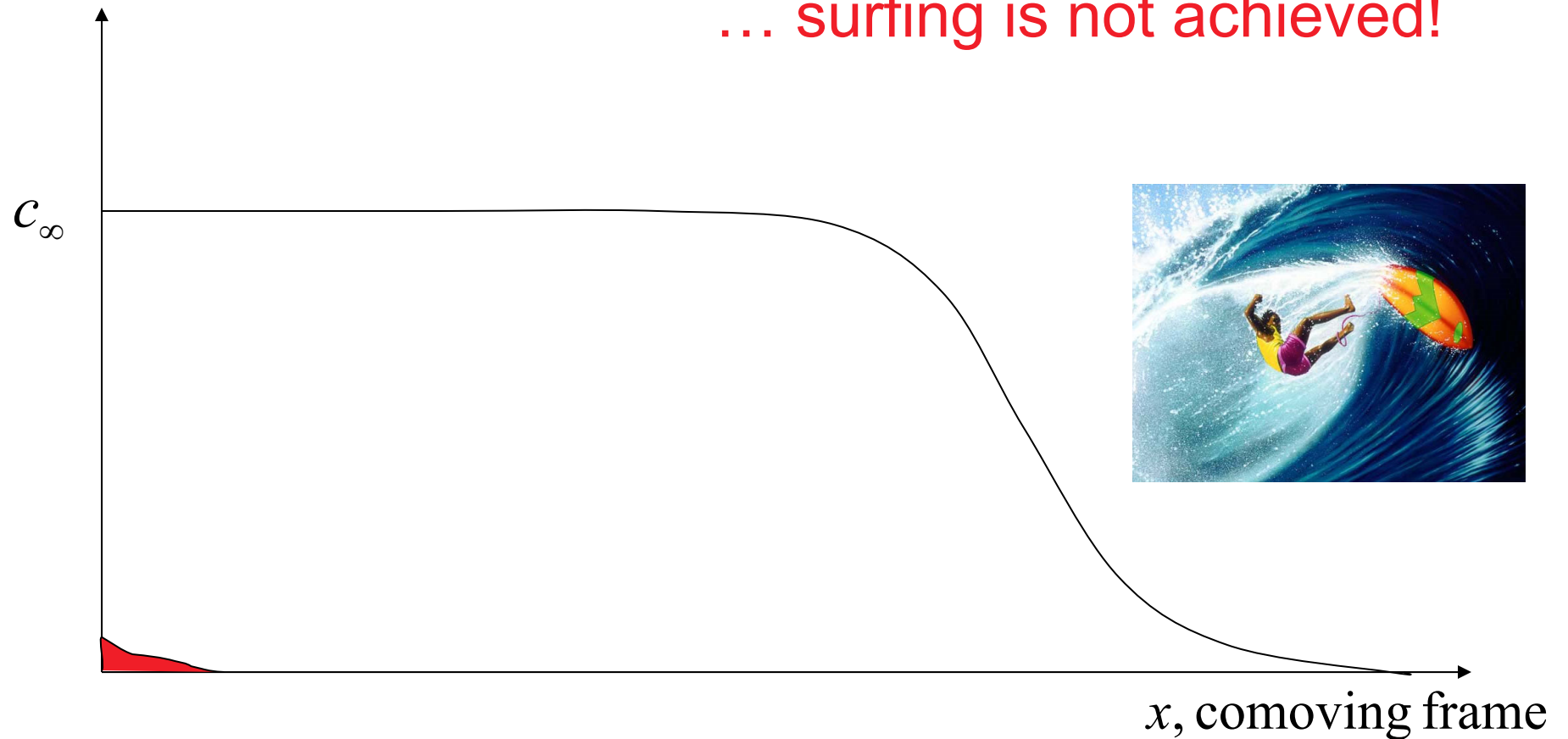
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Often however ...

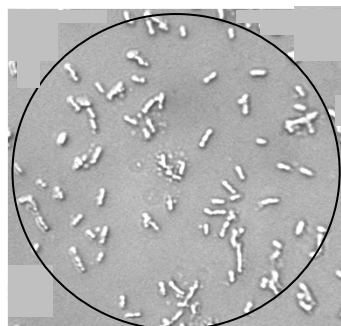
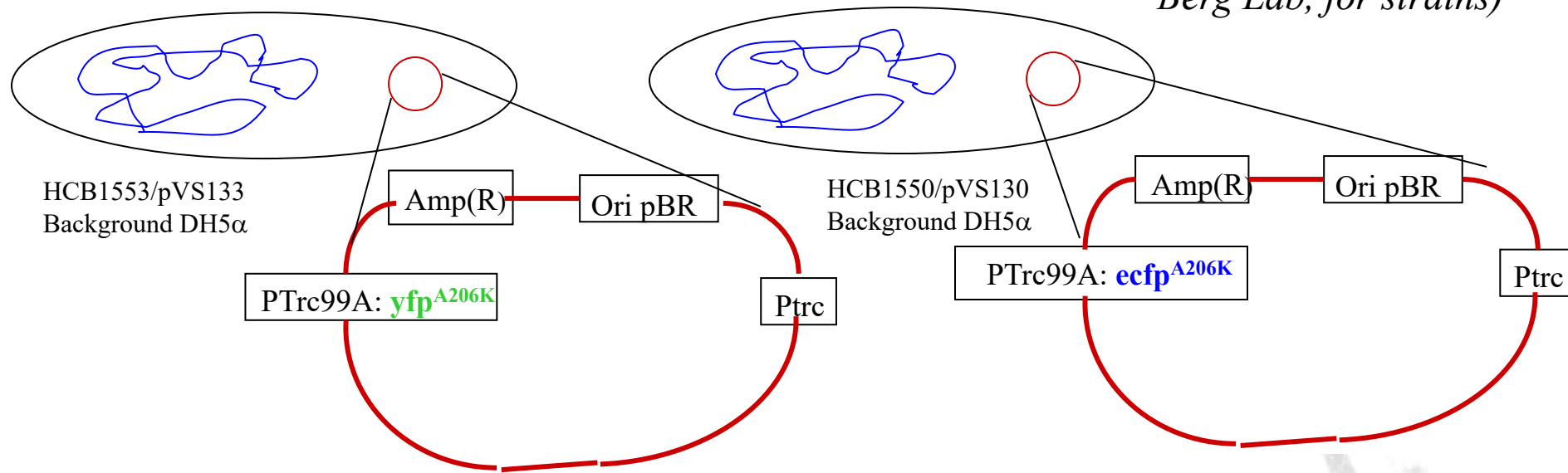
$c_s(x)$, steady state population density

... surfing is not achieved!

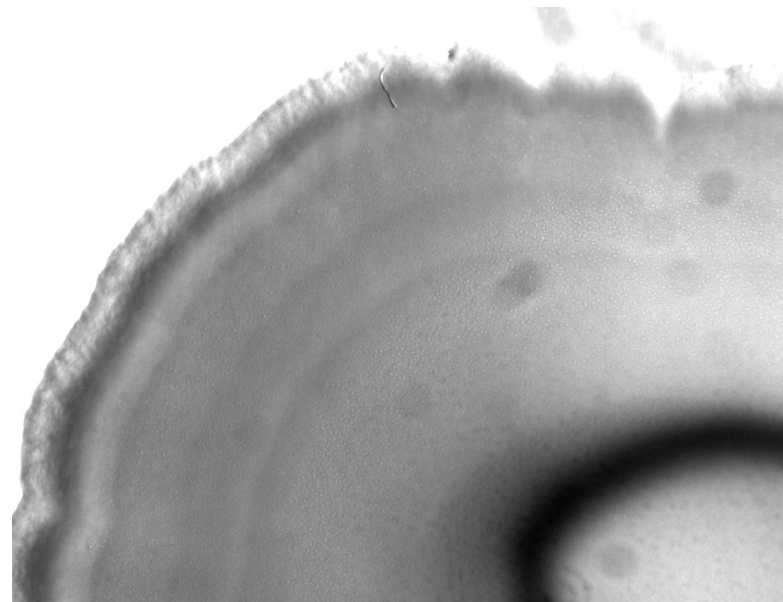


Genetic demxing in bacteria (*E. coli*)

(thanks to Tom Shimizu, Berg Lab, for strains)

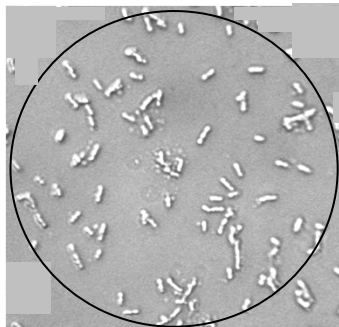
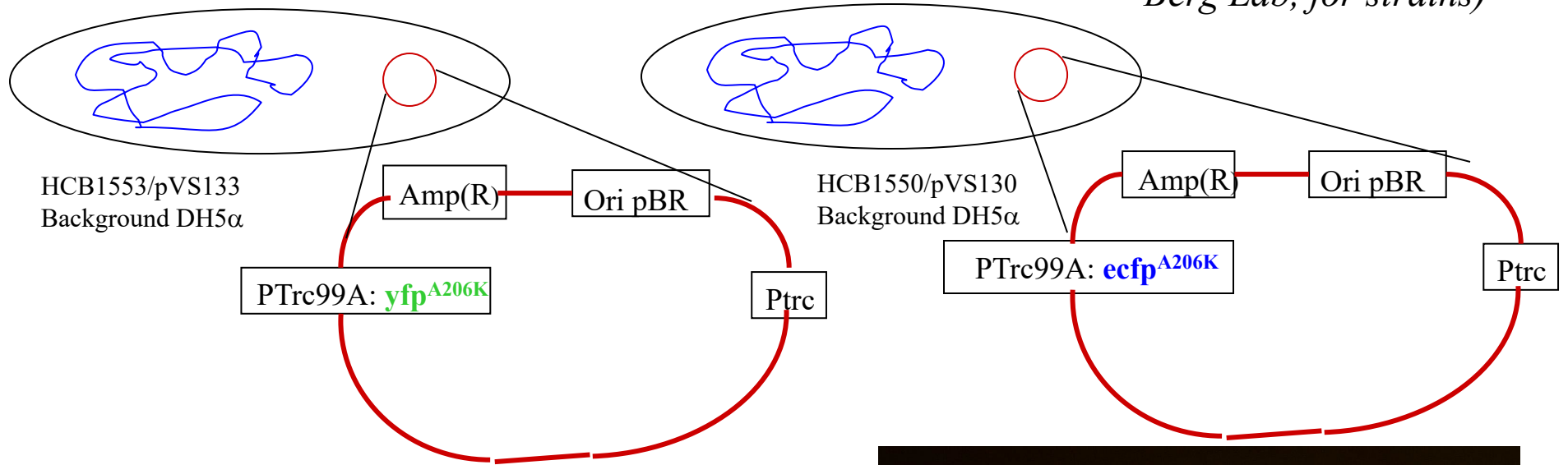


50-50 mixture,
1550/1553

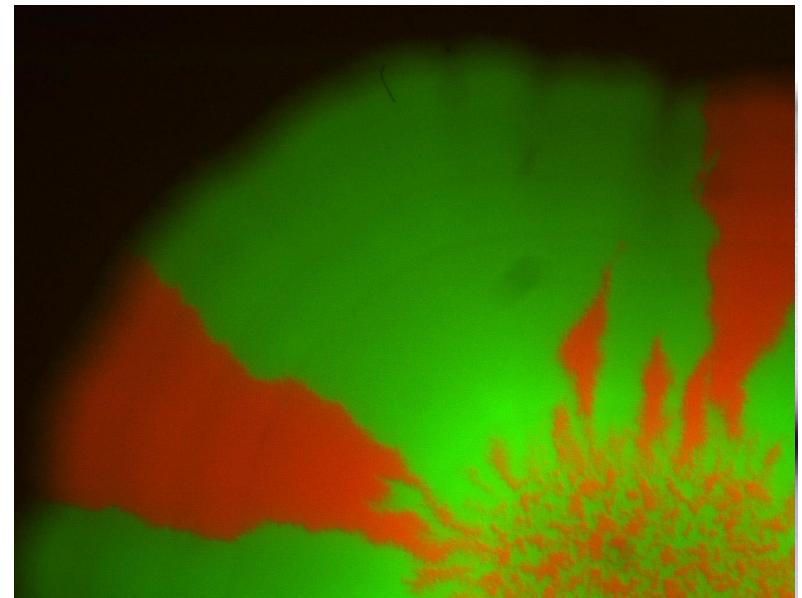


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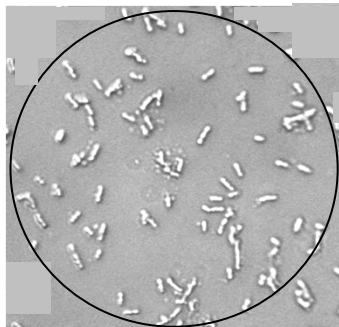
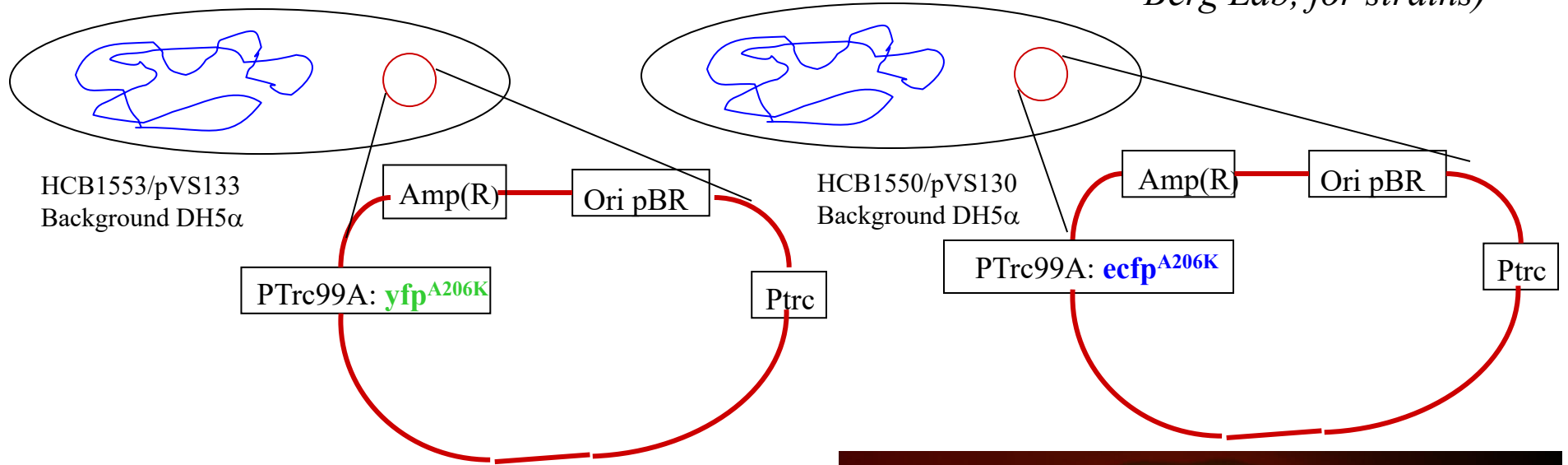
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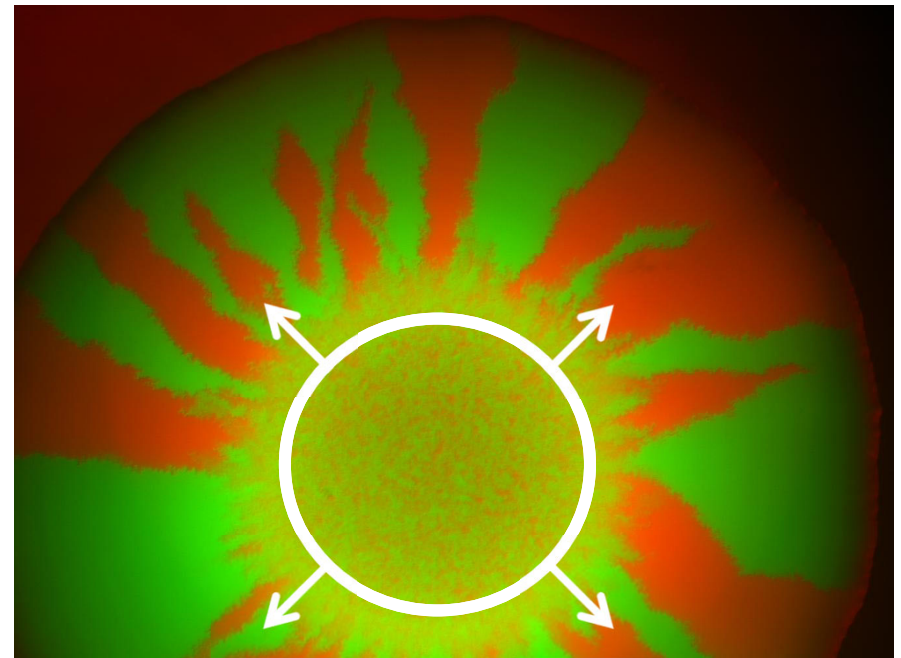
Cyan → Red

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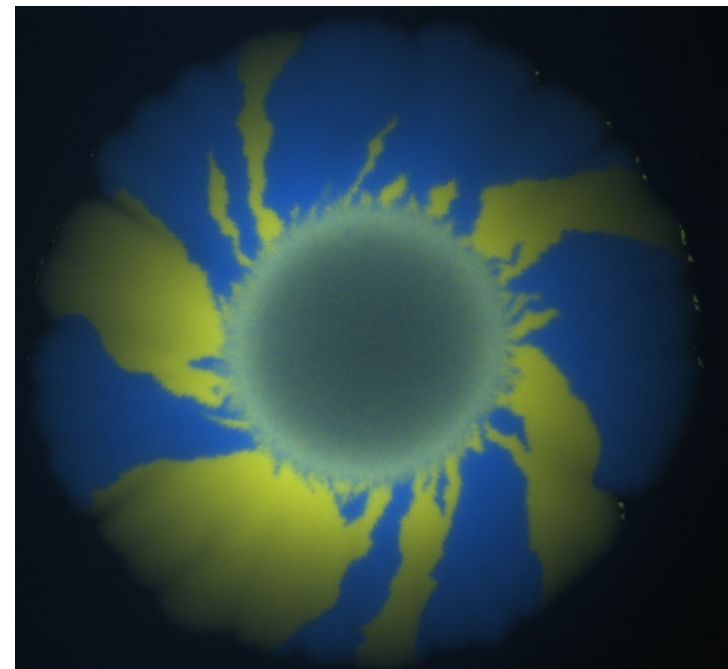
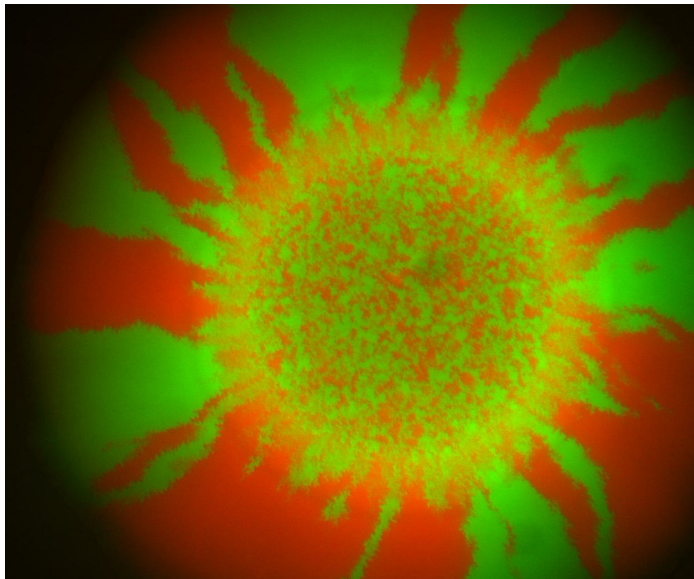
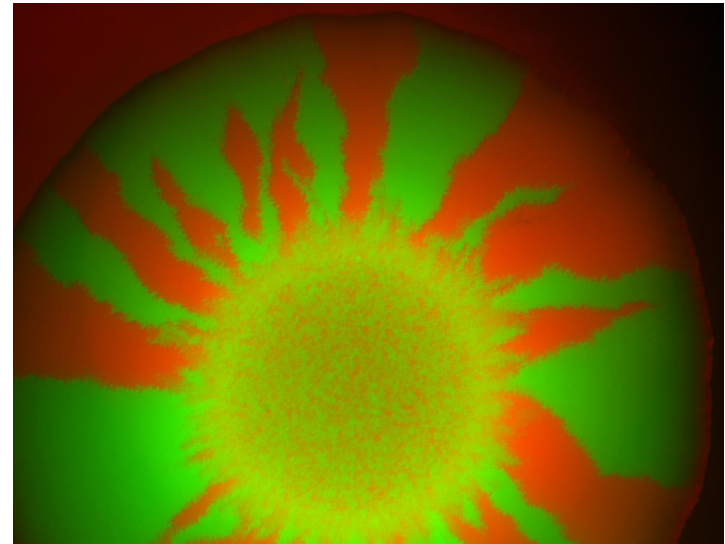
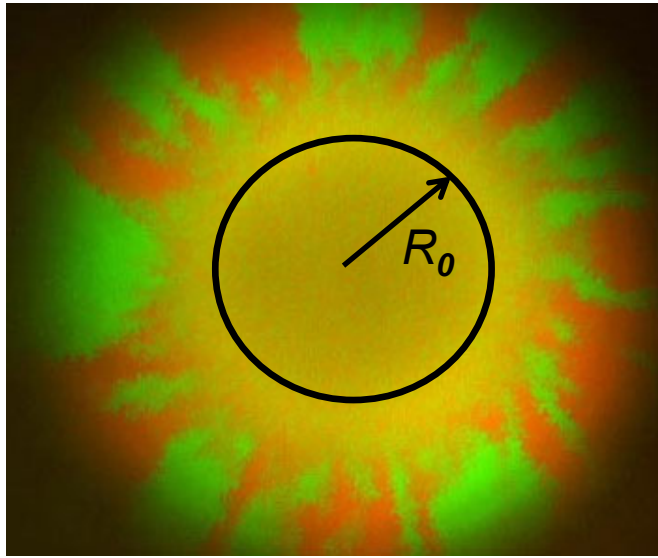


50-50 mixture,
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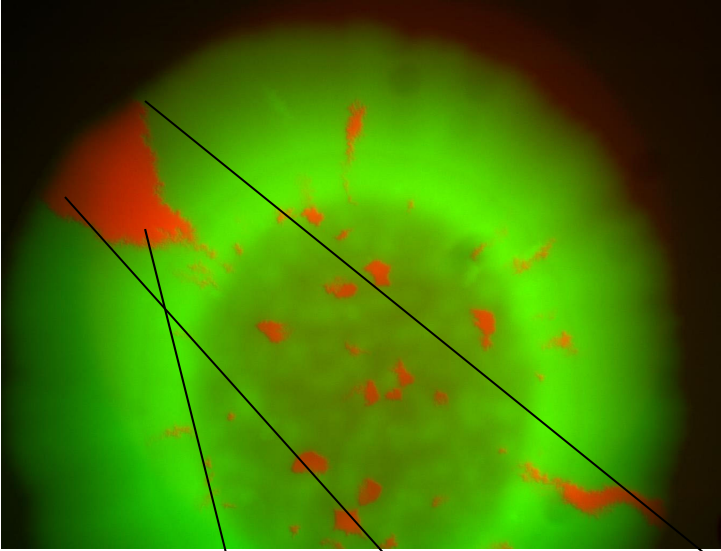
Cyan \rightarrow Red

What would happen if we could “replay the tape of life”?

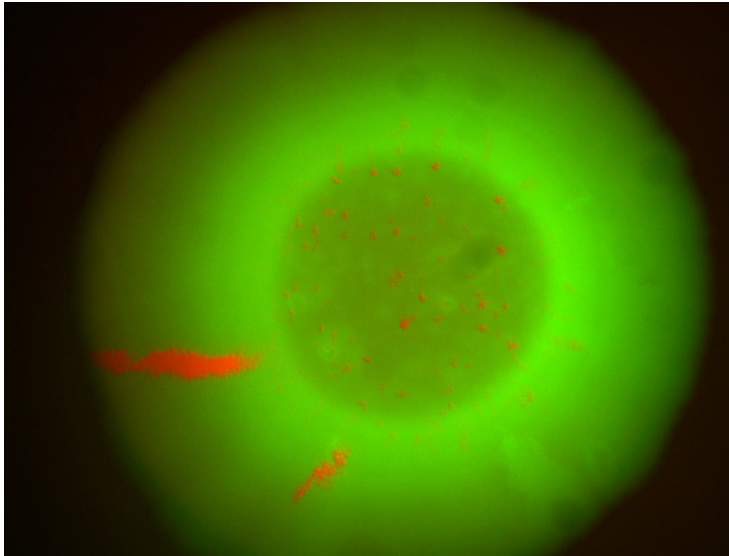


Chiral range expansions

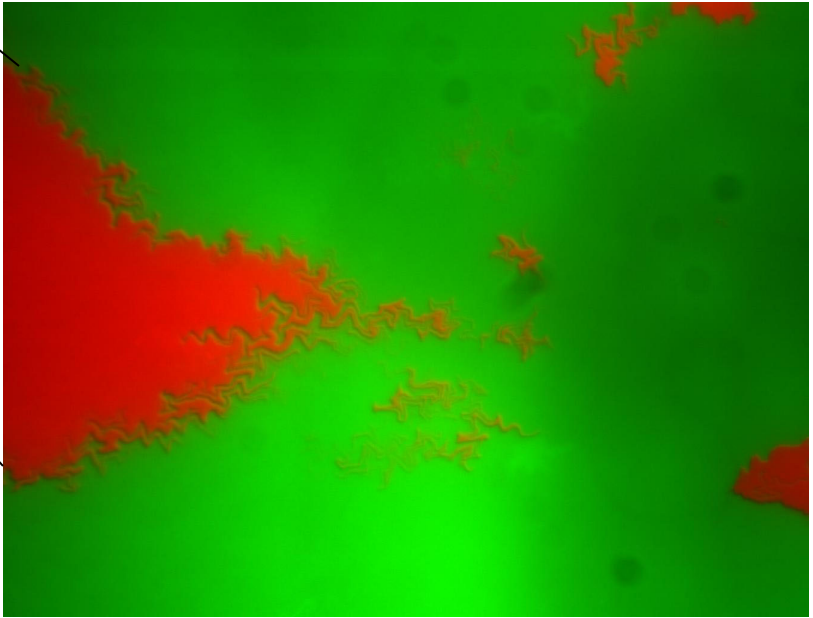
Gene surfing in the dilute limit: “survival of the luckiest”



95%-5% mixture, founder population ~ 500



98%-2% mixture, founder population ~ 5000



Questions & Break

But... life probably evolved first in a *liquid* environment

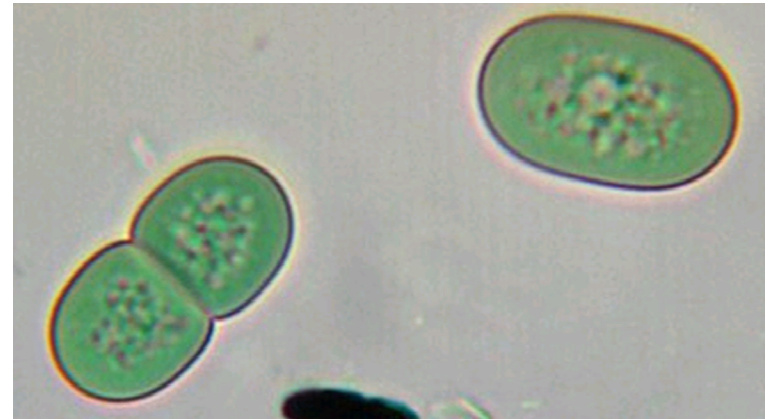
- ~2-3 billion years ago, water covered most of the earth

- Fossilized, oxygen-producing cyanobacteria have been dated at ~2.8-3.5 billion years ago.

- Oxygenic cyanobacteria transformed the atmosphere via photosynthesis

- Spatial growth and evolutionary competition took place at high Reynolds numbers

- These photosynthetic organisms can control their buoyancy to resist down welling currents and stay close to the ocean surface.



Cyanobacterium *Synechococcus*

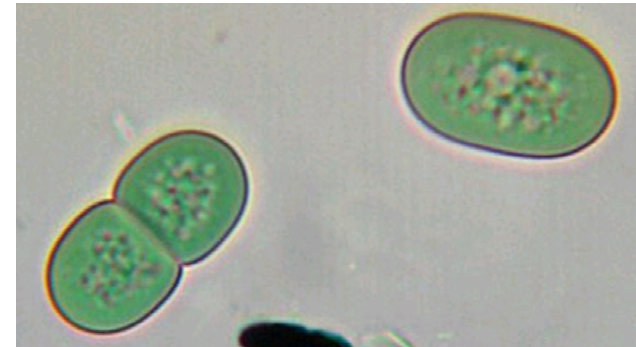
www.dr-raif-wagner.de/Blaualgen-englisch.htm



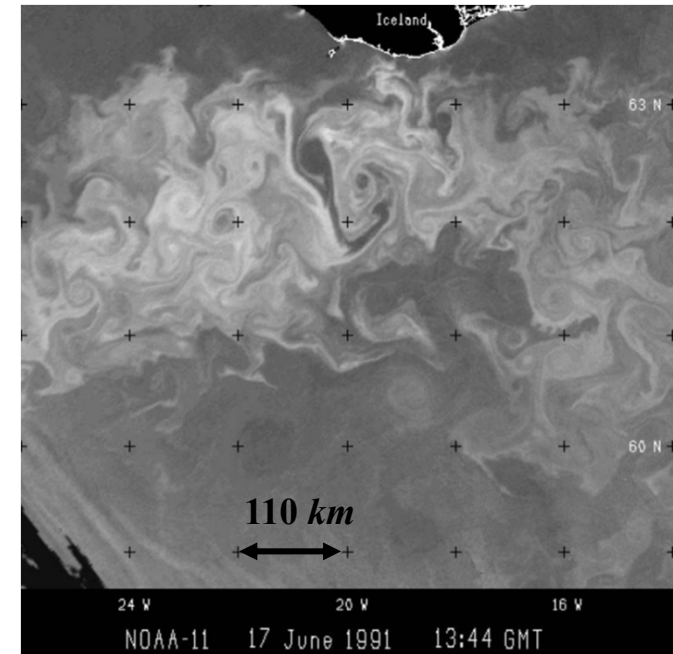
Bloom of cyanobacteria
in Lake Atitlán, Guatemala
NASA Earth observatory

Life probably evolved first in a *liquid* environment

- *~2-3 billion years ago, like today, water covered most of the earth*
- *Fossilized, oxygen-producing cyanobacteria have been dated at ~2 billion years ago.*
- *Oxygenic cyanobacteria transformed the atmosphere via photosynthesis*
- *Their spatial growth and evolutionary competition took place in liquid environments at both high and low Reynolds numbers*
- *These photosynthetic organisms control their height to resist down welling currents and stay close to the ocean or lake surface.*



Cyanobacterium *Synechococcus*



A. P. Martin, Prog. Oceanography **57**, 125 (2003)

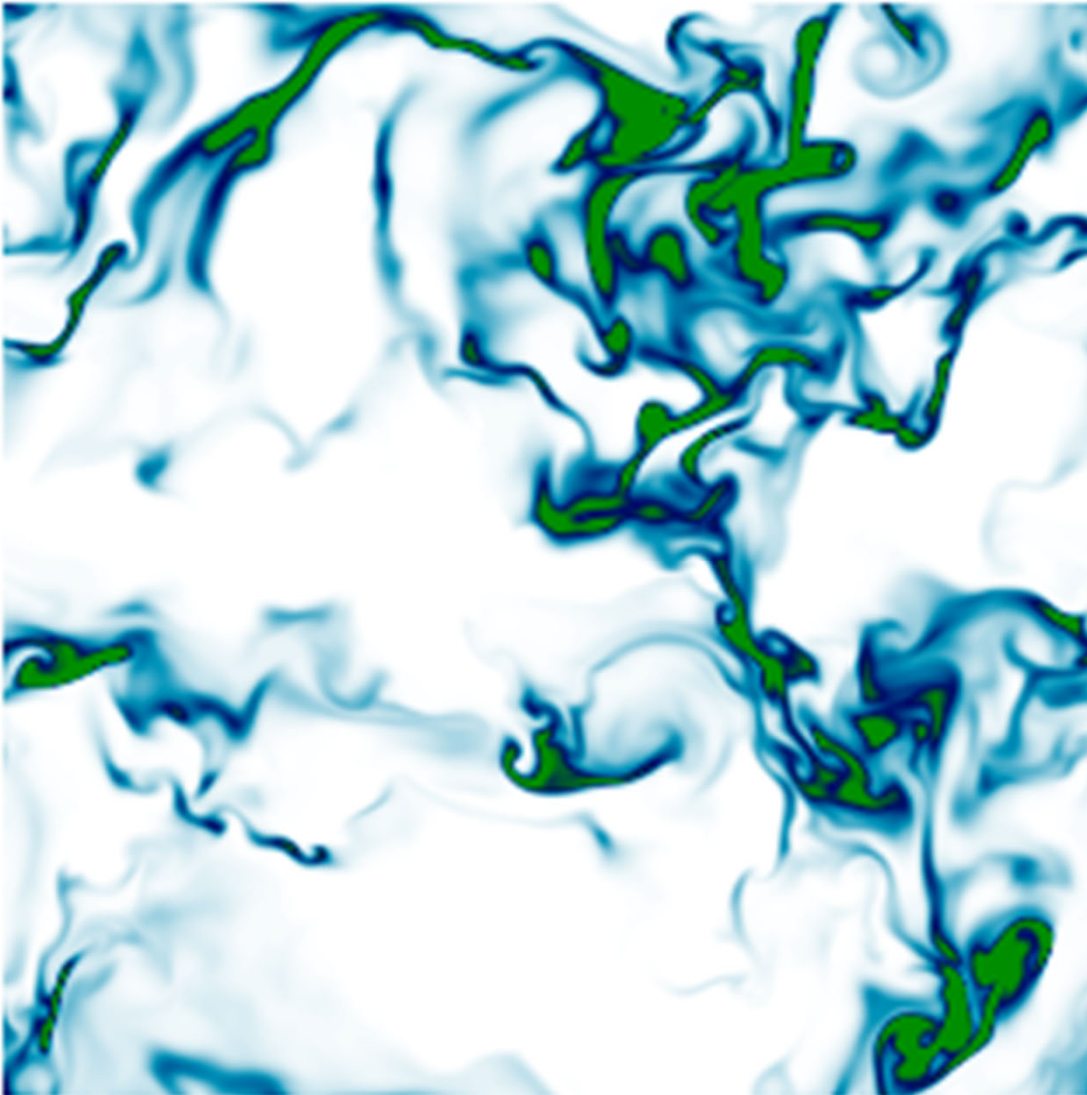
$$Re = LU / \nu = 10^8 - 10^9$$

Large eddy turnover time ≈ 50 days

Small eddy turnover time ≈ 5 minutes

Plankton doubling time $\approx 12-24$ hours

Buoyant population dynamics in Silico (Perlekar, Toschi, Benzi, drn)



$$\frac{\partial \vec{u}}{\partial t} + \vec{u} \cdot \vec{\nabla} \vec{u} = -\frac{1}{\rho} \vec{\nabla} p + \nu \nabla^2 \vec{u} + \vec{f}$$

project onto a 2d plane $\rightarrow \vec{\nabla} \cdot \vec{u}_{2d} \neq 0$

$$\frac{\partial c}{\partial t} + \nabla \cdot (\vec{u}_{2d} c) = D \nabla^2 c + \mu c(1 - c)$$

Reynolds number

$$Re = \frac{u_{\text{rms}} L}{\nu}$$

Schmidt number

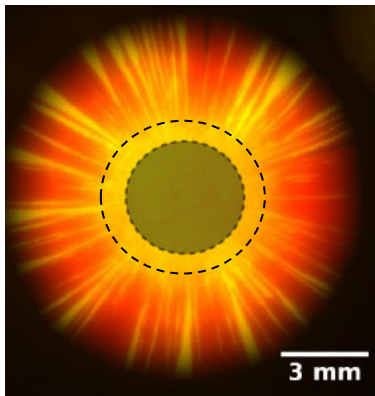
$$Sc = \frac{\nu}{D}$$

Doubling time/eddy turnover time

$$\tau_2 / \tau_{\text{eddy}} \sim 1 / (\mu \tau_{\text{eddy}})$$

Microorganisms grown on liquid but highly viscous substrates create their own flows (without pumps and syringes!)

Hard Agar

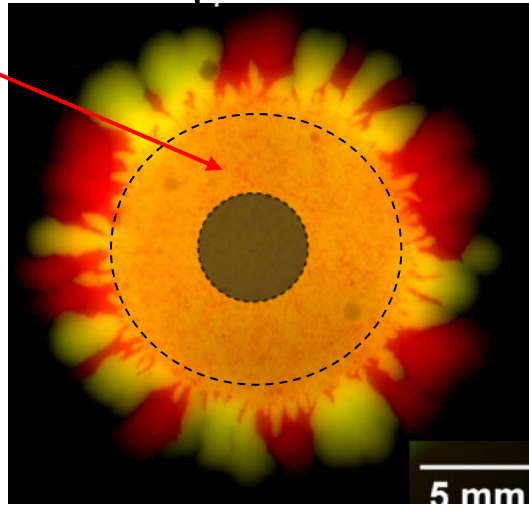


Genetic demixing of yeast on a 1% hard agar YPD plate (viscosity $\eta = \infty$)

Epoch of genetic demixing stretched out....



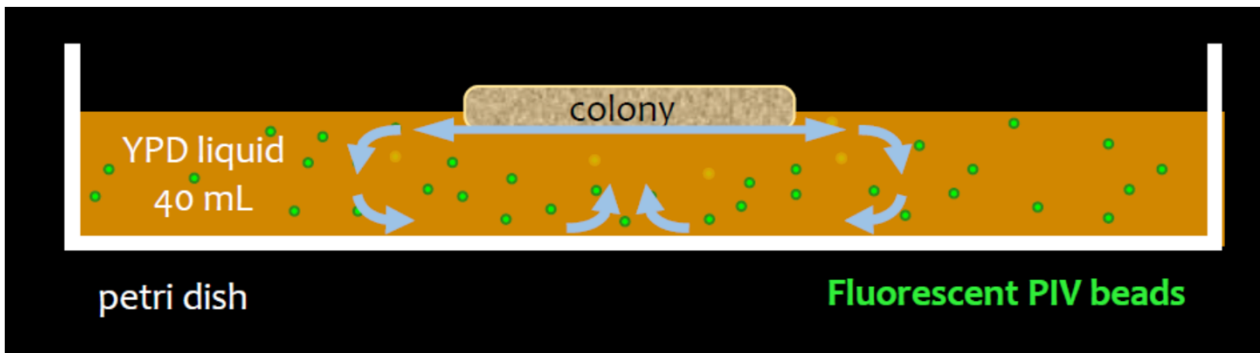
Liquid Media



Yeast on a liquid but highly viscous YPD media with 3% cellulose ($\eta \approx 600 \text{ Pa}\cdot\text{s}$)

Cellulose % (w/v)	Viscosity (Pa·s)
1.8	22 ± 3
2.0	51 ± 6
2.2	81 ± 9
2.4	120 ± 10
2.6	340 ± 50

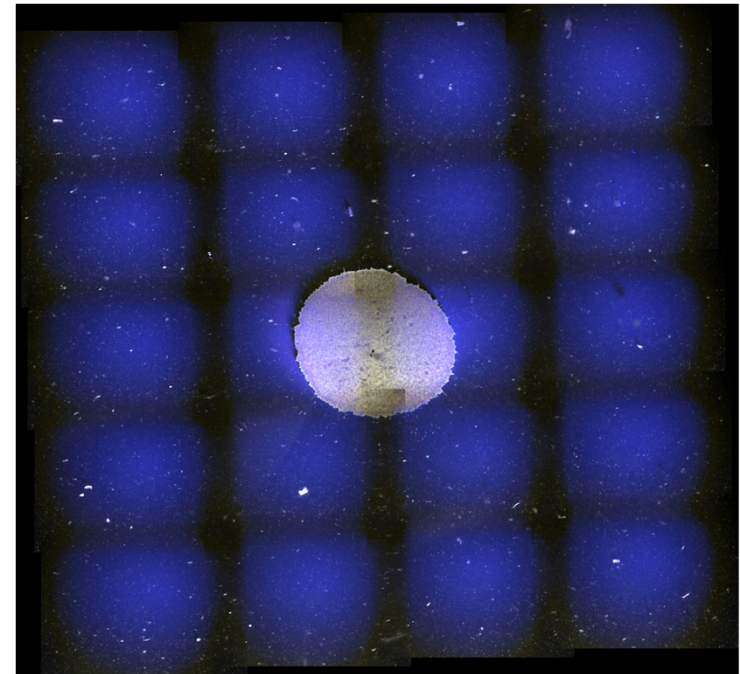
(the viscosity of water is $\eta \approx 10^{-3} \text{ Pa}\cdot\text{s}$; our viscosities are $10^4 - 10^5$ times larger)



The colony itself generates flows that dilate the growing cell mass radially!

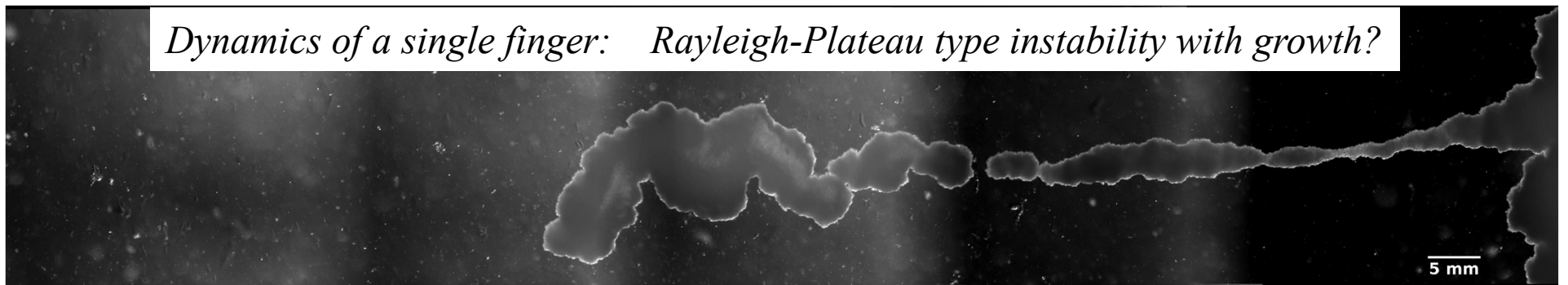
Enhancing the radial flow field...

*(moderate substrate
viscosity $\eta \approx 450 \text{ Pa}\cdot\text{s}$)*

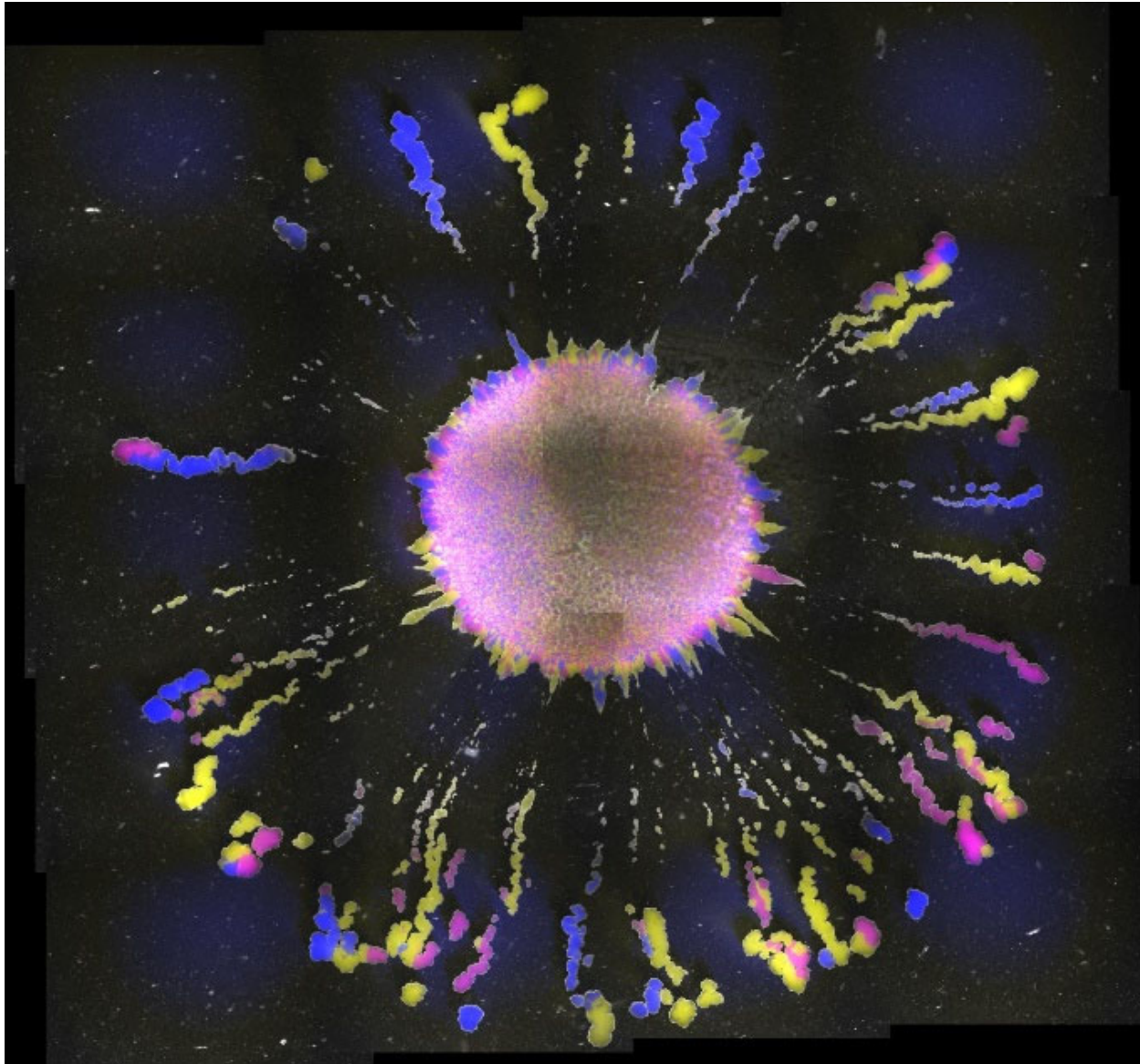


Liquid-like fingering instabilities

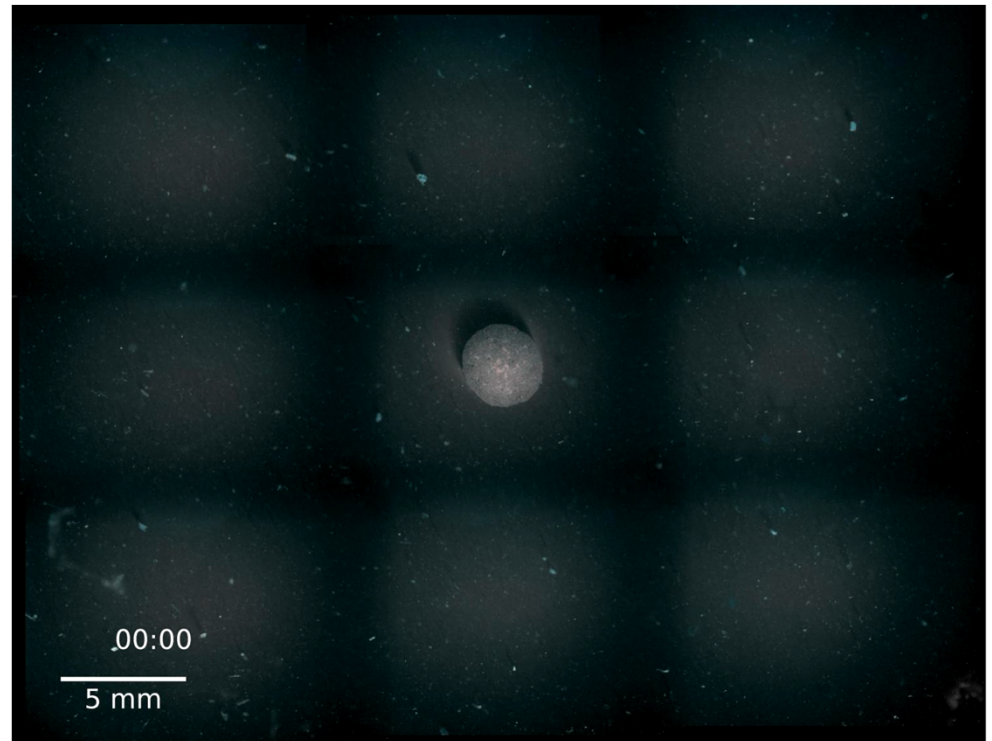
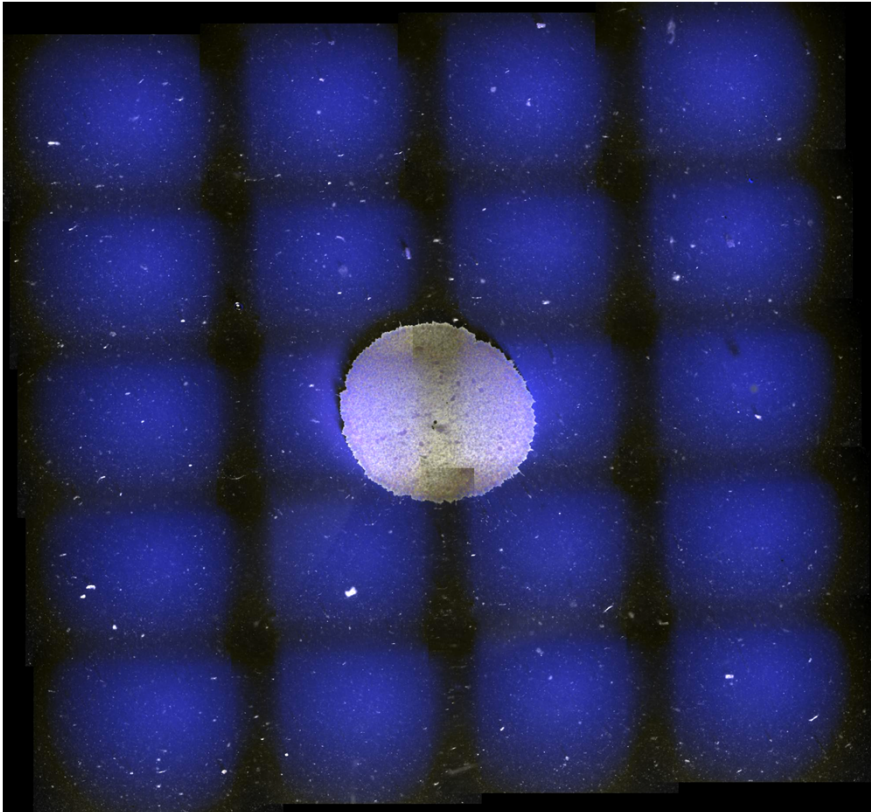
Dynamics of a single finger: Rayleigh-Plateau type instability with growth?



Three colors: genetic demixing
among *three* different genotypes



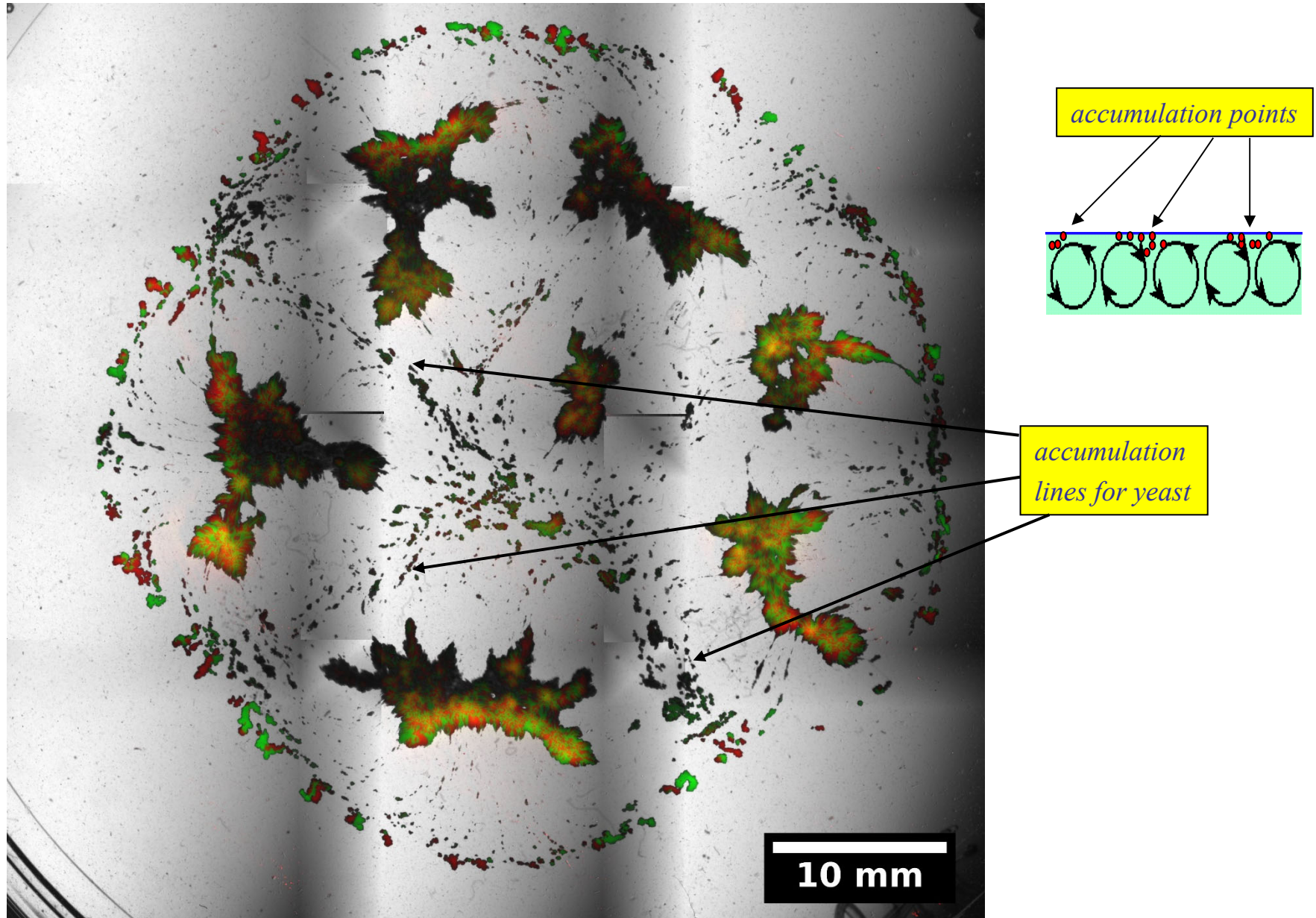
- Anomalous expansion of neutral *S. cerevisiae* strains on a liquid substrate (driven by a submerged vortex ring!)
S. Atis, B. Weinstein, A. Murray, drn



• *Range expansion floating on a high viscosity substrate → fingering with population genetics*

• *Range expansion floating on a lower viscosity substrate → fracture with population genetics*

At lower viscosities, submerged vortex rings fracture the colony – like plate tectonics & continental drift !



Entry #V0020

ROCKET YEAST

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Thank you!



<http://streetanatomy.com>