

Smoluchowski rate: $k = 4\pi D R c$

$\nearrow$ rate at which searchers find target
 $\swarrow$ $D_1 + D_2$
 $\downarrow$ $R_1 + R_2$
 $\nwarrow$ conc. of searcher particles

any reaction rate $k \leq k_{\text{smol}} = 4\pi D R c$

How can biology optimize speed of chemistry?

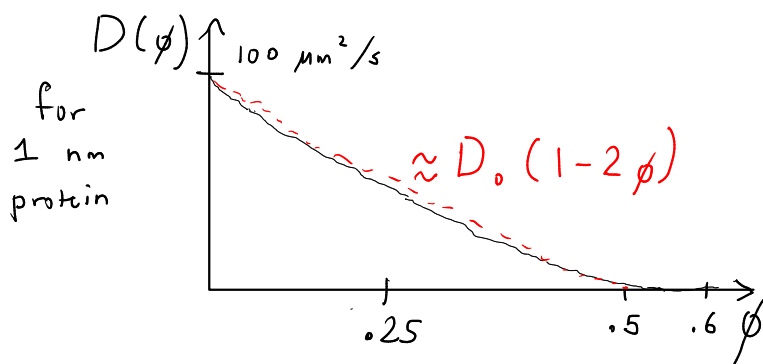
$\Rightarrow$ increasing c

$\Rightarrow$ cell more crowded

$\Rightarrow$ smaller diffusion const. D

ϕ = volume fraction of (non-water) stuff in cell (proteins, DNA/RNA, etc.)

natural range: $0.2 - 0.4$ in modern cells of ϕ



$$c(\phi) = \frac{\phi}{V_{\text{prot}}}$$

$\Leftarrow$

ϕ = dominated by proteins

$$\approx \frac{n V_{\text{prot}}}{V}$$

$V \leftarrow$ total cell volume

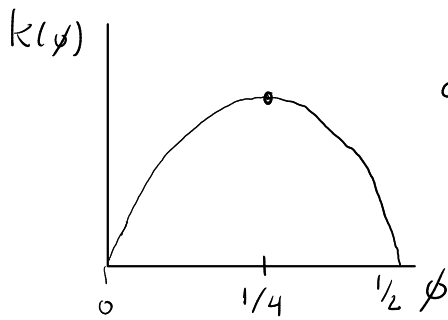
$n = \#$ proteins

$V_{\text{prot}} \approx$ Vol. per protein

$$\phi = c V_{\text{prot}} \quad c = \text{conc. of proteins}$$

$$\max k(\phi) = 4\pi D(\phi) R c(\phi)$$

$$\approx 4\pi D_0 (1-2\phi) R \phi V_{\text{prot}}^{-1}$$



$$\frac{dk}{d\phi} = 0 \Rightarrow \phi = \frac{1}{4} \text{ at } \max k(\phi)$$

estimate total conc. of proteins in a cell:

$$c \sim \frac{\phi}{V_{\text{prot}}} = \frac{1/4}{60 \text{ nm}^3} \leftarrow \text{typical prot. vol.}$$

$$\text{conc. unit} = \text{molar (M)} = \frac{N_A}{L} \begin{matrix} \nearrow 6.022 \times 10^{23} \\ \searrow \text{liter} \end{matrix}$$

typical total conc. of proteins $\sim c = 7 \text{ mM}$

cell volume of bacteria $\sim 1 \mu\text{m}^3$

$$\Rightarrow n = cV \sim 4 \times 10^6 \text{ proteins/cell}$$

actual E. coli values: $3 \times 10^6 - 10^7 \text{ proteins/cell}$

$$\text{optimal } k \approx 2.6 \times 10^6 \text{ s}^{-1}$$

$$\text{reaction timescales } \tau \sim 0.4 \mu\text{s}$$

broad lessons: any one protein $c \ll \text{mM}$

$$n \ll 10^6 \\ (\text{for } V \sim 1 \mu\text{m}^3)$$

one last point:

$$\phi = \frac{n_{\text{prot}} V_{\text{prot}} + n_{\text{bp}} V_{\text{bp}}}{\bar{V}}$$

n_{bp} = # of base
pairs of
genetic material

$\sim .005$ for E. coli

V_{bp} = volume per
bp