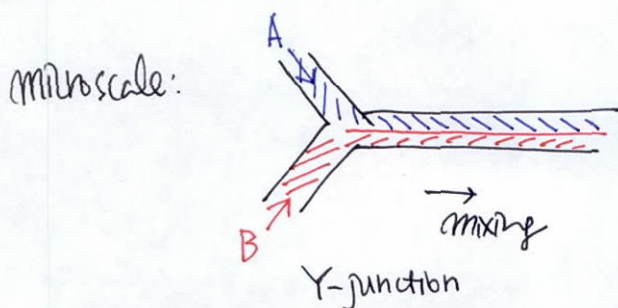
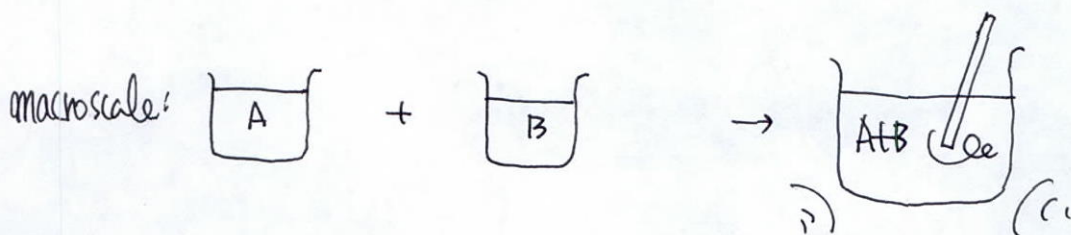
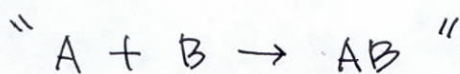


MIXING IN MICROFLUIDICS

- Lab-on-a-chip > biochemical reaction
- μ TAS

- biochemical analysis
- drug delivery
- sequencing / synthesis of nucleic acids
- cell activation
- enzyme reaction
- protein folding
- hazardous chemical detection



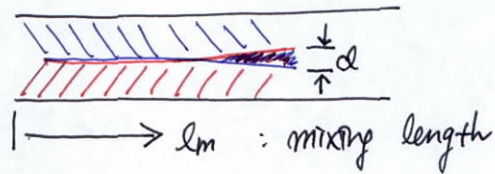
surface-to-volume ratio

$$\frac{A}{V} \sim \frac{l^2}{l^3} \sim \frac{1}{l} \quad \uparrow \text{ as } l \downarrow$$

Mass transfer mechanism of mixing

1. molecular diffusion - low Re
 2. eddy diffusion
 3. bulk diffusion
-) \sim turbulence

Simplest case



molecular diffusion

$$\frac{\partial c}{\partial t} = D \frac{\partial^2 c}{\partial y^2}$$

D : molecular diffusivity

scaling analysis

$$\frac{\Delta c}{\tau} \sim D \frac{\Delta c}{d^2}$$

$$d \sim \sqrt{D\tau}$$

$$\tau \sim \frac{l}{U}$$

$$d \sim \sqrt{D \cdot \frac{l}{U}}$$

$$l_m \sim \frac{U}{D} d^2$$

e.g. $U = 1 \text{ mm/s}$, d (channel width) = $100 \mu\text{m}$,

$D \approx 10^5 \text{ cm}^2/\text{s} = 10^{-9} \text{ m}^2/\text{s}$ (small molecules)

$10^{-7} \text{ cm}^2/\text{s}$ (large molecules, e.g. haemoglobin in water)

channel length for complete mixing

$$l_m \sim \frac{10^{-3}}{10^{-9}} (10^{-4})^2 \sim 10^{-2} \text{ m} \sim 1 \text{ cm}.$$

"too long for multi-step reactions"

* Peclet number. $Pe = \frac{Ud}{D} = \frac{\text{convective transport}}{\text{diffusive transport}}.$

$$l_m \sim Pe \cdot d$$

$$l_m \propto Pe.$$

* Chaotic advection / Lagrangian chaos

- laminar flow at low $Re \rightarrow$ chaotic particle trajectories
- velocity field $\left\{ \begin{array}{l} 2-D, \text{ time-dependent} \\ 3-D. \end{array} \right.$
- Analysis : hydrodynamics + nonlinear dynamics

* Mixing devices

- active mixers (moving parts, varying pressure gradient)
- passive mixers

* Evaluation of mixing

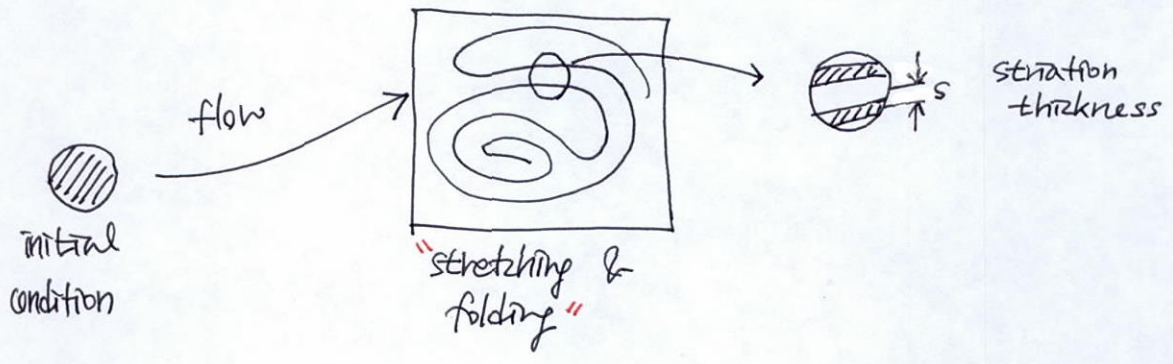
- observing color or intensity variations of a dye or pH indicator
- indicator $\left\{ \begin{array}{l} \text{bromothymol blue (pH)} \\ \text{fluorescein (angiography, fluorescent tag)} \\ \text{potassium permanganate (KMnO}_4, \text{ oxidizing agent)} \\ \text{rhodamine / uranine dyes} \\ \text{phenolphthalein (pH) : acid (colorless) } \rightarrow \text{ base (pinkish)} \\ \qquad \qquad \qquad \sim \text{pH 9} \end{array} \right.$

* Necessary condition for chaos :

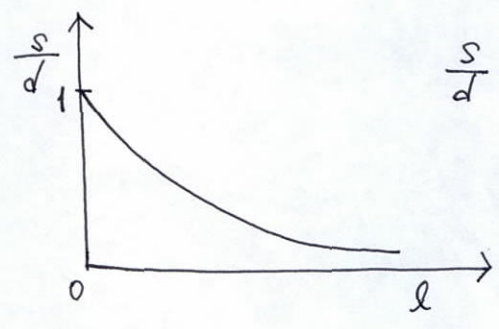
'crossing' of streamlines at different times.

~ two successive streamline portraits, say at t and $t + \Delta t$ for time-periodic 2-D flows, or at z and $z + \Delta z$ for spatially periodic flows, when superimposed, should show intersecting streamlines when projected onto the (x, y) -plane.

Basic processes during mixing (except molecular diffusion) by flow



In a static chaotic flow,



$$\frac{s}{d} = \exp\left(-\frac{l}{\lambda}\right)$$

λ : characteristic length determined by the geometry of trajectories in the chaotic flow

$l_{m}(d)$?

residence time $\tau_r = \frac{l}{U}$

diffusion time $\tau_d = \frac{s^2}{D} = \frac{d^2}{D} \exp\left(-\frac{2l}{\lambda}\right)$

$$\tau_r \sim \tau_d$$

$$\frac{l}{U} \sim \frac{d^2}{D} \exp\left(-\frac{2l}{\lambda}\right)$$

$$\frac{Ud}{D} \sim \frac{l}{d} \exp\left(\frac{2l}{\lambda}\right)$$

$$\ln(Pe) \sim \ln\left(\frac{l}{d}\right) + 2\frac{l}{\lambda}$$

if $\ln\left(\frac{l}{d}\right) \ll 2\frac{l}{\lambda}$

$$l_m \sim \lambda \ln(Pe)$$