## **I – Bioreactor Kinetics**



#### Introduction

- 1 Batch Reactor (BSTR)
- 2 Continuous Reactor (CSTR)
- 3 Plug Flow Reactor (PFR)
  - 3.1 Definition
  - 3.2 Material balances
  - 3.3 Kinetics
  - 3.3 Productivity
  - 3.4 PFR/CSTR comparison
  - 3.5 Discussion about PFR



# Plug flow photobioreactor









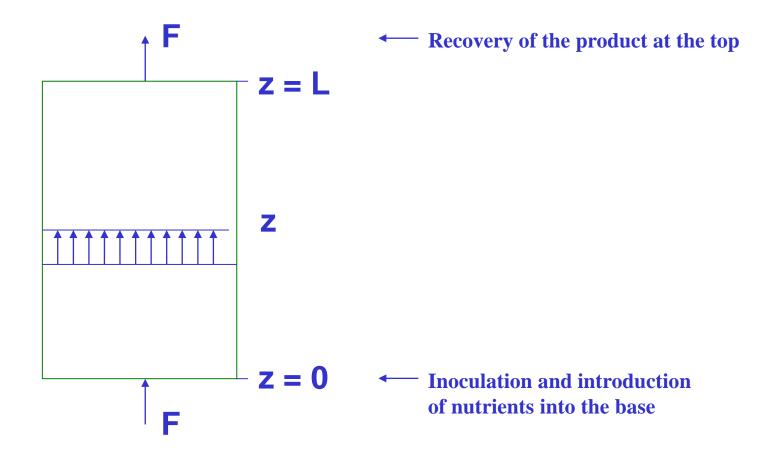
## 3.1 – Definition





Geometry: cylindrical column

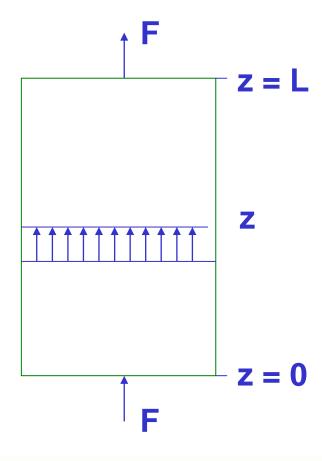
**Operation**: Continuous





#### 3.1 – Definition

Geometry: cylindrical column



**Operation**: Continuous

z – position on a vertical axis (m)

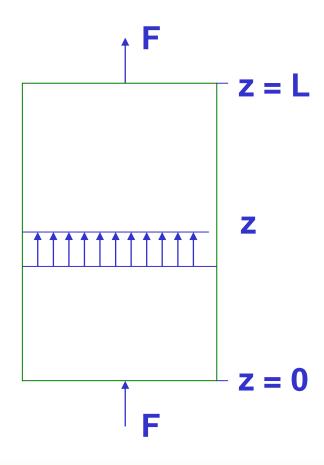
L - column height (m)

F - fluid flow rate in ascending flow (m<sup>3</sup>/h)



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Geometry: cylindrical column



**Operation**: Continuous

z – position on a vertical axis (m)

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F - fluid flow rate in ascending flow (m<sup>3</sup>/h)

A - cross section area (m<sup>2</sup>)

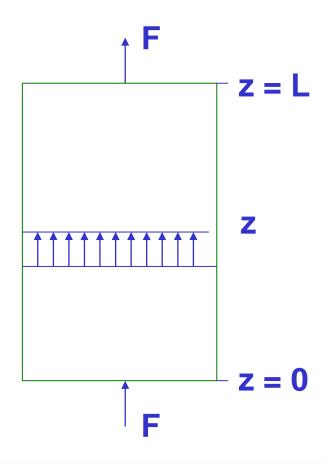
$$A = \frac{\pi \, \mathrm{d}^2}{4}$$

d - diameter of the cylindrical column (m)



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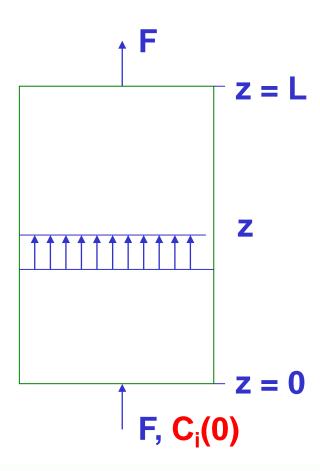
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$$V = AL - column volume (m^3)$$



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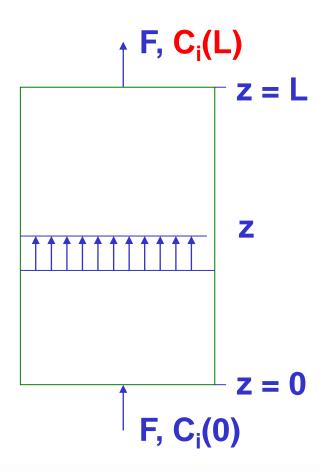


**Operation**: Continuous

Ci(0) - concentration of a generic i component at the base of the column (kg/m³)



Geometry: cylindrical column



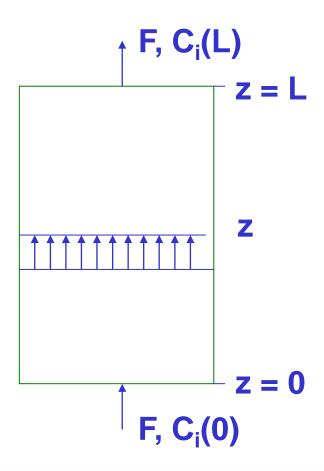
**Operation**: Continuous

Ci(0) - concentration of a generic i component at the base of the column (kg/m³)

Ci(L) - concentration of a generic i component at the top of the column (Kg/m³)



Geometry: cylindrical column



**Operation**: Continuous

Ci(0) - concentration of a generic i component at the base of the column (kg/m³)

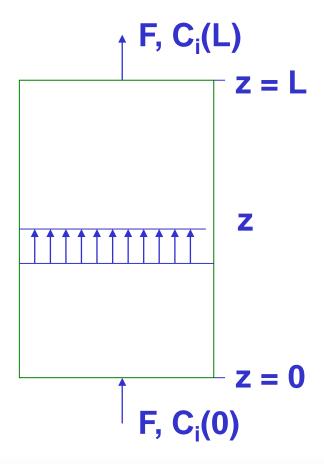
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Ci(z) - concentration of a generic i component in a z position of the column (Kg/m<sup>3</sup>)



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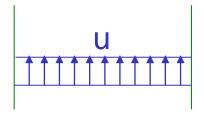
Ci(L) - concentration of a generic i component at the top of the column (Kg/m³)

Ci(z) - concentration of a generic i component in a z position of the column (Kg/m<sup>3</sup>)

u - axial velocity of the fluid inside the column (m/s)



## 3.1 – Definition

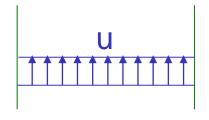


Velocity profile in plug flow = **CONSTANT** 

Therefore: all fluid elements move at the same velocity u.



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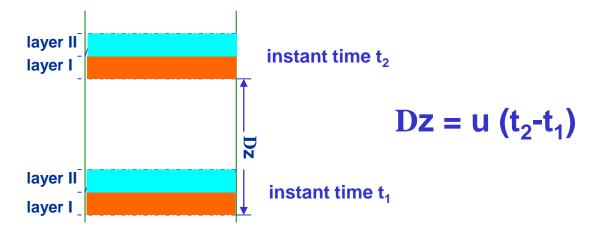
Therefore: all fluid elements move at the same velocity u.

$$Re = \frac{\rho u d}{\mu} > 2000$$

(Nº Reynolds)

- $\rho$  Specific mass of fluid (Kg/m<sup>3</sup>)
- u axial velocity of the fluid (m/s)
- d column diameter (m)
- $\mu$  Viscosity of the fluid (Pa.s)

Displacement analysis of 2 adjacent fluid layers



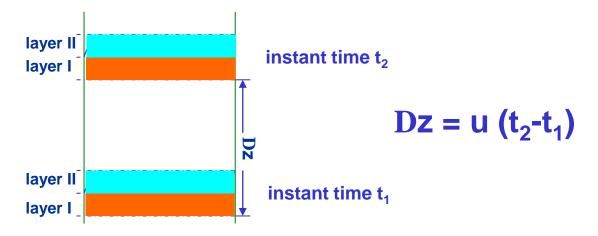
all fluid elements move at the same velocity u



Two adjacent fluid layer never mix



Displacement analysis of 2 adjacent fluid layers



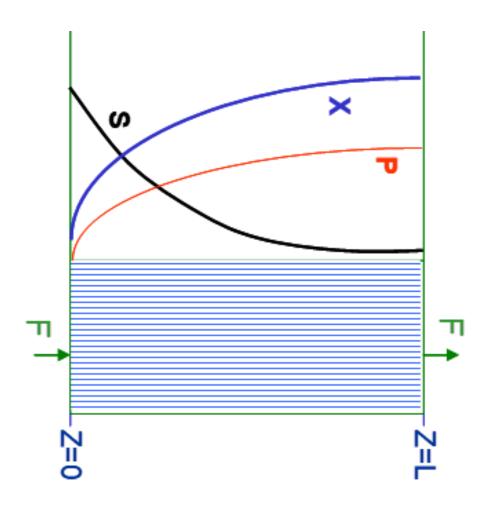
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Two adjacent fluid layer never mix

PFR: TOTAL SEGREGATION CSTR: PERFECT MIX

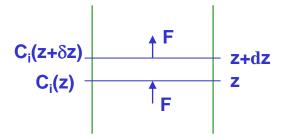






#### 3.2 – Material balances

Material balance to the infinitesimal section of the column with height dz a generic 'i' component



Mass of 'i' that enters z per unit of time

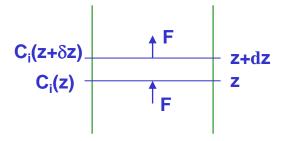
Mass of 'i' produced by reaction in Adz volume = per unit of time

Mass of "i" leaving z+dz per unit of time



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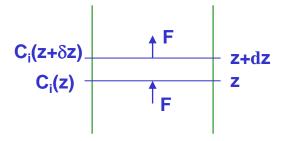
$$FC_i(z)$$

+ 
$$r_i(z)Adz = FC_i(z+dz)$$

Note: r<sub>i</sub> – volumetric rate of i production



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Mass of 'i' that enters z per unit of time

Mass of "i" leaving z+dz per unit of time

$$FC_i(z)$$
 +  $r_i(z)Adz$  =  $FC_i(z+dz)$ 

It's only true if dz

is infinitely small

Note: r<sub>i</sub> – volumetric rate of i production



$$F C_i(z) + r_i(z) A dz = F C_i(z + dz)$$

$$F C_i(z) + r_i(z) A dz = F C_i(z + dz)$$

$$\Leftrightarrow$$
 r<sub>i</sub> (z) A dz = F C<sub>i</sub> (z + dz) - F C<sub>i</sub> (z)

$$F C_i(z) + r_i(z) A dz = F C_i(z + dz)$$

$$\Leftrightarrow$$
  $r_i(z) A dz = F C_i(z + dz) - F C_i(z)$ 

$$\Leftrightarrow$$
  $r_i(z) = \frac{F}{A} \frac{C_i(z + dz) - C_i(z)}{dz}$ 

$$F C_i(z) + r_i(z) A dz = F C_i(z + dz)$$

 $\left(\mathbf{u} = \frac{\mathbf{F}}{\mathbf{A}}\right)$ 

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$$\Leftrightarrow r_{i}(z) = u \frac{C_{i}(z + dz) - FC_{i}(z)}{dz}$$



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$$\Leftrightarrow \lim_{dz \to 0} r_i(z) = u \lim_{dz \to 0} \frac{C_i(z + dz) - FC_i(z)}{dz}$$



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$$\Leftrightarrow r_i = u \frac{dC_i}{dz}$$



$$F C_i(z) + r_i(z) A dz = F C_i(z + dz)$$

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 r<sub>i</sub> (z) A dz = F C<sub>i</sub> (z + dz) - F C<sub>i</sub> (z)

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$$\Leftrightarrow r_i = u \frac{dC_i}{dz}$$

Eq. of material to component 'i' em PFR



### 3.3 – Kinetics

Example: product formation associated with growth (Type I)

Reaction: 
$$S \longrightarrow X + P$$

$$\mu = \frac{\mu_{\text{max}} S}{k_{\text{m}} + S} \qquad \text{(assuming Monod kinetics)}$$

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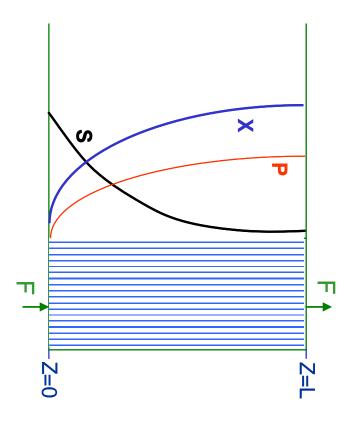
Material balances:  $u \frac{dX}{dz} = \mu X - k_d X - k_e X$ 
 $u \frac{dS}{dz} = -\frac{\mu X}{y_{XS}} - m_S X$ 
 $u \frac{dP}{dz} = \frac{\mu X}{y_{XP}}$ 

Analytical integration just for the case de  $\mu \cong \mu$ max (high excess of S)



#### 3.3 – Kinetics

Example: product formation associated with growth (Type I)



Typical Concentration Profile for Type I Product

- All concentrations depend on the 'z' position.
- Substrate concentration (S) decreases from bottom to top.
- Product (P) and biomass (X) concentrations increase from bottom to top.



## 3.4 – Productivity

Volumetric productivity of product

Prod = 
$$\frac{FP(z=L)}{V}$$
 =  $DP(z=L)$  g product l-1 h-1

F Flow rate (1/h)

P(z=L) conc. of product at the top of the column (at the exit of the reactor) (g/l)

V reactor volume (1)

## **Important note**

A design or optimization study should look for the maximum product concentration to occur at the top of the reactor.



**Objective:** convert So to  $S^* \rightarrow Volume needed?$ 





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### Material balance to the substrate:

$$FS_0 - r_s V - FS^* = 0$$

$$V = \frac{F(S_0 - S^*)}{r_s}$$



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#### Material balance to the substrate:

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$$u \frac{dS}{dz} = -r_s \iff u dS = -r_s dz$$
$$\iff -\frac{ds}{r_s} = \frac{dz}{u}$$

$$\Leftrightarrow -\int_{S_0}^{S^*} \frac{ds}{r_s} = \int_0^L \frac{dz}{u} = \frac{L - 0}{u} = \frac{L A}{u A} = \frac{V}{F}$$



$$u \frac{dS}{dz} = -r_s \iff u dS = -r_s dz$$
$$\Leftrightarrow -\frac{ds}{r_s} = \frac{dz}{u}$$

$$\Leftrightarrow -\int_{S_0}^{S^*} \frac{\mathrm{ds}}{\mathrm{r_s}} = \frac{V}{F}$$

$$\Leftrightarrow V = \frac{F(S_0 - S^*)}{\overline{r}_S}$$

$$\bar{r} = \frac{\int_{S^*}^{S_0} r_S(S) dS}{S_0 - S^*}$$



Objective: convert So to  $S^* \rightarrow Volume needed$ ?



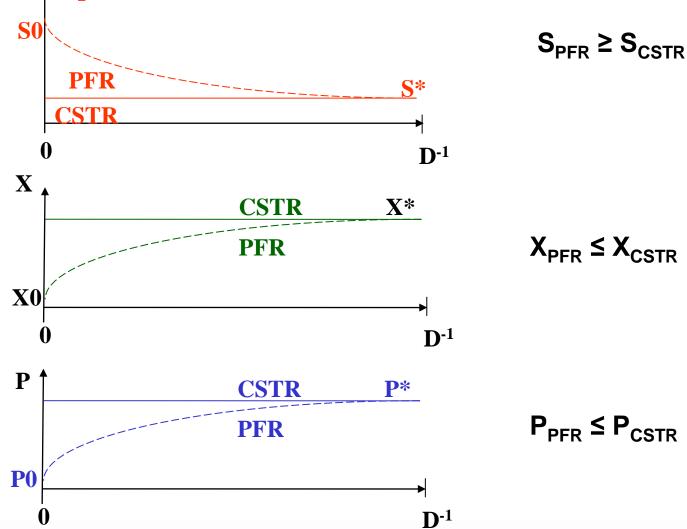
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$$V = \frac{F(S_0 - S^*)}{\overline{r}_s}$$





### 3.4 – Comparison of PFR with CSTR

Case 1: Negligible growth ( $X_{PFR} = X_{CSTR} = constant$  over time)

	CSTR $V = \frac{F(S_0 - S^*)}{r(S^*)}$		$\mathbf{PFR}$ $\mathbf{V} = \frac{F(S_0 - S^*)}{\bar{r}_S}$	
Kinetics	S <sub>CSTR</sub> P <sub>CSTR</sub>	< >	S <sub>PFR</sub> P <sub>PFR</sub>	
Order 0 $r_s = k_0$ (indepedent of S)	=		=	
Order 'n' $r_s = k S^n$			+++	
Michaelis-Menten $\mathbf{r}_s = \frac{r_{s \max S}}{K_m + S}$			+++	
Inhibition by S (S オ r <sub>s</sub> 凶)	+++			
Inhibition by produt (P オ rs 凶)			+++	

Note: the signs "+++" refer to the best reactor and "---" to the worst reactor



Case 2: significant cell growth (i.e. autocatalytic kinetics)

$$V = \frac{F(S_0 - S^*)}{r(S^*)}$$

$$V = \frac{F(S_0 - S^*)}{\overline{r}_S}$$

$$X_{CSTR}$$
 >  $X_{PFR}$ 

Autocatalitic

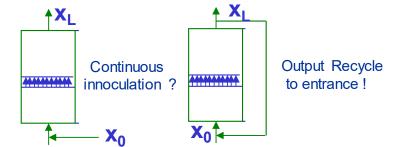
$$r_s = v_s X$$

- .: Biocatalysis with cells (bacteria, fungi, animal cell lines) CSTR tends to be more productive!!!!!!!!
- .: Biocatalysis with enzymes PFR tends to be more productive!!!!!!!!



### 3.5 – Discussion about PFR

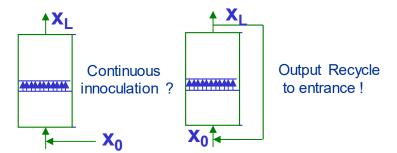
1. How to do Innoculation?



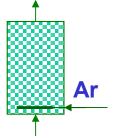


#### 3.5 – Discussion about PFR

1. How to do Innoculation?



2. How to make areation?

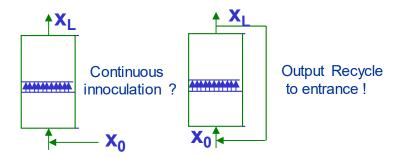


Bubbling at the bottom of the reactor promotes mixing.
PFR tends to CSTR

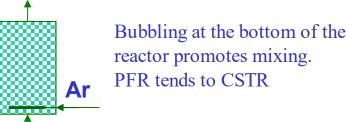


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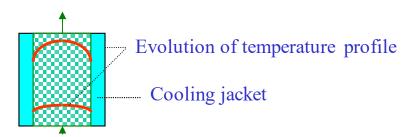


2. How to make areation?



3. How to control temperature?

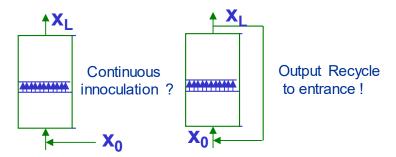
heat transport less efficient than in CSTR



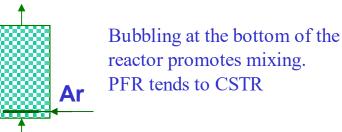


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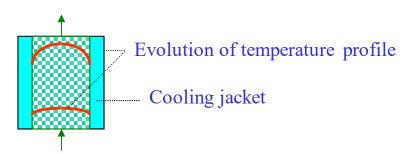
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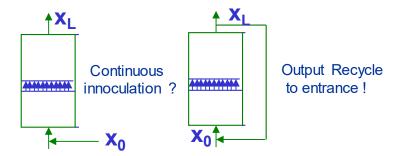
4. How to control pH?



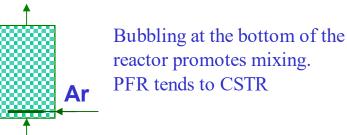


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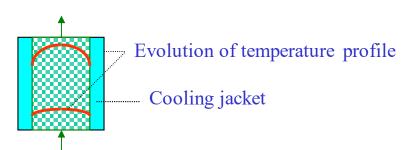


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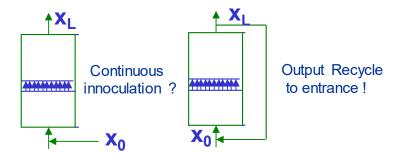


- 4. How to control pH?
- 5. How to control the concentration of any component?

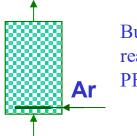


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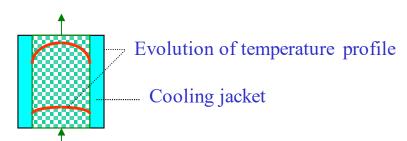
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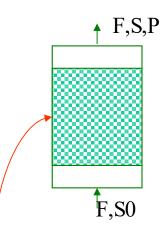
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PFR with cells in suspension (and remarkable cell growth) is an impractical construct. It can, however, occur in practice in association with other bioreactors (example: plug flow associated with CSTRs)



#### 3.5 – Discussion about PFR

Exception: tubular bioreactor with immobilized cells (or enzymes)



solid support
Cells grow adherent to
solid support or
'incarcerated' in a
polymer matrix

- Very stable cultures that remain viable for long periods of time (months, years)
- Cells at rest (low maintenance)
- High cell density (much higher than cells in suspension). Cells grow adherent and form biofilms
- After a growth phase, cell density remains constant over time (new cells simply replace the dying cells)
- Higher dilution rates because washout cannot occur

## μ negligible

⇒kinetically favorable to the PFR regarding the CSTR