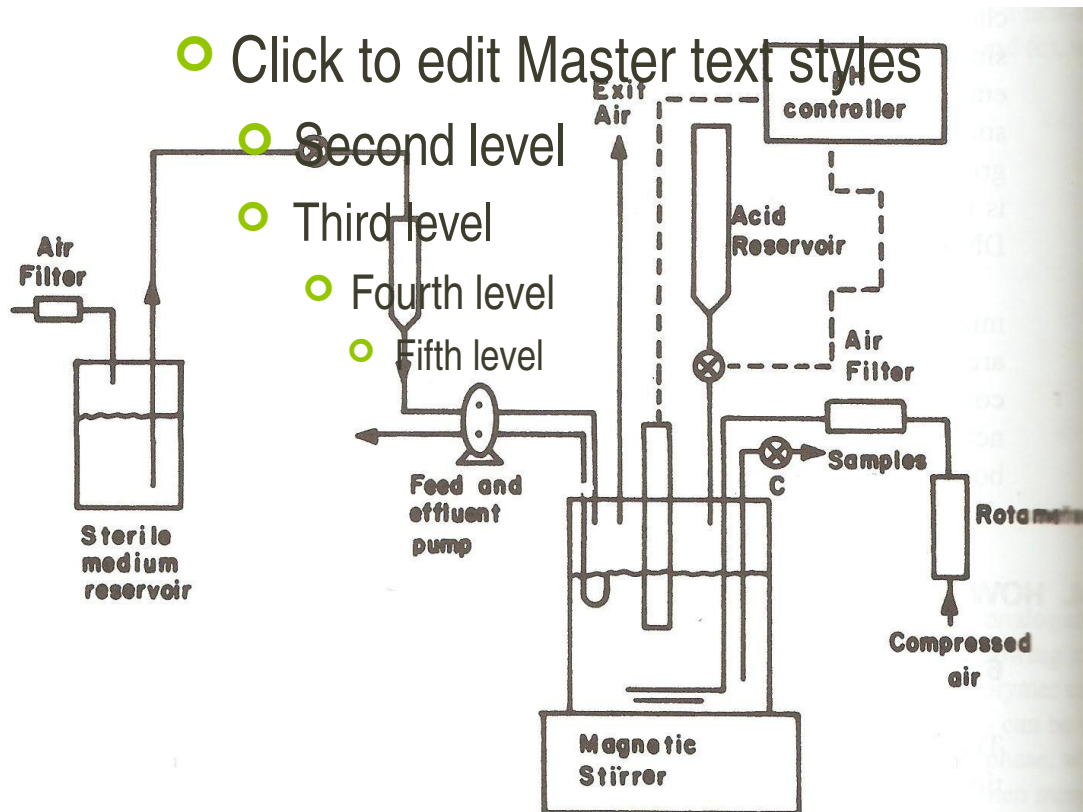


Načini obratovanja bioreaktorjev

Kontinuirni bioproces



Tipi kontinuirnih obratovanj

Glede na metodo nadzora:

- ● **Kemostat** - reguliran na osnovi nadzora koncentracije limitnega hranila
- ● **Turbidostat** - reguliran na osnovi nadzora biomase z uporabo optične gostote (fotoelektrična celica)
- ● **Biostat** - reguliran na osnovi sistema za nadzor biomase, ki ne temelji na optični gostoti (npr. proizvodnja CO₂)

Uporaba kontinuirnih procesov

- **Industrija**
- Biološke čistilne naprave
- Proizvodnja „single-cell“ proteinov
- Kontinuirna proizvodnja piva
- Kontinuirna proizvodnja amino kislin
- Kontinuirna proizvodnja organskih kislin
- Kontinuirna proizvodnja etanola

Uporaba kontinuirnih procesov

○ Raziskave

○ Fiziološke in biokemijske študije za nadzor hitrosti rasti

○ Vpliv dejavnikov okolja/ procesnih parametrov na rast in tvorbo produkta

Indukcija, represija, hitrost rasti, vpliv temperature, pH itd.

○ Mikrobna ekologija

- Izbor populacij, ki rastejo počasi
- Interakcije žrtev-plenilec
- Kompetitivnost (npr. plasmidi +/-)

○ Kinetične študije

Izračun rastnih konstant, podatki o fermentacijah

Snovne bilance – kontinuirni proces

- za biomaso (X):

$$\frac{dX}{dt} = \mu X - \frac{FX}{V} = \mu X - D X$$

- za substrat (S):

$$\frac{dS}{dt} = \frac{F S_v}{V} + r_s - \frac{FS}{V} = D(S_v - S) + r_s$$

- za produkt (P):

$$\frac{dP}{dt} = r_p - \frac{FP}{V} = r_p - D P$$

Kemostat

- Stacionarno stanje

- Biomasa:

$$0 = \mu X - D X$$

$$\mu = D$$

- Substrat:

$$0 = D(S_v - S) + r_s$$

$$X = Y_{x/s} (S_v - S)$$

$$S = \frac{\mu K_s}{\mu_{\max} - \mu}$$

$$S = \frac{DK_s}{\mu_{\max} - D}$$

Kemostat – tvorba produkta

$$D(P_F - P) + Y_{P/X} \cdot \mu \cdot X = 0$$

Če je PF = 0

$$P = \frac{Y_{P/X} \mu X}{D}$$

Uporaba kontinuirnih procesov

$$V_R \cdot \frac{dS}{dt} = F \cdot S_F - F \cdot S - V_R \cdot \mu_g \cdot X \cdot \frac{1}{Y_{X/S}^M} - V_R \cdot q_P \cdot X \cdot \frac{1}{Y_{P/S}}$$

$$\frac{\mu_g}{Y_{X/S}^M}$$

...maksimalni izkoristek X/S

Če ni tvorbe produkta + stacionarno stanje:

$$D \cdot (S_F - S) = \mu_g \cdot X \cdot \frac{1}{Y_{X/S}^M}$$

$$D = \mu_g - k_d = \mu$$

$$D \cdot (S_F - S) - (D + k_d) \cdot X \cdot \frac{1}{Y_{X/S}^M} = 0 \quad /: X$$

$$D \left(\frac{S_F - S}{X} \right) - \frac{D}{Y_{X/S}^M} - \frac{k_d}{Y_{X/S}^M} = 0 \quad /: D$$

$$\frac{1}{Y_{X/S}^{AP}} = \frac{1}{Y_{X/S}^M} + \frac{k_d}{Y_{X/S}^M \cdot D} = \frac{1}{Y_{X/S}^M} + \frac{m_s}{D}$$

$$m_s = \frac{k_d}{Y_{X/S}^M}$$

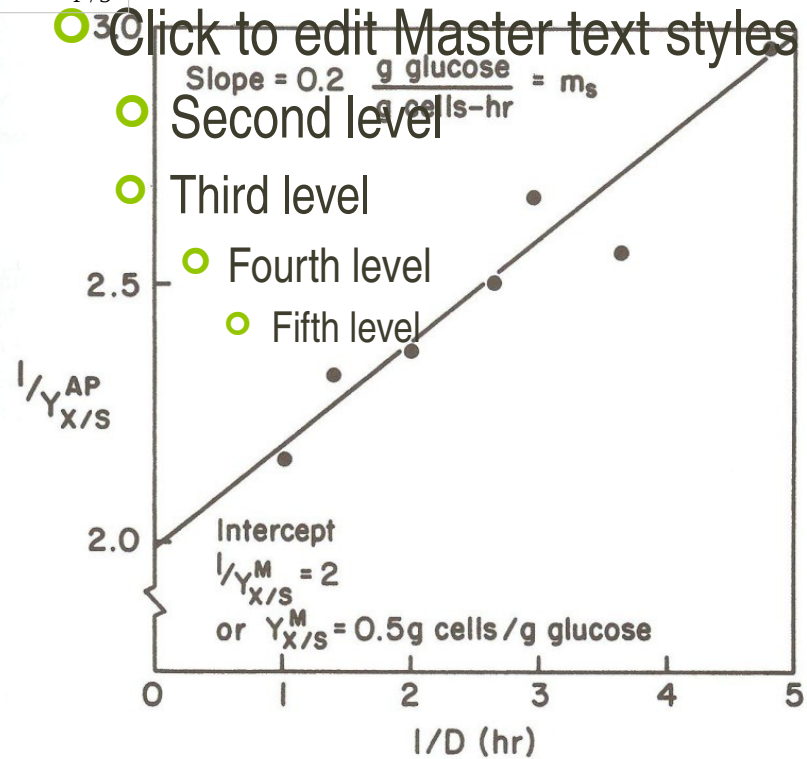


Figure 6.19. Graphical approach to estimating $Y_{X/S}^M$ and m_s for chemostat data for *E. coli* growing on glucose as the limiting nutrient.

Uporaba kontinuirnih procesov

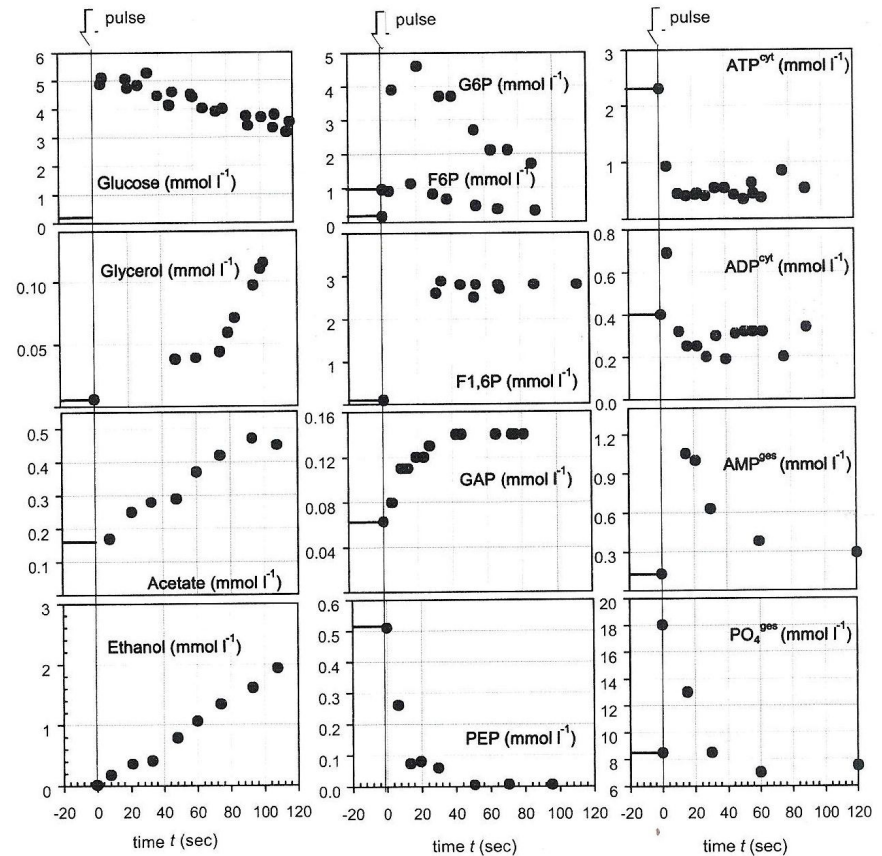
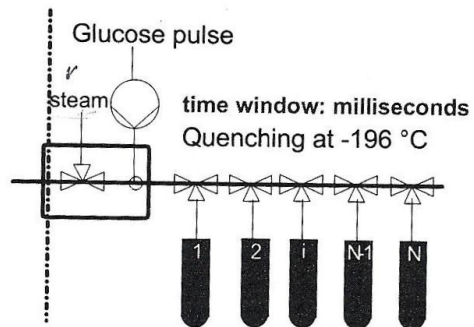
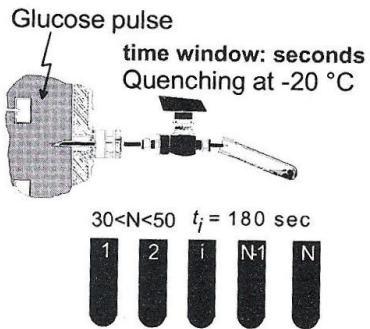
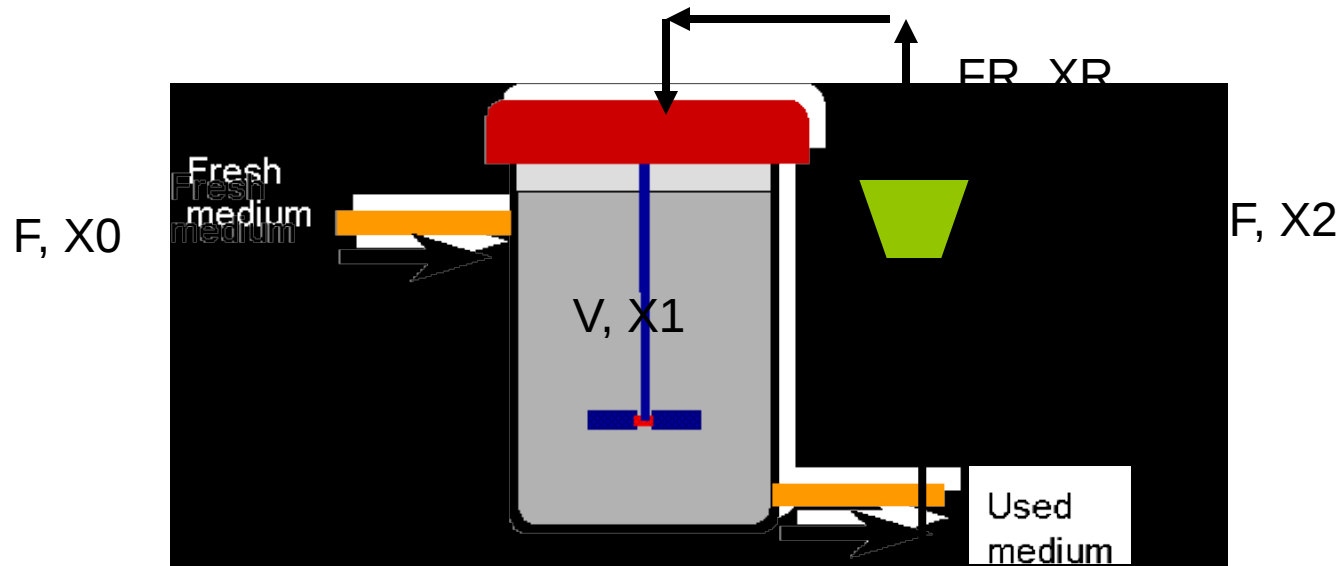


Figure 4. Changes in the concentration of extracellular products and substrates (left side), intracellular metabolites (middle) and intracellular co-metabolites (right hand side) a glucose pulse at $t=0$ sec.

Kemostat z reciklom celic



F – pretok napajalne raztopine

V – volumen reaktorja

X1 – koncentracija biomase v reaktorju

X2 - koncentracija biomase v iztoku

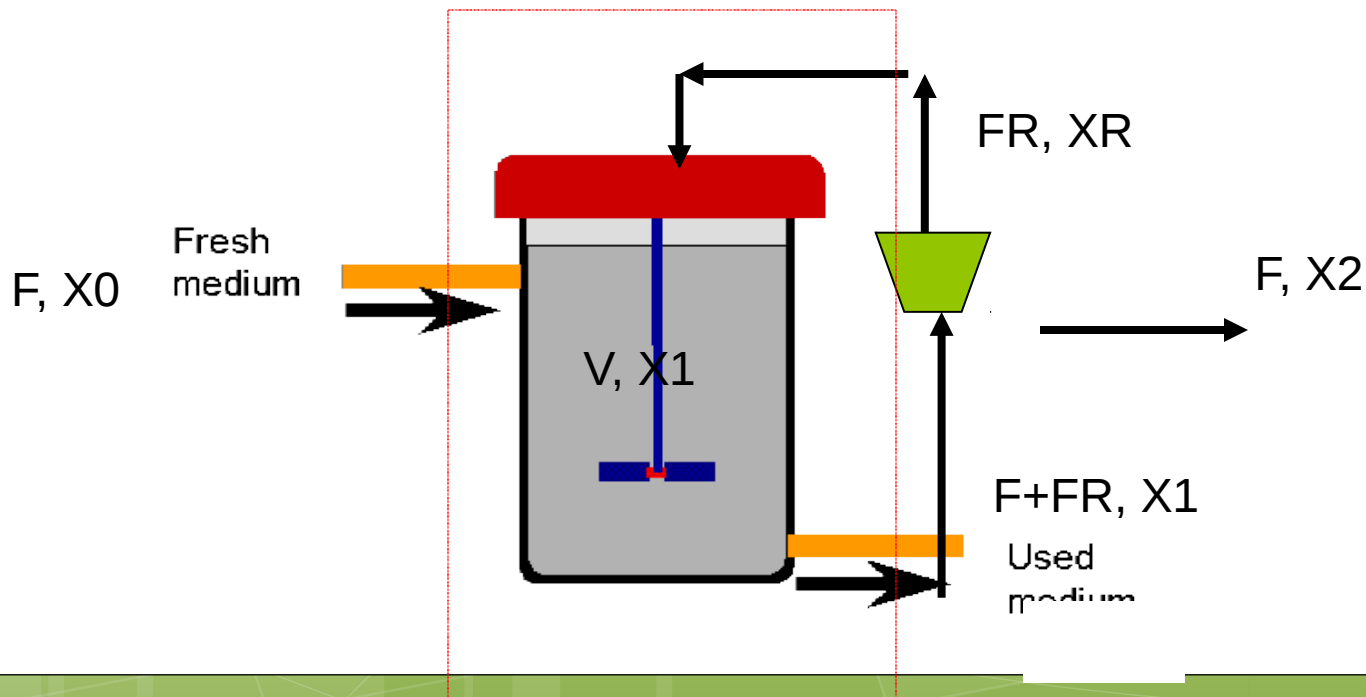
XR - koncentracija biomase v reciklu

FR – pretok recikla

Kemostat z reciklom celic

Snovna bilanca za biomaso:

$$F X_0 + F_R X_R - (F + F_R) X_1 + V \mu X_1 = \frac{dX}{dt} V$$



Kemostat z reciklom celic

Definicije

$$\alpha = FR/F$$

$$C = XR / X1$$

Izpeljava

- $F + FR = (1 + \alpha)F$

- $FRXR = \alpha CFX1$

$$FR = F\alpha$$

$$XR = CX1$$

$$FRXR = \alpha CFX1$$

$$F X_0 + FR XR - (F + FR) X_1 + V\mu X_1 = \frac{dX_1}{dt} V$$

$$F X_0 + \alpha CFX_1 - (1 + \alpha)F X_1 + V\mu X_1 = \frac{dX_1}{dt} V$$

Kemostat z reciklom celic

- Predpostavke

- Stacionarno stanje:

$$= 0 \frac{dX_1}{dt}$$

- Sterilni vtok: $X_0 = 0$

$$(\alpha C - 1 - \alpha)F + V\mu = 0$$

e je $D = F/V$, velja za recikel:

$$\mu = D(1 + \alpha(1 - C))$$

e je $C > 1$ (konc. celic), potem je $\alpha(1 - C) < 0$

in je $\mu < D$

Kemostat z
reciklom
lahko
deluje pri
 $D > \mu_{\max}$

Bilanca za substrat- Recikel

$$FS_0 + \alpha FS - V \frac{\mu X_1}{Y_{X/S}} - (1 + \alpha) FS = V \frac{dS}{dt}$$

- V stacionarnem stanju in menjavi D za μ :

$$X_1 = \frac{D}{\mu} Y_{X/S} (S_0 - S) = \frac{Y_{X/S} (S_0 - S)}{(1 + \alpha - \alpha C)}$$

Bilanca za substrat- Recikel

- Upoštevamo kinetiko Monoda

$$S = \frac{K_s D(1 + \alpha - \alpha C)}{\mu_{\max} - D(1 + \alpha - \alpha C)}$$

$$X_1 = \frac{Y_{X/S}}{(1 + \alpha - \alpha C)} \left[S_0 - \frac{K_s D(1 + \alpha - \alpha C)}{\mu_{\max} - D(1 + \alpha - \alpha C)} \right]$$

Šaržni proces z dohranjevanjem

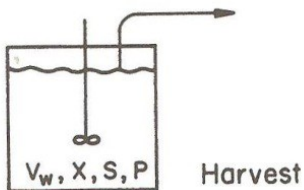
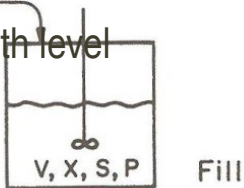
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Second level

Third level

Fourth level

Fifth level



$$\frac{d(CV)}{dt} = V \frac{dC}{dt} + C \frac{dV}{dt} = V \frac{dC}{dt} + CF_{in}(t)$$

$$\frac{F_{in}(t)}{V(t)} = D$$

$$\frac{d(XV)}{dt} = \mu X V$$

$$\frac{d(SV)}{dt} = F S V + r_s V$$

$$\frac{d(PV)}{dt} = r_p V$$