STABILIZATION OF CHEMOSTATS WITH MORTALITY AND SUBSTRATE DYNAMICS

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PART I

The simplest chemostat model

The chemostat: A microbial bioreactor



The chemostats are used to

- produce antibiotics and some proteins
- Ireat water (degradation of pollutant)
- oproduce biomass

4 ...

The simplest chemostat is modeled by:

$$\begin{cases} \dot{X} = (p_0 \mu(S) - \boldsymbol{b} - D)X\\ \dot{S} = D(S_{\text{in}} - S) - \mu(S)X. \end{cases}$$

(1)

Symbols	Amounts
X	Microbial concentration or biomass
S	Concentration of the substrate (nutrient)
$S_{\sf in}$	Inlet concentration of the nutrient
$\mu(S)$	Specific growth rate
D	Dilution rate
b	Mortality rate
$p_0 = 1/k$	Yield factor

The parameters X, S, D, b, p_0 are positive and $\mu \in C^1(\mathbb{R}_{\geq 0}, \mathbb{R}_{\geq 0})$ with $\mu(0) = 0$.

The goal in the chemostat is to maintain the system in a non washout steady state.

An equilibrium point of the chemostat model (1) is (X^*, S^*) such that

$$p_0\mu(S^*) = b + D^*,$$

 $X^* = rac{D^*(S_{
m in} - S^*)}{\mu(S^*)}.$

Given $D^* > 0$, it may exist many equilibrium points

Given $D^* > 0$ and an equilibrium (X^*, S^*) , the linearization of the system (1) is

$$\begin{pmatrix} \dot{X} \\ \dot{S} \end{pmatrix} = \begin{pmatrix} 0 & p_0 \mu'(S^*) X^* \\ -(b+D^*)/p_0 & -D^* - \mu'(S^*) X^* \end{pmatrix} \begin{pmatrix} X \\ S \end{pmatrix}.$$
 (2)

The characteristic polynomial is:

$$P(s) = s^{2} + (D^{*} + \mu'(S^{*})X^{*})s + (b + D^{*})\mu'(S^{*})X^{*}.$$
(3)

The system is unstable if $\mu'(S^*) < 0$.

When considering the Monod kinetics

$$\mu(S) := \frac{\mu_{\max}S}{K+S},\tag{4}$$



Theorem 1 (Dali-Youcef, Rapaport, and Sari 2022)

There exists a unique positive equilibrium point of (1) which is globally asymptotically stable and exponentially stable.

When, considering the Haldane kinetics

$$\mu(S) := \frac{\mu_{\max}S}{K + S + aS^2},\tag{5}$$



Theorem 2

The system (1) admits two positive equilibrium points $(X^*, S^*), (X^{**}, S^{**})$ such that

 \succ (X^*, S^*) is unstable.

> (X^{**}, S^{**}) is exponentially stable.

Unstable open-loop chemostat model



Figure: The phase diagram for the open-loop chemostat model (1) with Haldane equation

Theorem 3

Assume that

$$p_0\mu(S) > b, \quad \forall S \in [S^*, S_{in}].$$
 (6)

Then for every $\delta>0$ and $\alpha\in[0,1)$ the feedback law defined as

$$D(X,S) = \frac{D^*\mu(S)X}{\mu(S^*)X^*} + \frac{\delta b}{(\mu(S^*))^{1+\alpha}} \begin{cases} |\mu(S) - \mu(S^*)|^{1+\alpha}, & \text{if } S \le S^* \\ 0, & \text{if } S > S^* \end{cases}$$
(7)

achieves global asymptotic stabilization of (X^*, S^*) . Moreover, if $\alpha > 0$ then the feedback law (7) also achieves local exponential stabilization of (X^*, S^*) .

Stable closed-loop model



Figure: The phase diagram for the closed-loop chemostat model (1) with (7).

Is the sufficient condition (6) also necessary to ensure GAS of system (1)?

Theorem 4

Assume that there exists a constant $\overline{S} \in (S^*, S_{in})$ such that

μ

$$p_0\mu(\overline{S}) < b \tag{8}$$
$$T(S) \le 0, \quad \forall S \in [\overline{S}, S_{in}]. \tag{9}$$

Then, there is no feedback law $D(X,S) \ge 0$ that achieves global stabilization of the equilibrium (X^*, S^*) of (1).

PART II

An age structured chemostat model with substrate dynamics

Consider the following age structured chemostat model with substrate dynamics

$$\frac{\partial f}{\partial t}(t,a) + \frac{\partial f}{\partial a}(t,a) = -(\beta(a) + D(t))f(t,a), \tag{10}$$

$$f(t,0) = \mu(S(t)) \int_0^{+\infty} k(a) f(t,a) da,$$
(11)

$$\dot{S}(t) = D(t)(S_{\text{in}} - S(t)) - \mu(S(t)) \int_0^{+\infty} q(a)f(t,a)da,$$
(12)

where f(t,a) > 0 is the distribution function of the microbial population in the chemostat at time $t \ge 0$ and age $a \ge 0$.

We derived the following ODE chemostat model

$$\dot{S} = D(S_{\rm in} - S) - \mu(S)X \tag{13a}$$

$$\dot{X} = q_0 \mu(S) Y - (b+D) X \tag{13b}$$

$$\dot{Y} = p_0 \mu(S) Y + \gamma X - (b+D) Y, \qquad (13c)$$

where

$$X(t) = \int_0^{+\infty} q(a)f(t,a)da, \quad Y(t) = \int_0^{+\infty} k(a)f(t,a)da.$$
 (14)

Unstable open-loop equilibrium



Figure: The phase diagram for the chemostat model (13) with Haldane kinetics.

(A) There exists a constant $\varphi > 1$ such that for $S \in [S^*, S_{in}]$:

$$\begin{pmatrix} b - \frac{\lambda(b+D^*)}{2} \end{pmatrix} \left(\frac{\mu(S^*)}{\mu(S)} - 1 \right) + (1+\lambda) \frac{\lambda^2 \varphi(b+D^*)^2}{4D^*} \left(\frac{\mu(S^*)}{\mu(S)} - 1 \right)^2 < D^* \left(1 - \frac{1}{4(1+\lambda)\varphi} \right)$$

Theorem 5

Under assumption (A) the feedback law

$$D(X,S) = \frac{D^*\mu(S)X}{\mu(S^*)X^*} + \frac{\delta}{(\mu(S^*))^{1+\alpha}} \begin{cases} |\mu(S) - \mu(S^*)|^{1+\alpha}, & \text{if } S \le S^* \\ 0, & \text{if } S > S^* \end{cases}$$
(15)

achieves global asymptotic stabilization of the equilibrium. Moreover, if $\alpha > 0$, then the feedback law (15) achieves in addition local exponential stabilization of the equilibrium.



Figure: The phase diagram of the closed-loop system (13) with (15)

- Consider S_{in} and/or the volume V to be depending on t
- 2 Consider another reproduction rate μ than Monod and Haldane
- Investigate the stabilization of the PDE-ODE model (10)-(11)-(12)

Thank you for your attention