

Optimum dilution rate for cell productivity with substrate inhibition.  
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Operating condition:

$$s_f := 200$$

Model parameters:

$$\mu_m := 0.3 \quad K := 50 \quad K_i := 0.01 \quad A := 0.004 \quad B := 0.001$$

$$\mu(s) := \frac{\mu_m \cdot s}{K + s + K_i \cdot s^2} \quad Y(s) := A + B \cdot s$$

$$s := 0 .. s_f$$

Maximum specific growth rate  $\mu$ :  $s := 10$  ... initial guess

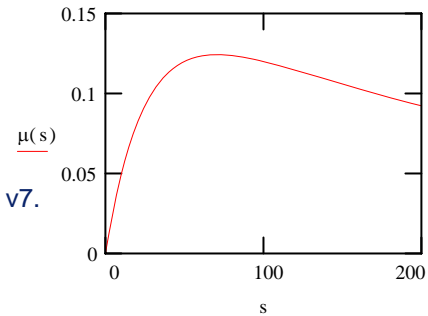
$$\text{Given} \quad \frac{d}{ds} \mu(s) = 0 \quad s := \text{Find}(s) \quad s = 70.711$$

$$\mu(s) = 0.124$$

An initial guess of  $s=0$  is o.k. in Mathcad v5, but no good in v7.

Note that the "root" function does not seem to be as accurate:

$$s := 0 \quad s := \text{root}\left(\frac{d}{ds} \mu(s), s\right) \quad s = 51.301 \quad \mu(s) = 0.121$$



Initial guesses:

$$x := 1 \quad s := 0$$

Given Steady-state equations:

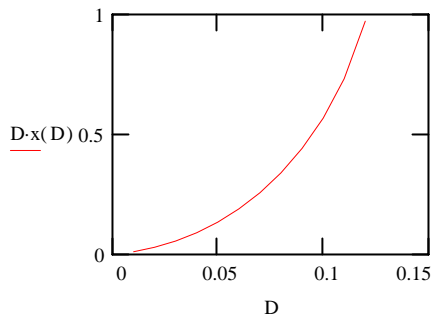
$$dx/dt: \quad 0 = (\mu(s) - D) \cdot x$$

$$ds/dt: \quad 0 = D \cdot (s_f - s) - \frac{1}{Y(s)} \cdot \mu(s) \cdot x$$

$$\text{ans}(D) := \text{Find}(x, s) \quad x(D) := \text{ans}(D)_0$$

$$\text{An example:} \quad \text{ans}(0.1) = \begin{pmatrix} 5.683 \\ 29.289 \end{pmatrix}$$

Dependence of  $D \cdot x$  on  $D$ :  $D := 0.01, 0.02 .. 0.124$



Note that cell productivity increases monotonically with  $D$ , right up to maximum  $\mu$ , at which point cell washout occurs. The maximum cell productivity occurs at:  $D := 0.124$   
The maximum cell productivity is:  $D \cdot x(D) = 1.155$