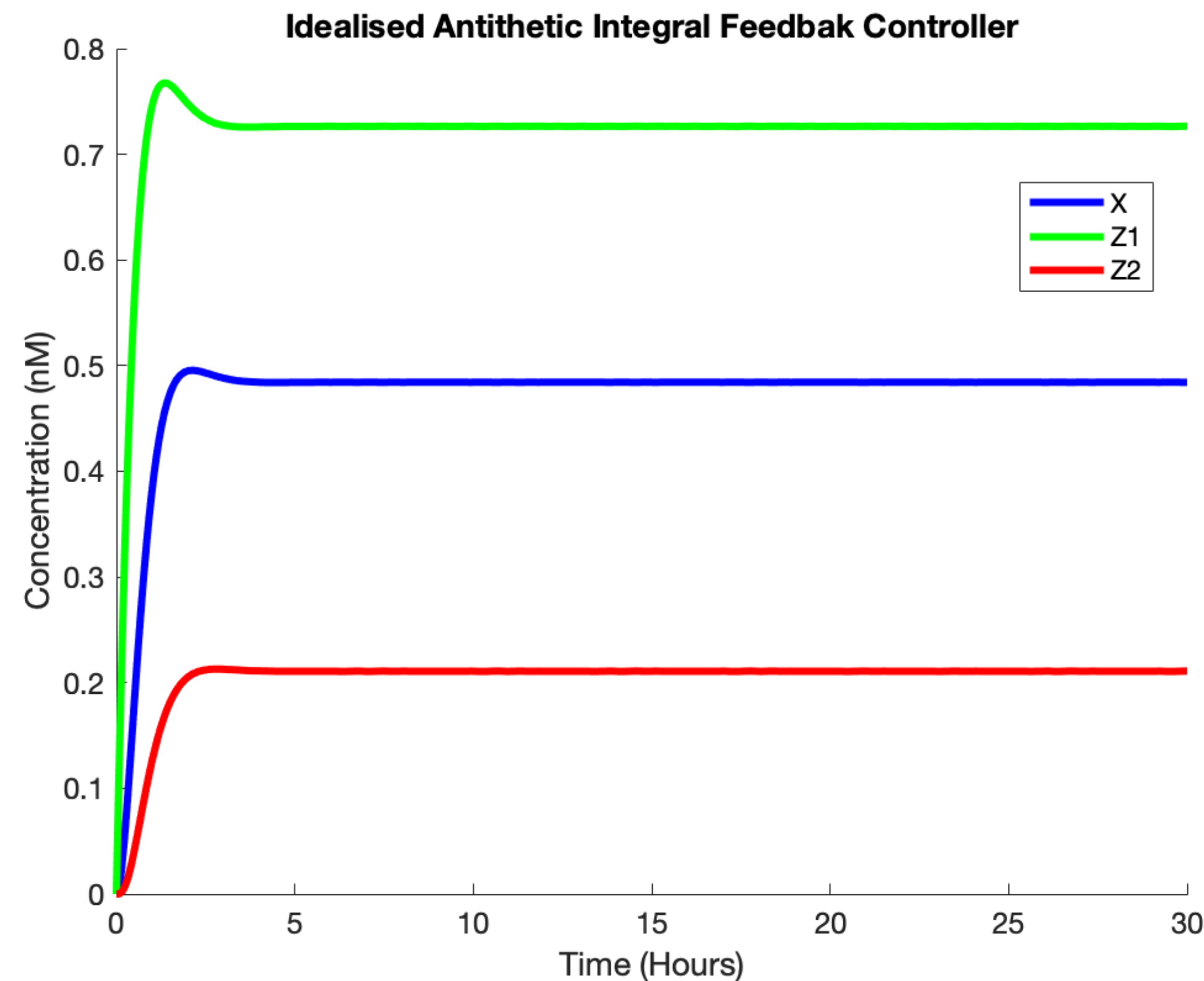
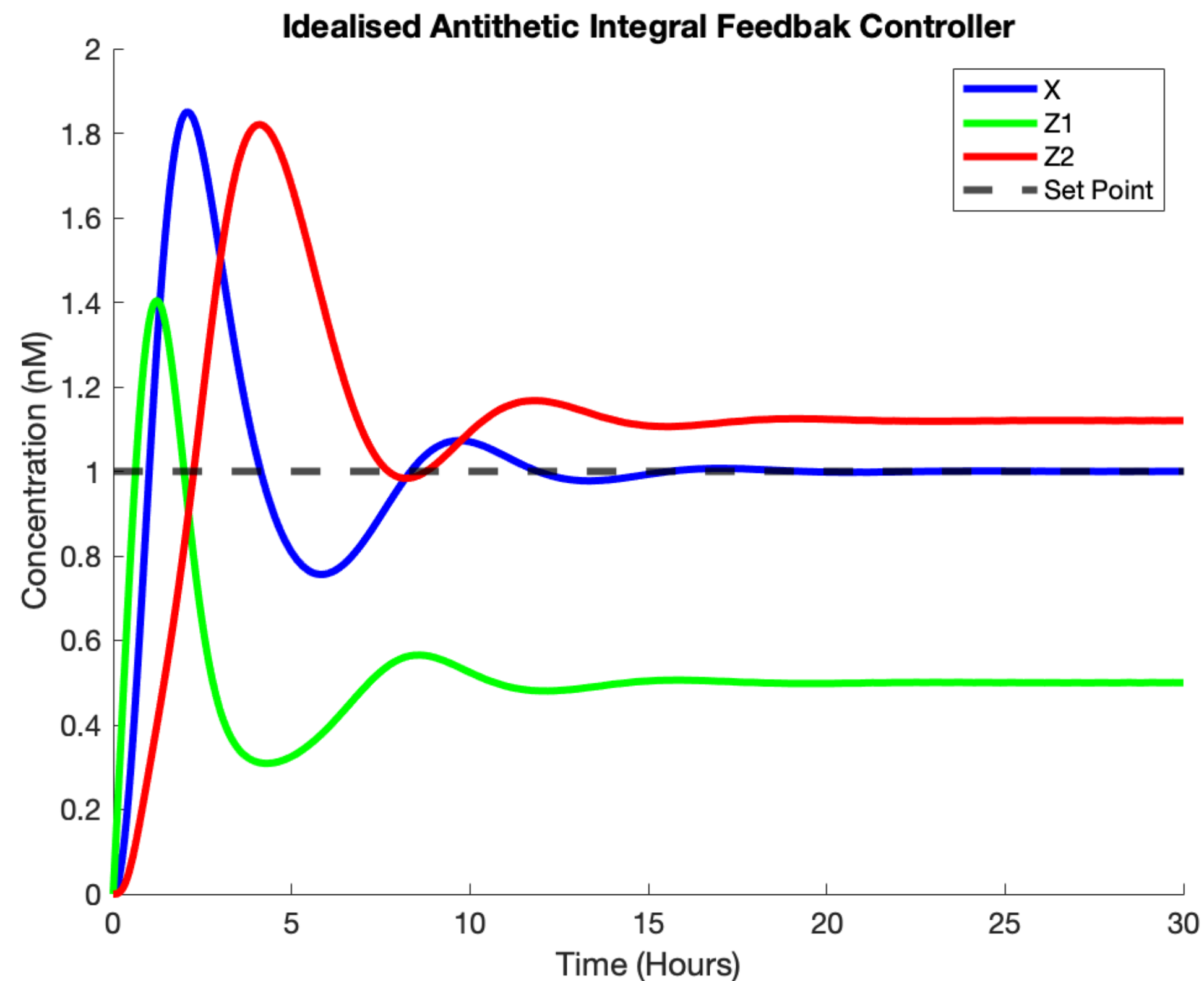


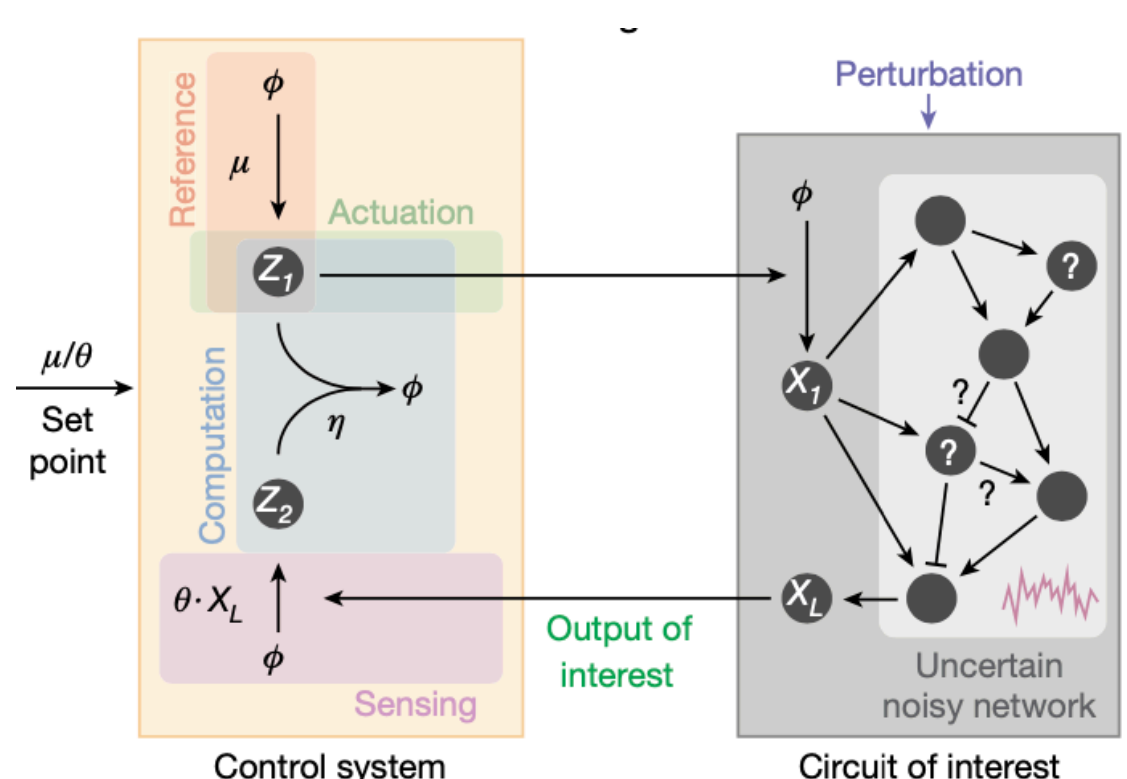
Antithetic

$$\frac{dZ_1}{dt} = \mu - \eta Z_1 Z_2 - \delta Z_1$$
$$\frac{dZ_2}{dt} = \theta X - \eta Z_1 Z_2 - \delta Z_2$$
$$\frac{dX}{dt} = k Z_1 - (\gamma + \delta) X$$
$$\text{setpoint} = \frac{\mu}{\theta}$$



$$\mu = 0.028 \text{ nMmin}^{-1}; \eta = 0.05 \text{ nM}^{-1}\text{min}^{-1}; \theta = 0.028 \text{ min}^{-1}$$
$$k = 0.028 \text{ min}^{-1}; \gamma = 0.014 \text{ min}^{-1}; \delta = 0 \text{ min}^{-1}$$

$$\mu = 0.028 \text{ nMmin}^{-1}; \eta = 0.05 \text{ nM}^{-1}\text{min}^{-1}; \theta = 0.028 \text{ min}^{-1}$$
$$k = 0.028 \text{ min}^{-1}; \gamma = 0.014 \text{ min}^{-1}; \delta = 0.028 \text{ min}^{-1}$$



Antithetic with Damping

$$\frac{dZ_1}{dt} = \mu - \eta Z_1 Z_2 - \delta Z_1$$

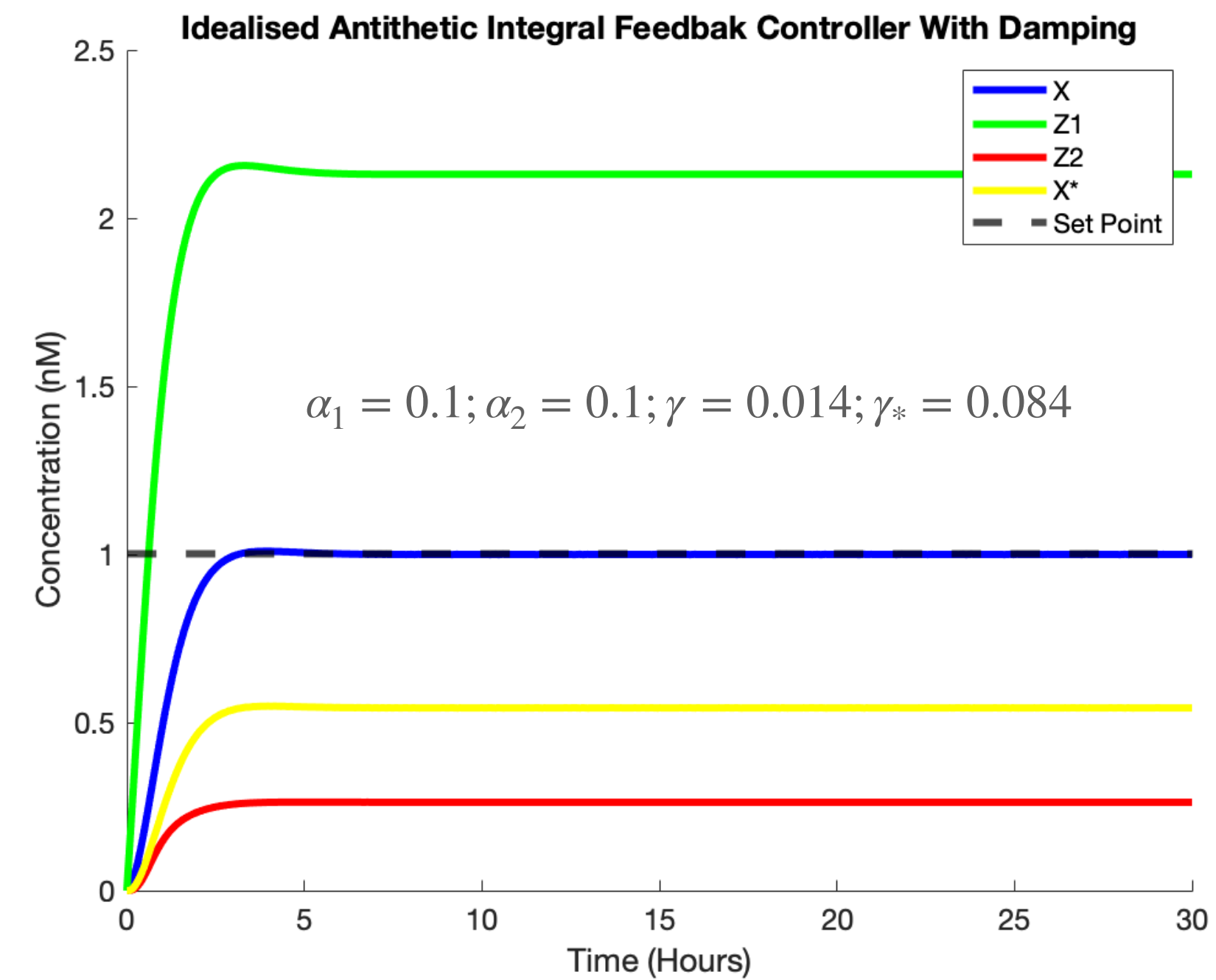
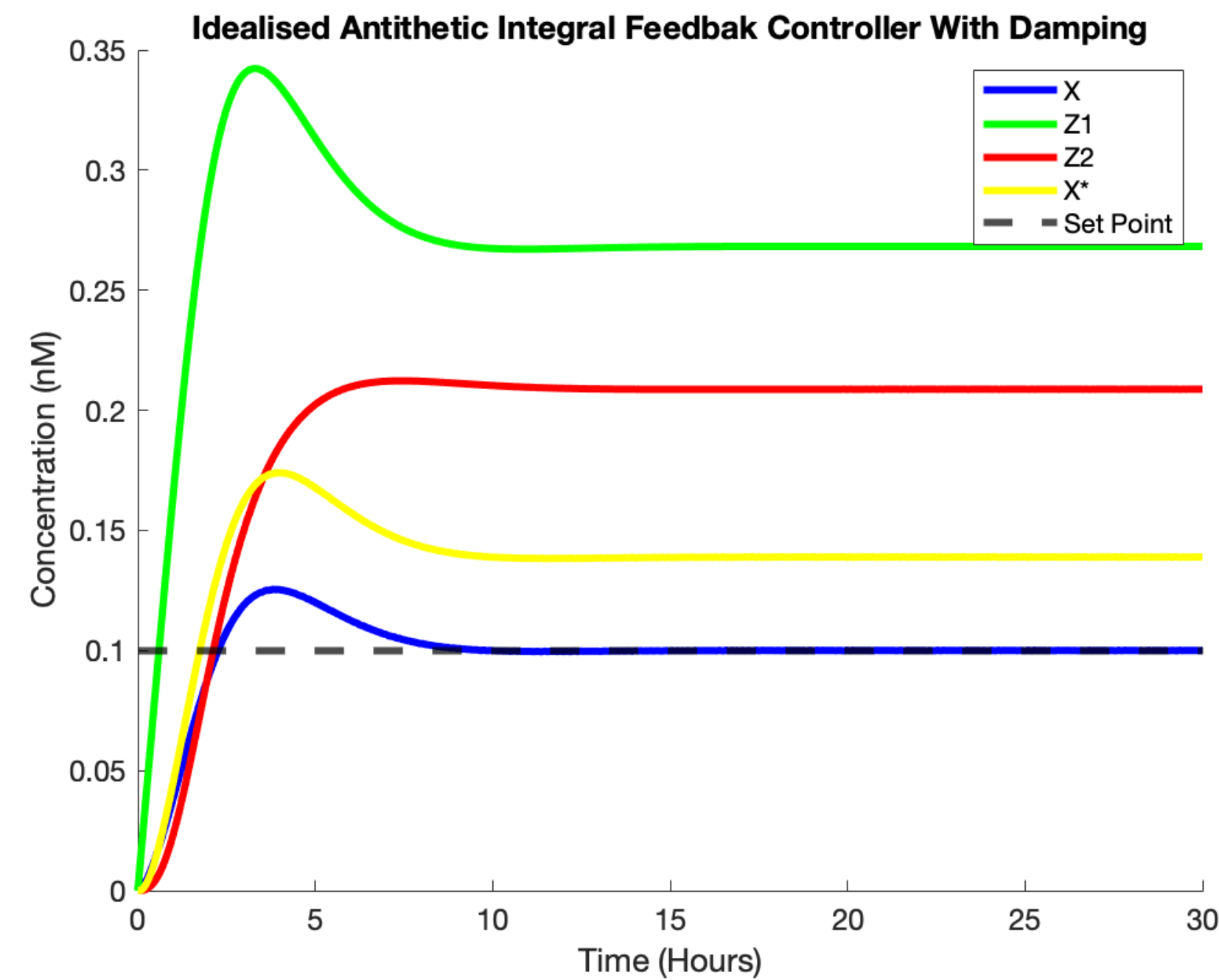
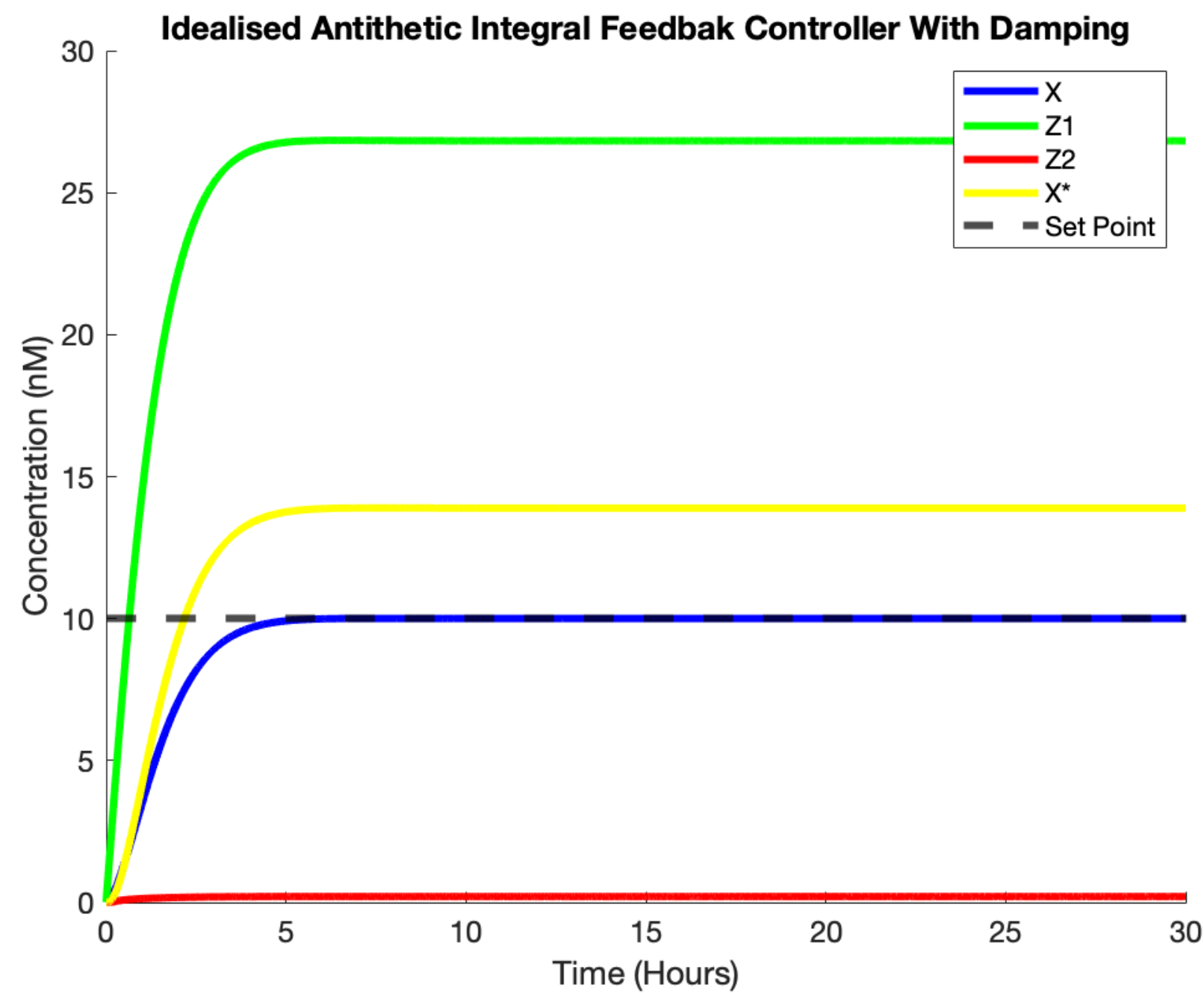
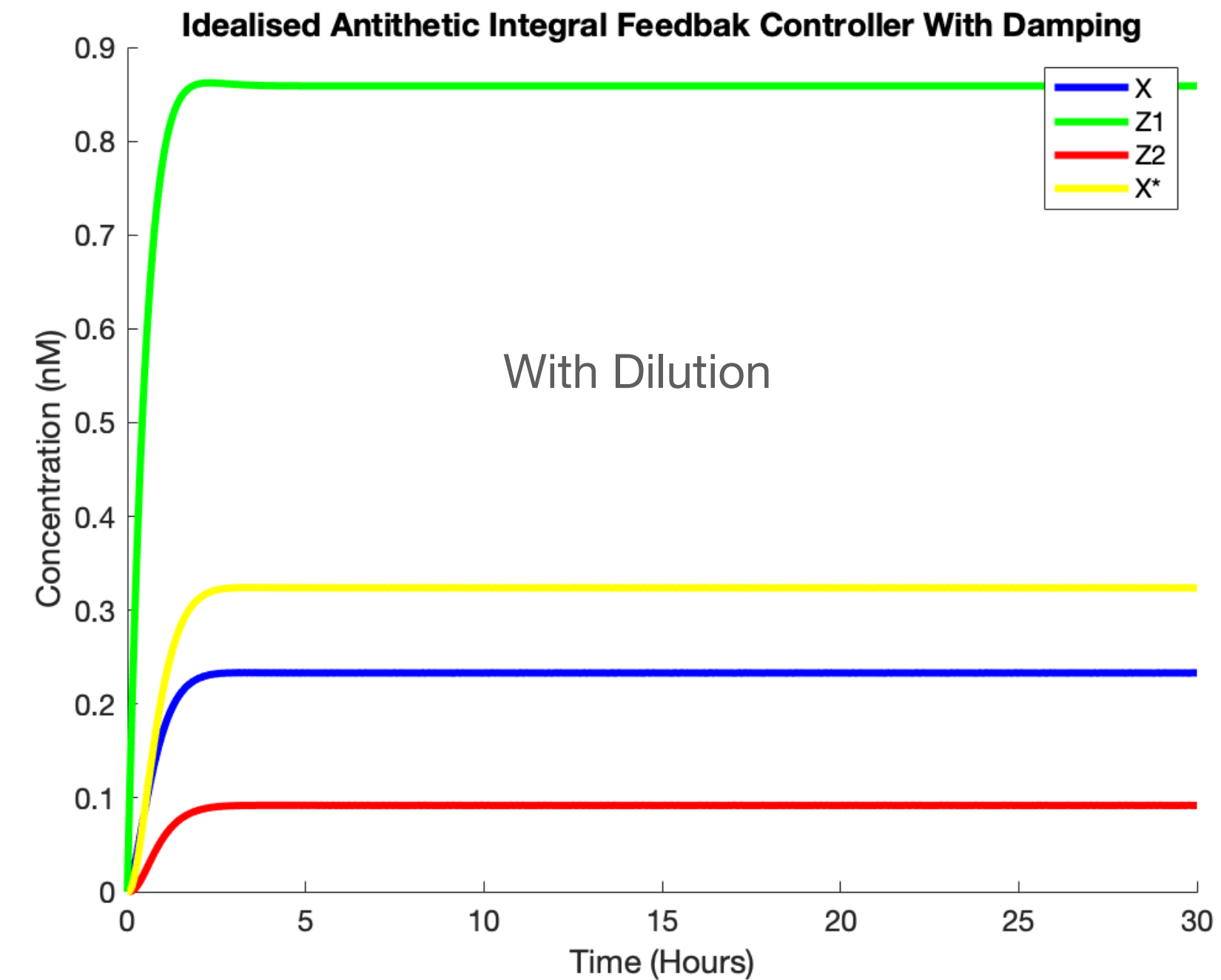
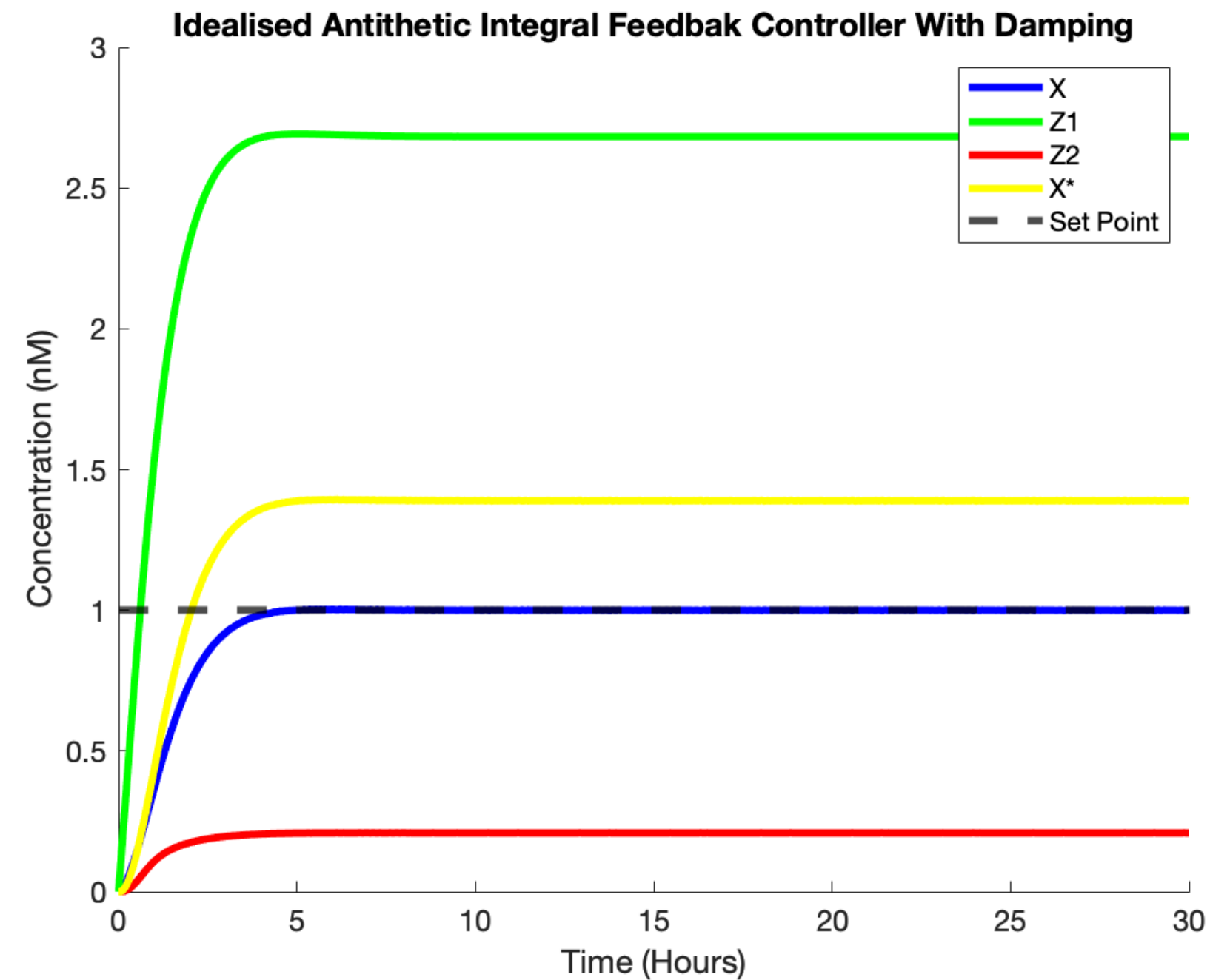
$$\frac{dZ_2}{dt} = \theta X - \eta Z_1 Z_2 - \delta Z_2$$

$$\frac{dX}{dt} = k Z_1 - (\gamma + \delta) X - \alpha_1 X + \alpha_2 X^*$$

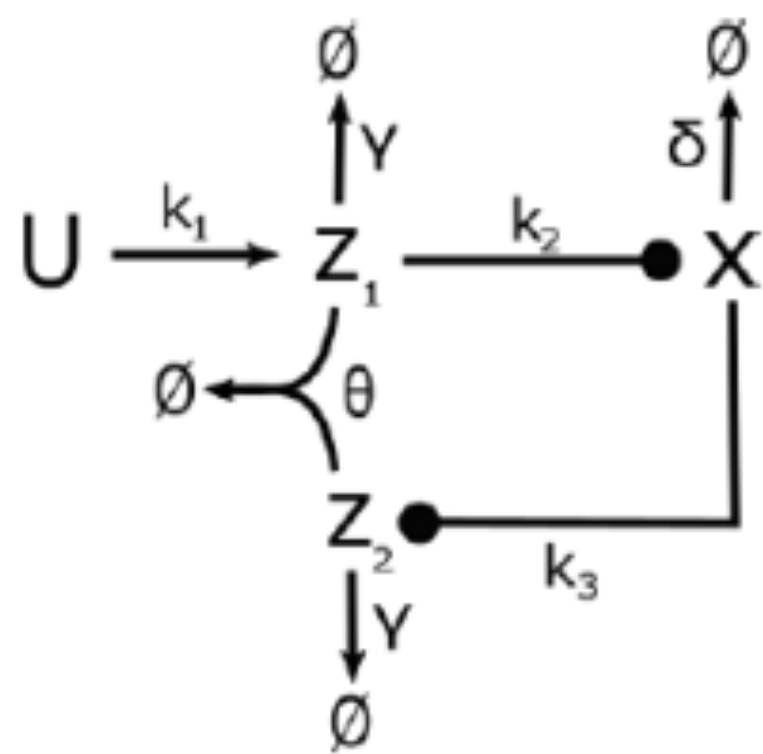
$$\frac{dX^*}{dt} = \alpha_1 X - \alpha_2 X^* - \gamma_* X^*$$

$$\text{setpoint} = \frac{\mu}{\theta}$$

$$\alpha_1 = 0.2; \alpha_2 = 0.1; \gamma = 0.014; \gamma_* = 0.044$$



Antithetic sRNA Implementation



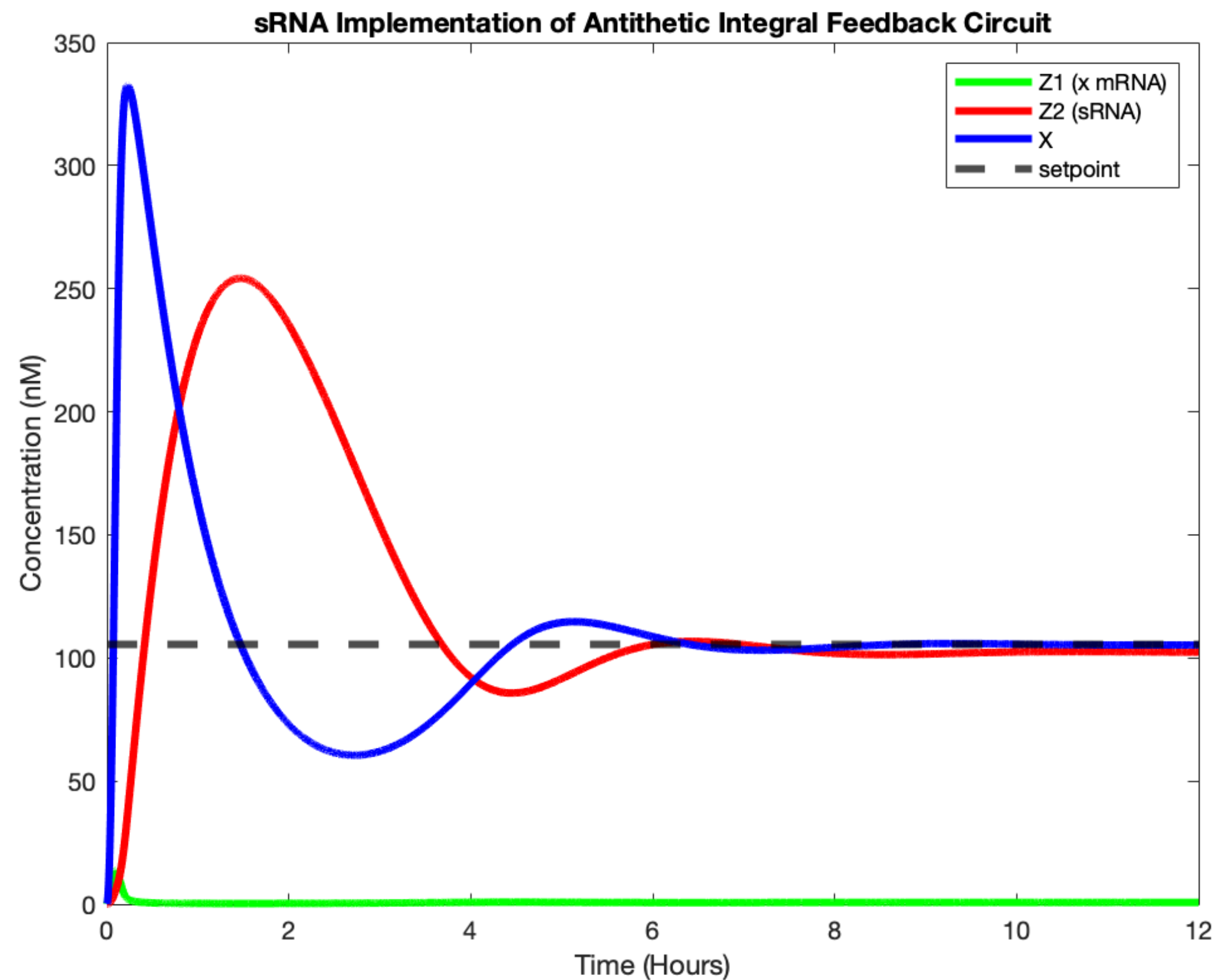
$$\frac{dZ_1}{dt} = \tau_1 + \frac{k_1 U}{K_U + U} - \gamma_m Z_1 - \theta Z_1 Z_2$$

$$\frac{dZ_2}{dt} = \tau_2 + \frac{k_3 X}{K_X + X} - \gamma_s Z_2 - \theta Z_1 Z_2$$

$$\frac{dX}{dt} = \tau_3 C + k_2 Z_1 - \delta X$$

$$\frac{dC}{dt} = \theta Z_1 Z_2 - \delta_C C$$

Without Dilution/Degradation



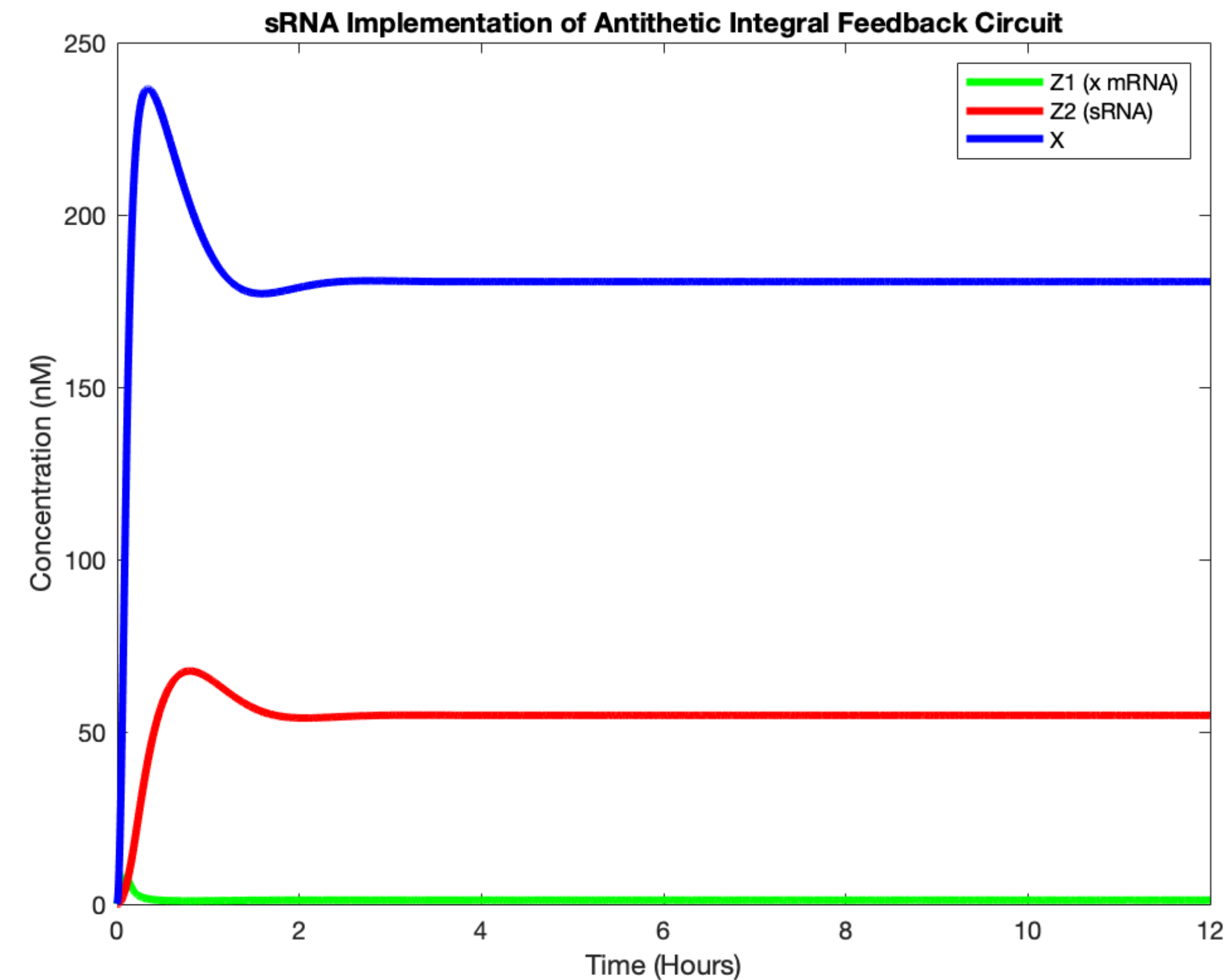
$$\tau_1 = 0; \tau_2 = 0; \tau_3 = 0; \gamma_m = 0; \gamma_s = 0$$

$$k_1 = 0.1 \text{ nMs}^{-1}; k_2 = 0.06 \text{ s}^{-1}; k_3 = 1.5 \text{ nMs}^{-1}$$

$$\theta = 0.05 \text{ nM}^{-1} \text{ s}^{-1}; \delta = 0.00039 \text{ s}^{-1}; \delta_C = 0.0041 \text{ s}^{-1}$$

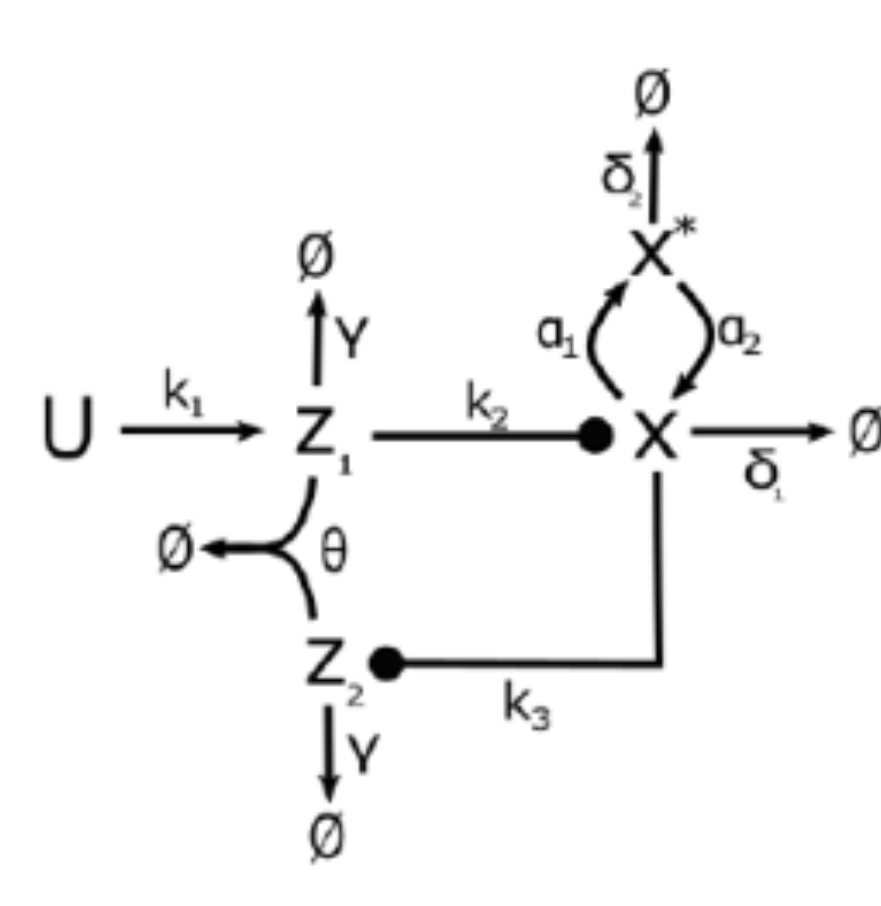
$$K_U = 178000 \text{ nM}; K_X = 2600 \text{ nM}$$

With Dilution/Degradation



$$\gamma_m = 0.0041 \text{ s}^{-1}; \gamma_s = 0.0008 \text{ s}^{-1}$$

Antithetic sRNA Implementation with Damping



$$\frac{dZ_1}{dt} = \tau_1 + \frac{k_1 U}{K_U + U} - \gamma_m Z_1 - \theta Z_1 Z_2$$

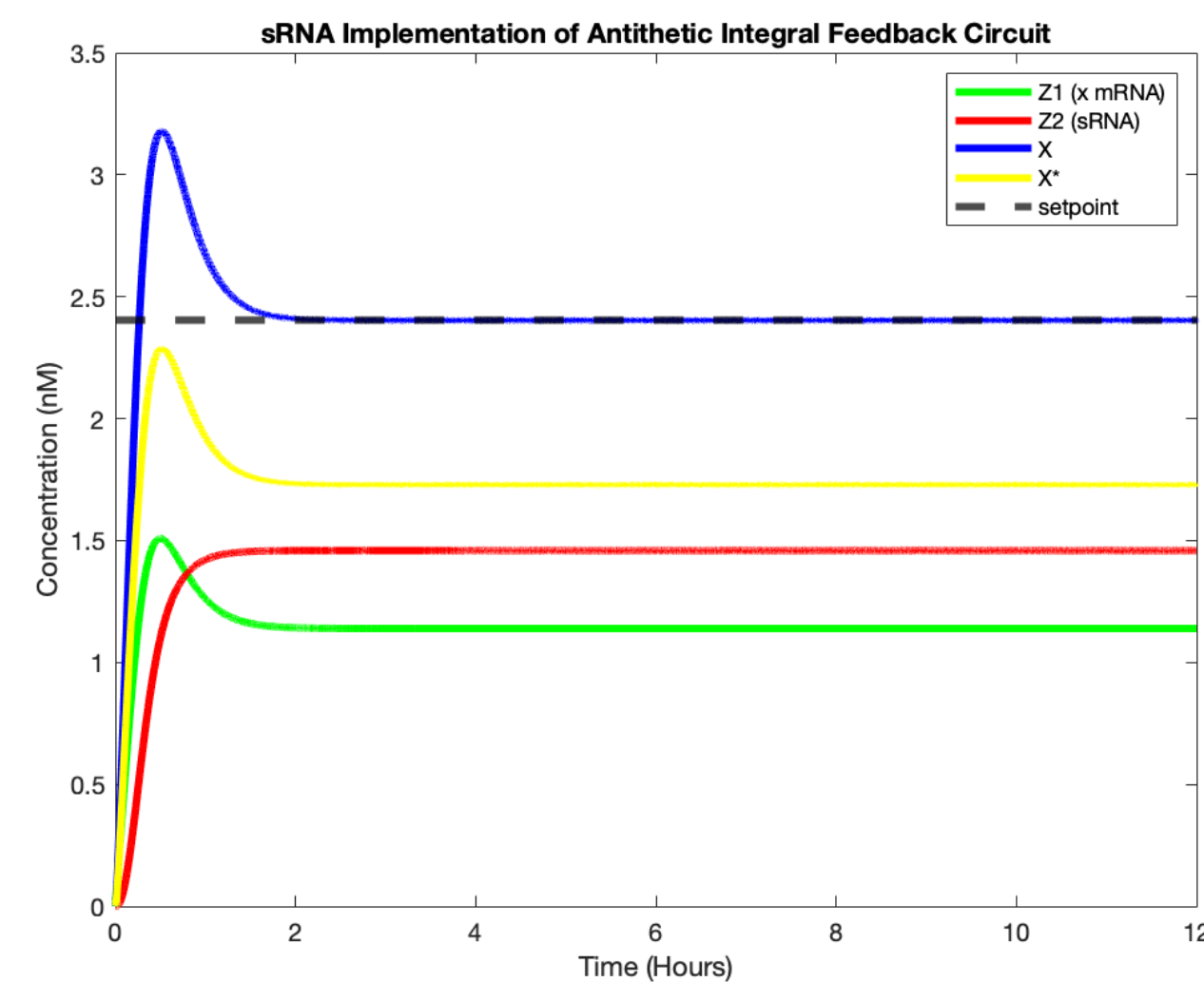
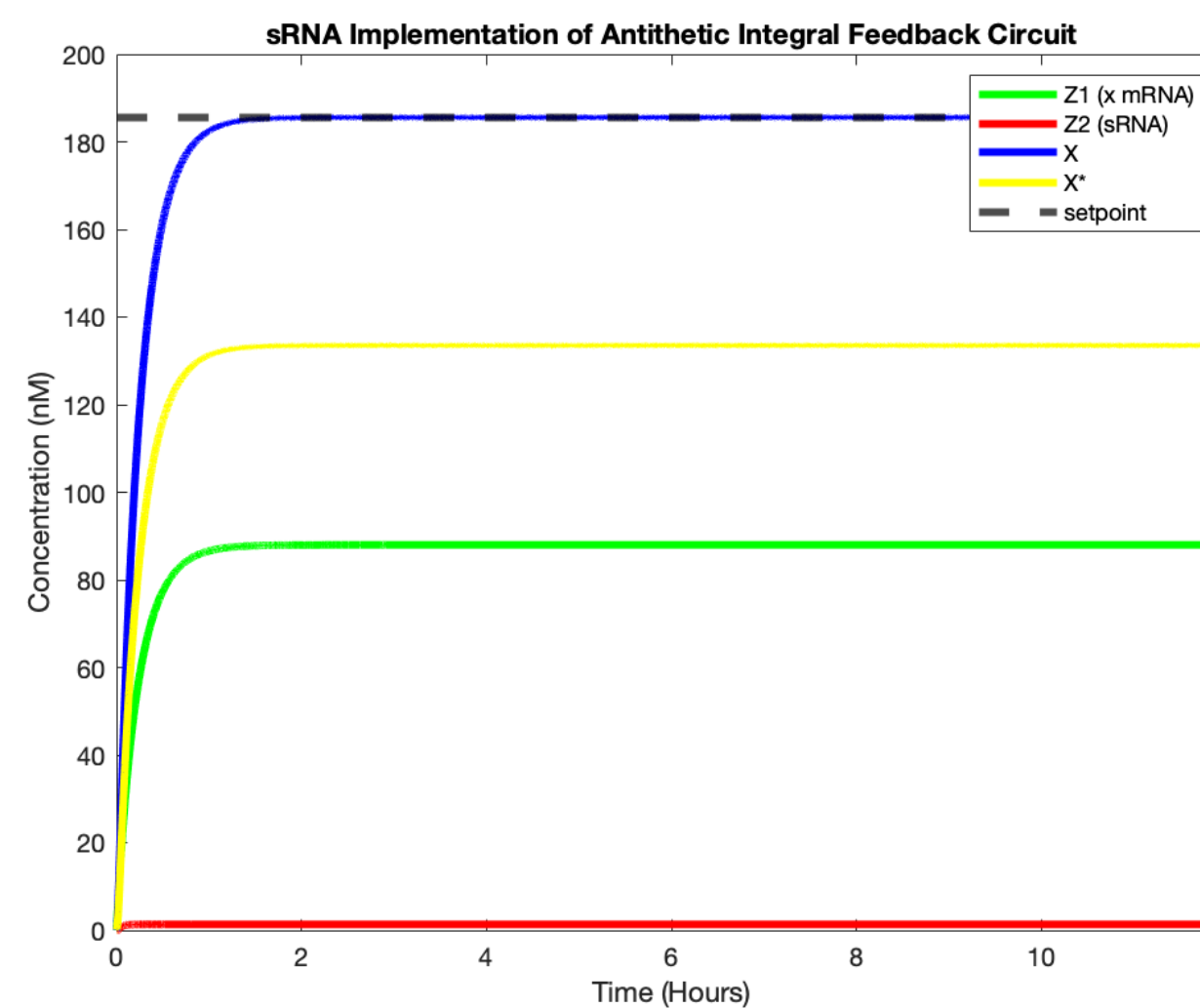
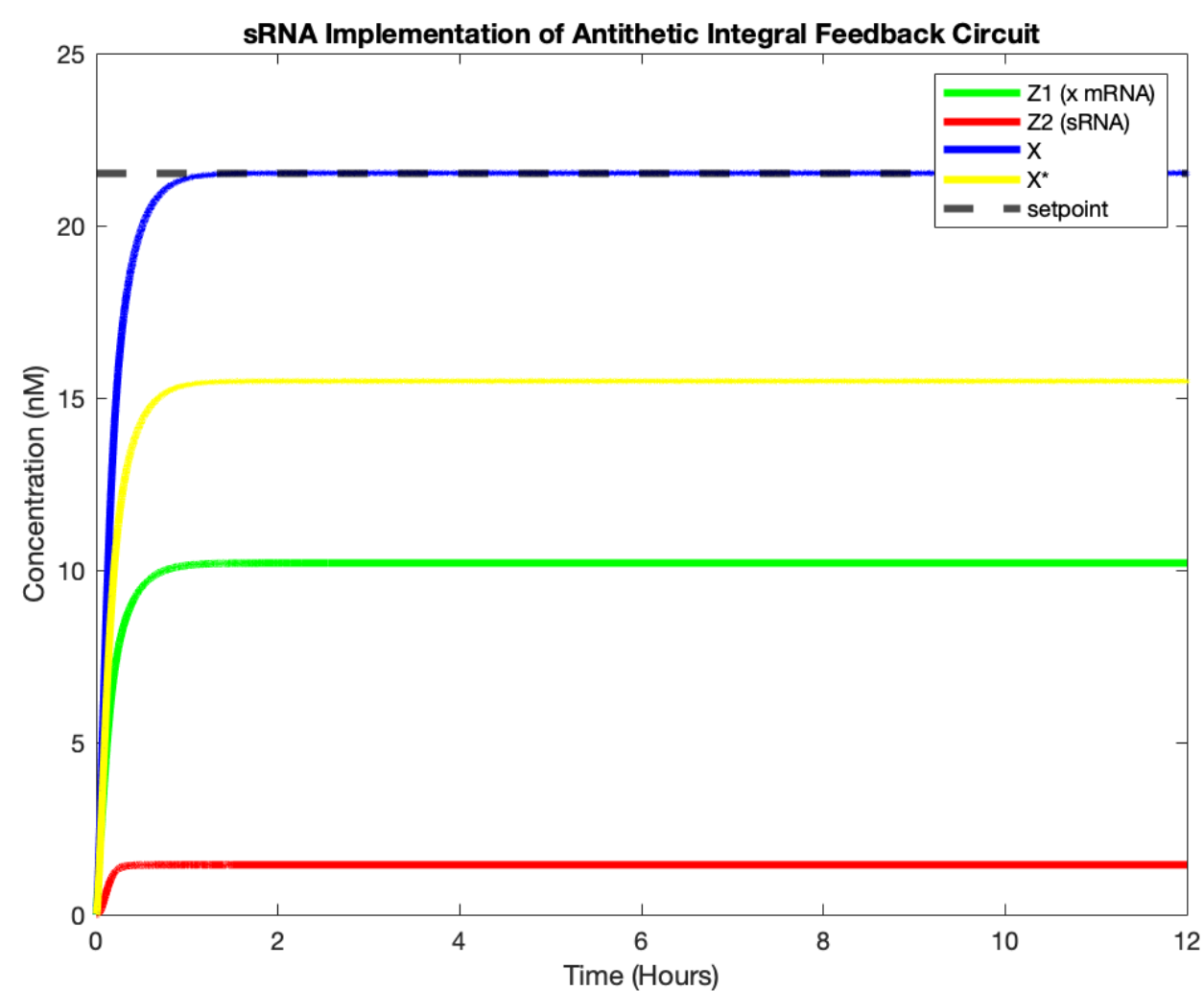
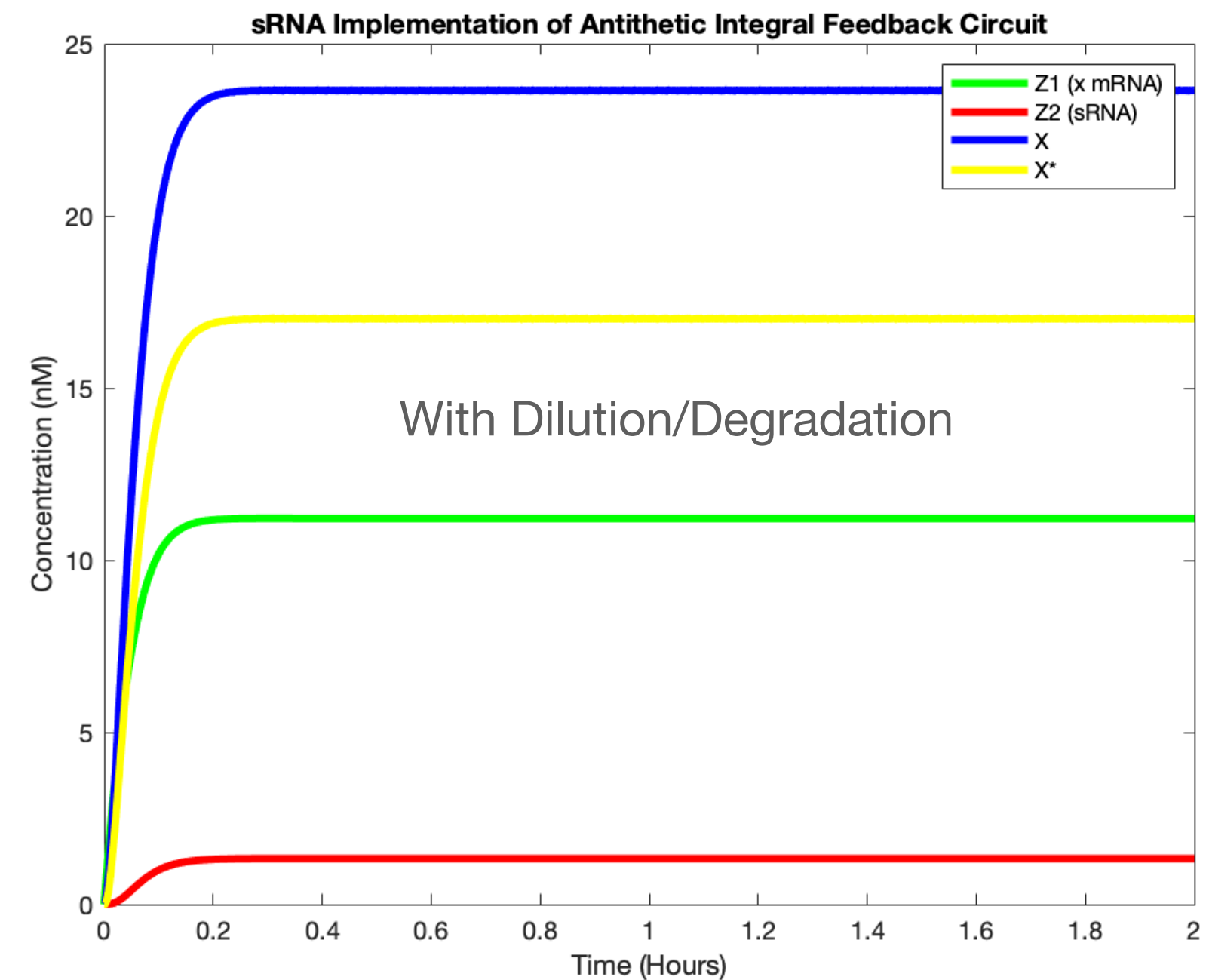
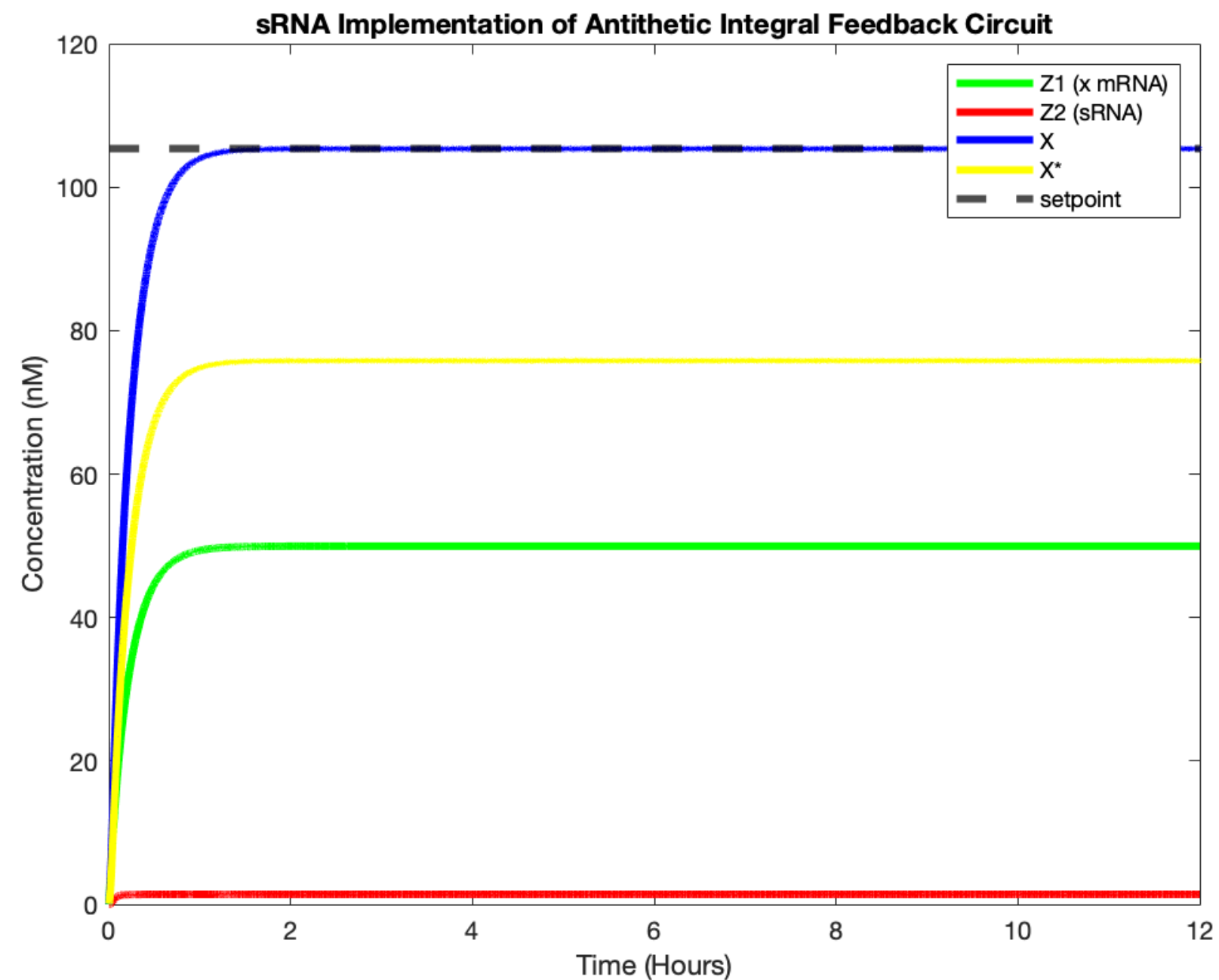
$$\frac{dZ_2}{dt} = \tau_2 + \frac{k_3 X}{K_X + X} - \gamma_s Z_2 - \theta Z_1 Z_2$$

$$\frac{dX}{dt} = \tau_3 C + k_2 Z_1 - \delta X - \alpha_1 X + \alpha_2 X^*$$

$$\frac{dC}{dt} = \theta Z_1 Z_2 - \delta_C C$$

$$\frac{dX^*}{dt} = \alpha_1 X - \alpha_2 X^* - \delta_* X^*$$

$$\alpha_1 = 0.1 \text{ s}^{-1}; \alpha_2 = 0.1 \text{ s}^{-1}; \delta = 0.00039 \text{ s}^{-1}; \delta_* = 0.039 \text{ s}^{-1}$$



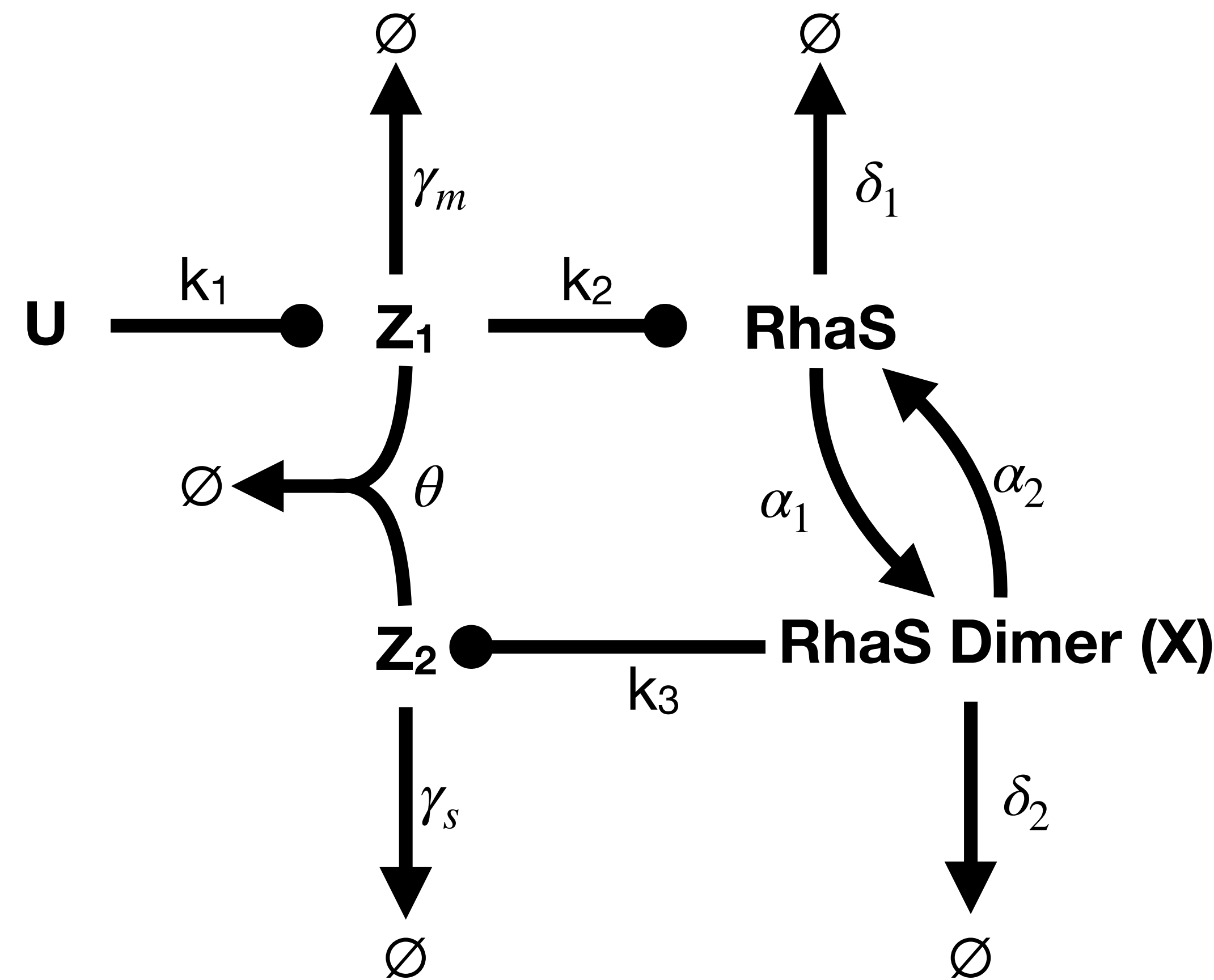
Can tune this to make it work by increasing alpha 1 but always stops working at some point and leads to more X* than X in the system

sRNA RhaS PI Controller

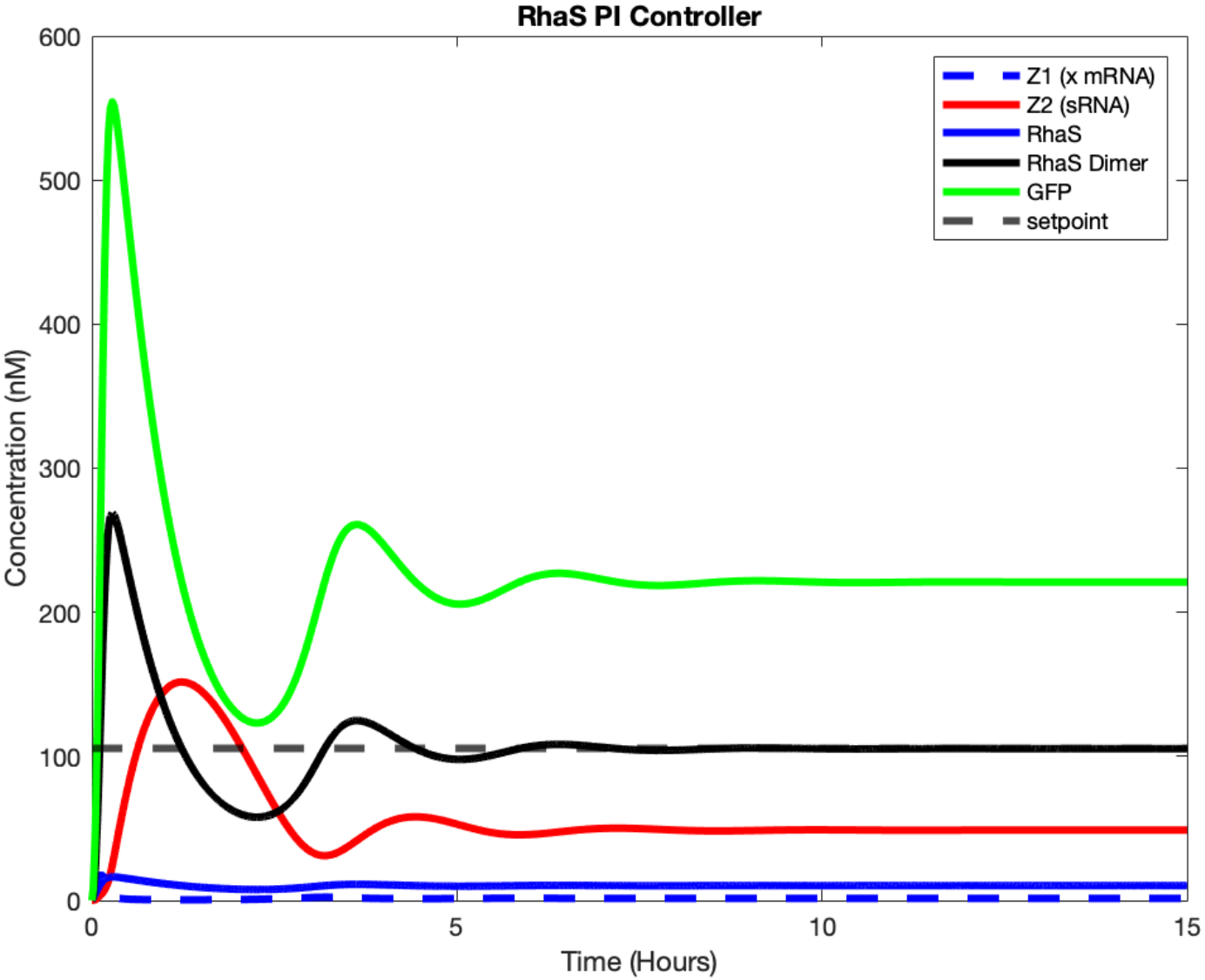
$$\frac{dZ_1}{dt} = \frac{k_1 U}{K_U + U} - \gamma_m Z_1 - \theta Z_1 Z_2 \qquad \frac{dZ_2}{dt} = \frac{k_3 X}{K_X + X} - \gamma_s Z_2 - \theta Z_1 Z_2$$

$$\frac{d\text{RhaS}}{dt} = k_2 Z_1 - \delta_1 \text{RhaS} - 2\alpha_1 \text{RhaS}^2 + 2\alpha_2 X$$

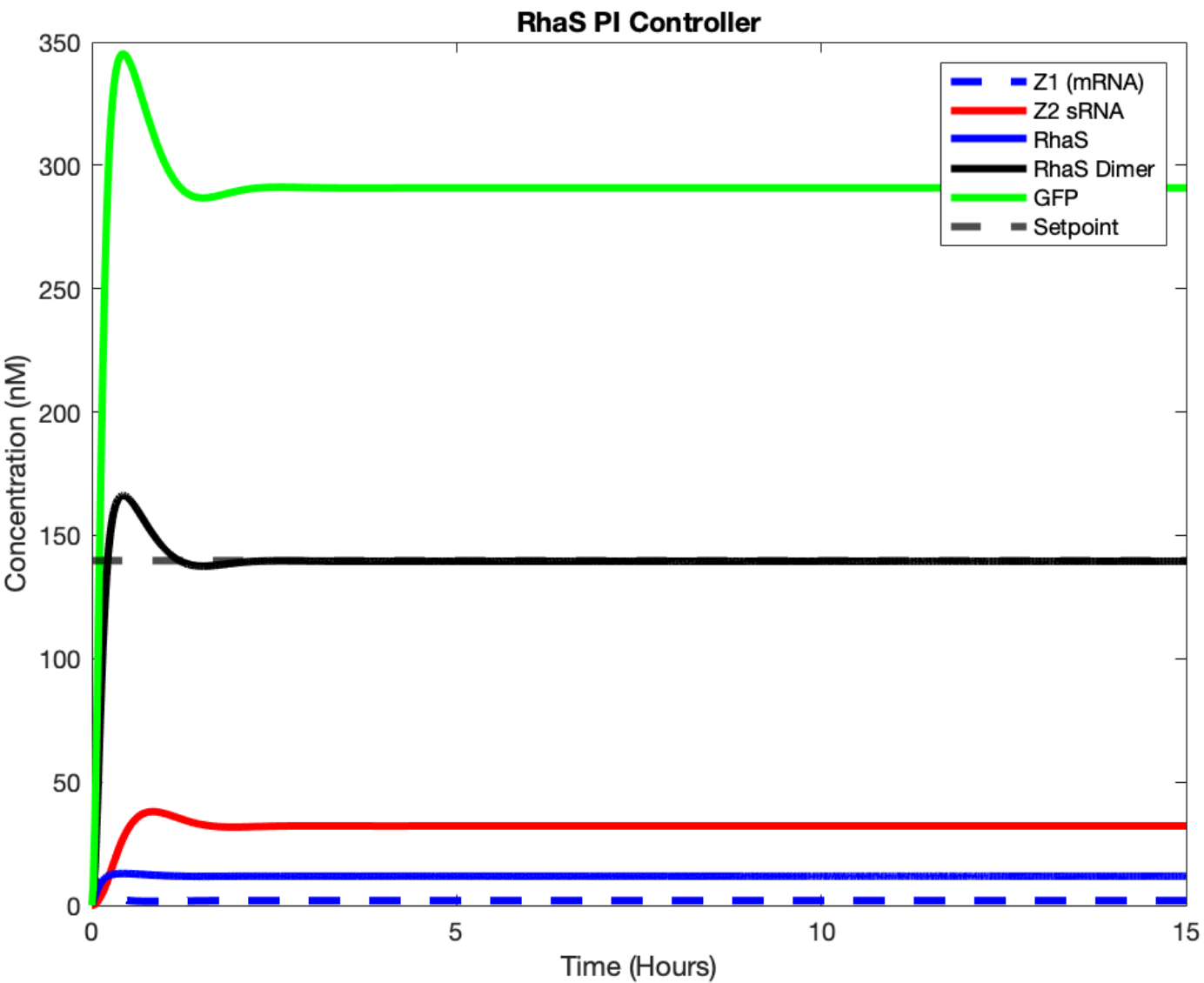
$$\frac{dX}{dt} = \alpha_1 \text{RhaS}^2 - \alpha_2 X - \delta_2 X$$



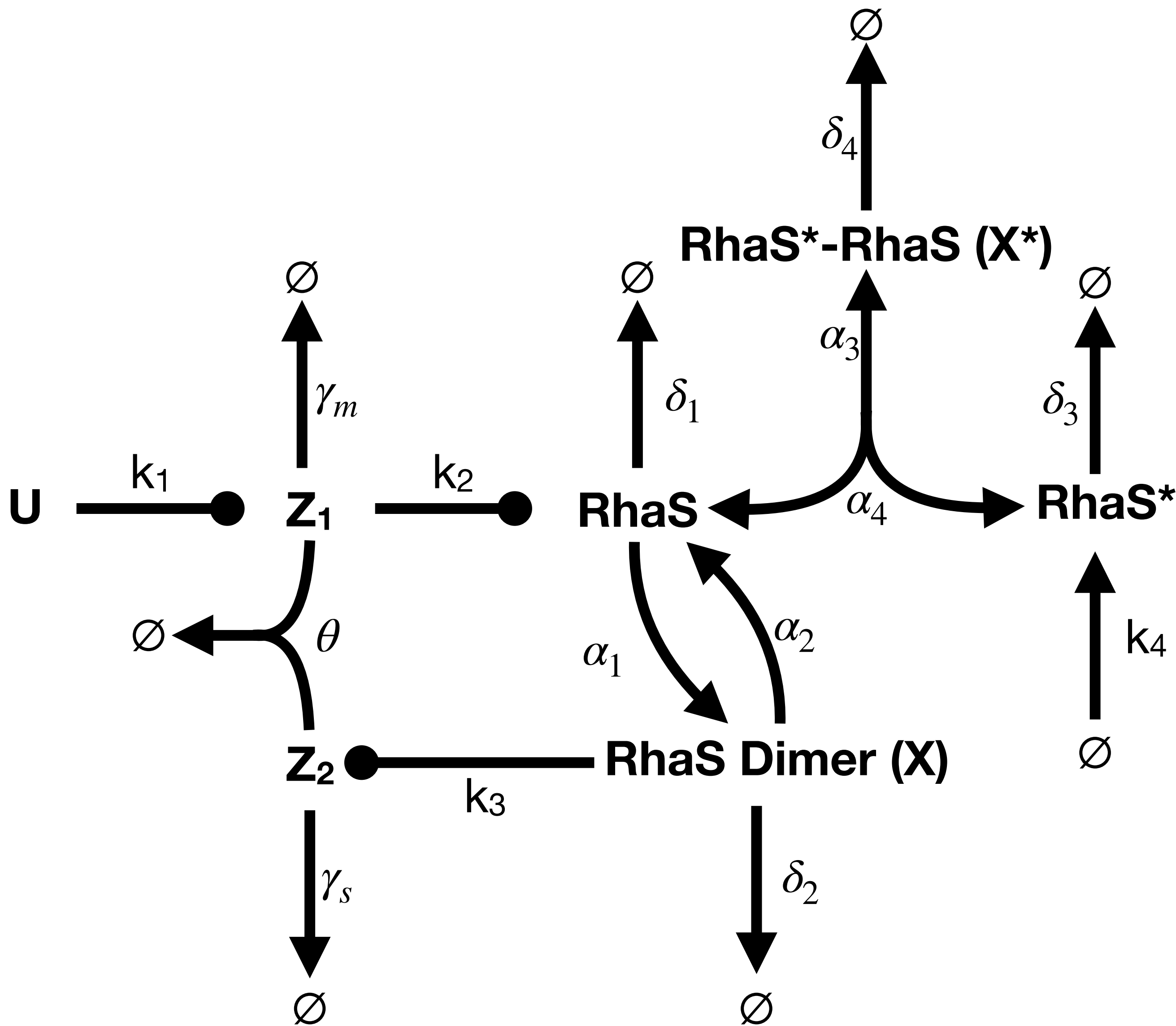
No Dilution/
degradation of Z1 and
Z2



Dilution/degradation
of Z1 and Z2



sRNA RhaS PID Controller



$$\frac{dZ_1}{dt} = \frac{k_1 U}{K_U + U} - \gamma_m Z_1 - \theta Z_1 Z_2$$

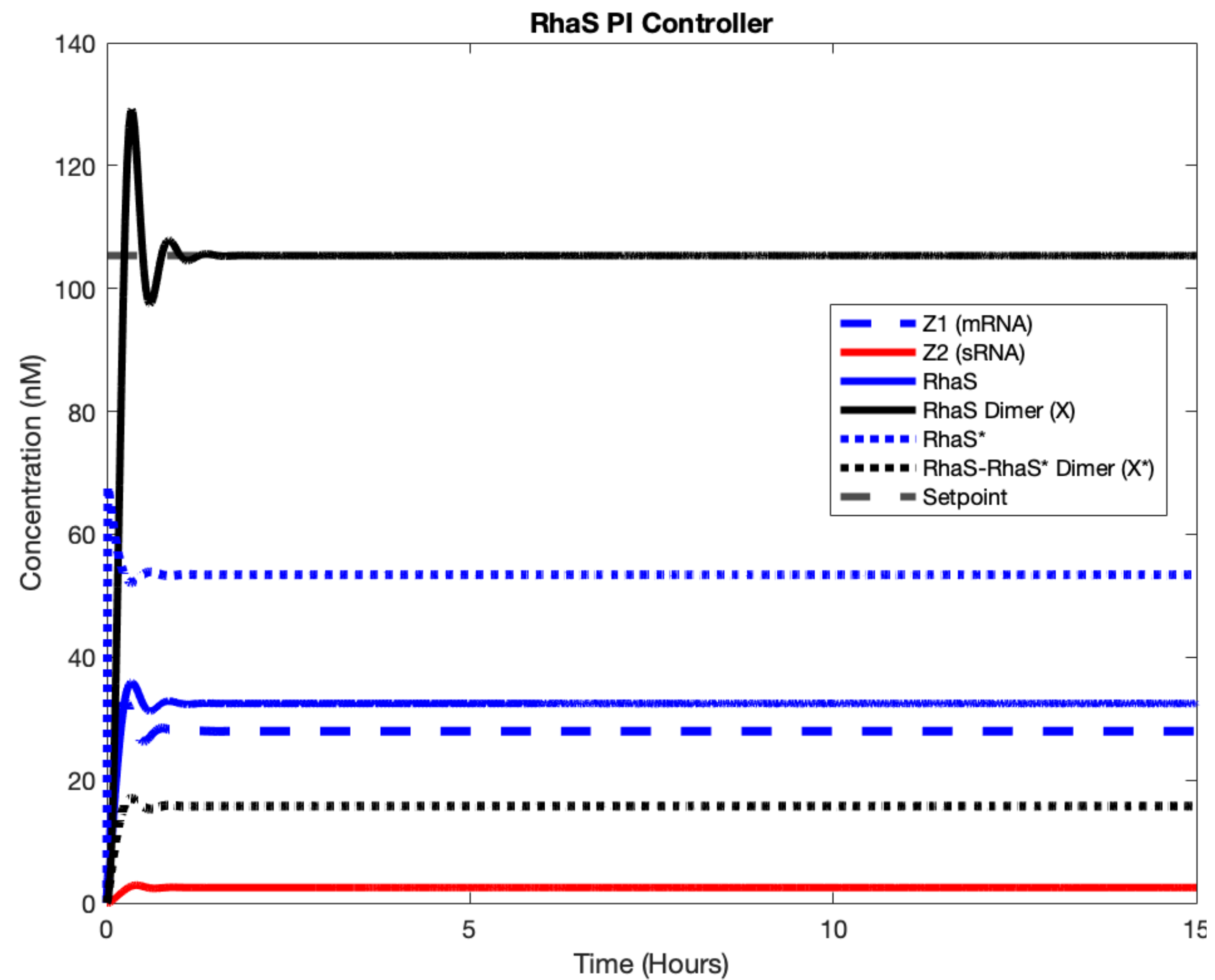
$$\frac{dZ_2}{dt} = \frac{k_3 X}{K_X + X} - \gamma_s Z_2 - \theta Z_1 Z_2$$

$$\frac{d\text{RhaS}}{dt} = k_2 Z_1 - \delta_1 \text{RhaS} - 2\alpha_1 \text{RhaS}^2 + 2\alpha_2 X - \alpha_3 \text{RhaS RhaS}^* + \alpha_4 X^*$$

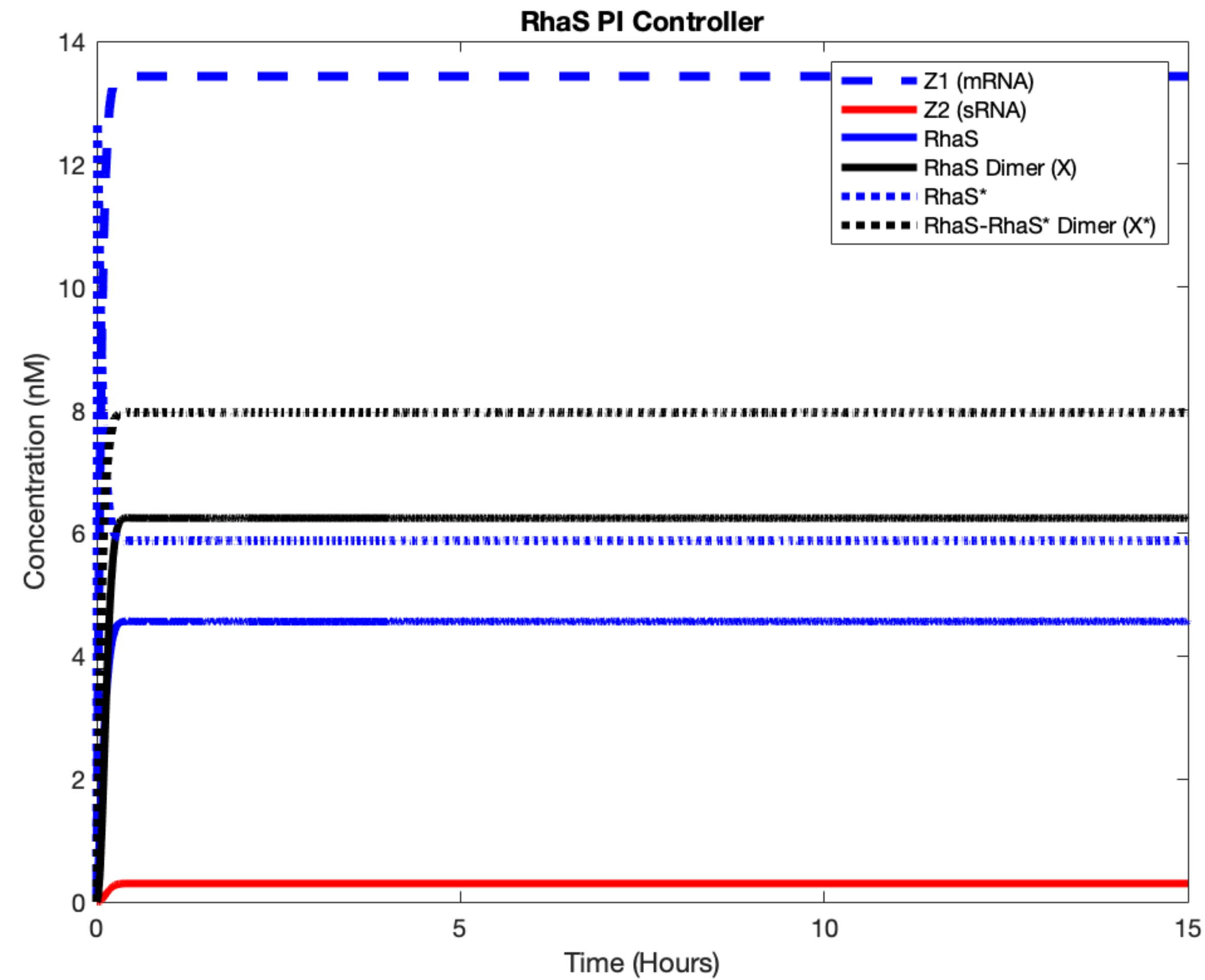
$$\frac{d\text{RhaS}^*}{dt} = k_4 - \delta_3 \text{RhaS}^* - \alpha_3 \text{RhaS RhaS}^* + \alpha_4 X^*$$

$$\frac{dX}{dt} = \alpha_1 \text{RhaS}^2 - \alpha_2 X - \delta_2 X$$

$$\frac{dX^*}{dt} = \alpha_3 \text{RhaS RhaS}^* - \alpha_4 X^* - \delta_4 X^*$$



Gammas = 0
Closest I got



Gammas non 0

Still a bit confused as to how degradation of the mixed dimer would work



Change this so some RhaS is released on degradation of the dimer

A Possible solution to the GFP problem is to have a TF being controlled by Z2 that produces the Z1 mRNA

And then have the buffering act upon that transcription factor rather than the output