

# Enzyme rhythms in model linear\_chain\_dilution - spontaneous oscillations

Model name: linear\_chain\_dilution

o Optimisation problem

- Protein turnover time  $1.8e+03$  s = 30 min
- Perturbed parameter(s) : x1
- Perturbation frequency  $f$  : 0.0167/s (period 60 s)
- Scored quantity: v4
- State-averaged fitness
- Posttranslational rhythms allowed
- Standard frequency considered  $f$  : 0.0167/s (period 60 s)

o Model properties:

- inactive\_enzymes: 0
- balanced\_reference\_state: 1
- consider\_external\_rhythm: 1
- adaptive\_rhythm: 0
- spontaneous\_rhythm: 1
- spontaneous\_rhythm\_at\_omega: 1
- has\_spontaneous\_rhythm\_and\_inactive\_enzymes: 0

o Beneficial autonomous oscillation found

- Maximum principal synergy found (in tested range) at frequency  $f = 0.0001$ /s (period  $1e+04$  s)
- Maximum fitness found (in tested range) at frequency  $f = 0.0001$ /s (period  $1e+04$  s)

o Fitness changes after external perturbation at frequency  $f = 0.0167$ /s

- Change by perturbation alone (xx):  $-3e-13$

o Autonomous oscillations?

- Maximally autonomous oscillations (in tested range) at  $f = 0.0001$ , principal synergy 0.0014
- Beneficial autonomous oscillations found at frequency  $f = 0.0167$ /s (principal synergy = 0.00139)
- Predicted fitness change (autonomous, 2nd order, amplitude below 1/2 of mean) at frequency  $f = 0.0167$ : 0.0863
- Predicted maximal fitness change (autonomous, numeric opt, full amplitude constraints) at frequency  $f = 0.0001$ : 0.726

WARNING: an external rhythm is given and an autonomous rhythm has been found

o Numerical calculation (responsive,  $f = 0.0167$ )

- Fitness change (fitness-averaged):  $-3.73e-05$
- Fitness change (state-averaged):  $-1.29e-05$

o Numerical calculation (autonomous rhythm, amplitude below 1/2 of mean,  $f = 0.0167$ )

- Fitness change (fitness-averaged) :  $2.04e-06$
- Fitness change (state-averaged):  $2.28e-06$

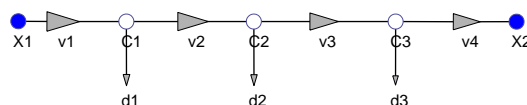


Figure 1: Network and reference flux

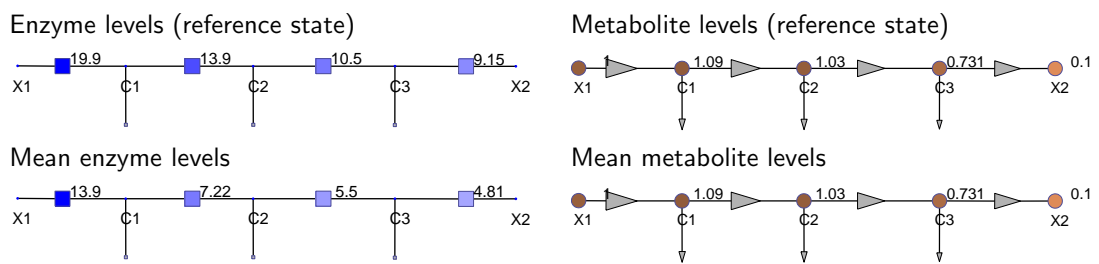


Figure 2: Reference state (top) and mean state during oscillation (bottom).

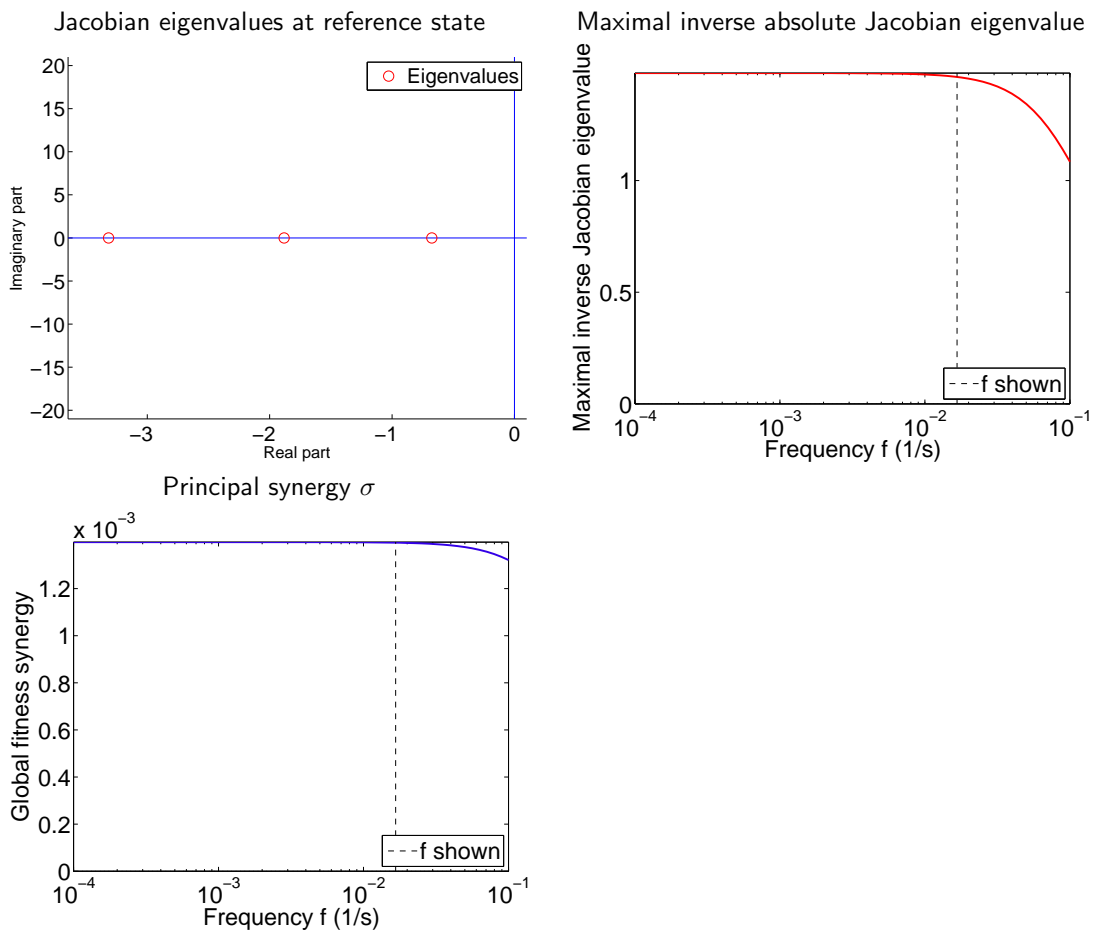
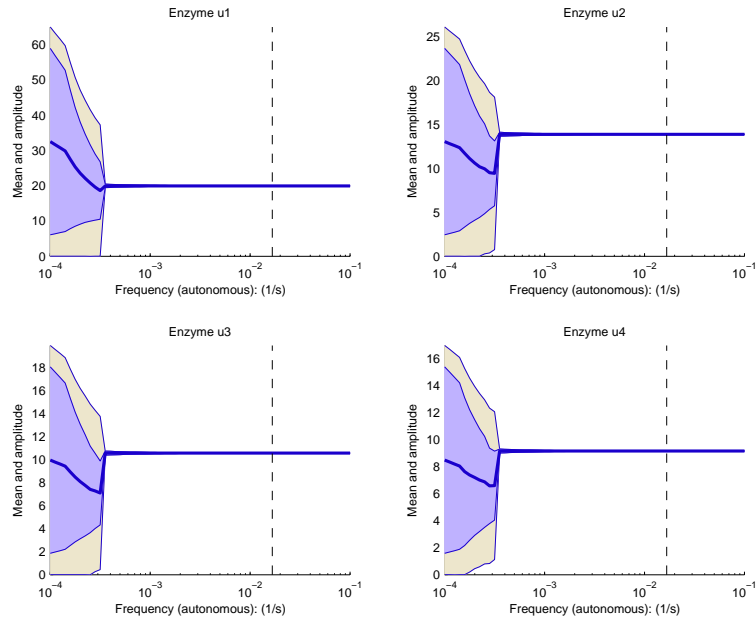
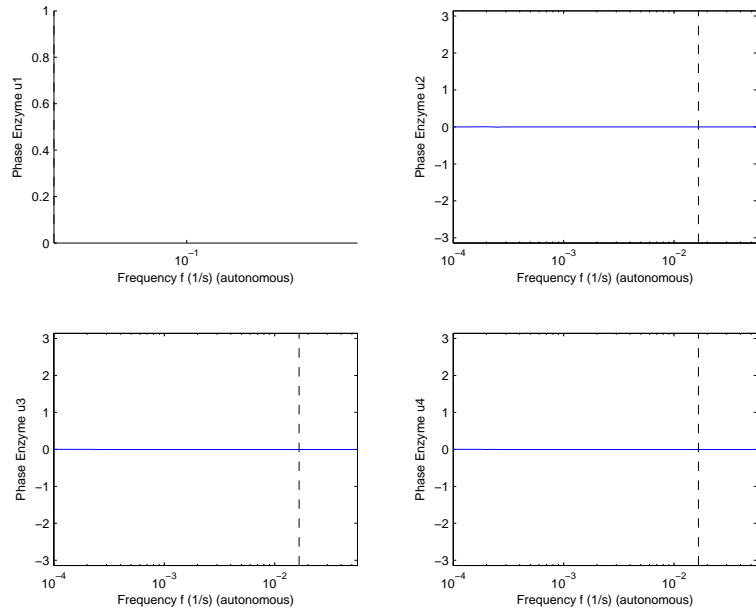


Figure 3: Control analysis: fitness curvatures. Left: Frequency-dependent fitness curvature eigenvalues. Right: relative sizes and phases of the individual enzyme levels (components of the leading fitness curvature eigenvector).

### Protein level and enzyme activity (mean and amplitude)



### Phase angles $[0, 2\pi]$



### Fitness change

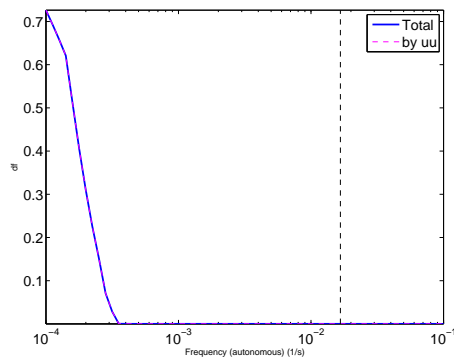


Figure 4: Self-promoting oscillations. Left: amplitudes of protein levels (blue) and modification (grey). Right: phase shifts.

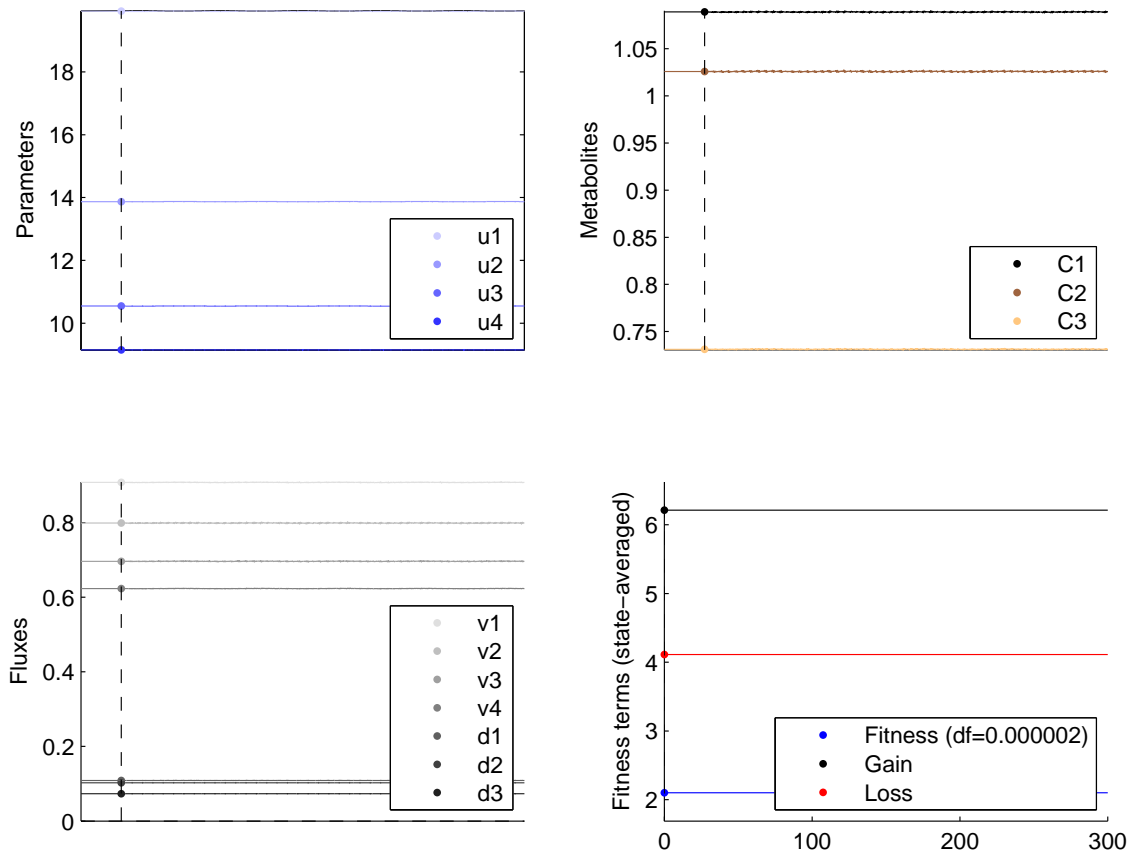


Figure 5: Numerical calculations: spontaneous oscillations. Perturbation frequency see first page.

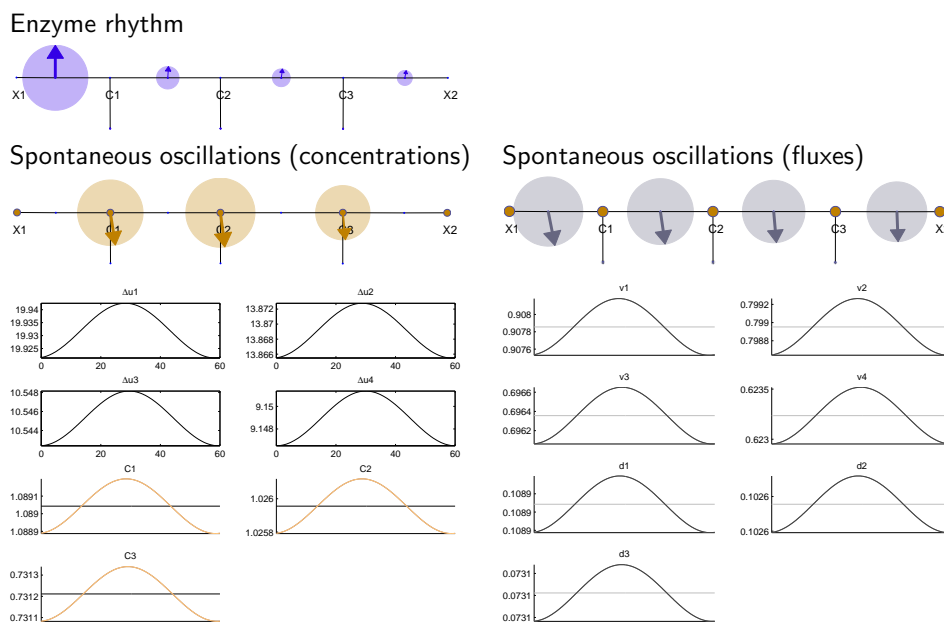


Figure 6: Spontaneous oscillations (local expansion; arrows: absolute changes). Perturbation frequency see first page.

Spontaneous oscillations

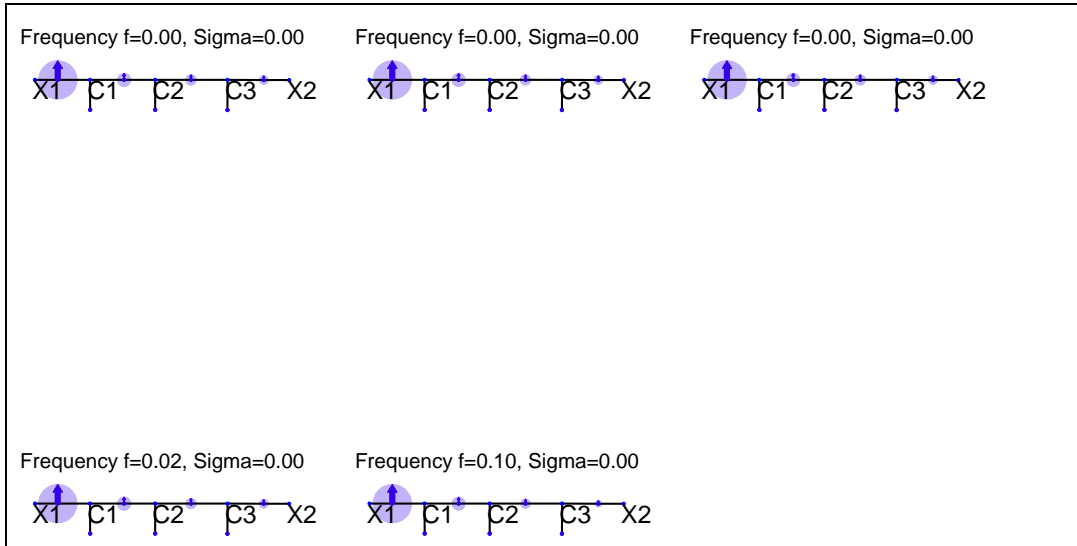
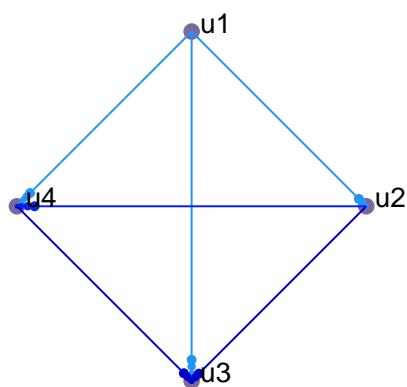
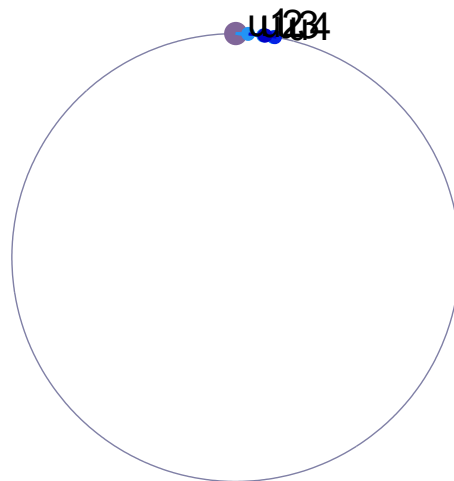


Figure 7: Spontaneous oscillations (or tendencies towards them) for various circular frequencies  $\omega$ . If the maximal fitness curvatures  $\lambda$  is positive, the rhythm is beneficial (local expansion; arrows: absolute changes).

Synergies



Synergy phase plot



Synergies on network

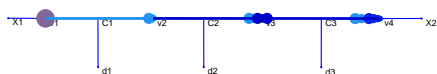


Figure 8: Periodic economic potentials and direct enzyme values.