

## **Additional file 2**

# **Dynamic modeling of yeast meiotic initiation**

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**Table S1 - The *rim11* knockout model**

$$(2) \frac{dpUme6}{dt} = -u_{ume6} \cdot pUme6$$

$$(3) \frac{dpSok2}{dt} = p_{sok2} \cdot \frac{c_{sok2}}{c_{sok2} + Ime1} \cdot (1 - pSok2) - u_{sok2} \cdot pSok2$$

$$(4) \frac{dIme1}{dt} = s_{ime1} \cdot \frac{c_{ime1}}{c_{ime1} + pSok2} - d_{ime1} \cdot Ime1 - d_{ime1} \cdot Ime2 \cdot \frac{Ime1}{c_1 + Ime1}$$

$$(5) \frac{dpIme1}{dt} = -d_{pime1} \cdot pIme1$$

$$(6) \frac{dIme2}{dt} = s_{ime2} \cdot pUme6 \cdot pIme1 + s_{ime2} \cdot \frac{Ime2^5}{c_2^5 + Ime2^5} - d_{ime2} \cdot \frac{Ime2}{c_3 + Ime2}$$

**Table S2 - The *ume6* knockout model**

$$\begin{aligned}
 (1) \quad & \frac{dRim11}{dt} = u_{rim11} \cdot (1 - Rim11) - p_{rim11} \cdot Rim11 \\
 (3) \quad & \frac{dpSok2}{dt} = p_{sok2} \cdot \frac{c_{sok2}}{c_{sok2} + Ime1} \cdot (1 - pSok2) - u_{sok2} \cdot pSok2 \\
 (4) \quad & \frac{dIme1}{dt} = s_{ime1} \cdot \frac{c_{ime1}}{c_{ime1} + pSok2} - p_{ime1} \cdot Rim11 \cdot Ime1 - d_{ime1} \cdot Ime1 - d_{ime1} \cdot Ime2 \cdot \frac{Ime1}{c_1 + Ime1} \\
 (5) \quad & \frac{dpIme1}{dt} = p_{ime1} \cdot Rim11 \cdot Ime1 - d_{pime1} \cdot pIme1 \\
 (6) \quad & \frac{dIme2}{dt} = s_{ime2} \cdot \frac{Ime2^5}{c_2^5 + Ime2^5} - d_{ime2} \cdot \frac{Ime2}{c_3 + Ime2}
 \end{aligned}$$

**Table S3 - The *sok2* knockout model**

- (1)  $\frac{dRim11}{dt} = u_{rim11} \cdot (1 - Rim11) - p_{rim11} \cdot Rim11$
- (2)  $\frac{dpUme6}{dt} = p_{ume6} \cdot Rim11 \cdot (1 - pUme6) - u_{ume6} \cdot pUme6$
- (4)  $\frac{dIme1}{dt} = s_{ime1} - p_{ime1} \cdot Rim11 \cdot Ime1 - d_{ime1} \cdot Ime1 - d_{ime1} \cdot Ime2 \cdot \frac{Ime1}{c_1 + Ime1}$
- (5)  $\frac{dpIme1}{dt} = p_{ime1} \cdot Rim11 \cdot Ime1 - d_{pime1} \cdot pIme1$
- (6)  $\frac{dIme2}{dt} = s_{ime2} \cdot pUme6 \cdot pIme1 + s_{ime2} \cdot \frac{Ime2^5}{c_2^5 + Ime2^5} - d_{ime2} \cdot \frac{Ime2}{c_3 + Ime2}$

**Table S4 - The *ime1* knockout model**

$$\begin{aligned} (1) \quad & \frac{dRim11}{dt} = u_{rim11} \cdot (1 - Rim11) - p_{rim11} \cdot Rim11 \\ (2) \quad & \frac{dpUme6}{dt} = p_{ume6} \cdot Rim11 \cdot (1 - pUme6) - u_{ume6} \cdot pUme6 \\ (3) \quad & \frac{dpSok2}{dt} = p_{sok2} \cdot (1 - pSok2) - u_{sok2} \cdot pSok2 \\ (6) \quad & \frac{dIme2}{dt} = s_{ime2} \cdot \frac{Ime2^5}{c_2^5 + Ime2^5} - d_{ime2} \cdot \frac{Ime2}{c_3 + Ime2} \end{aligned}$$

**Table S5 - The *ime2* knockout model**

- (1)  $\frac{dRim11}{dt} = u_{rim11} \cdot (1 - Rim11) - p_{rim11} \cdot Rim11$
- (2)  $\frac{dpUme6}{dt} = p_{ume6} \cdot Rim11 \cdot (1 - pUme6) - u_{ume6} \cdot pUme6$
- (3)  $\frac{dpSok2}{dt} = p_{sok2} \cdot \frac{c_{sok2}}{c_{sok2} + Ime1} \cdot (1 - pSok2) - u_{sok2} \cdot pSok2$
- (4)  $\frac{dIme1}{dt} = s_{ime1} \cdot \frac{c_{ime1}}{c_{ime1} + pSok2} - p_{ime1} \cdot Rim11 \cdot Ime1 - d_{ime1} \cdot Ime1$
- (5)  $\frac{dpIme1}{dt} = p_{ime1} \cdot Rim11 \cdot Ime1 - d_{pime1} \cdot pIme1$

**Table S6 - The pSok2-Ime1 feedback knockout model**

- (1)  $\frac{dRim11}{dt} = u_{rim11} \cdot (1 - Rim11) - p_{rim11} \cdot Rim11$
- (2)  $\frac{dpUme6}{dt} = p_{ume6} \cdot Rim11 \cdot (1 - pUme6) - u_{ume6} \cdot pUme6$
- (3)  $\frac{dpSok2}{dt} = p_{sok2} \cdot (1 - pSok2) - u_{sok2} \cdot pSok2$
- (4)  $\frac{dIme1}{dt} = s_{ime1} - p_{ime1} \cdot Rim11 \cdot Ime1 - d_{ime1} \cdot Ime1 - d_{ime1} \cdot Ime2 \cdot \frac{Ime1}{c_1 + Ime1}$
- (5)  $\frac{dpIme1}{dt} = p_{ime1} \cdot Rim11 \cdot Ime1 - d_{pime1} \cdot pIme1$
- (6)  $\frac{dIme2}{dt} = s_{ime2} \cdot pUme6 \cdot pIme1 + s_{ime2} \cdot \frac{Ime2^5}{c_2^5 + Ime2^5} - d_{ime2} \cdot \frac{Ime2}{c_3 + Ime2}$

**Table S7 - The Ime2-Ime1 feedback knockout model**

$$\begin{aligned}
 (1) \quad & \frac{dRim11}{dt} = u_{rim11} \cdot (1 - Rim11) - p_{rim11} \cdot Rim11 \\
 (2) \quad & \frac{dpUme6}{dt} = p_{ume6} \cdot Rim11 \cdot (1 - pUme6) - u_{ume6} \cdot pUme6 \\
 (3) \quad & \frac{dpSok2}{dt} = p_{sok2} \cdot \frac{c_{sok2}}{c_{sok2} + Ime1} \cdot (1 - pSok2) - u_{sok2} \cdot pSok2 \\
 (4) \quad & \frac{dIme1}{dt} = s_{ime1} \cdot \frac{c_{ime1}}{c_{ime1} + pSok2} - p_{ime1} \cdot Rim11 \cdot Ime1 - d_{ime1} \cdot Ime1 \\
 (5) \quad & \frac{dpIme1}{dt} = p_{ime1} \cdot Rim11 \cdot Ime1 - d_{pime1} \cdot pIme1 \\
 (6) \quad & \frac{dIme2}{dt} = s_{ime2} \cdot pUme6 \cdot pIme1 + s_{ime2} \cdot \frac{Ime2^5}{c_2^5 + Ime2^5} - d_{ime2} \cdot \frac{Ime2}{c_3 + Ime2}
 \end{aligned}$$



**Table S8 - The Ime2 feedback knockout model**

$$\begin{aligned}
 (1) \quad & \frac{dRim11}{dt} = u_{rim11} \cdot (1 - Rim11) - p_{rim11} \cdot Rim11 \\
 (2) \quad & \frac{dpUme6}{dt} = p_{ume6} \cdot Rim11 \cdot (1 - pUme6) - u_{ume6} \cdot pUme6 \\
 (3) \quad & \frac{dpSok2}{dt} = p_{sok2} \cdot \frac{c_{sok2}}{c_{sok2} + Ime1} \cdot (1 - pSok2) - u_{sok2} \cdot pSok2 \\
 (4) \quad & \frac{dIme1}{dt} = s_{ime1} \cdot \frac{c_{ime1}}{c_{ime1} + pSok2} - p_{ime1} \cdot Rim11 \cdot Ime1 - d_{ime1} \cdot Ime1 - d_{ime1} \cdot Ime2 \cdot \frac{Ime1}{c_1 + Ime1} \\
 (5) \quad & \frac{dpIme1}{dt} = p_{ime1} \cdot Rim11 \cdot Ime1 - d_{pime1} \cdot pIme1 \\
 (6) \quad & \frac{dIme2}{dt} = s_{ime2} \cdot pUme6 \cdot pIme1 - d_{ime2} \cdot \frac{Ime2}{c_3 + Ime2}
 \end{aligned}$$

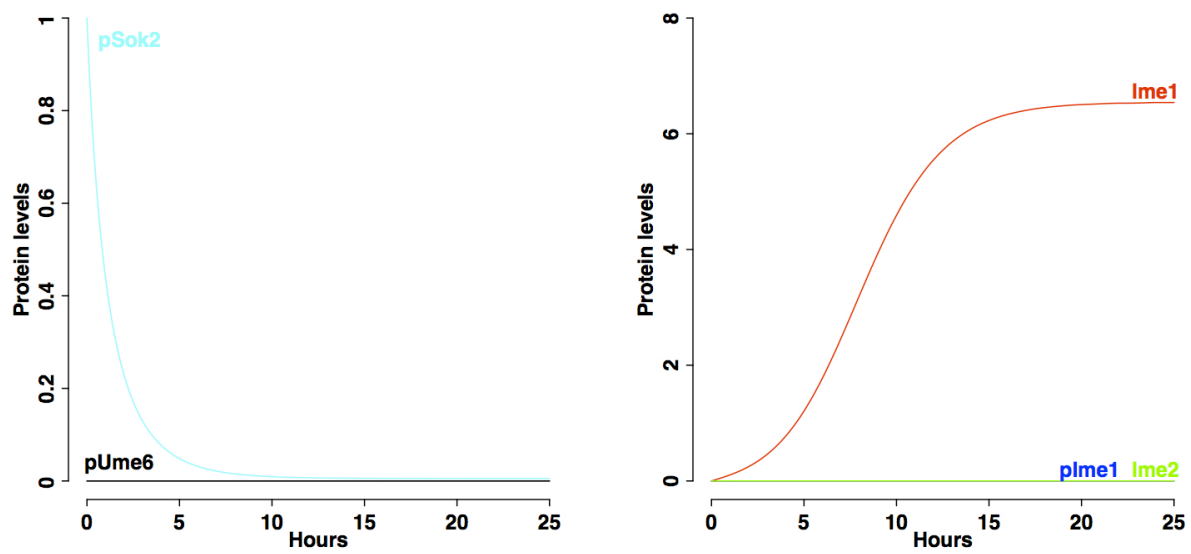


Figure S1 - Numerical simulations of the *rim11* knockout model

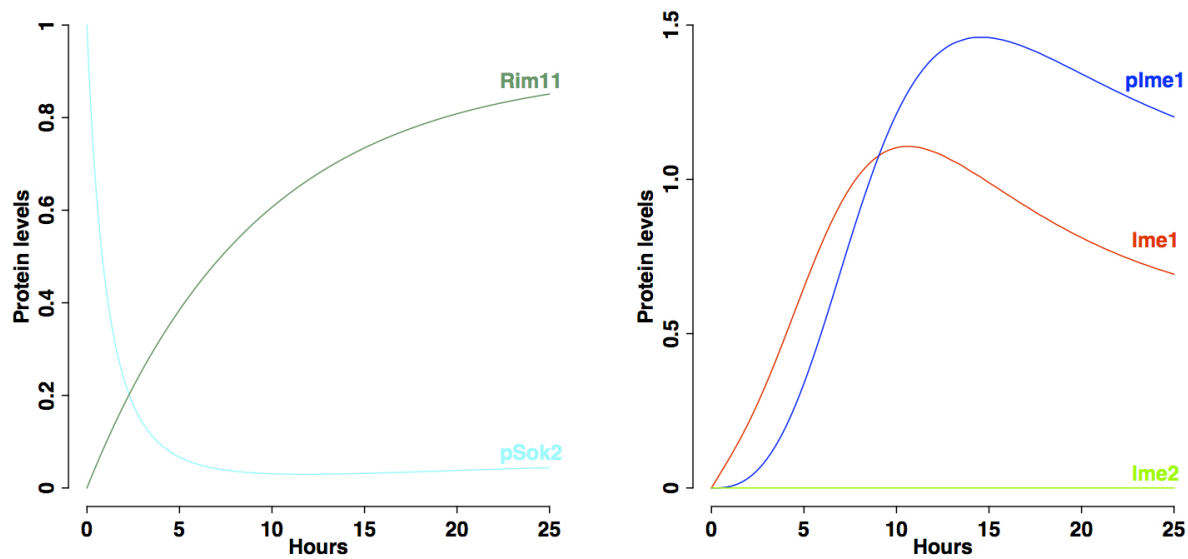
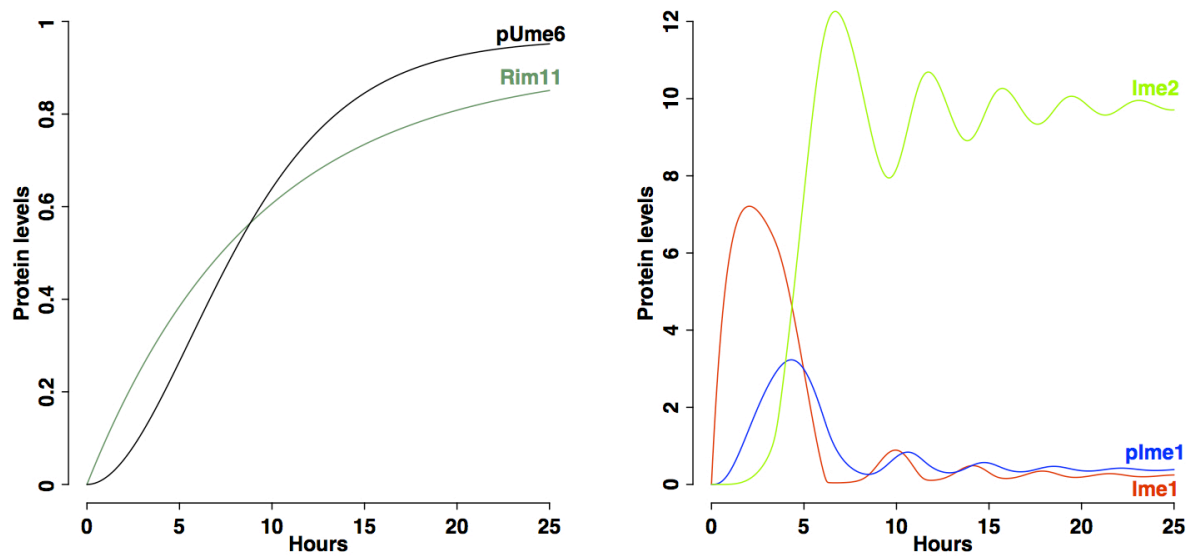


Figure S2 - Numerical simulations of the *ume6* knockout model



**Figure S3 - Numerical simulations of the sok2 knockout model**

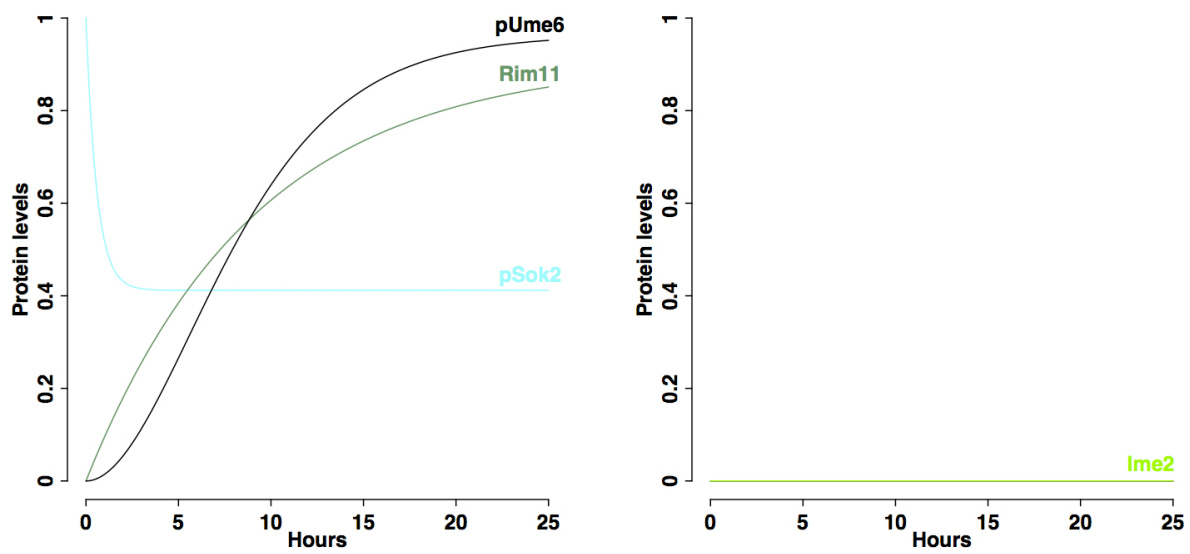


Figure S4 - Numerical simulations of the *ime1* knockout model

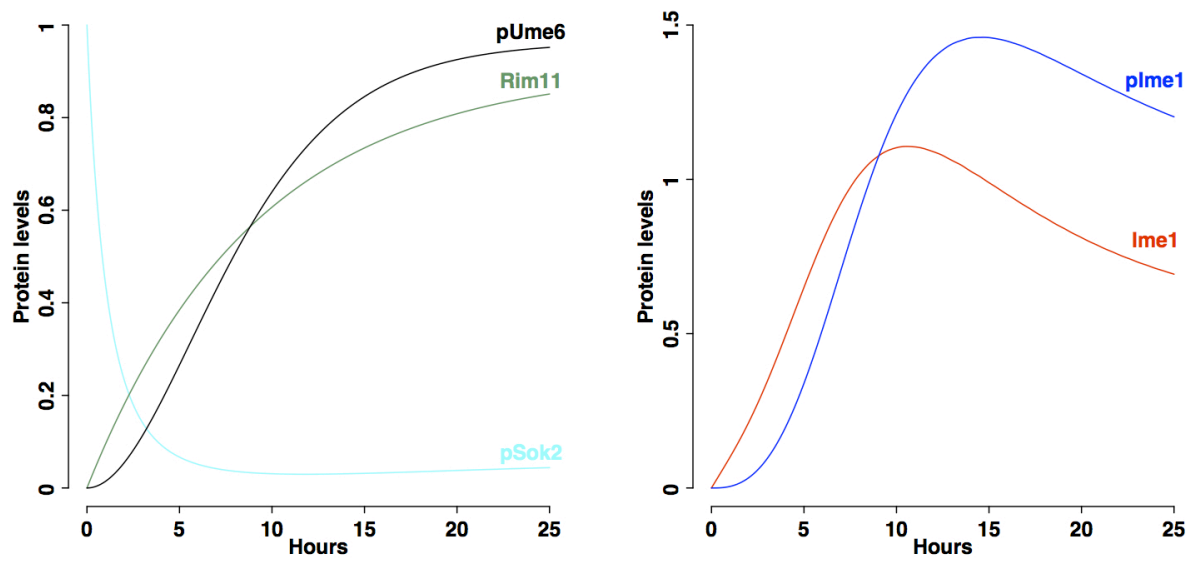
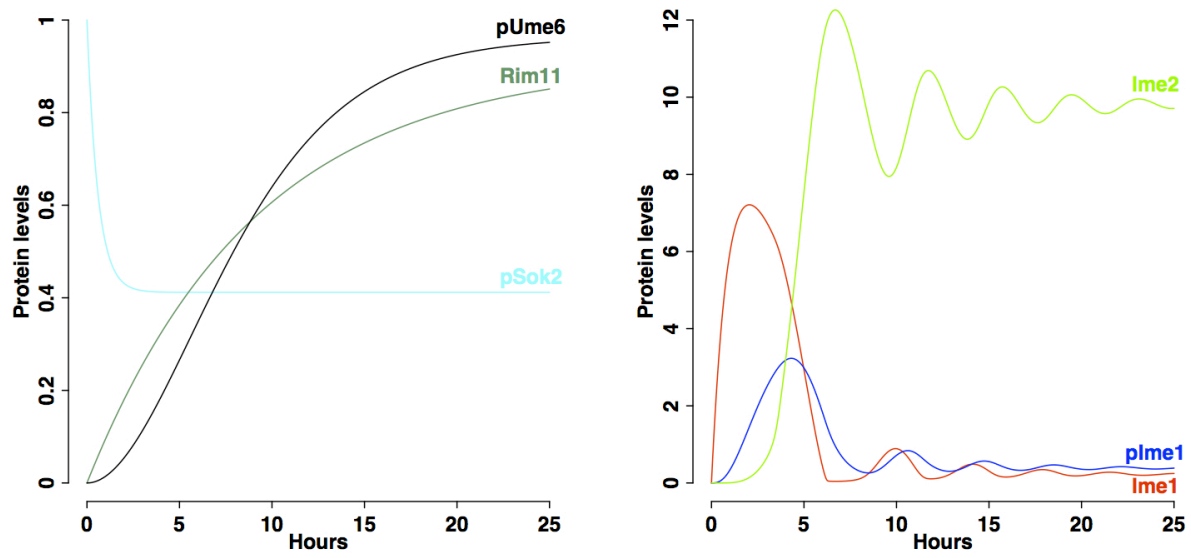
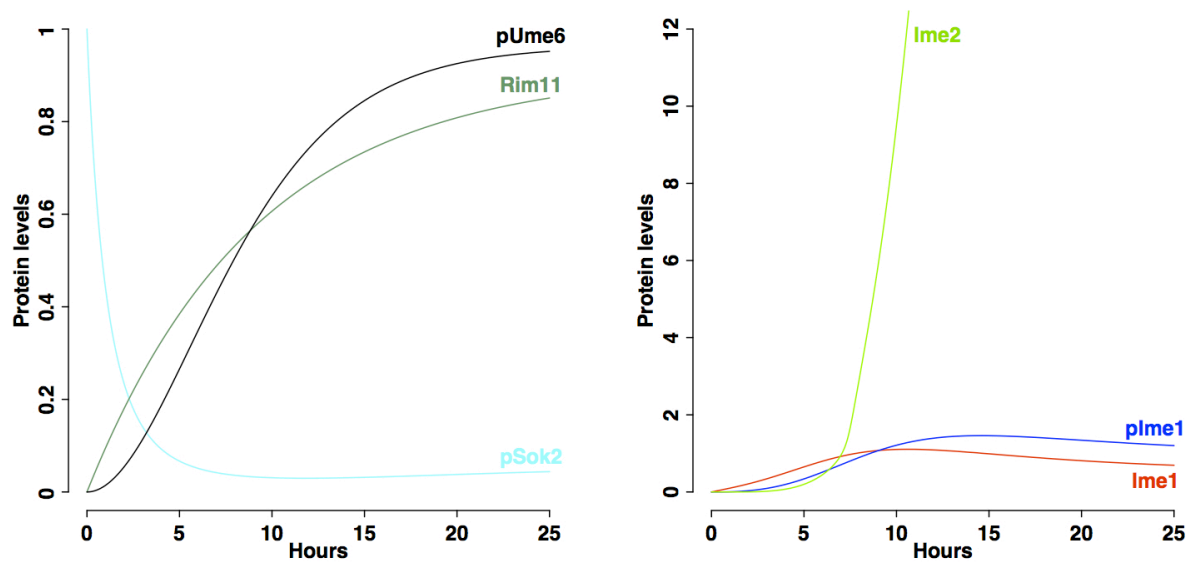


Figure S5 - Numerical simulations of the *ime2* knockout model

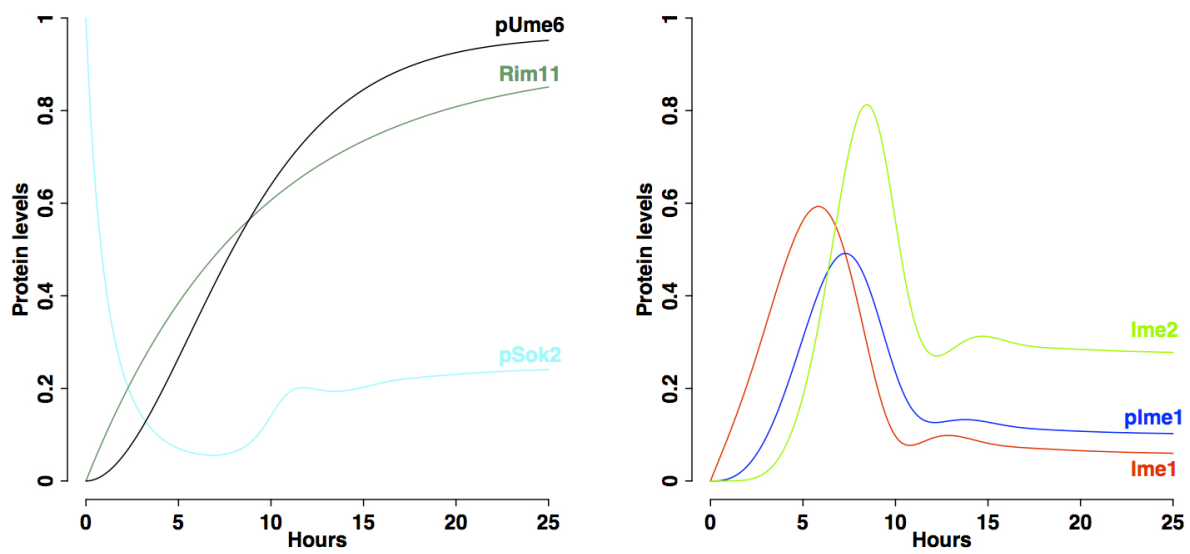


**Figure S6 - Numerical simulations of the pSok2-lme1 feedback knockout model**

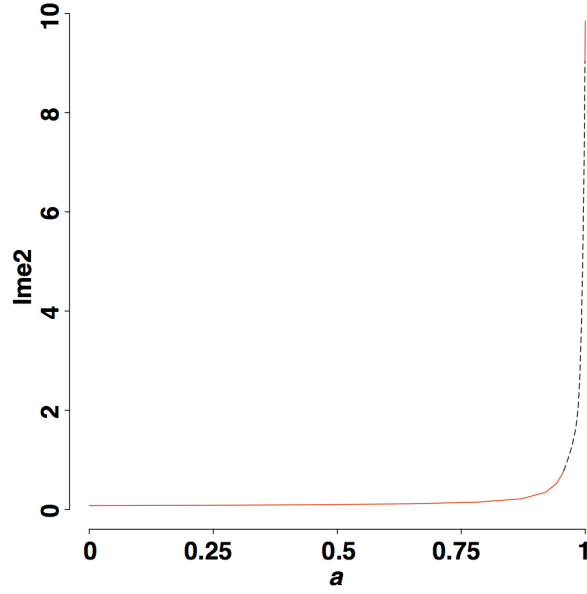


**Figure S7 - Numerical simulations of the lme2-lme1 feedback knockout model**





**Figure S8 - Numerical simulations of the lme2 feedback knockout model**



**Figure S9 - Bifurcation analysis of PKA**

Steady state value of Ime2 as a function of PKA. The change in PKA activity is represented by simultaneously varying  $p_{rim11}$  and  $p_{sok2}$ , phosphorylation rates of Rim11 and Sok2, respectively. This is achieved by varying  $a$ , where  $p_{rim11} = 0.1 - 0.1a$  and  $p_{sok2} = 7 - 7a$ . Red segments represent stable steady states, whereas black segments trace unstable steady states.