

Jornada Matemática SPM/CIM em Epidemiologia Teórica
Coimbra, 16 de Janeiro de 2010

Stochastic models of infection dynamics

[**Ana Nunes**]

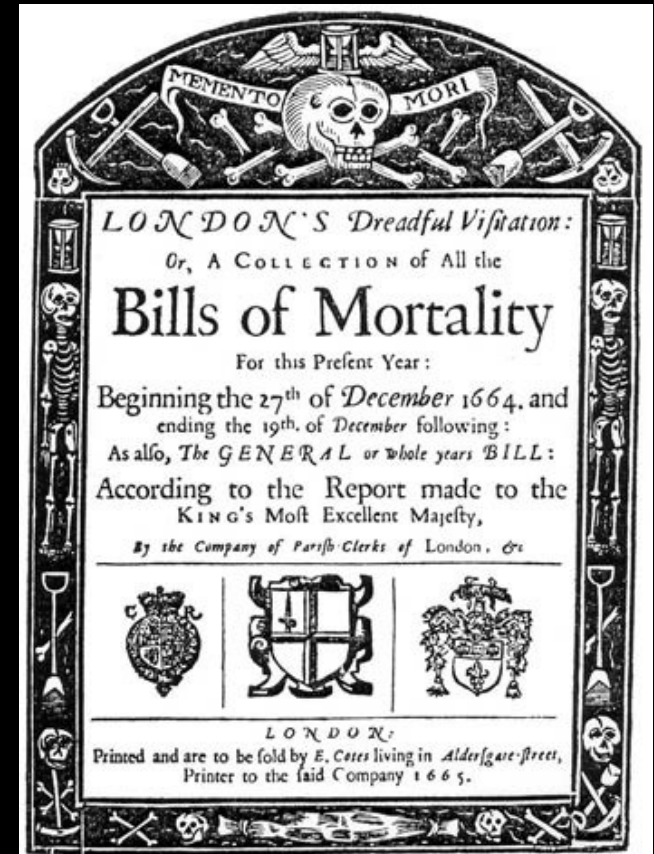
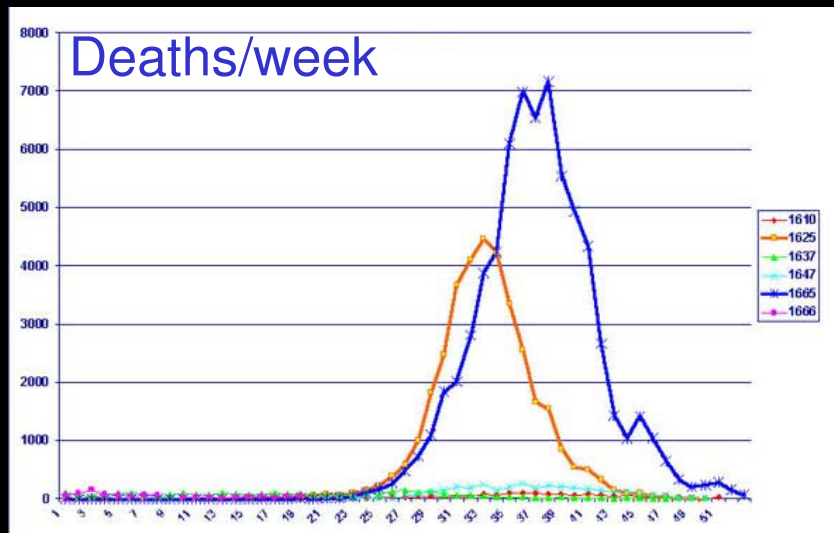


Departamento de Física, FCUL



Outline

- Deterministic models and phenomenology
- Resonant amplification of stochastic fluctuations



- The effect of correlations
- Conclusions and perspectives

Deterministic models & phenomenology

SIR dynamics

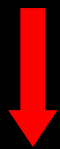
Population N , compartments S, I, R

$$S' = -b S I$$

$$I' = b S I - g I$$

$$R' = g I$$

Kermack – McKendrick, 1927

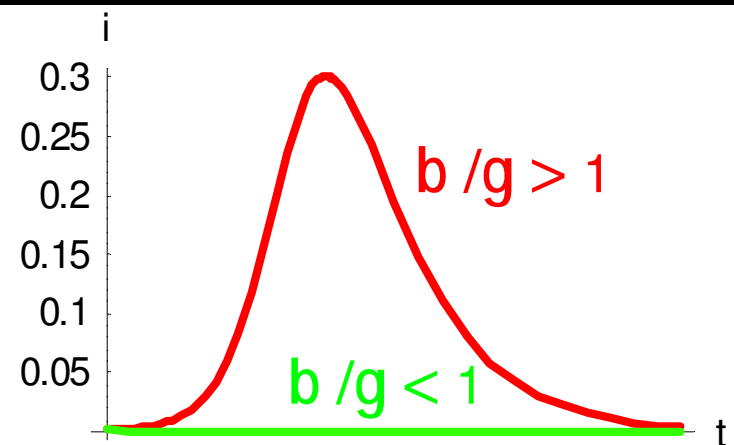


N constant, $S+I+R=N$, $N \gg 1$

$$s' = -b s i$$

$$i' = b s i - g i$$

s, i densities



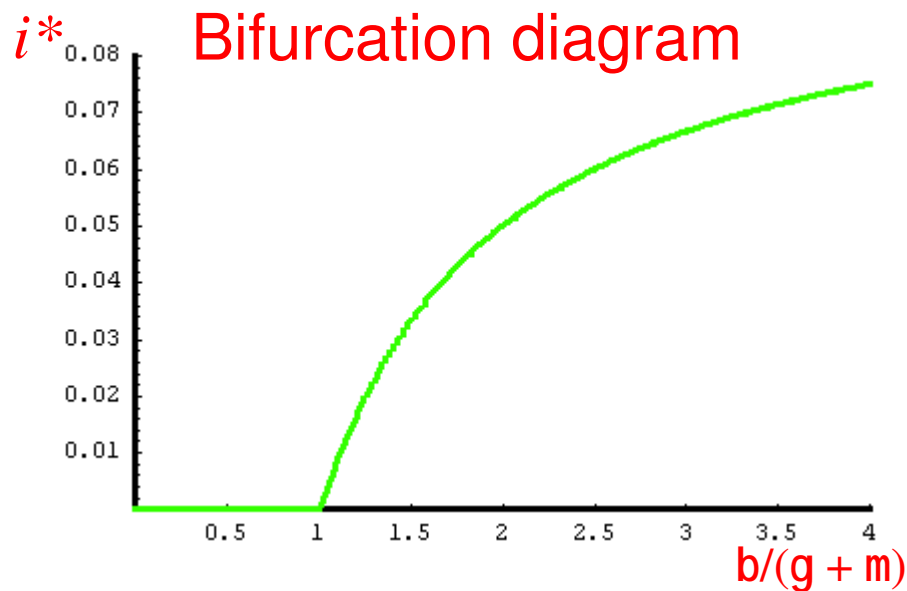
Deterministic models & phenomenology

SIR(S) dynamics

$$\begin{aligned} s' &= m(1-s) - b s i \\ i' &= b s i - (g+m) i \end{aligned}$$

$$\begin{aligned} s' &= m(1-s-i) - b s i \\ i' &= b s i - g i \end{aligned}$$

m birth/death or immunity waning rate

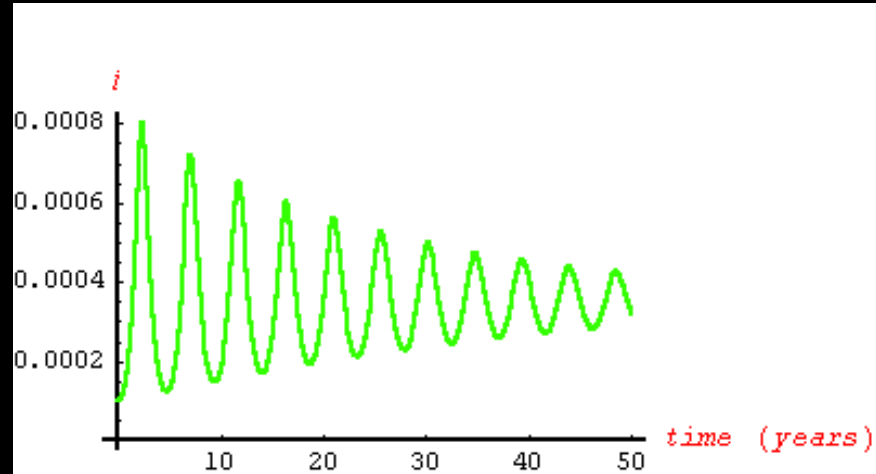


Epidemic/endemic threshold
 $R_0 = 1$,

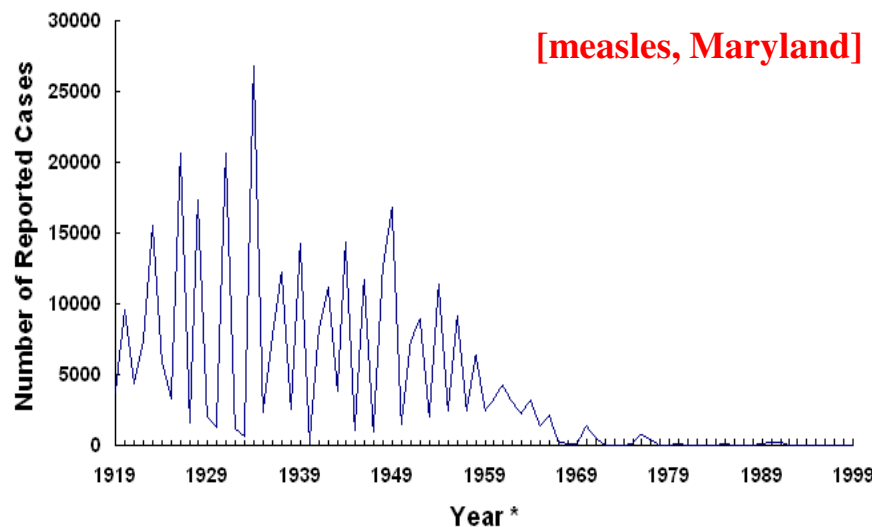
$$R_0 = b/(g+m) \text{ - dSIR,}$$
$$R_0 = b/g \text{ - SIRS,}$$

Deterministic models & phenomenology

SIR(S) dynamics



Damped oscillations



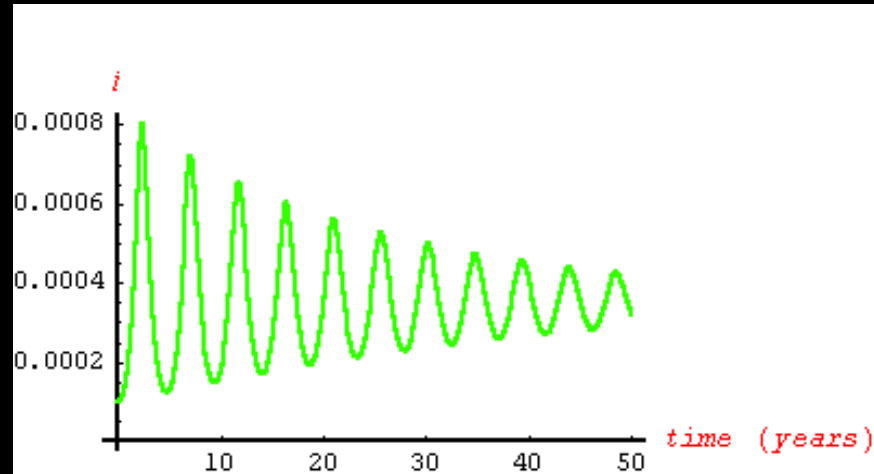
Recurrent epidemics

Diverse incidence patterns

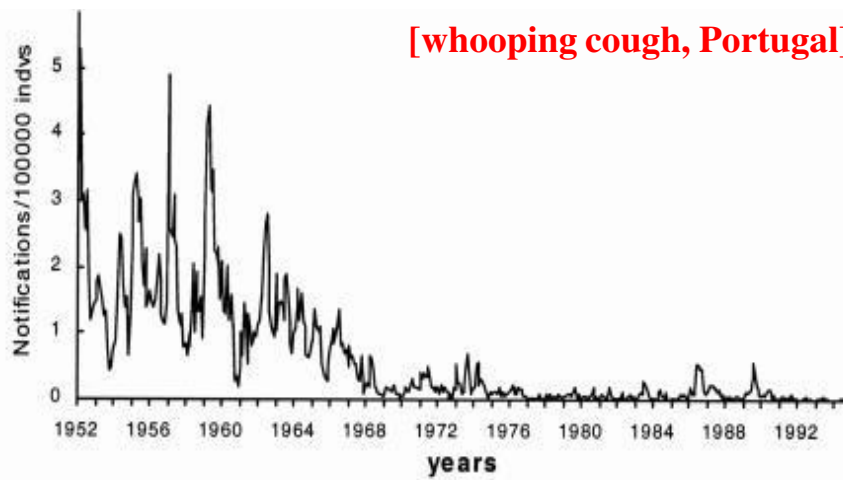
Extinctions

Deterministic models & phenomenology

SIR(S) dynamics



[whooping cough, Portugal]



Damped oscillations

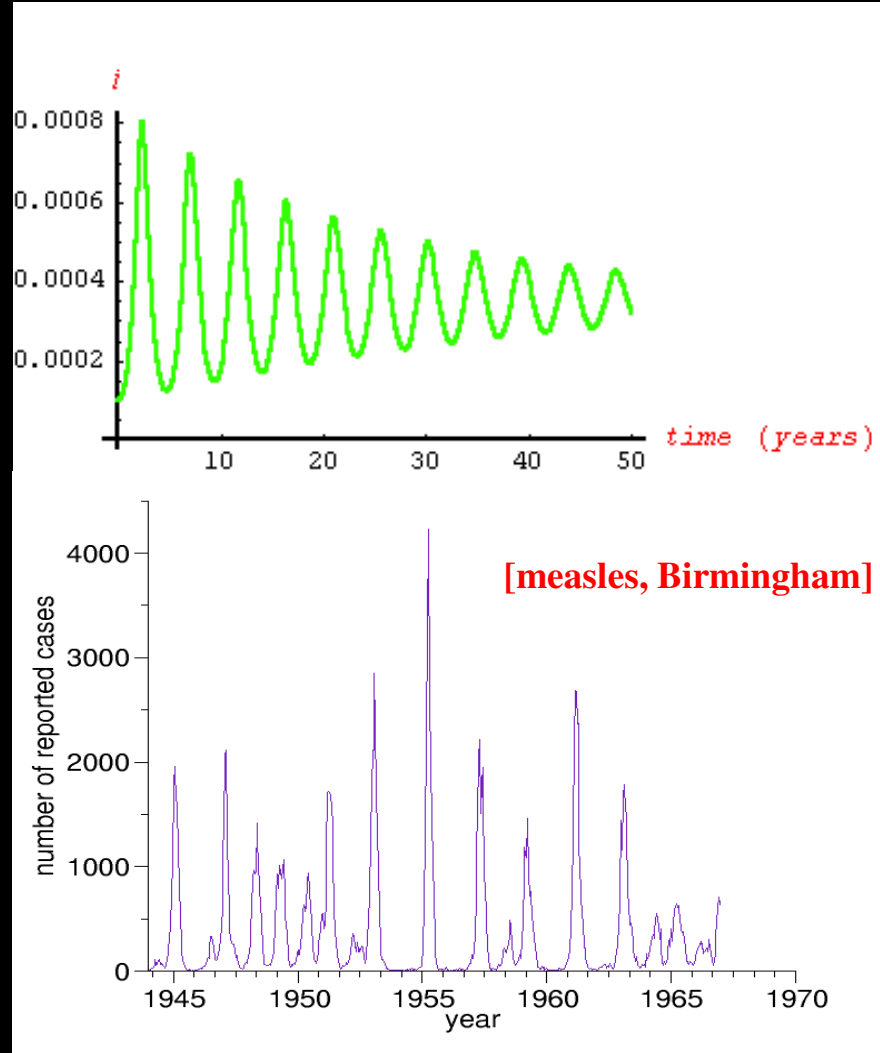
Recurrent epidemics

Diverse incidence patterns

Extinctions

Deterministic models & phenomenology

SIR(S) dynamics



Damped oscillations

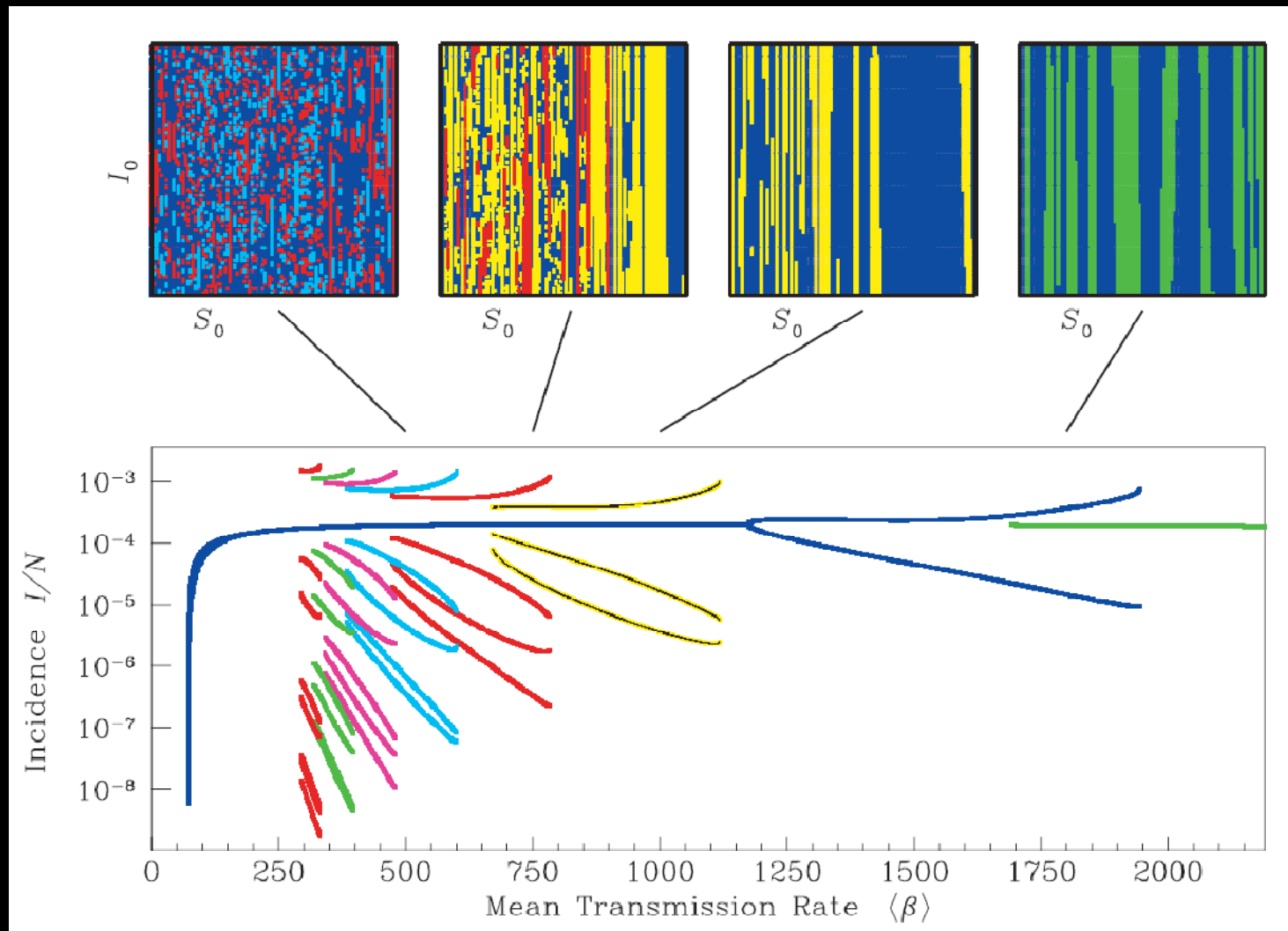
Recurrent epidemics

Diverse incidence patterns

Extinctions

Deterministic models & phenomenology

Periodic forcing

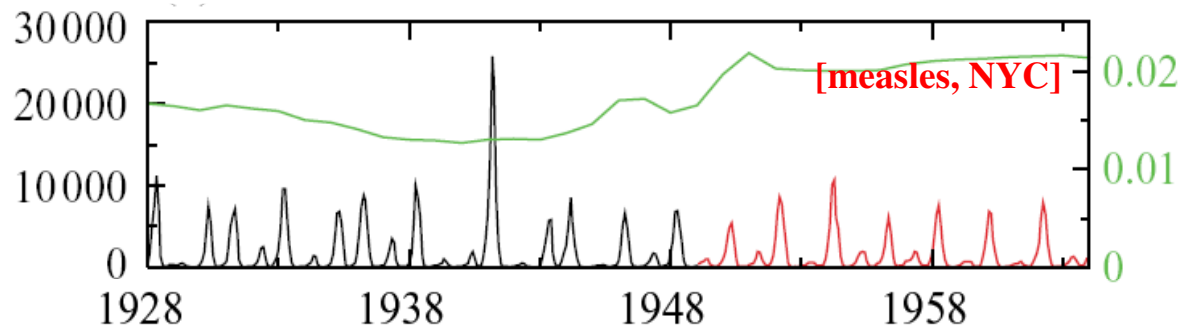


[1] Earn et al., Science 287, 667-670 (2000)

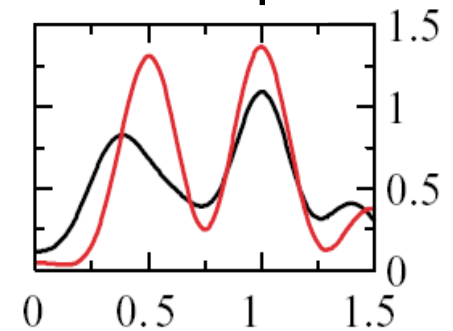
Deterministic models & phenomenology

Periodic forcing

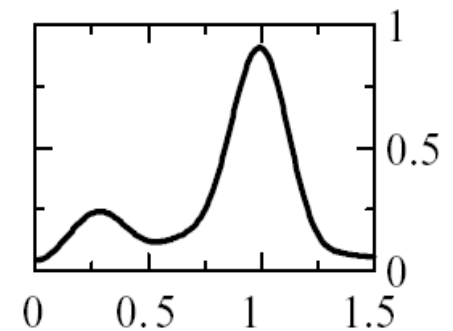
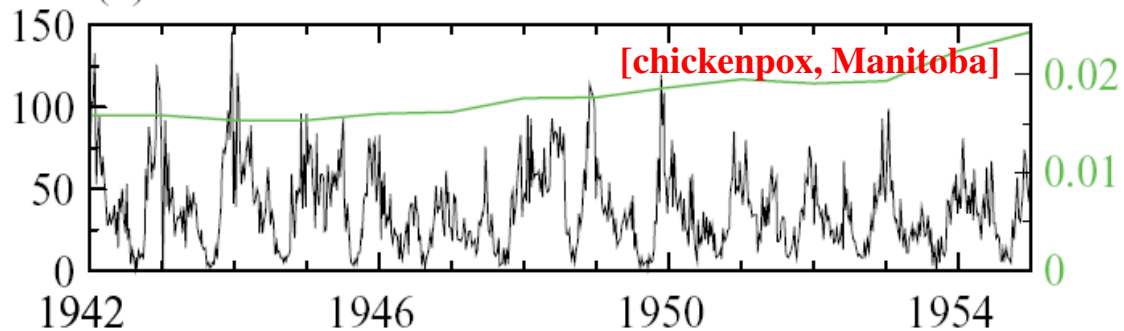
Time series



Power Spectra



(b)

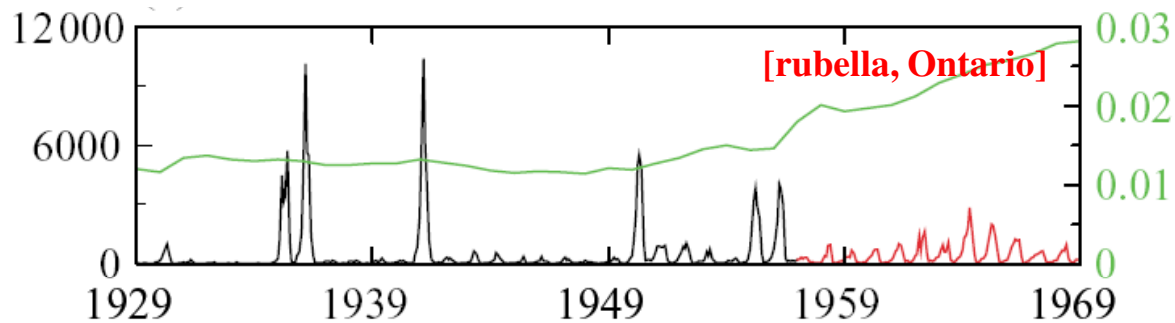


[2] Bauch and D. J. D. Earn, Proc. R. Soc. Lond. B 270, 1573-1578 (2003)

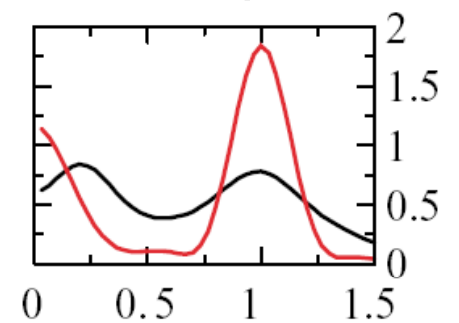
Deterministic models & phenomenology

Periodic forcing

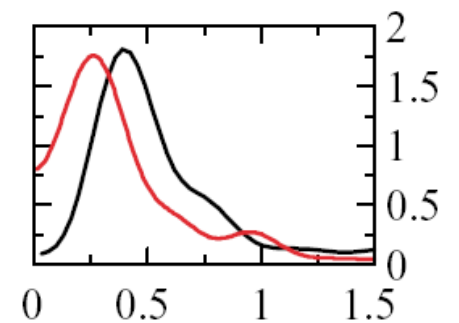
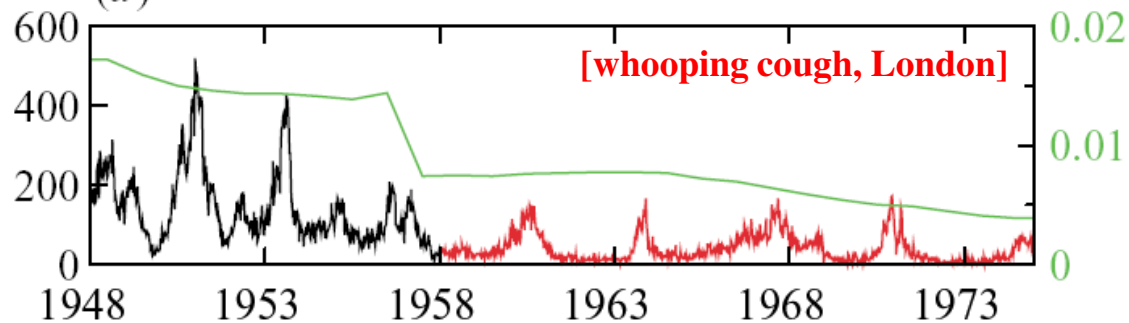
Time series



Power Spectra



(d)



[2] Bauch and D. J. D. Earn, Proc. R. Soc. Lond. B 270, 1573-1578 (2003)

Deterministic models & phenomenology

Periodic forcing

Seasonality versus stochasticity debate:

- Forced nonlinear dynamics explains the data, and stochasticity has a secondary role (switching between attractors and sustaining small amplitude fluctuations around the deterministic system's equilibrium)
- Many data records cannot be understood in a deterministic framework, fluctuations can be dominant

Resonant amplification of stochastic fluctuations

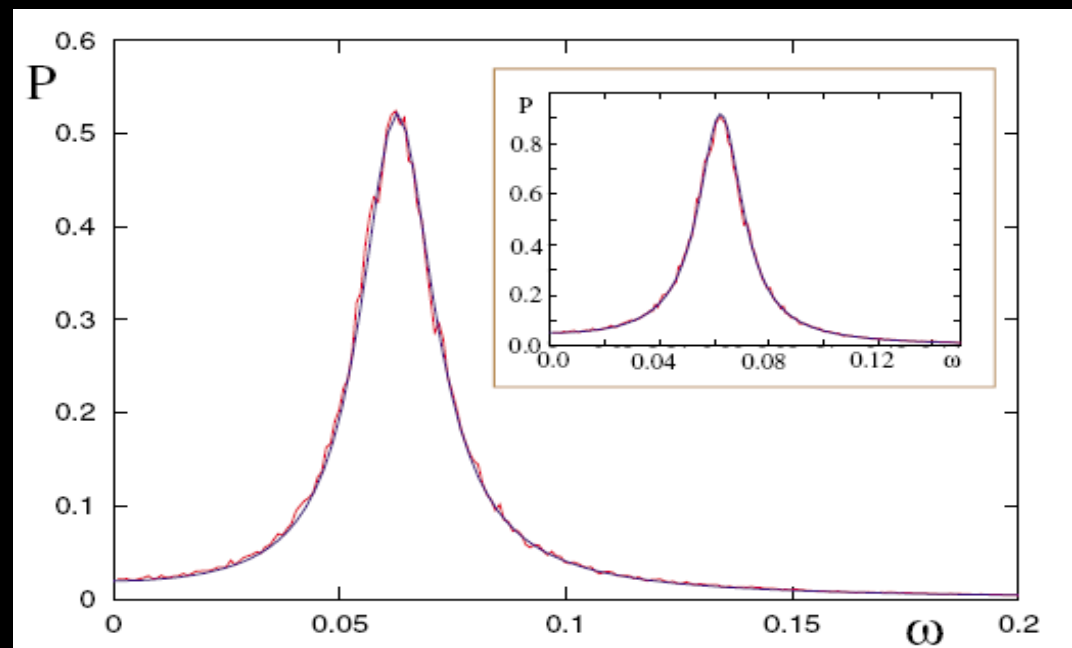
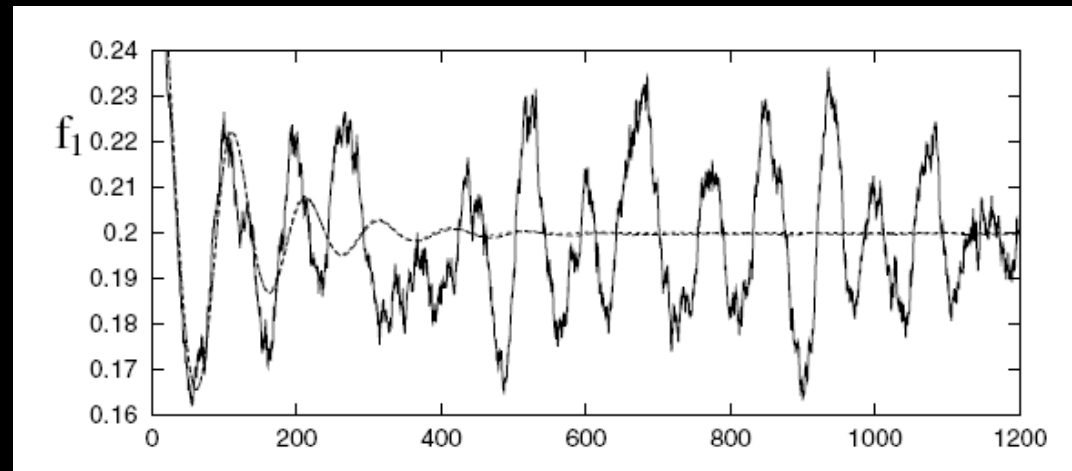
Predator-prey dynamics

[3] A J McKane and T J Newman,
Phys. Rev. Lett. 94 (2005), 218102

Mean field and average densities exhibit oscillatory transients.

$$P(\omega) = \frac{\alpha + \beta\omega^2}{[(\omega^2 - \Omega_0^2)^2 + \Gamma^2\omega^2]}$$

Sustained oscillatory patterns arise through resonant amplification of internal noise



Resonant amplification of stochastic fluctuations

Stochastic infection dynamics in a well mixed population,
 $N = S + I + R$

	Process	Rate
Infection	$S + I \rightarrow I + I$	$\lambda S I / N$
Recovery	$I \rightarrow R$	$d I$
Loss of immunity	$R \rightarrow S$	R

[4] D Alonso, A J McKane and M Pascual, J R Soc Interface 4, 575-82 (2007)

Resonant amplification of stochastic fluctuations

Master Equation

$$\frac{dP(\sigma; t)}{dt} = \sum_{\sigma' \neq \sigma} T(\sigma|\sigma')P(\sigma'; t) - \sum_{\sigma' \neq \sigma} T(\sigma'|\sigma)P(\sigma; t), \quad \sigma = (S, I)$$

$$T(S - 1, I + 1|S, I) = \lambda S I / N,$$

$$T(S, I - 1|S, I) = \delta I,$$

$$T(S + 1, I|S, I) = \gamma (N - S - I)$$

van Kampen's system size expansion

$$S = N\phi + \sqrt{N}x_0$$

$$I = N\psi + \sqrt{N}x_1$$

Resonant amplification of stochastic fluctuations

van Kampen's system size expansion

$$\begin{aligned}\phi' &= d(1 - \phi - \psi) - b\phi\psi \\ \psi' &= b\phi\psi - g\psi\end{aligned}$$

Leading order terms
MF equations for the densities

Next to leading order

Linear Fokker-Planck or Langevin equations for the normalized fluctuations

$$\dot{x}_i(t) = \sum_{j=1}^2 J_{ij} x_j(t) + L_i(t), \quad i = 1, 2$$

Resonant amplification of stochastic fluctuations

Fluctuation power spectrum around the MF equilibrium

$$P_i(\omega) \equiv \langle |\tilde{x}_i(\omega)|^2 \rangle = \sum_{j,k} M_{ik}^{-1}(\omega) B_{kj} M_{ji}^{-1}(-\omega)$$

$$M_{ik}(\omega) = i\omega\delta_{ik} - J_{ik} \text{ and } \langle \tilde{L}_i(\omega) \tilde{L}_j(\omega') \rangle = B_{ij} \delta(\omega + \omega')$$

$$P_S = \frac{B_{11}(J_{12}^2 + \omega^2)}{(D - \omega^2)^2 + T^2\omega^2} ,$$

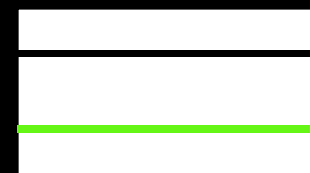
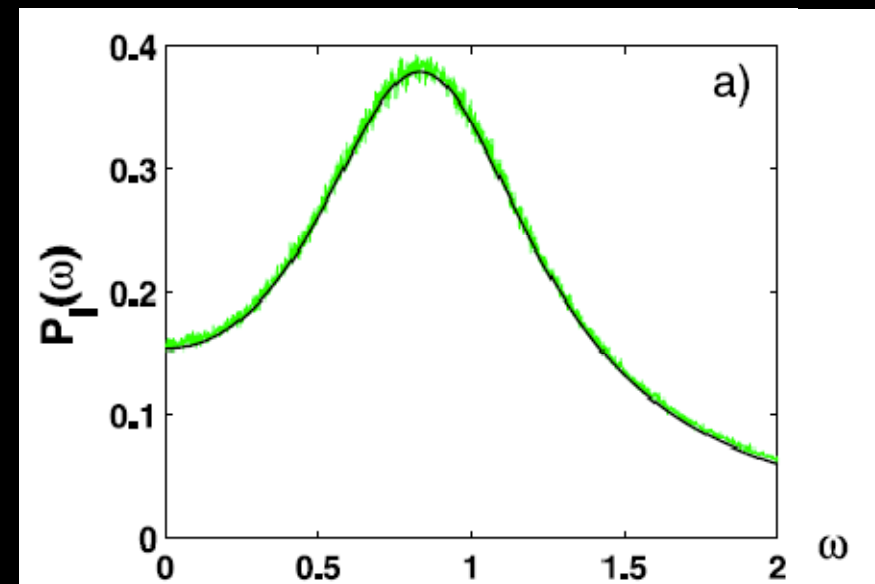
$$P_I = \frac{B_{11}(J_{11}^2 + J_{11}J_{21} + J_{21}^2 + \omega^2)}{(D - \omega^2)^2 + T^2\omega^2}$$

Resonant amplification of stochastic fluctuations

Power spectrum of the incidence fluctuations for the stochastic SIRS model

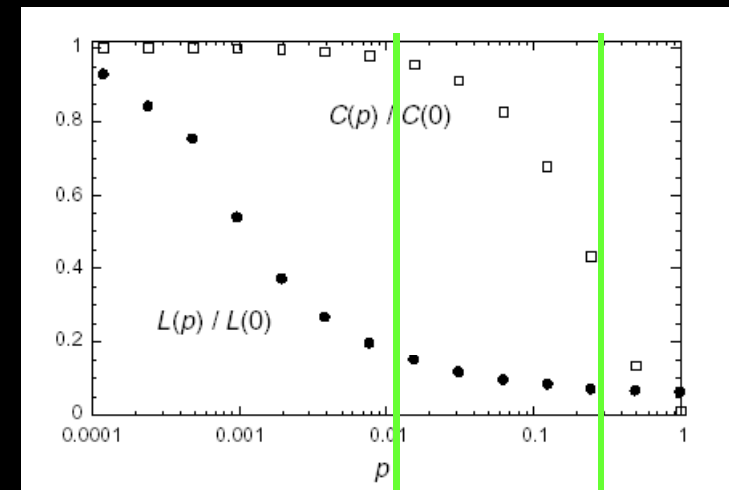
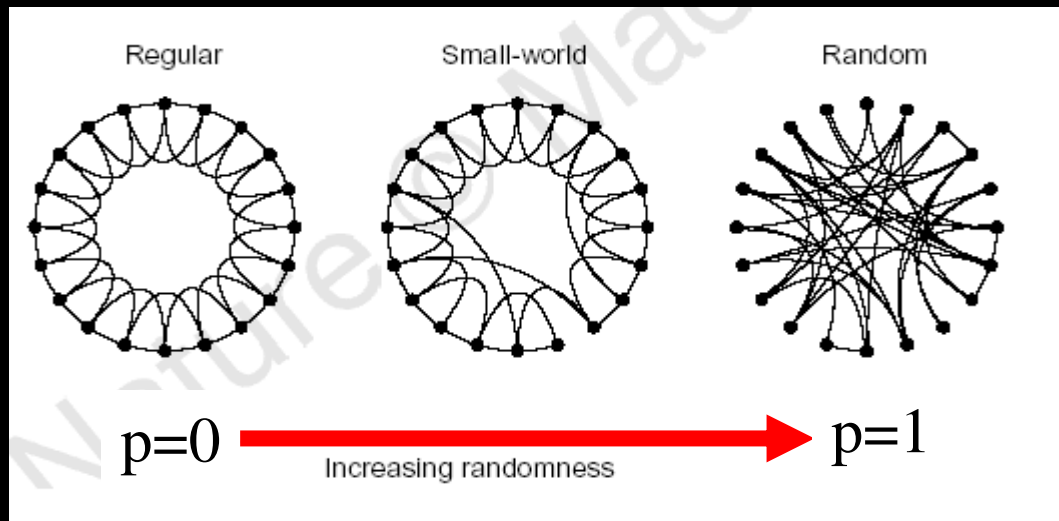
$$\lambda = 10, \quad \beta = 0.1, \quad N = 10^6$$

Finite N , sustained noisy oscillations



analytical
simulations

The effect of correlations in space



D Watts e S Strogatz, Nature 393 (1998), 440-442

SW

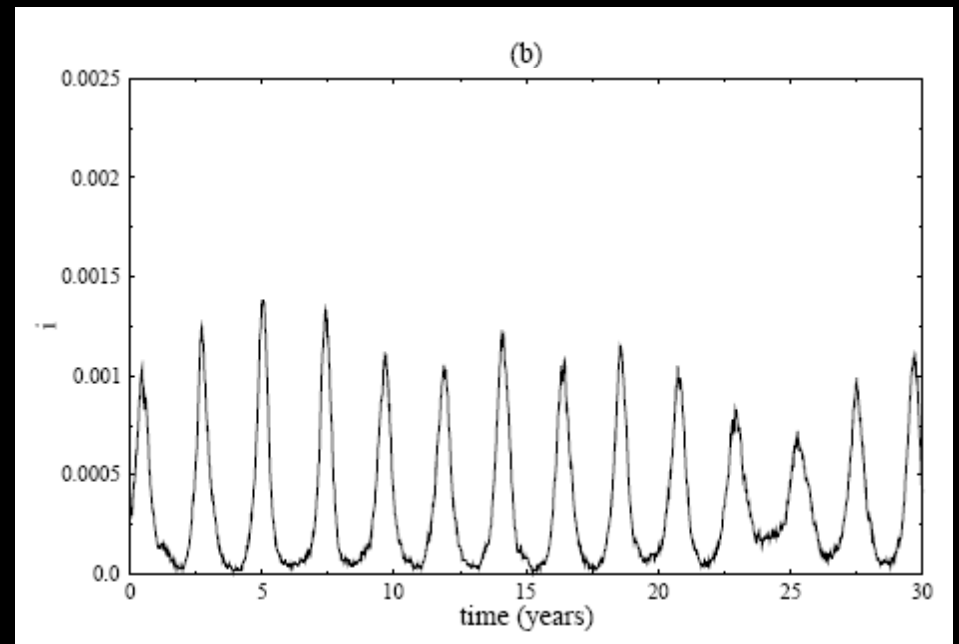
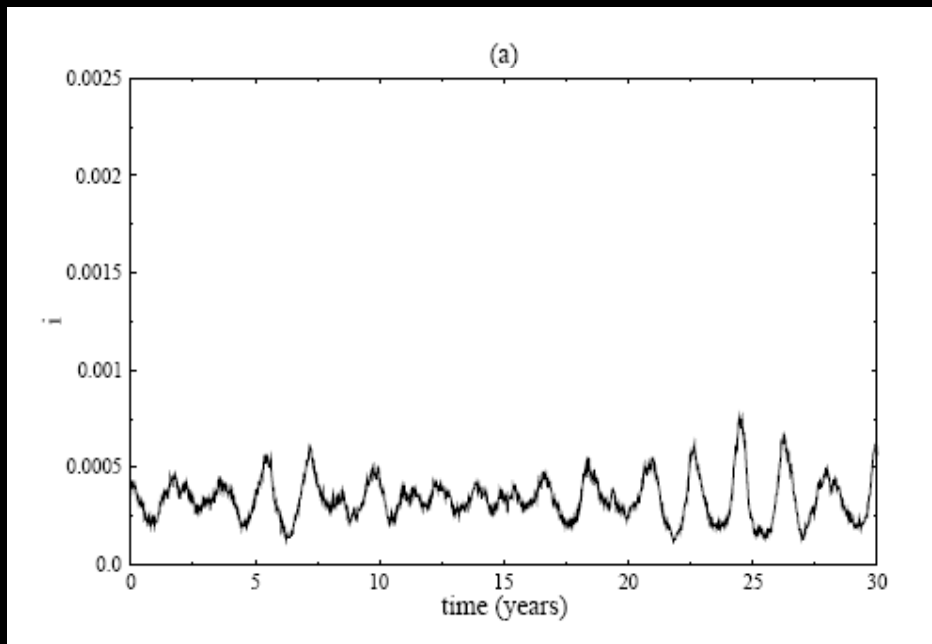
Simulations on small world networks may be used to assess the effect of spatial correlations on the stochastic fluctuations

The effect of correlations in space

SEIR stochastic simulations, **measles**, $N=10^6$

a) $p=1$

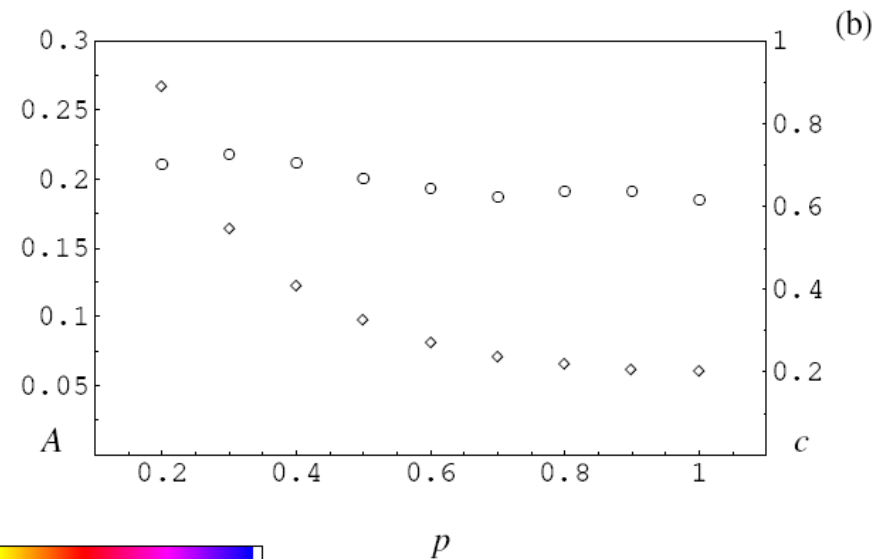
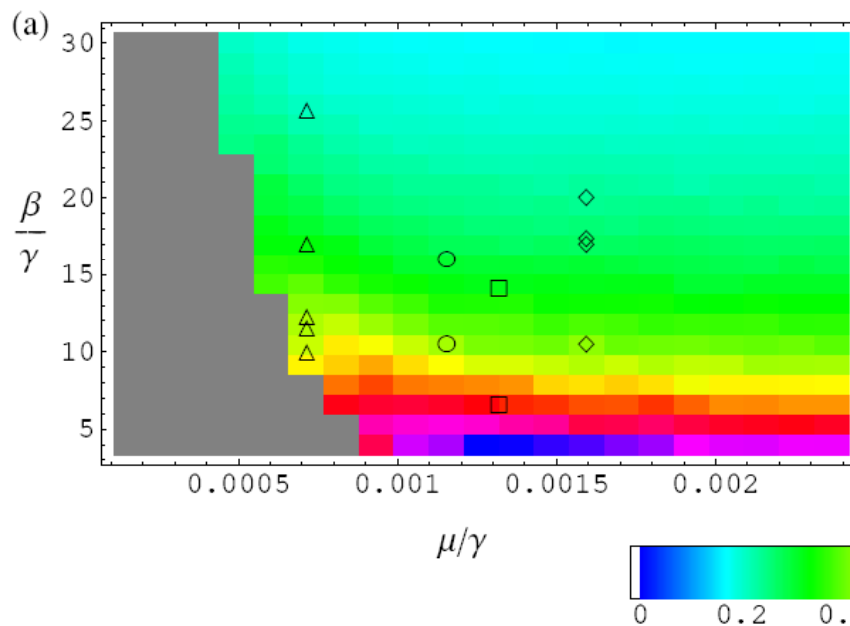
b) $p=0.2$



The effect of correlations in space

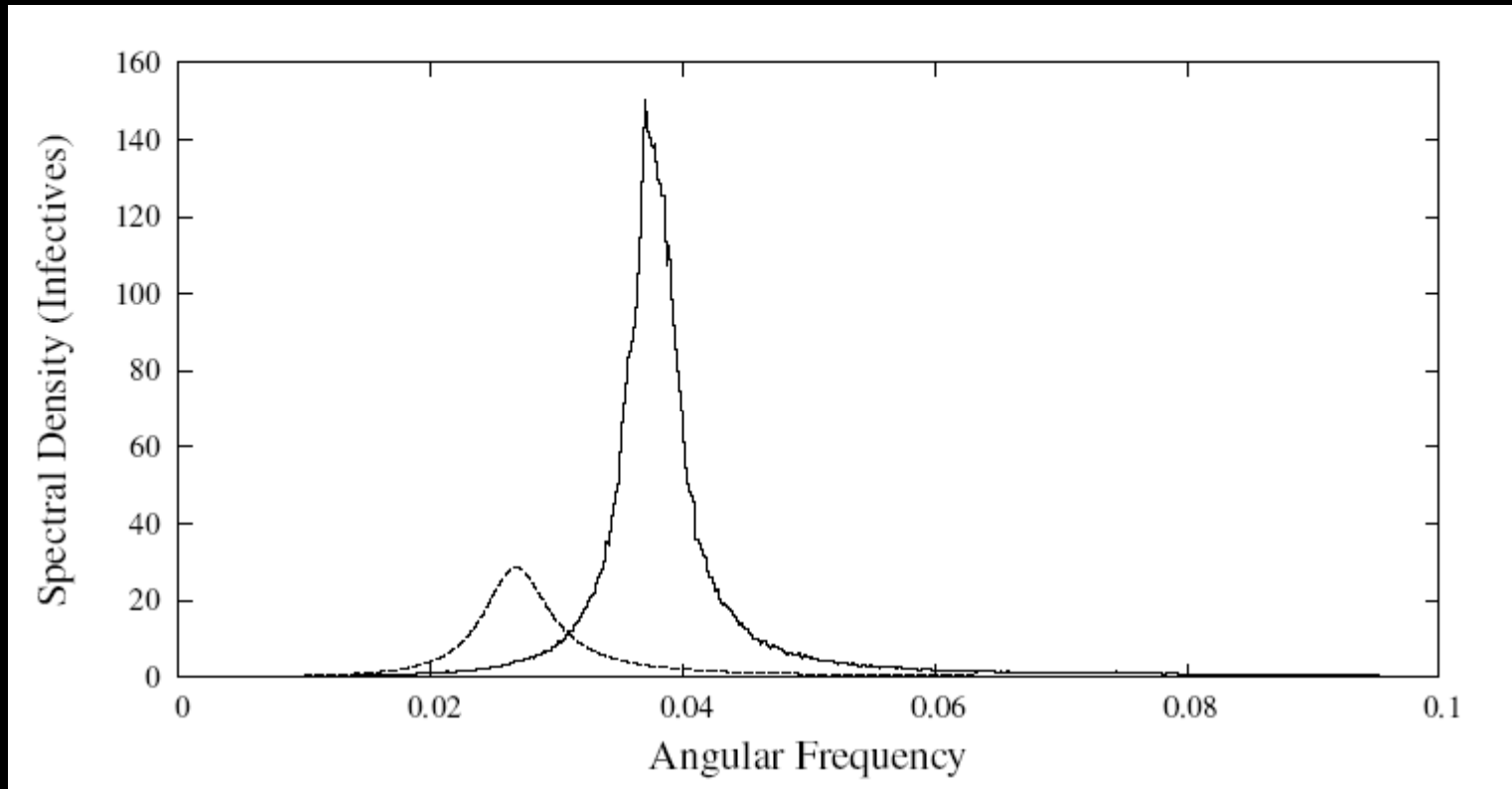
$p=0.6$

Amplification and coherence



Symbols correspond to published parameter values for childhood infectious diseases: measles (triangles), chickenpox (circle), rubella (squares) and whooping cough (diamonds), $N=10^6$

The effect of correlations in time



Fluctuation power spectrum for a stochastic dSIR model with the two extreme types of recovery period distribution: constant recovery rate (dashed), and fixed infectious period (full).

The effect of correlations

analytical description

Pair approximation for SIRS (k neighbours)

The rates of change of the pair densities $[xy]$, $x, y \in \{s, i, r\}$, depend on $[xy]$ e $[xyz]$

The standard pair approximation...

triplets are formed by uncorrelated pairs

$$\Sigma [asi] = (k-1)([as][si])/[s], \dots$$

... closes the equations for the pair densities

$$[si]' = g[ir] - (\lambda + d) [si] + \lambda (\Sigma [ssi] - \Sigma [isi]), \dots$$

and the pair densities determine the singlet densities

$$[s] = [si] + [ss] + [sr], \dots$$

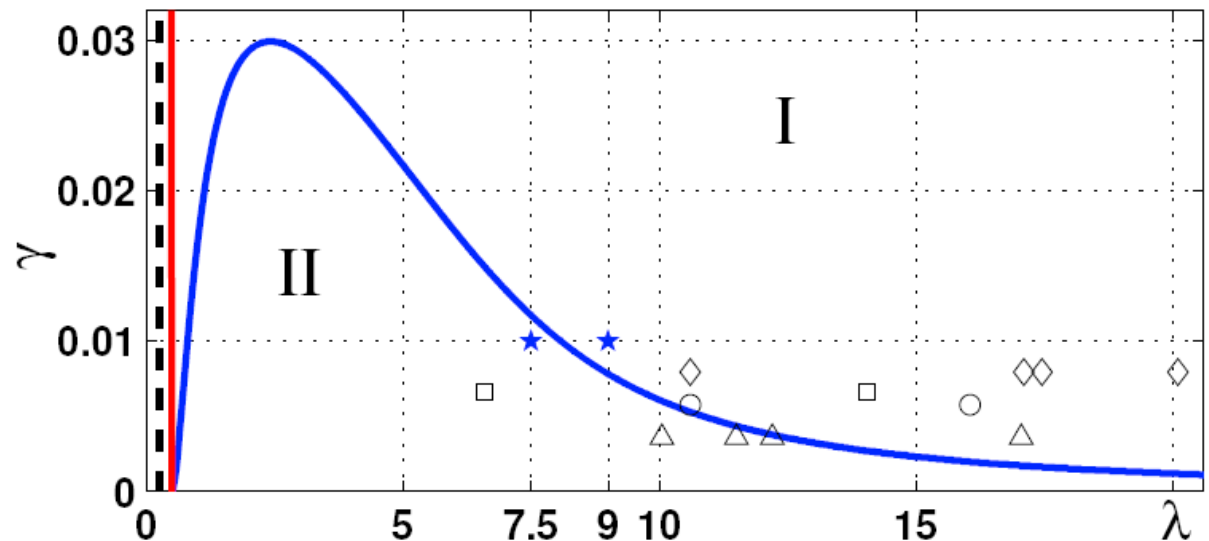
The effect of correlations

analytical description

$$\begin{aligned} \frac{d s}{d t} &= \gamma (1 - i - s) - k \lambda s i , \\ \frac{d i}{d t} &= k \lambda s i - \delta i , \\ \frac{d s i}{d t} &= \gamma r i - (\lambda + \delta) s i + \frac{(k - 1) \lambda s i}{s} (s - s r - 2 s i) , \\ \frac{d s r}{d t} &= \delta s i + \gamma (1 - s - i - r i - 2 s r) - \frac{(k - 1) \lambda s i s r}{s} , \\ \frac{d r i}{d t} &= \delta (i - s i) - (\gamma + 2 \delta) r i + \frac{(k - 1) \lambda s i s r}{s} . \end{aligned}$$

k is the fixed degree of the RRG

Bifurcation diagram
 $k=4$



The effect of correlations

analytical description

Pair approximation for stochastic SIRS

N individuals, $kN/2$ pairs

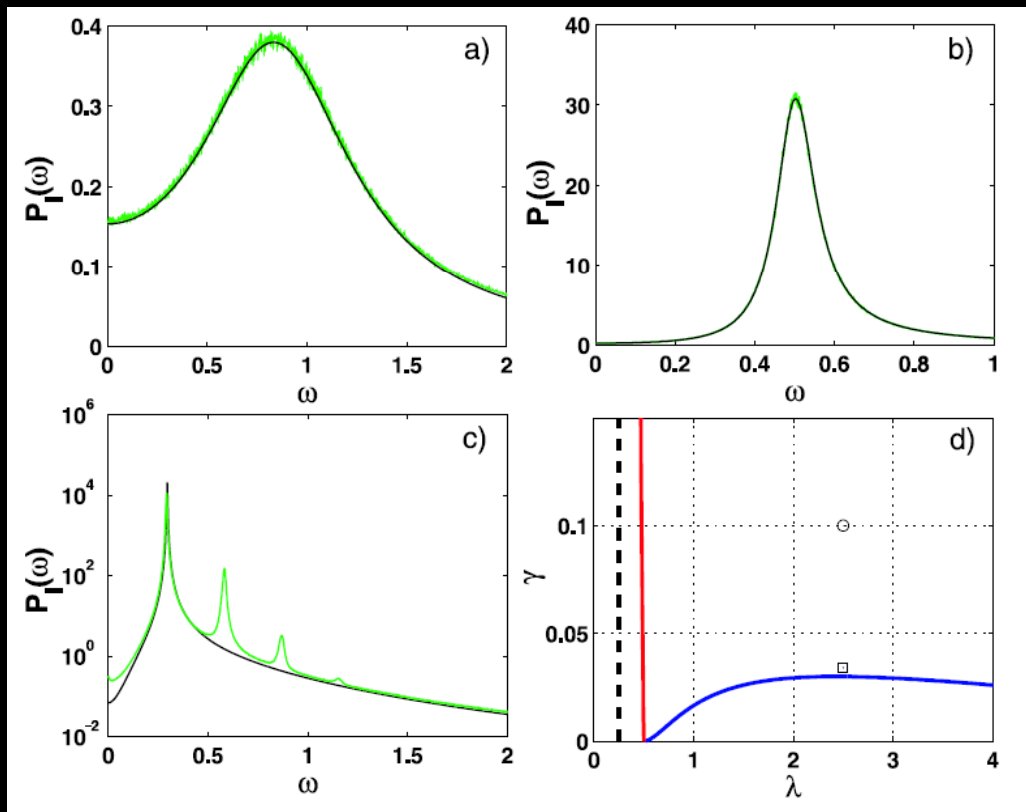
	Process	Rate
Infection	$SI \rightarrow II$	λSI
Recovery	$I \rightarrow R$	$d I$
Loss of immunity	$R \rightarrow S$	R

Fluctuation power spectra around the equilibrium can be computed using van Kampen's expansion as before

The effect of correlations

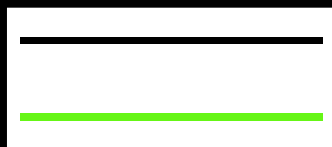
analytical description

Pair approximation for stochastic SIRS



Amplitude and coherence of the resonant fluctuations are much enhanced in the PA.

Harmonic peaks show up close to the boundary of the oscillatory phase.

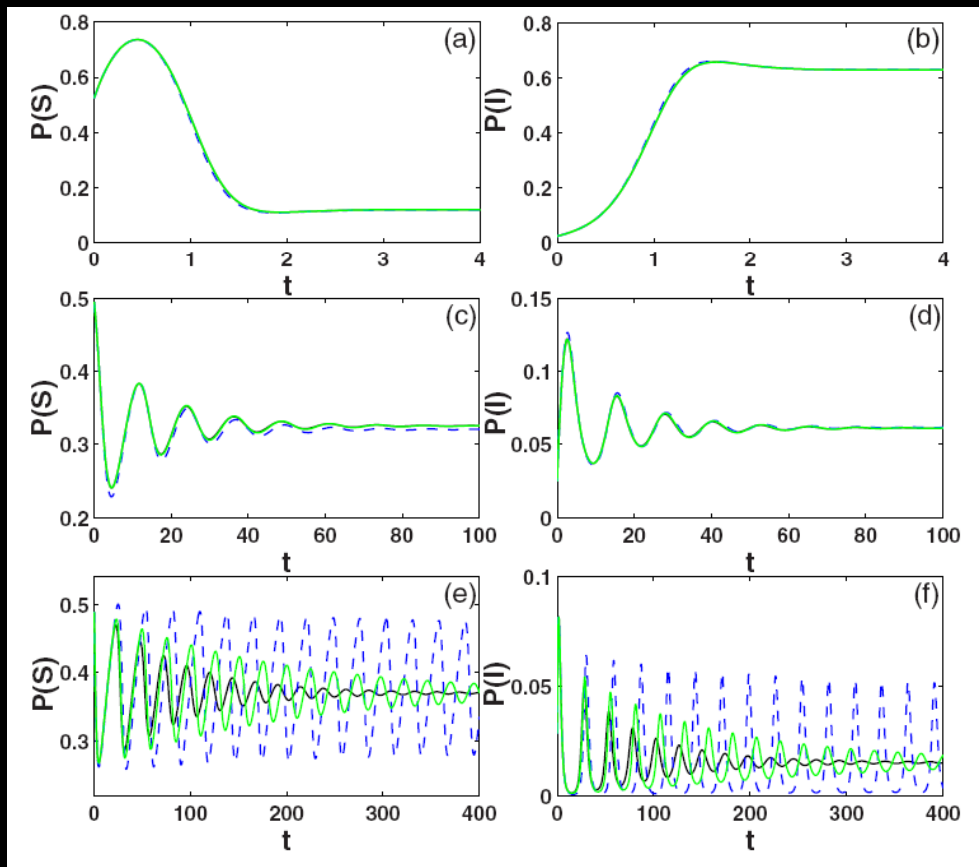


analytic
simulations $N=10^6$

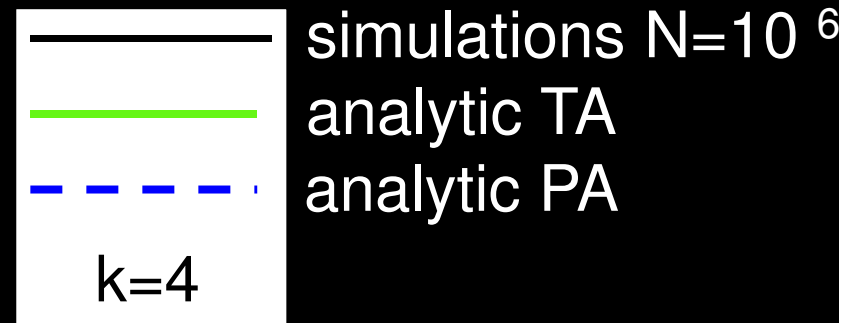
Rozhnova & Nunes,
Phys. Rev. E 79, 041922 (2009)

The effect of correlations analytical description

Beyond the pair approximation: SIRS on a k-RRG



Higher order cluster approximations are needed to describe some regimes of the dynamics on networks

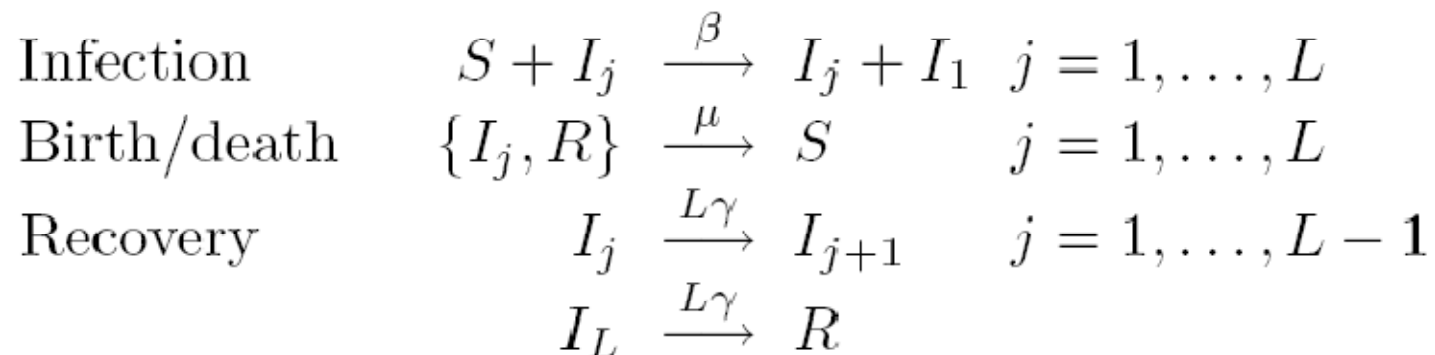


Rozhnova & Nunes,
Phys. Rev. E 80, 051915 (2009)

The effect of correlations

analytical description

Stochastic d-SIR with distributed infectious periods

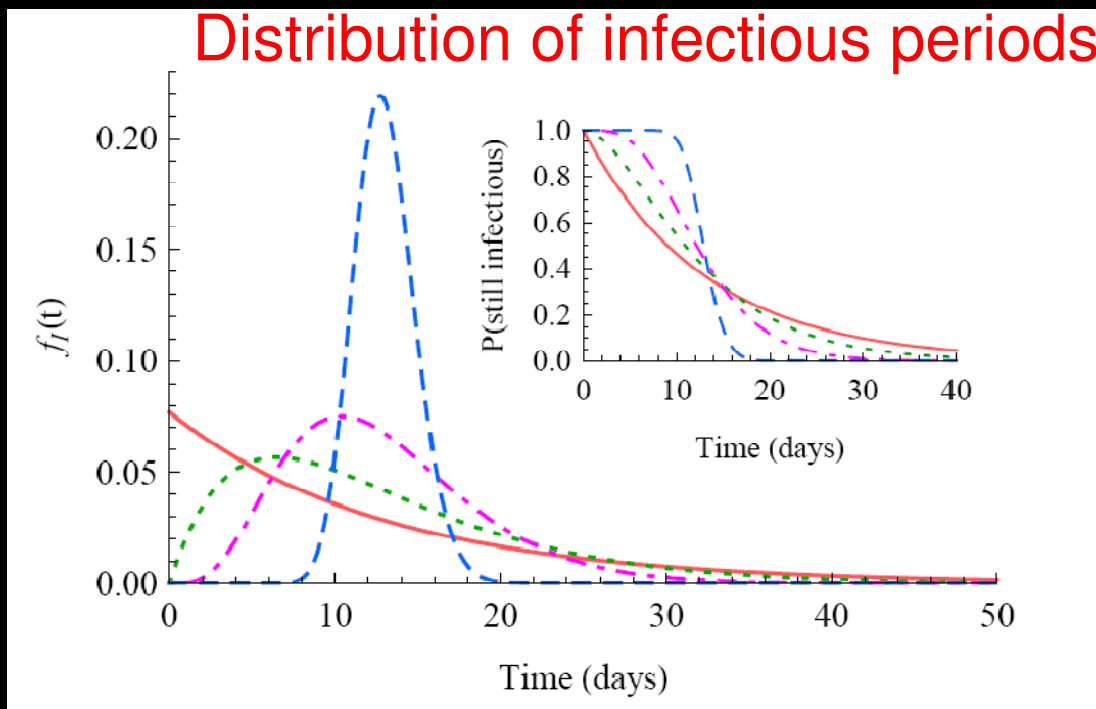


Gamma distributed recovery profiles that interpolate between exponentially distributed ($L=1$) and constant recovery period as $L \rightarrow \infty$

The effect of correlations

analytical description

Distribution of infectious periods

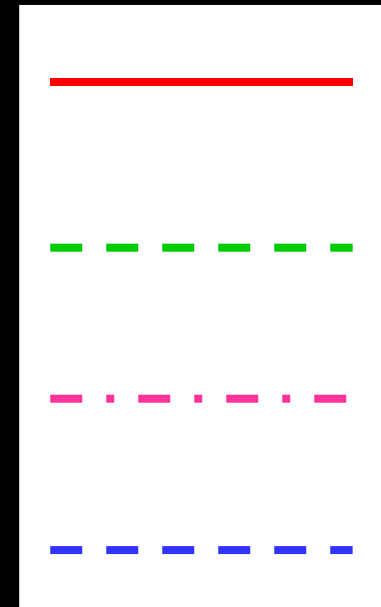


$L=1$

$L=2$

$L=5$

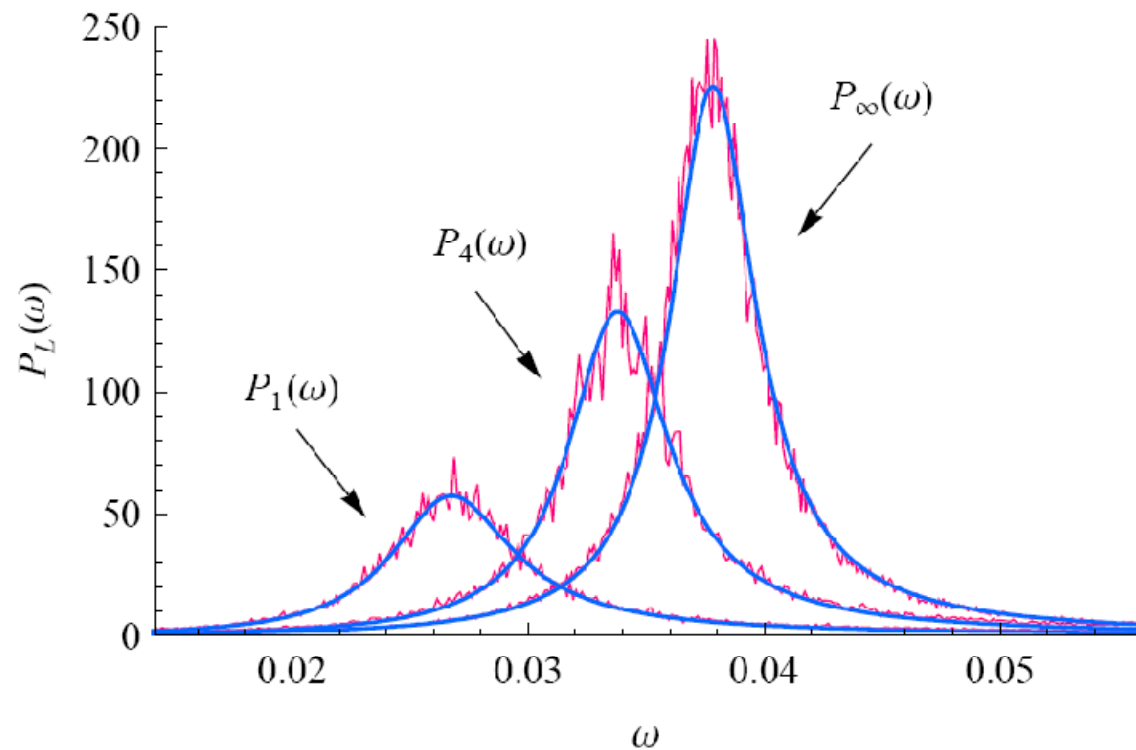
$L=50$



For each L , fluctuation power spectra around the equilibrium can be computed using van Kampen's expansion as before

The effect of correlations

analytical description



The power spectrum can be obtained in closed form in the limit $L \rightarrow \infty$

The peak frequency is very sensitive to the recovery profile

— analytic
— simulations $N=10^6$

Black et al.,
Phys. Rev. E 80, 021922 (2009)

Conclusions & Perspectives

In infection dynamics, intrinsic stochasticity combined with correlations can give rise in finite systems to sustained noisy oscillations compatible with realistic patterns of recurrent epidemics

The power spectrum of the fluctuations around the endemic equilibrium can be computed analytically

The interplay between fluctuations and forcing is not yet well understood

Contact network structure and stochasticity can be treated analytically with cluster approximations and system size expansions

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