

# MeccanicaMente

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-Afferenze: INdAM (GNFM), UMI (Giunta AI), INFN, ARxiv, CMCC, Sant'Anna, PhD-AI

-Mansioni Editoriali: Neural Networks (Elsevier), Nature Sci. Rep., Proc. Royal Society

-Grants in essere: MAECI (Italy-Israel), PRIN (Mat07), PNRR (Foundation AI) [←expected...]

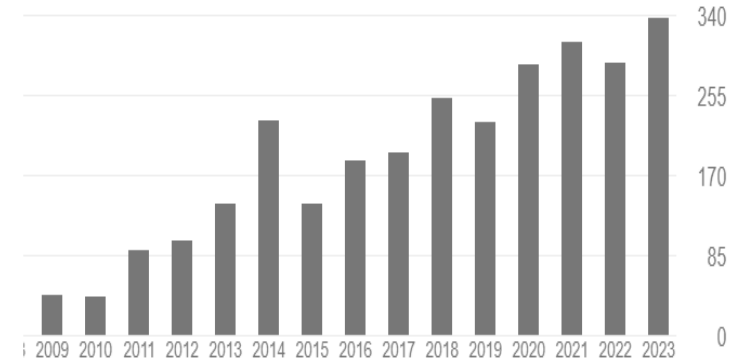
-Pubblicazioni : >100 in Meccanica Statistica ad ampio spettro

Problemi diretti:

- Neural Networks and Learning Machines
- Spin Glasses and Random Networks
- Reaction Kinetics & Biological Computing
- Mathematical models & methods

Problemi inversi:

- Stochastic Processes in Quantitative Biology (mainly immune response)
- HD Statistical Inference in Quantitative Biology (mainly cancerogenesis)
- Machine Learning per Quantitative Biology (genomics & trajectory reconstruction)



Ministero degli Affari Esteri e della Cooperazione Internazionale

# Structure of the talk

We use a simple physical model to inspect the techniques (mean field Ising model, Curie-Weiss)

- 1) Physicist approach to ferromagnetic phase transitions
- 2) Statistical mechanical description (probabilistic methods)
- 3) Analytical mechanical description (PDE techniques)

From mean-field Ising to mean field neural networks:

- Hopfield model for pattern recognition
- Hopfield model for machine learning



# 1 Mean field phase transitions: heuristics

Oscillatori di Kuramoto: [video](#)  
Spin di Curie-Weiss: [video](#)

We use the *Free Energy F*

$$\xrightarrow{\text{red arrow}} \underline{F = E - TS}$$

$$E = H(\sigma|J, h) = -\frac{1}{N} \sum_{i < j}^{N, N} J_{ij} \sigma_i \sigma_j - h \sum_i^N \sigma_i = -N \left( J \frac{m^2}{2} + hm \right)$$

and, with the choice  $J_{ij} = J > 0$ , in the large  $N \rightarrow \infty$  limit, we write

$$m = \frac{1}{N} \sum_i^N \sigma_i$$

$$\mathcal{P}(\sigma_1, \sigma_2, \dots, \sigma_N) = \prod_{i=1}^N \mathcal{P}(\sigma_i),$$

$$\mathcal{P}(\sigma_i) = \left( \frac{1+M}{2} \right) \delta(\sigma_i - 1) + \left( \frac{1-M}{2} \right) \delta(\sigma_i + 1),$$

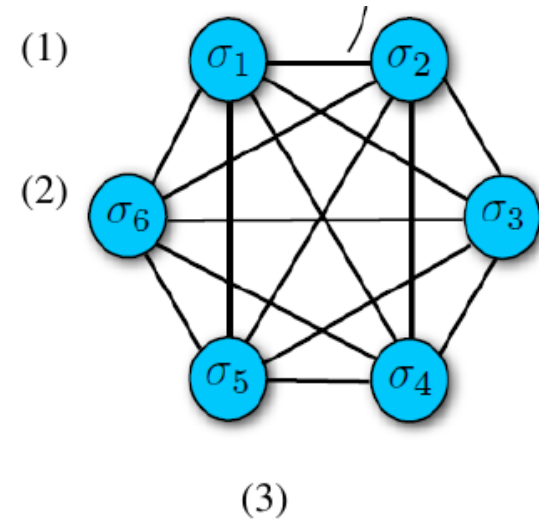
$$\langle \sigma \rangle = \left[ \left( \frac{1+M}{2} \right) \cdot (+1) + \left( \frac{1-M}{2} \right) \cdot (-1) \right] = M, \quad \leftarrow \lim_{N \rightarrow \infty} \mathcal{P}(m) = \delta(m - M)$$

$$S(\sigma) = - \sum_{\sigma_i = \pm 1} \mathcal{P}(\sigma_i) \ln \mathcal{P}(\sigma_i) = -N \left[ \left( \frac{1+M}{2} \right) \ln \left( \frac{1+M}{2} \right) + \left( \frac{1-M}{2} \right) \ln \left( \frac{1-M}{2} \right) \right]$$

• Control parameter:  $T, h, J$ .

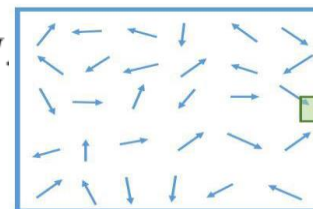
• Order parameter:  $m = \frac{1}{N} \sum_i^N \sigma_i$ , whose mean is  $\langle m \rangle = M$ .

furthermore, keep in mind that the temperature obeys a random walk (i.e. Fourier PDE).



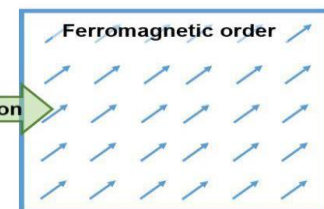
**T alta (vince Smax)**

**T bassa (vince Emin)**



Magnetization = 0

Phase transition



Magnetization = M

thus we can write  $F = F(M|T, h, J)$  as

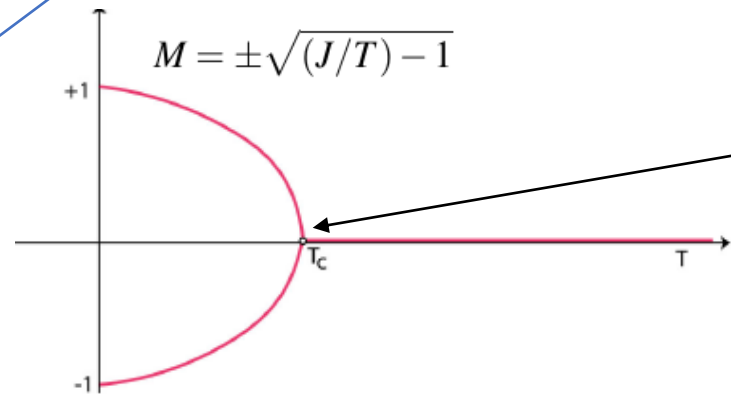
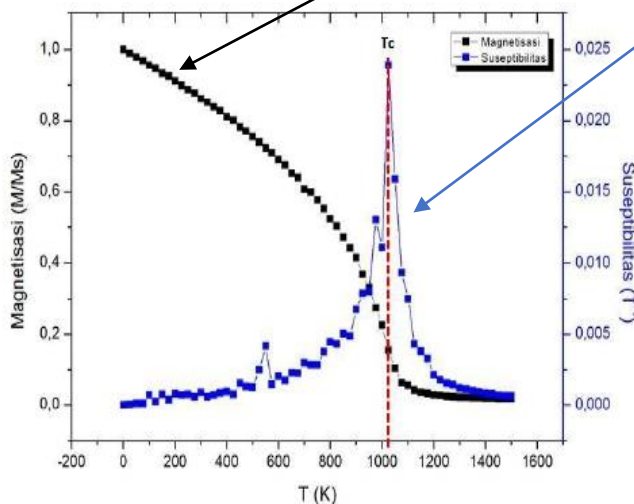
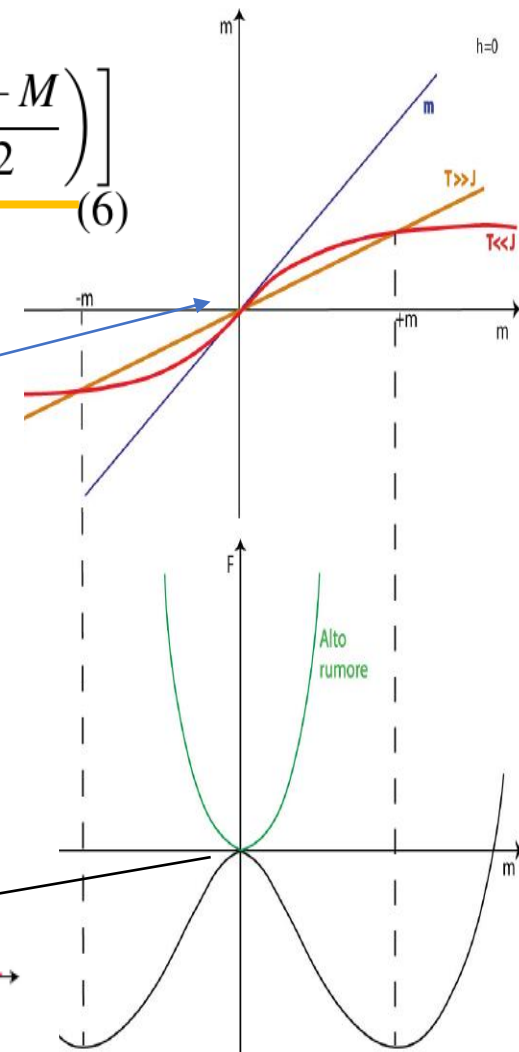
$$F = -N \left[ \left( \frac{J}{2} M^2 + hM \right) - T \left( \frac{1+M}{2} \ln \left( \frac{1+M}{2} \right) + \left( \frac{1-M}{2} \right) \ln \left( \frac{1-M}{2} \right) \right) \right] \quad (6)$$

and impose Thermodynamics by requiring (remember  $F = E - TS$ )

$$\frac{dF}{dM} = 0 \quad \& \quad \frac{d^2F}{dM^2} \geq 0 \quad (7)$$

$$\frac{dF}{dM} = 0 \quad \rightarrow \quad M = \tanh \left( \frac{JM + h}{T} \right) \quad (8)$$

$$\frac{d^2F}{dM^2} = \chi \quad \rightarrow \quad \chi = \frac{C}{T - T_c} \quad (9)$$



We expand for small  $M$  at  $h = 0$  to see spontaneously broken symmetries  
 Memo:  $\tanh(x) \sim x - x^3/3 + O(x^5)$

Figure 3. Graph of magnetization value [13]

$$M = \tanh \left( \frac{J}{T} M \right) \sim \frac{J}{T} M - \left( \frac{J}{T} \right)^3 \frac{M^3}{3} \rightarrow M \left( \frac{J}{T} - 1 \right) \sim CM^3 \rightarrow M \propto \pm \sqrt{\frac{J}{T} - 1}$$

# Shannon Entropy = Boltzmann entropy?

$$S[\sigma] = \ln \Omega, \quad \Omega = \binom{N}{N_A} = \binom{N}{N_B} \quad (10)$$

$$= \ln \left( \frac{N!}{N_A! N_B!} \right) = \ln N! - \ln N_A! - \ln N_B! \quad (11)$$

$$\sim N \ln N - N_A \ln N_A - N_B \ln N_B - (N - N_A - N_B) = \quad (12)$$

$$= (N_A + N_B) \ln N - N_A \ln N_A - N_B \ln N_B \quad (13)$$

$$= -N_A \ln \left( \frac{N_A}{N} \right) - N_B \ln \left( \frac{N_B}{N} \right) = -N_A \ln \rho_A - N_B \ln \rho_B \quad (14)$$

by which, calling  $\rho_A = N_A/N$  and  $\rho_B = N_B/N$ , the intensive entropy reads

$$\frac{S[\sigma]}{N} = -\rho_A \ln \rho_A - \rho_B \ln \rho_B = \rho \ln \rho + (1 - \rho) \ln(1 - \rho), \quad (15)$$

$$\frac{S[\sigma]}{N} = -\left(\frac{1+M}{2}\right) \ln\left(\frac{1+M}{2}\right) - \left(\frac{1-M}{2}\right) \ln\left(\frac{1-M}{2}\right). \quad (16)$$

**When the Entropy is extremized?**  $\rho = 1/2 \rightarrow M = 0$ .

whose extremization (as expected) returns

$$\frac{d}{d\rho} \frac{S[\sigma]}{N} = 0 \rightarrow -\ln \rho - \frac{\rho}{\rho} + \ln(1 - \rho) + \frac{(1 - \rho)}{(1 - \rho)} = -\ln \left( \frac{\rho}{1 - \rho} \right) \rightarrow \rho = 1/2.$$

namely  $M = 0$ .



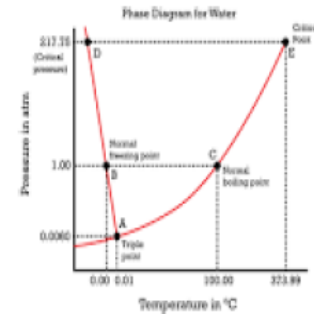
# Statistical Mechanics in a nutshell



## Statistical mechanics has its genesis in Physics (late 800/early 900)

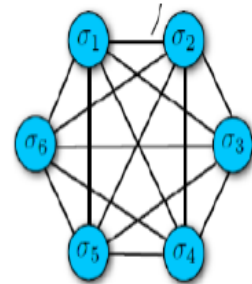
- It was born as an alternative route (w.r.t system's of ODE) to capture thermodynamic properties of large ensembles of (interacting) units
- Grounded on probabilistic techniques reveals «emerging capabilities»
- Focuses on few effective degrees of freedom (control and order parameters)

alla ricerca delle proprietà emergenti....



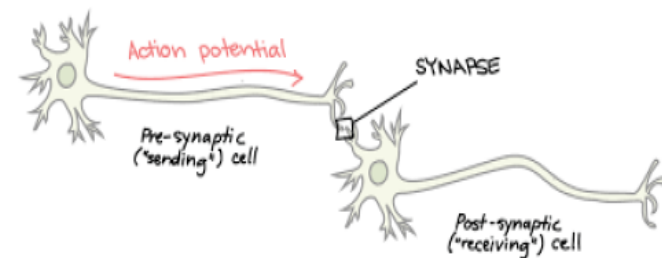
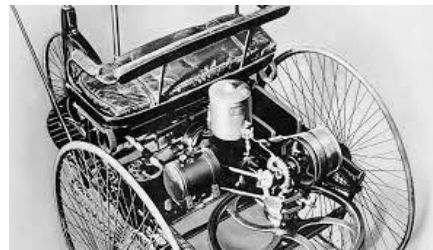
## It has then «exported» in Neural Networks (1985/2000, AGS-Theory)

- Basic Reference: discrete statistical mechanics (spin systems) [two-state neurons]
- Simple models: paramagnetic  $\rightarrow$  ferromagnetic transitions (Curie-Weiss)
- Complex models: spin glasses & Parisi Theory [synapses are both positive and negative]



(almost-deterministic) PROTOCOL:

- 1) Define the cost-function (i.e. the Hamiltonian in Physics).
- 2) Define control and order parameters
- 3) Introduce the free energy
- 4) Extremize the free energy  $F/A$  w.r.t. the order parameters to work out phase diagrams in the control parameters.



FREE ENERGY = ENERGY MINUS TEMPERATURE TIMES ENTROPY (not so different from ML...)

meccanicamente....

## 2 F in Statistical Mechanics: Mom. Gen. Func.

$$\longrightarrow E = H(\sigma|J,h) = -\frac{J}{2N} \sum_{i,j}^{N,N} \sigma_i \sigma_j - h \sum_i^N \sigma_i = -N \left( J \frac{m^2}{2} + hm \right) \quad (17)$$

$$\mathcal{P}(\{\sigma\}) = \frac{1}{Z(T,J,h)} e^{-\frac{H(\sigma|J,h)}{T}} = \frac{e^{\frac{J}{2N} \sum_{i,j}^{N,N} \sigma_i \sigma_j + h \sum_i^N \sigma_i}}{\sum_{\sigma}^{2^N} e^{\frac{J}{2N} \sum_{i,j}^{N,N} \sigma_i \sigma_j + h \sum_i^N \sigma_i}}$$

---

$$\begin{aligned} F(m|T,J,h) &= -\frac{T}{N} \ln \sum_{\sigma}^{2^N} \exp \left( -\frac{H(\sigma|J,h)}{T} \right) \\ &= -\frac{T}{N} \ln \sum_{\sigma}^{2^N} \exp \left( \frac{JN}{2} m^2 + hNm \right) \end{aligned} \quad (18)$$

- First moment: magnetization  $\frac{dF}{dh} = -\langle m \rangle$
- Second moment: susceptibility  $\frac{d^2F}{dh^2} = \chi = N (\langle m^2 \rangle - \langle m \rangle^2)$

$$\langle m \rangle = \frac{\sum_{\sigma}^{2^N} m e^{-H/T}}{\sum_{\sigma}^{2^N} e^{-H/T}}$$

$$m = \frac{1}{N} \sum_i^N \sigma_i$$



### 3 Symmetry breaking: Guerra's interpolation

Interpolation  $\lim_{N \rightarrow \infty} \mathcal{P}(m) = \delta(m - M)$  cum grano salis...

$$F(t) = -\frac{T}{N} \ln \sum_{\sigma} \exp \left[ (t) \left( \frac{J}{TN} \sum_{i < j}^{N,N} \sigma_i \sigma_j \right) + (1-t) \left( \frac{JM}{T} \sum_{i=1}^N \sigma_i \right) + h \sum_{i=1}^N \sigma_i \right] \quad (13)$$

Solution via Fundamental Theorem of Calculus

$$F(T, J, h) = F(t = 1) = \underline{F(t = 0)} + \int_0^1 \dot{F}(t') dt' \quad (14)$$

Thus we are left to two calculations to do, the *one-body* (i.e.  $F(t = 0)$ ) and the *streaming* (i.e.  $d_t F(t)$ ).

Let us start with the former, the one-body term:

$$F(t = 0) = -\frac{T}{N} \ln \sum_{\sigma} \exp \left[ \left( \frac{JM}{T} \sum_{i=1}^N \sigma_i \right) + h \sum_{i=1}^N \sigma_i \right] \quad (15)$$

$$= -\frac{T}{N} \ln \sum_{\sigma} \exp \left[ \left( \frac{JM + h}{T} \right) \sum_{i=1}^N \sigma_i \right] \quad (16)$$

$$= -\frac{T}{N} \ln \sum_{\sigma} \prod_{i=1}^N \exp \left[ \left( \frac{JM + h}{T} \right) \sigma_i \right] \quad (17)$$

$$= -T \ln \left[ 2 \cosh \left( \frac{JM + h}{T} \right) \right] \quad (18)$$

Let us focus now on the streaming, namely the evaluation of  $\int_0^1 \dot{F}(\mathbf{t}) dt$ .  
 Before proceeding note that  $m = \frac{1}{N} \sum_i \sigma_i$  thus

- where it is simple to evaluate (e.g. in the high T limit)  $\frac{1}{\sqrt{N}} \sum_i^N \sigma_i \sim Z[0, 1]$
- thus  $\lim_{N \rightarrow \infty} \mathcal{P}(m) = \delta(m - M) \quad m = \frac{1}{N} \sum_i^N \sigma_i$
- or equivalently  $\lim_{N \rightarrow \infty} \langle (m - M)^2 \rangle = 0$  that means  $\langle m^2 \rangle - 2M\langle m \rangle = -M^2$

Let us start now with the derivative of the interpolating free energy, that reads

$$F(t) = -\frac{T}{N} \ln \sum_{\sigma} 2^N \exp \left[ (t) \left( \frac{J}{TN} \sum_{i < j}^{N,N} \sigma_i \sigma_j \right) + (1-t) \left( \frac{JM}{T} \sum_{i=1}^N \sigma_i \right) + h \sum_{i=1}^N \sigma_i \right] \quad (19)$$

hence its t-derivative reads (in the thermodynamic limit  $N \rightarrow \infty$ )

$$\frac{dF}{dt} = -\frac{J}{2} \left( \langle m^2 \rangle - 2M\langle m \rangle \right) = \frac{JM^2}{2} \quad (20)$$

that does no longer depends on t and, thus, its integral in  $t \in (0, 1)$  is just multiplication by one.

Pasting the two pieces together, we can write

$$F(m|T, J, h) = F(t=1) = F(t=0) + \int_0^1 \dot{F}(t') dt' \quad (21)$$

$$= -T \ln \left[ 2 \cosh \left( \frac{JM+h}{T} \right) \right] + \frac{JM^2}{2} \quad (22)$$

Thus we can finally extremize the latter to obtain

$$\frac{dF}{dM} = 0 \quad \rightarrow \quad \boxed{\tanh \left( \frac{JM+h}{T} \right) = M.} \quad (23)$$

### What can we say on the Susceptibility? Gaussian Fluctuation Theory

Memo:

$$Z = \sum_{\sigma}^{2^N} \exp \left( t \left( \frac{Nm^2}{2T} \right) + (1-t) \frac{NMm}{T} \right)$$

- We first rescale the magnetization properly, i.e.  $\tilde{m} = \sqrt{N}(m - M)$
- We then prove that  $\frac{d}{dt} \langle f(\sigma) \rangle = \frac{1}{2T} (\langle f(\sigma) \tilde{m}^2 \rangle - \langle f(\sigma) \rangle \langle \tilde{m}^2 \rangle)$
- use it with the choice  $f(\sigma) = \tilde{m}^2 \rightarrow \frac{d}{dt} \langle \tilde{m}^2 \rangle = \frac{1}{2T} \langle \tilde{m}^2 \rangle^2$
- the solution of the ODE is

$$\boxed{\langle \tilde{m}^2 \rangle = \frac{C}{1 - t \frac{J}{T}} \rightarrow_{t=1} \frac{C'}{T - J}}$$

## 4 Symmetry breaking as bifurcation: PDE approach

It is useful to introduce  $A(T, J, h) = -\frac{F(T, J, h)}{T}$  such that

$$A(T, J, h) = \frac{1}{N} \ln \sum_{\sigma} \exp \left( \left( \frac{J}{T} \frac{N}{2} m^2 + \frac{h}{T} Nm \right) \right) \quad (24)$$

and pave the *Mechanical Analogy* (note: these are all Joules by Legendre)

$$A(t, x) = \frac{1}{N} \ln \sum_{\sigma} \exp \left( -t \frac{N}{2} m^2 + x Nm \right) \quad (25)$$

such that it is trivial to check that

$$\frac{\partial A}{\partial t} = \frac{1}{N} \frac{1}{Z} \sum_{\sigma} \left( -\frac{1}{2} Nm^2 \right) e^{-H/T} = -\frac{1}{2} \langle m^2 \rangle, \quad (26)$$

$$\frac{\partial A}{\partial x} = \frac{1}{N} \frac{1}{Z} \sum_{\sigma} (m) e^{-H/T} = \langle m \rangle. \quad (27)$$

As a consequence, if I introduce a potential  $V_N(t, x) = \frac{1}{2} (\langle m \rangle^2 - \langle m^2 \rangle)$  by construction the following HJ PDE holds (even at finite volumes)

$$\frac{\partial A}{\partial t} + \frac{1}{2} \left( \frac{\partial A}{\partial x} \right)^2 + V_N(t, x) = 0. \quad (28)$$

$$\longrightarrow A(t, x) = \frac{1}{N} \ln \sum_{\sigma} \exp \left( -t \frac{N}{2} m^2 + xNm \right) \quad (23)$$

$$\longrightarrow \frac{\partial A}{\partial t} + \frac{1}{2} \left( \frac{\partial A}{\partial x} \right)^2 + V_N(t, x) = 0. \quad (26)$$

Now, as  $\lim_{N \rightarrow \infty} V_N(t, x) = 0$ , in the thermodynamic limit we are left with a Galilean motion, where the *velocity* of the unitary mass is  $\langle m \rangle$  and the related trajectory reads as  $x(t) = x_0 + \langle m \rangle t$ .

The solution of the HJ PDE is the Cauchy condition  $A(t = 0, x = x_0)$  plus the integral of the Lagrangian  $\mathcal{L}$  over time (and the Lagrangian is just the kinetic term), namely

$$A(t, x) = A(t = 0, x = x_0) + \int_0^t \mathcal{L} dt \quad (27)$$

We have thus to perform these two evaluations, that read the former

$$A(t = 0, x = x_0) = \frac{1}{N} \ln \sum_{\sigma} \exp(x_0 Nm) = \frac{1}{N} \ln \sum_{\sigma} \exp \left( x_0 \sum_i^N \sigma_i \right) \quad (28)$$

$$= \ln 2 + \ln \cosh(x_0) = \ln 2 + \ln \cosh(x(t) - \langle m \rangle t) \quad (29)$$



the latter

$$\int_0^t \mathcal{L} dt = \frac{1}{2} \langle m^2 \rangle t \quad (30)$$

Merging the two together, we can write

$$A(t, x) = \ln 2 + \ln \cosh(x(t) - \langle m \rangle t) + \frac{1}{2} \langle m^2 \rangle t, \quad (31)$$

$$A\left(t = -\frac{J}{T}, x = \frac{h}{T}\right) = \ln 2 + \ln \cosh\left(\frac{h}{T} + \frac{J}{T} \langle m \rangle\right) - \frac{J}{2T} \langle m^2 \rangle \quad (32)$$

namely

$$A(T, J, h) = \ln 2 + \ln \cosh\left(\frac{h + JM}{T}\right) - \frac{J}{2T} M^2, \quad (33)$$

$$d_M A = 0 \rightarrow M = \tanh\left(\frac{h + JM}{T}\right). \quad (34)$$

...è il principio di Minima Azione nel formalismo della Meccanica Analitica!

If we work already in the TD limit we can write a Riemann-Hopf PDE for the magnetization

$$\frac{\partial A}{\partial x} = \frac{1}{N} \frac{1}{Z} \sum_{\sigma}^{2^N} \langle m \rangle e^{-H/T} = \langle m \rangle.$$

$$\frac{\partial}{\partial x} \left( \frac{\partial A}{\partial t} + \frac{1}{2} \left( \frac{\partial A}{\partial x} \right)^2 \right) = 0 \rightarrow \boxed{\frac{\partial \langle m \rangle}{\partial t} + \langle m \rangle \frac{\partial \langle m \rangle}{\partial x} = 0} \quad (37)$$

that is the harmonic oscillator of a Hopf (or Hopf-Lax) bifurcation. Indeed we can inspect mass conservation to see that

$$\rho(x)dx = \rho(y)dy$$

and, by the Galilean equation of motion we can write (at  $h = 0$ )

$$x = y + \langle m \rangle t = y + \tanh\left(\frac{J}{T} \langle m \rangle\right) t$$

such that the Jacobian reads

$$\frac{dx}{dy} = 1 + \partial_y \langle m \rangle t = 1 + (1 - \tanh^2\left(\frac{J}{T} \langle m \rangle\right)) t$$

thus for the mass density at a generic point  $x$  we get, from the high temperature (where  $\langle m \rangle = 0$ )

$$\rho(x) = \rho(y) \frac{1}{1 + (1 - \tanh^2\left(\frac{J}{T} \langle m \rangle\right)) t} = \rho(y) \frac{1}{1 + t} = \rho(y) \frac{1}{1 - J/T}$$

namely there is a pole at  $T_c = J$ , where the spin-flip symmetry becomes spontaneously broken (and, accordingly, a shock happens in this dual representation).

## 4 Relativity beyond Statistical Reductionism

$$A(t, x) = \frac{1}{N} \ln \sum_{\sigma} \exp \left( -t \frac{N}{2} m^2 + xNm \right) \quad (38)$$

There are two points in the previous PDE approach that *smell relativistic* namely

- the velocity is the magnetization, that is bounded by 1 (and in Classical Mechanics velocity is not bounded).
- The metric tensor is Lorenzian rather than Euclidean (i.e.  $-t, +x$  rather than  $+t, +x$ )

the whole suggests to move from  $E = -\frac{1}{2}m^2$  to  $E = \sqrt{1+m^2} = 1 - \frac{1}{2}m^2 + O(m^4)$ .

- This is the analogous of considering solely binary collisions when using particles
- This is deeply related to the fact that if forces are linear, then energies are quadratic forms

L'energia totale relativistica  $E$ , nella teoria della relatività ristretta, è data dall'Energia a riposo  $E_0 = mc^2$  a cui va aggiunta l'Energia cinetica relativistica  $K = (\gamma - 1) mc^2$  :

$$E = E_0 + K = mc^2 + (\gamma - 1) mc^2 = \gamma mc^2 = \sqrt{p^2 c^2 + m^2 c^4}$$

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# Quantifying heterogeneity to drug response in cancer-stroma kinetics

Francesco Alemanno<sup>ab,1</sup>, Marta Cavo<sup>a,1</sup>, Donatella Delle Cave<sup>c</sup>, Alberto Fachechi<sup>d</sup>, Riccardo Rizzo<sup>a</sup>, Eliana D'Amone<sup>a</sup>, Giuseppe Gigli<sup>ab</sup>, Enza Lonardo<sup>c</sup>, Adriano Barra<sup>be,2</sup>, and Loretta L. del Mercato<sup>a,2</sup>

Edited by Michele Castellana, Institut Curie, Paris, France; received December 13, 2021; accepted February 4, 2023, by Editorial Board Member Mehran Kardar



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# Probing Single-Cell Fermentation Fluxes and Exchange Networks via pH-Sensing Hybrid Nanofibers

Valentina Onesto, Stefania Forciniti, Francesco Alemanno, Krishnadev Narayanankutty, Anil Chandra, Saumya Prasad, Amalia Azzariti, Giuseppe Gigli, Adriano Barra, Andrea De Martino, Daniele De Martino,\* and Loretta L. del Mercato\*

Cite This: *ACS Nano* 2023, 17, 3313–3323

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# Fully Automated Computational Approach for Precisely Measuring Organelle Acidification with Optical pH Sensors

Anil Chandra, Saumya Prasad, Francesco Alemanno, Maria De Luca, Riccardo Rizzo, Roberta Romano, Giuseppe Gigli, Cecilia Bucci, Adriano Barra,\* and Loretta L. del Mercato\*

Cite This: *ACS Appl. Mater. Interfaces* 2022, 14, 18133–18149

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OPEN

# Cancer-driven dynamics of immune cells in a microfluidic environment

SUBJECT AREAS:  
COMPUTATIONAL BIOPHYSICS

Elena Agliari<sup>1</sup>, Elena Biselli<sup>1</sup>, Adele De Ninno<sup>2</sup>, Giovanna Schiavoni<sup>2</sup>, Lucia Gabriele<sup>3</sup>, Anna Gerardino<sup>4</sup>, Fabrizio Mitrilli<sup>3</sup>, Adriano Barra<sup>1</sup> & Luca Ravinno<sup>4</sup>

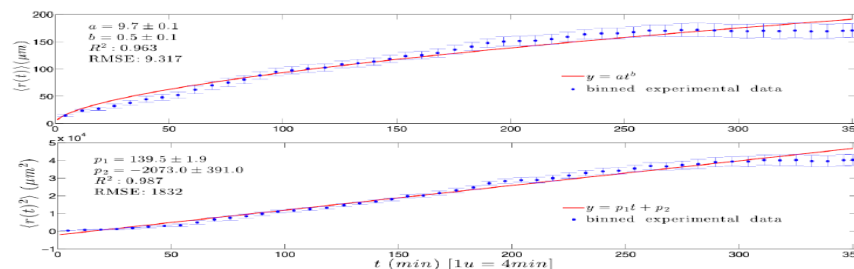
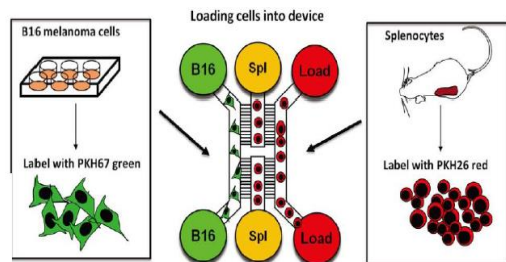
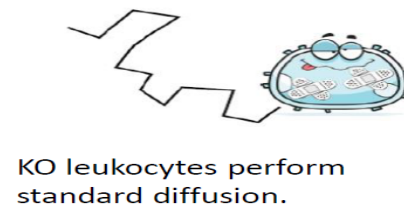


Figure 9 | Mean displacement  $\langle r(t) \rangle$  (upper panel) and mean squared displacement  $\langle r^2(t) \rangle$  (lower panel) for KO splenocytes. Experimental (binned data) with standard errors are compared with best fits (solid line) whose coefficients are properly shown.



# OK (WT) leukocytes perform ballistic motion: they perceive the chemoattractant gradient.

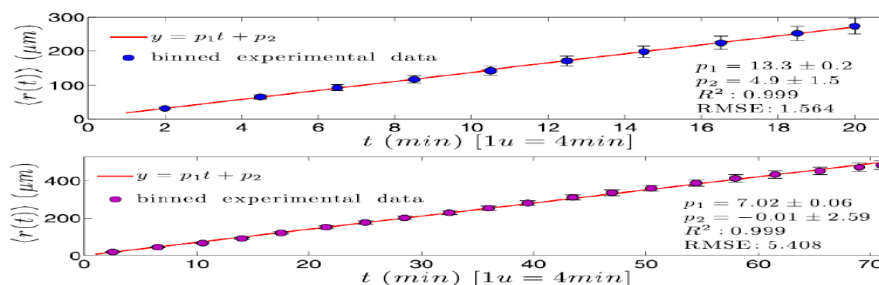
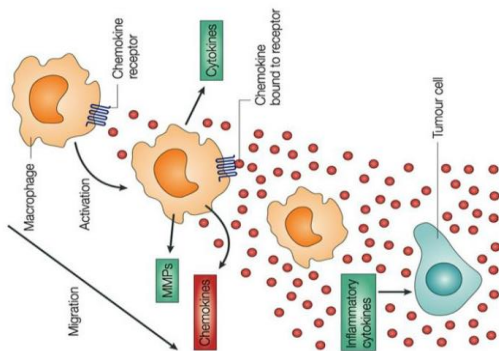
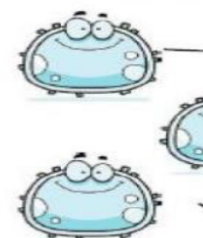
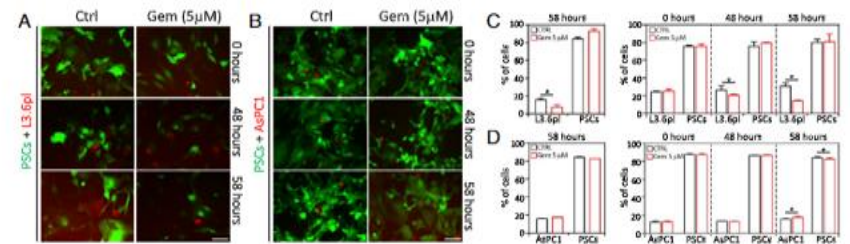
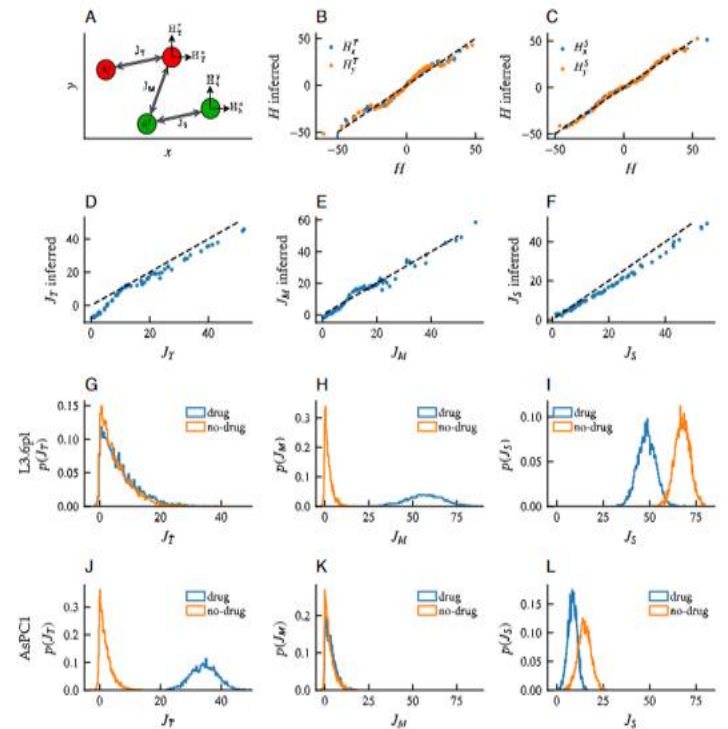
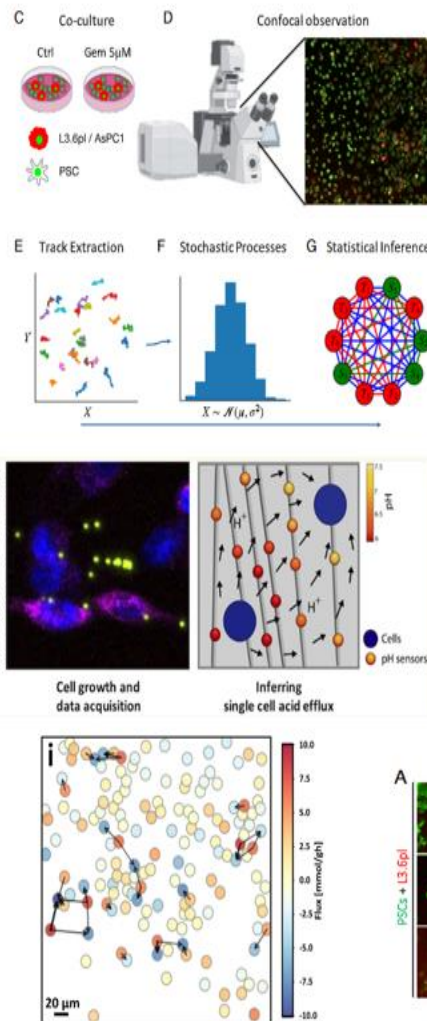
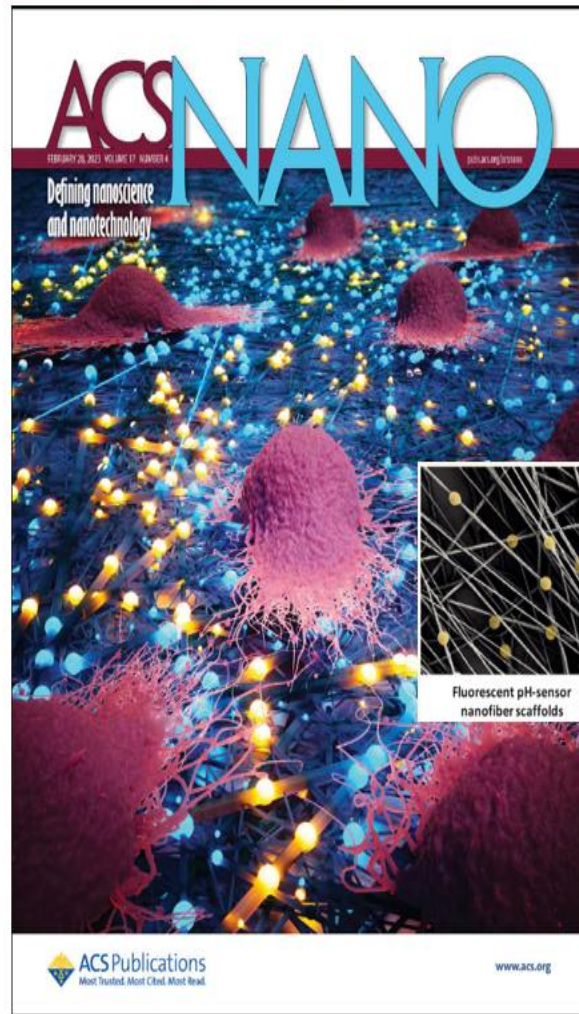


Figure 16 |  $\langle r(t) \rangle$  versus  $t$  for WT-PRE splenocytes (upper panel) and for WT-POST splenocytes (lower panel). As expected, the mean displacement grows linearly with time. Binned data (•) with standard errors are compared with best fit (solid line), whose coefficients are also reported.



## Theoretical Division [TD] (Dipartimento di Matematica e Fisica, Università del Salento)





# une petite parenthèse historique: is Boltzmann = Shannon for these systems?

Vedremo che partendo dalle definizione di entropia data da Boltzmann arriviamo a quella di Shannon.

Il numero di configurazioni possibili è  $\Omega = \frac{N!}{N_A!N_B!}$  da cui possiamo definire l'entropia come

$$S \sim \ln \Omega = \ln N! - \ln N_A! - \ln N_B!$$

Per  $N \rightarrow \infty$  usiamo Stirling ed otteniamo

$$S \sim N \ln N - N_A \ln N_A - N_B \ln N_B + (-N + N_A + N_B).$$

A questo punto aggiungiamo la quantità nulla  $[(N_A + N_B)\ln N - N \ln N]$ . Quindi possiamo riscrivere

$$S \sim N_A \ln \rho_A - N_B \ln \rho_B$$

dove abbiamo introdotto le densità  $\rho_A = N_A/N$  e  $\rho_B = N_B/N$ . Adesso dividiamo tutto per  $N$  (per avere l'espressione intensiva) e troviamo

$$s = \frac{S}{N} = -\rho_A \ln \rho_A - \rho_B \ln \rho_B$$

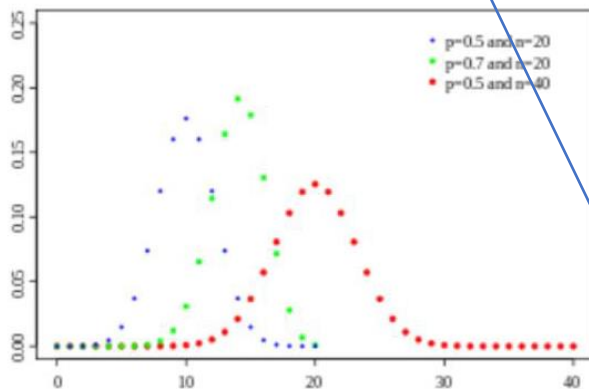
Un altro concetto importante:

QUANTITA' INTENSIVE VS QUANTITA' ESTENSIVE

$$\mu = \sum_{x=0}^n x P_{\text{bin}}(x) = np$$

$$\sigma = \sqrt{np(1-p)}$$

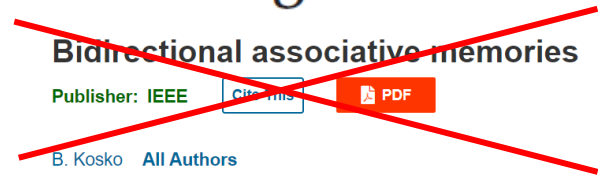
E quindi troviamo che  $\rho_{eq} = \frac{1}{2}$ .



i.e.  $M=0$

# From Pavlov Conditioning to Hebb Learning

What do we need?



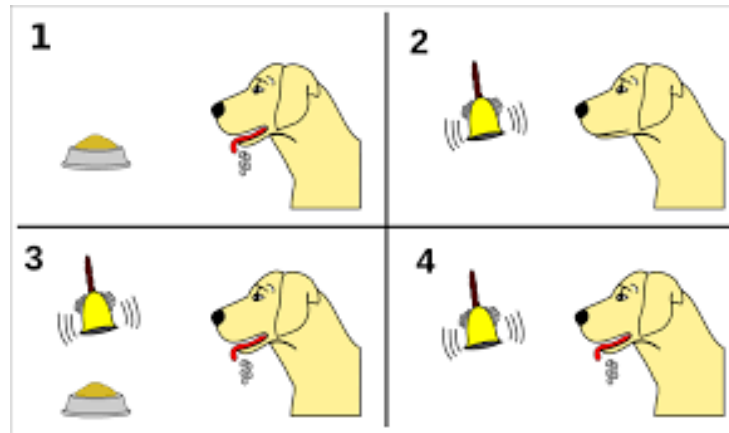
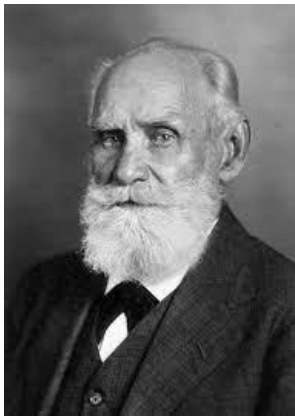
- 1) A reference model in glassy statistical mechanics:  
Switch from serial processing (standard AGS theory) to parallel processing (modern theory)

Why standard Hopfield model can not be the correct reference?

- 1) A toy model to show that «*neurons that fire together wire together*» (1-bit prescription)

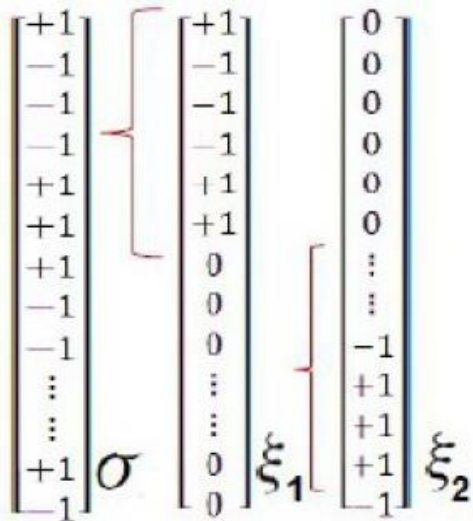
Guerra scheme for 1-bit Classical Conditioning

- 1) The generalization from bits to informative patterns (e.g. ringing bell and food for dogs)



- 2) A proof that such a generalization converges to the Hebbian kernel

# Multi-tasking associative networks: patterns with diluted entries



$$\langle H \rangle = E = - \sum_{\mu} m_{\mu}^2, \quad \text{emergent parallel processing is the most convenient}$$

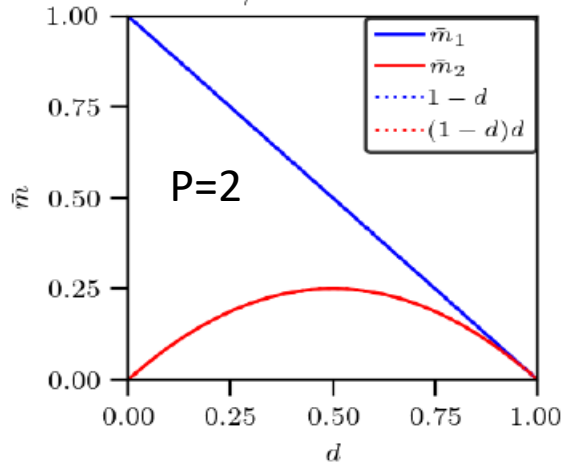
$$E_{hp1} = -m_1^2 - m_2^2 = -\left(\frac{1}{2}\right)^2 + \text{random} = -.25,$$

$$E_{hp2} = -m_1^2 - m_2^2 = -\left(\frac{1}{2}\right)^2 - \left(\frac{1}{2}\right)^2 = -.50.$$

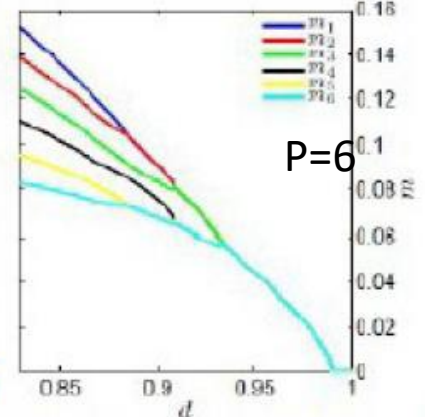
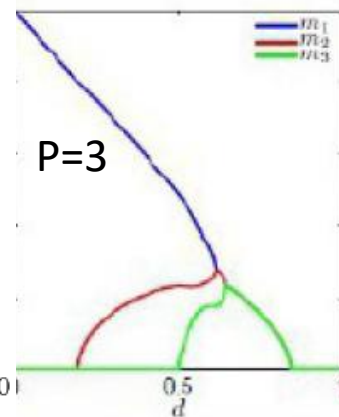
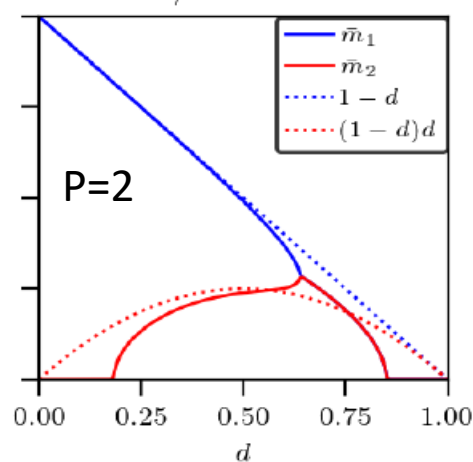
$$m_1 = d(1-d) \tanh(\beta m_1) + \frac{(1-d)^2}{2} \{ \tanh[\beta(m_1 + m_2)] + \tanh[\beta(m_1 - m_2)] \}, \quad (2.6)$$

$$m_2 = d(1-d) \tanh(\beta m_1) + \frac{(1-d)^2}{2} \{ \tanh[\beta(m_1 + m_2)] - \tanh[\beta(m_1 - m_2)] \} \quad (2.7)$$

$\beta = 1000.0$



$\beta = 6.66$





## Multitasking Associative Networks

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(Received 25 February 2012; published 26 December 2012)

## Extensive Parallel Processing on Scale-Free Networks

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## Retrieval Capabilities of Hierarchical Networks: From Dyson to Hopfield

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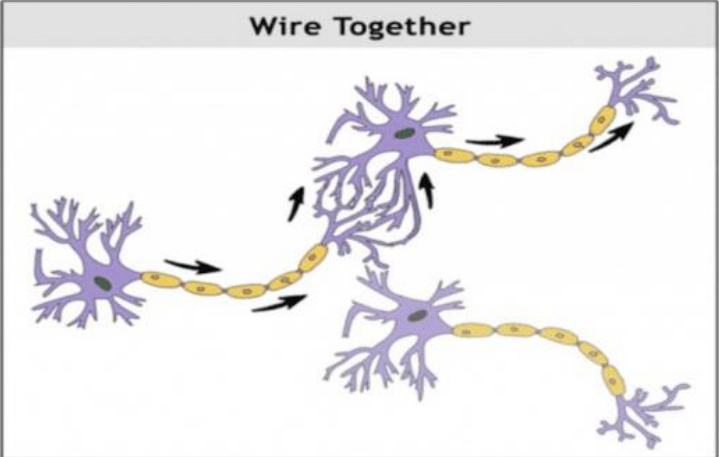
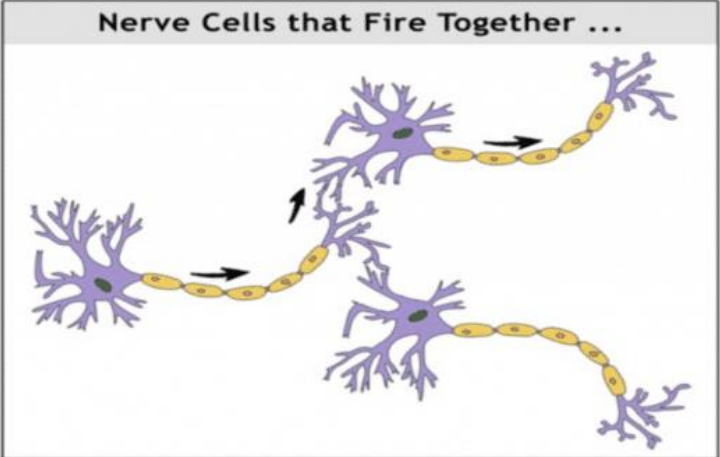
(Received 9 July 2014; published 16 January 2015)

Pavlov's Classical Conditioning	Hebb's learning
Correlations between concepts	Correlations between pairs of neurons
Patterns of bits	Single bits
Neutral stimuli associated with unconditioned stimuli	simultaneous activation of cells increases the synaptic strength.
Synapses store the correlations between concepts	Synaptic modification by correlated activity

} **Different Degrees of information**

} **Learning by association**

} **Synaptic modification**



## Neural and Synaptic Dynamics

The evolution equation of the expectation value of the spins and synaptic matrix over time:

$$\bar{J}_{ij} := \sum_{\sigma \in \Sigma_N} \sum_{J \in \mathcal{J}_N} \mathcal{P}_N(\sigma, J, t) J_{ij}, \quad \bar{\sigma}_i := \sum_{\sigma \in \Sigma_N} \sum_{J \in \mathcal{J}_N} \mathcal{P}_N(\sigma, J, t) \sigma_i$$

### Dynamical equations

$$\tau \frac{d\bar{\sigma}_i}{dt} = \sum_{J, J' \in \mathcal{J}_N} \sum_{\sigma, \sigma' \in \Sigma_N} \sigma_i \left[ \mathcal{P}_N(\sigma', J', t) w_{N, \beta}(J', \sigma' \rightarrow J, \sigma) - \mathcal{P}_N(\sigma, J, t) w_{N, \beta}(J, \sigma \rightarrow J', \sigma') \right]$$

$$\tau' \frac{d\bar{J}_{ij}}{dt} = \sum_{J, J' \in \mathcal{J}_N} \sum_{\sigma, \sigma' \in \Sigma_N} J_{ij} \left[ \mathcal{P}_N(\sigma', J', t) w_{N, \beta}(J', \sigma' \rightarrow J, \sigma) - \mathcal{P}_N(\sigma, J, t) w_{N, \beta}(J, \sigma \rightarrow J', \sigma') \right]$$

Where  $\tau$  and  $\tau'$  are respectively the typical timescale for the neural and synaptic dynamics

After some tedious calculations we get

*Note: self-consistency at equilibrium (lhs=0).*

$$\tau \frac{d\bar{\sigma}_i}{dt} = -\bar{\sigma}_i + \tanh \left( \beta \sum_j \bar{J}_{ij} \bar{\sigma}_j + \beta h_i \right), \quad \tau' \frac{d\bar{J}_{ij}}{dt} = -\bar{J}_{ij} + \tanh(\beta) \bar{\sigma}_i \bar{\sigma}_j$$

# Dynamical equations

$$\tau \frac{d\bar{\sigma}_i}{dt} = \sum_{J, J' \in \mathcal{J}_N} \sum_{\sigma, \sigma' \in \Sigma_N} \sigma_i \left[ \mathcal{P}_N(\sigma', J', t) w_{N, \beta}(J', \sigma' \rightarrow J, \sigma) - \mathcal{P}_N(\sigma, J, t) w_{N, \beta}(J, \sigma \rightarrow J', \sigma') \right]$$

$$\tau' \frac{d\bar{J}_{ij}}{dt} = \sum_{J, J' \in \mathcal{J}_N} \sum_{\sigma, \sigma' \in \Sigma_N} J_{ij} \left[ \mathcal{P}_N(\sigma', J', t) w_{N, \beta}(J', \sigma' \rightarrow J, \sigma) - \mathcal{P}_N(\sigma, J, t) w_{N, \beta}(J, \sigma \rightarrow J', \sigma') \right]$$

Where  $\tau$  and  $\tau'$  are respectively the typical timescale for the neural and synaptic dynamics

After some tedious calculations we get

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## Discrete dynamical equations

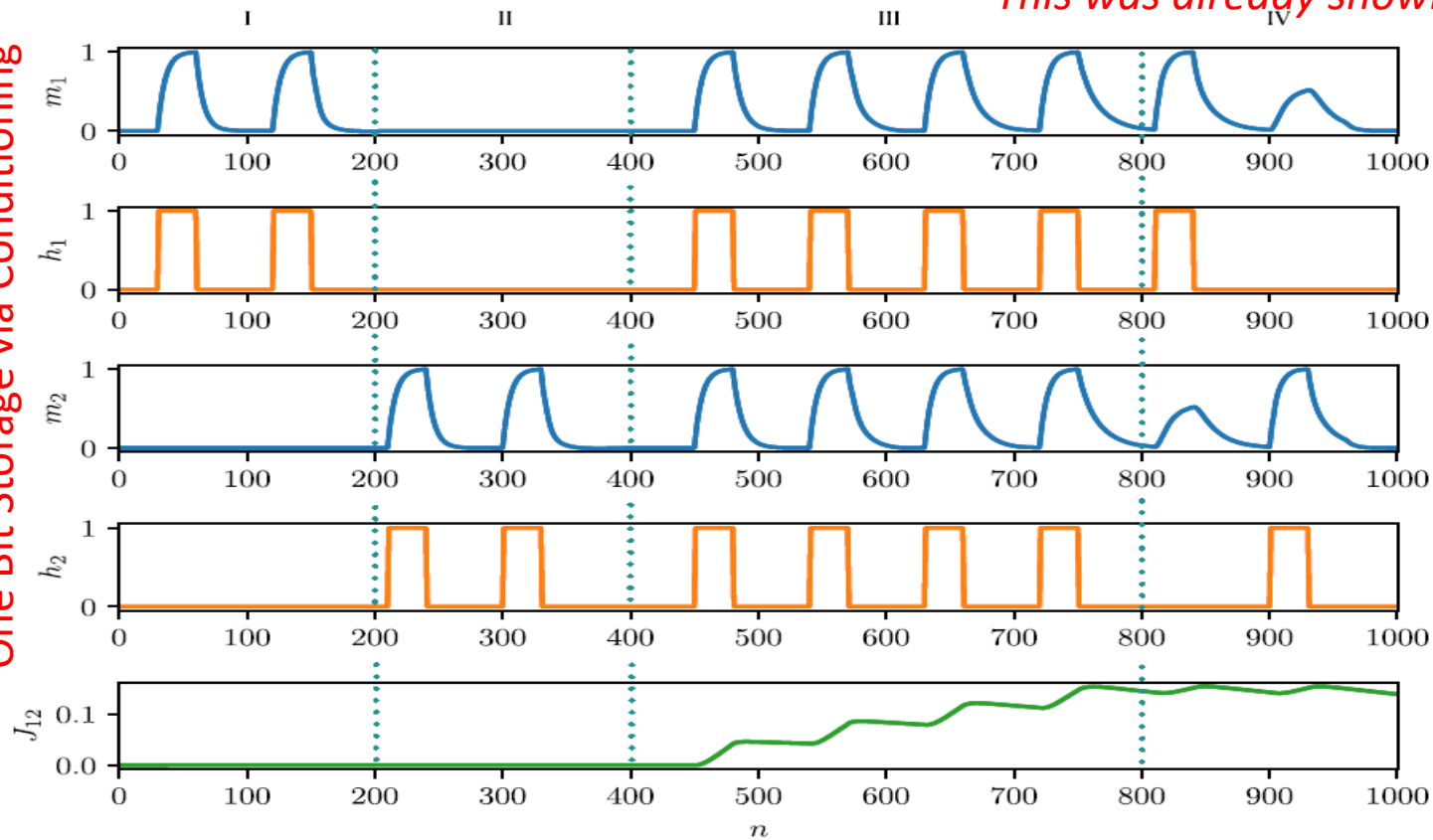
$$\bar{\sigma}_i^{(n+1)} = \bar{\sigma}_i^{(n)} \left( 1 - \frac{\delta t}{\tau} \right) + \frac{\delta t}{\tau} \tanh \left( \beta \sum_{i \neq j} \bar{J}_{ij}^{(n)} \bar{\sigma}_j^{(n)} + u \beta h_i^{(n)} \right) \quad \bar{J}_{ij}^{(n+1)} = \bar{J}_{ij}^{(n)} \left( 1 - \frac{\delta t}{\tau'} \right) + \frac{\delta t}{\tau'} \bar{\sigma}_j^{(n)} \bar{\sigma}_i^{(n)} \tanh \beta,$$

We can choose  $\delta t = c\tau$ ,  $c \in (0, 1]$  such that the dynamics depends on  $c$  and  $\tau/\tau'$

$$\tau \frac{d\bar{\sigma}_i}{dt} = -\bar{\sigma}_i + \tanh \left( \beta \sum_j \langle J_{ij} \rangle \bar{\sigma}_j + \beta h_i \right) \quad \tau' \frac{d\bar{J}_{ij}}{dt} = -\bar{J}_{ij} + \tanh(\beta) \bar{\sigma}_i \bar{\sigma}_j.$$

*This was already shown by Guerra in 1990.*

One Bit Storage via Conditioning



$$\tau \sim 1 \text{ ms}$$

$$\frac{\tau}{\tau'} \ll 1$$

$$\tau' \sim h$$

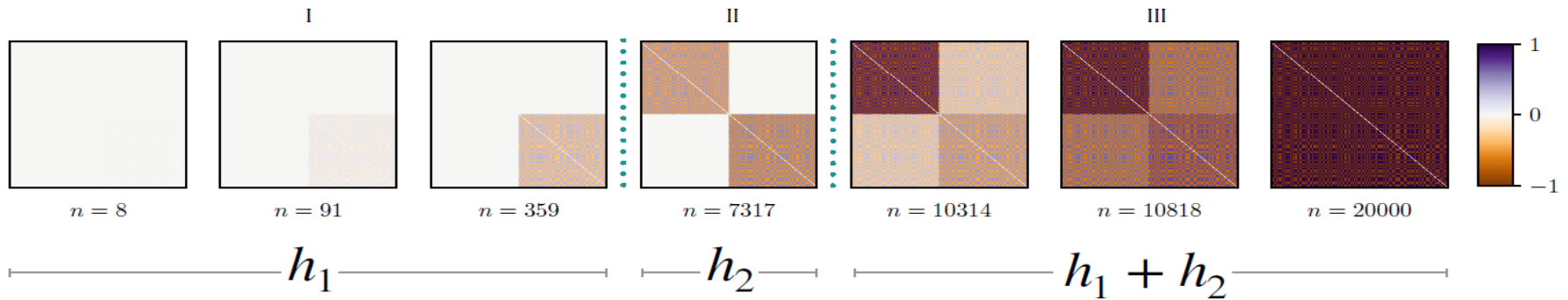
At the single-bit level Hebb's prescription is recovered. Now we move to concepts (whose length is  $N/2$ ) to inspect Pavlov!

$$\xi^1 = \left( \xi_1^1, \xi_2^1, \dots, \xi_{\frac{N}{2}}^1, 0, 0, \dots, 0 \right),$$

$$\xi^2 = \left( 0, 0, \dots, 0, \xi_{\frac{N}{2}+1}^2, \xi_{\frac{N}{2}+2}^2, \dots, \xi_N^2 \right).$$



# Emergence of concept's correlations by Classical Conditioning



**II:**

$$J(\{\xi^1, \xi^2\}) = \frac{1}{2} \times \begin{bmatrix} J^1 & \mathbf{0} \\ \mathbf{0} & J^2 \end{bmatrix}, \quad J_{ij}(\{\xi^1, \xi^2\}) = \frac{1}{2} \times \begin{cases} \xi_i^1 \xi_j^1 & (i, j) \in B_1 \\ \xi_i^2 \xi_j^2 & (i, j) \in B_2 \\ 0 & (i, j) \vee (j, i) \in B_{mix} \end{cases} .$$

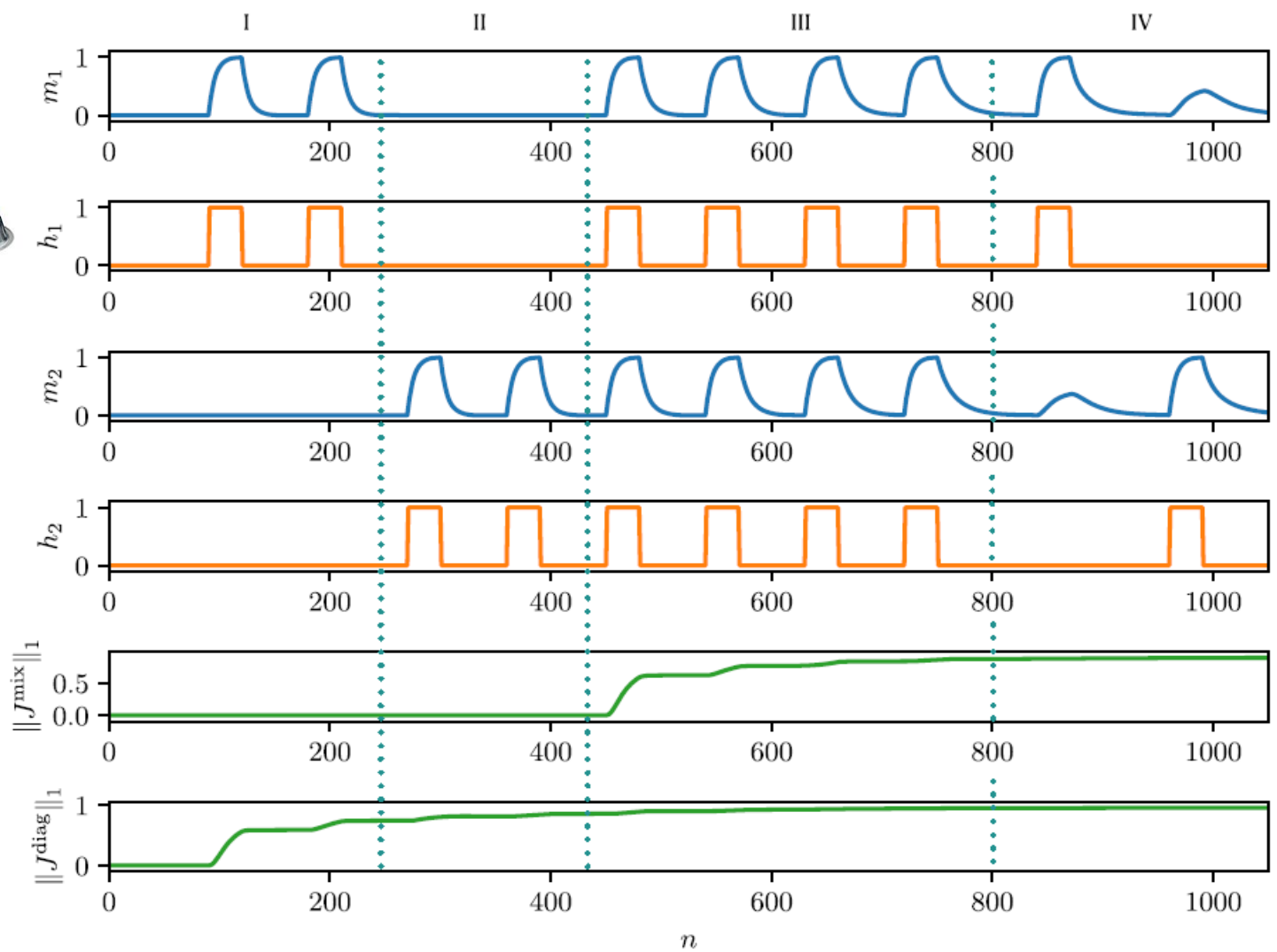
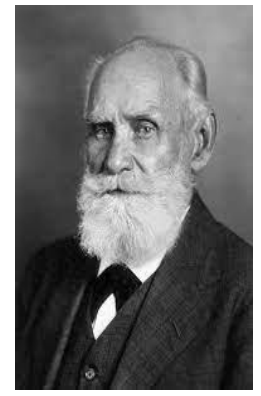
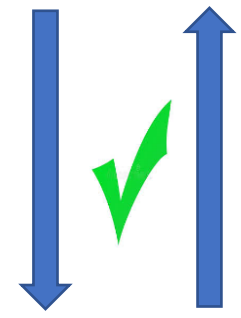
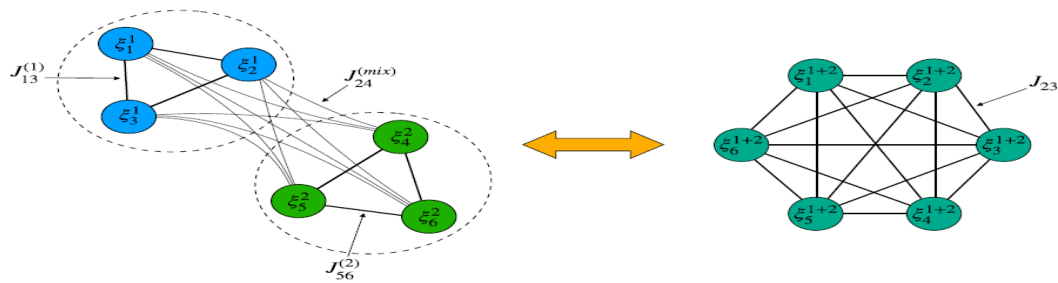
$$B_1 = [1, N/2] \times [1, N/2], B_2 = [N/2 + 1, N] \times [N/2 + 1, N], B_{mix} = [1, N/2] \times [N/2 + 1, N] .$$

**III:**

$$J(\{\xi^1, \xi^2\}) = \frac{1}{2} \times \begin{bmatrix} J^1 & J^{(mix)} \\ J^{(mix)} & J^2 \end{bmatrix}, \quad J_{ij}(\xi^{1+2}) = \begin{cases} \xi_i^1 \xi_j^1 & (i, j) \in B_1 \\ \xi_i^2 \xi_j^2 & (i, j) \in B_2 \\ \xi_i^1 \xi_j^2 & (i, j) \in B_{mix} \\ \xi_j^1 \xi_i^2 & (j, i) \in B_{mix} \end{cases} .$$

The synaptic matrix converges to the temporal mean of the stimuli  $\bar{J}_{ij}(t) \approx \frac{1}{t} \int_0^t \xi_i(t') \xi_j(t') dt'$





# Point Three: Classical Conditioning among two concepts. Guerra's argument on multi-tasking nets.

$$q_{\text{diag}}^{(n)} := \frac{1}{2} \sum_{k=1}^2 \frac{\sum_{(i,j) \in B_k} J_{ij}^k J_{ij}^{(n)}}{\sum_{(l,m) \in B_k} J_{lm}^k J_{lm}^k},$$

$$q_{\text{mix}}^{(n)} := \frac{\sum_{(i,j) \in B_{\text{mix}}} J_{ij}^{\text{mix}} J_{ij}^{(n)}}{\sum_{(l,m) \in B_{\text{mix}}} J_{lm}^{\text{mix}} J_{lm}^{\text{mix}}}.$$

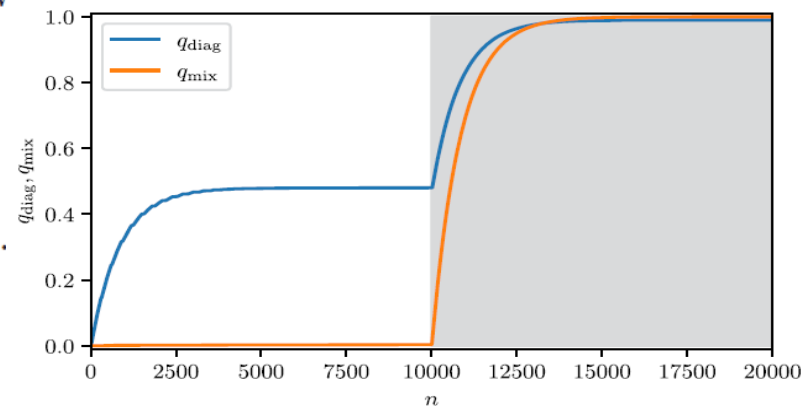
As for the off-diagonal blocks, since there are no correlations among the related neurons, they contain no information. Next, we present both stimuli simultaneously: the overall stimulation provided to the network is

$$\xi^{1+2} = \xi^1 + \xi^2 = (\xi_1^1, \xi_2^1, \dots, \xi_{\frac{N}{2}}^1, \xi_{\frac{N}{2}+1}^2, \xi_{\frac{N}{2}+2}^2, \dots, \xi_N^2),$$

and it is retained long enough to allow synapses to be plastic. synaptic matrix converges to

$$J(\xi^{1+2}) = \begin{pmatrix} J^1 & J^{\text{mix}} \\ J^{\text{mix}} & J^2 \end{pmatrix},$$

$$J_{ij}(\xi^{1+2}) = \begin{cases} \xi_i^1 \xi_j^1 & (i, j) \in B_1 \\ \xi_i^2 \xi_j^2 & (i, j) \in B_2 \\ \xi_i^1 \xi_j^2 & (i, j) \in B_{\text{mix}} \\ \xi_j^1 \xi_i^2 & (j, i) \in B_{\text{mix}}. \end{cases}$$



(3.8)

$$\xi^1 = (a, b, 0, 0)$$

$$\xi^2 = (0, 0, c, d)$$

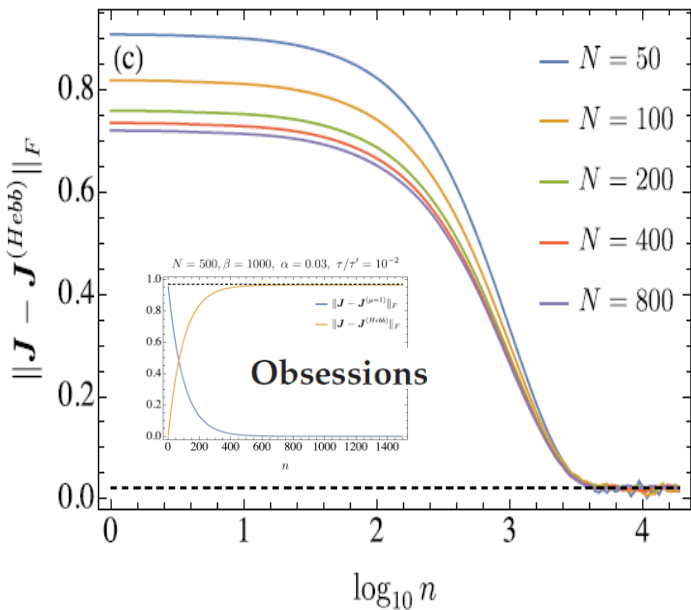
$$\xi^C = (a, b, c, d)$$

$$J_{\text{Hebb}} = \xi_i^C \xi_j^C \longrightarrow$$

$$\begin{bmatrix} 0 & ab & ac & ad \\ ba & 0 & bc & bd \\ ca & cb & 0 & cd \\ da & db & dc & 0 \end{bmatrix}$$



$\beta = 100, \alpha = 0.06, \tau/\tau' = 10^{-3}$



# Neural Computation

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## From Pavlov Conditioning to Hebb Learning

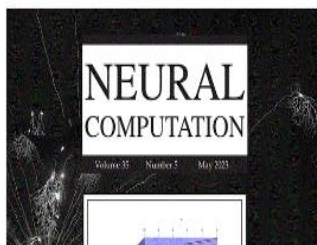
In Special Collection: CogNet

Elena Agliari, Miriam Aquaro, Adriano Barra, Alberto Fachechi, Chiara Marullo

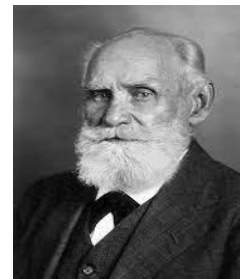
Author and Article Information

Neural Computation (2023) 35 (5): 930–957.

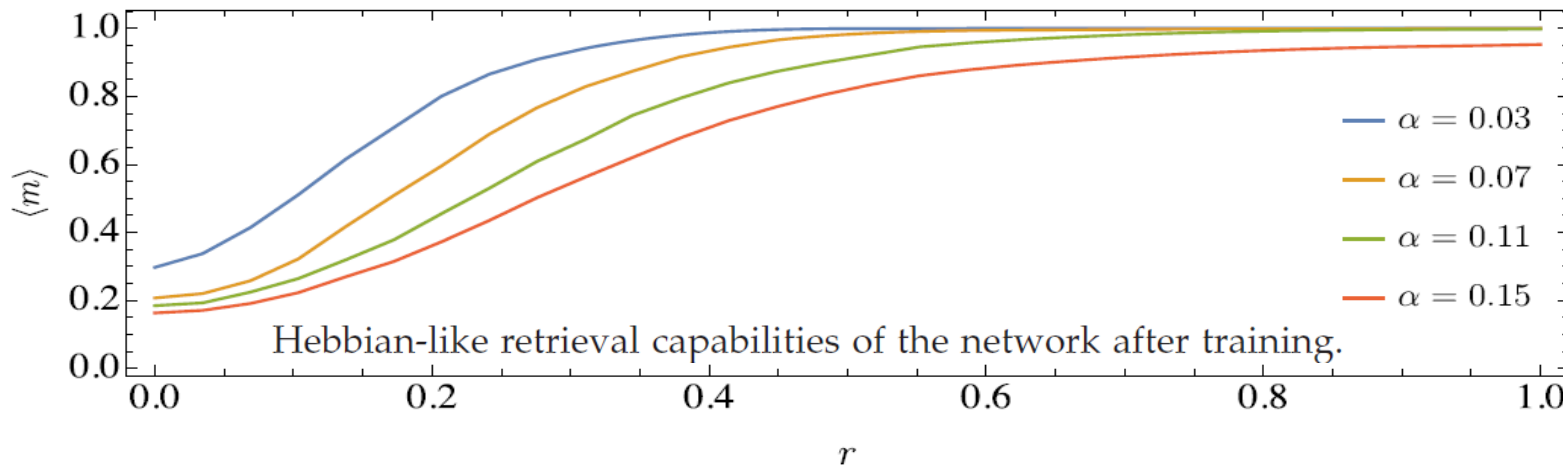
[https://doi.org/10.1162/neco\\_a\\_01578](https://doi.org/10.1162/neco_a_01578)



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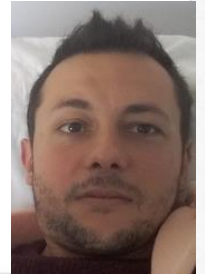


$N = 150, \alpha = 0.05, \tau/\tau' = 10^{-3}$



Hebbian-like retrieval capabilities of the network after training.

# acknowledgement



Brain-inspired ULtra-fast & ULtra-sharp machines for AI-assisted health-care  
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**THANK YOU FOR YOUR GENTLE ATTENTION!**

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