Neural Fields: from vision to consciousness A prospective mathematical perspective

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Outline

Introduction

Mathematical study of the NF equations

Visual orientation

Stochastic Neural Fields

Consciousness

Conclusion

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Introduction

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Predictive mathematical models for Neuroscience

 The goal of mathematical neuroscience is to develop predictive, falsifiable theories (Karl Popper).



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Predictive mathematical models for Neuroscience

- The goal of mathematical neuroscience is to develop predictive, falsifiable theories (Karl Popper).
- One of the most famous ones is found in the work of H. Wilson and J. Cowan (1972-1973)



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Work of H. Wilson and J. Cowan



Neural masses (Biophysical Journal 1972) Neural fields (Kybernetik 1973)

EXCITATORY AND INHIBITORY INTERACTIONS IN

INTERACTIONS IN

LOCALIZED POPULATIONS

OF MODEL NEURONS

HUGH R. WILSON and JACK D. COWAN

From the Department of Theoretical Biology, The University of Chicago, Chicago, Illinois 60637 (inster copy)

4. D. COWAN

> A Mathematical Theory of the Functional Dynamics of Cortical and Thalamic Nervous Tissue

> > H. R. Wilson and J. D. Cowan Department of Theoretical Biology. The University of Chicago, Chicago, 111.. USA

> > > Received: January 14. 1973

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The neural mass equations

E(t) (resp. I(t)) average membrane potential of excitatory (resp. inhibitory) cells at t,

$$\tau \frac{dE}{dt} = -E + J_{ee}S_E(E(t)) - J_{ie}S_I(I(t)) + I_{ext}^E(t)$$

$$\tau' \frac{dI}{dt} = -I + J_{ei}S_E(E(t)) - J_{ii}S_I(I(t)) + I_{ext}^I(t)$$

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The celebrated Wilson and Cowan equations This is the voltage-based neural mass model

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History

 Starts with the work of Wilson and Cowan, 1973, and Amari, 1977,

Followed by many attempts by theoreticians and mathematicians to characterize the solutions, i.e. Bressloff, Coombes, Ermentrout and their followers Atay, Chow, Faugeras, beim Graben, Guo, Gutkin, Hutt, Laing, Pinto, Potthast, Troy, Veltz....

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Functional analysis framework

Functional spaces: Banach, Hilbert, Sobolev

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- Fixed point theorems
- Spectral Theory
- Bifurcation Theory

Functional analytic setting

$$\begin{cases} \dot{\mathbf{V}}(\mathbf{X},t) = -\mathbf{V}(\mathbf{X},t) + [\mathbf{J}(t) \cdot \mathbf{S}(\lambda \mathbf{V})](\mathbf{X}) + \mathbf{I}_{\text{ext}}(\mathbf{X},t) \\ \mathbf{V}(\cdot,0) = \mathbf{V}_{0}(\cdot) \end{cases}$$

- V and V₀ are *p*-dimensional vector functions defined on a set Ω, an open bounded/unbounded piece of ℝ^{2,3}, or a more complicated space. They represent the average membrane potential of the neural field.
- They both belong to some functional space \mathcal{F}
- J(t) is a linear operator F → F (connectivity kernel). It describes the intensity of the projections from masses at X' to masses at X.

$$\left[\mathbf{J}(t) \cdot \mathbf{U}\right](\mathbf{X}) = \int_{\Omega} \mathbf{J}(\mathbf{X}, \mathbf{X}', t) \mathbf{U}(\mathbf{X}') d\mathbf{X}'$$

Functional analytic setting

$$\dot{\mathbf{V}}(\mathbf{X},t) = -\mathbf{V}(\mathbf{X},t) + \left[\mathbf{J}(t)\cdot\mathbf{S}(\lambda\mathbf{V})
ight](\mathbf{X}) + \mathbf{I}_{\mathrm{ext}}(\mathbf{X},t)$$

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- **S** is a sigmoidal function (regular and bounded),
- \blacktriangleright λ determines its stiffness.
- **X**, in Ω, represents space, features, ...

Choice of \mathcal{F}

Three criteria

- 1. The problem should be well-posed
- 2. Its biological relevance
- 3. Allow numerical computations
- Hilbert space: $\mathcal{F} = \mathbf{L}^2(\Omega, \mathbb{R}^p)$
- More spatial regularity can be imposed

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Choice of $\boldsymbol{\mathsf{J}}$

► If $\mathbf{J}(\cdot, \cdot, t) \in \mathbf{L}^2(\Omega \times \Omega, \mathbb{R}^{p \times p})$ for all t > 0 then $\mathbf{J}(t)$ defines a continuous linear operator from \mathcal{F} to \mathcal{F}

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Cauchy problem

$$\begin{array}{rcl} \bullet & \text{ODE defined on } \mathcal{F} \\ \left\{ \begin{array}{l} \frac{d\mathbf{V}}{dt} &= -\mathbf{V} + \mathbf{R}(t,\mathbf{V}) \\ \mathbf{V}(0) &= \mathbf{V}_0 \in \mathcal{F} \end{array} \right. \end{array}$$

where

$$\mathsf{R}(t,\mathsf{V}) = \mathsf{J}(t) \cdot \mathsf{S}(\lambda \mathsf{V}) + \mathsf{I}_{\mathrm{ext}}(t)$$

Proposition (O.F, F. Grimbert, J.-J. Slotine, SIAM J. Appl. Math., 2008) If

- 1. J is in $\mathcal{C}\left(\mathbb{R}^+; \mathbf{L}^2\left(\Omega \times \Omega, \mathbb{R}^{p \times p}\right)\right)$ and $\|\mathbf{J}\|_{\mathcal{F}} \leq J, t \geq 0$
- 2. $I_{ext} \in \mathcal{C}(\mathbb{R}^+; \mathcal{F})$

then $\forall V_0 \in \mathcal{F}$ there is a unique solution V, defined on \mathbb{R}^+ and continuously differentiable of the Cauchy problem Related work: R. Potthast, P. beim Graben, Mathematical Methods in the

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Applied Sciences, 2009

Boundedness, absolute stability

▶ If $\|\mathbf{I}_{ext}(t)\|_{\mathcal{F}} \leq I_{ext}$ for all $t \geq 0$ the solution is bounded for all $\mathbf{V}_0 \in \mathcal{F}$.

- If λρ(J_s) < 1, J_s = (J + J^{*})/2, every solution is globally asymptotically stable, in particular unique
- O.F., R. Veltz, F. Grimbert, Neural Computation, 2009

- $\blacktriangleright\,$ Assume $I_{\rm ext}$ and J do not depend upon time
- Study the equilibrium states: bumps, persistent states, stationary solutions

Solve:

$$\mathbf{V} = \mathbf{J} \cdot \mathbf{S}(\lambda \mathbf{V}) + \mathbf{I}_{\text{ext}} := F(\mathbf{V}, \lambda)$$

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Questions:

- How many?
- Manifold of solutions when varying the parameters

Define $\mathcal{B}_{\lambda} = \{ \mathbf{V} \mid F(\mathbf{V}, \lambda) = 0 \}$, note \mathbf{V}_{λ}^{f} the solutions.

Proposition (R. Veltz and O.F., SIAM J. Applied Dynamical Systems, 2010)

1.
$$\left\|\mathbf{V}_{\lambda}^{f}-\mathbf{V}_{0}^{f}\right\|_{\mathcal{F}}\leq B(\lambda)$$



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Define $\mathcal{B}_{\lambda} = \{ \mathbf{V} \mid F(\mathbf{V}, \lambda) = 0 \}$, note \mathbf{V}_{λ}^{f} the solutions.

Proposition (R. Veltz and O.F., SIAM J. Applied Dynamical Systems, 2010)

2. If $|\mathcal{B}_{\lambda}|$ is finite then it must be odd (Leray-Schauder degree theory)



Define $\mathcal{B}_{\lambda} = \{ \mathbf{V} \mid F(\mathbf{V}, \lambda) = 0 \}$, note \mathbf{V}_{λ}^{f} the solutions.



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Visual Orientation sensitivity or how symmetries make things more complicated



- V1 is organized as a set of orientation sensitive columns and hypercolumns (D. Hubel and T. Wiesel, 1974)
- Periodic preferred orientation (PO) map



From E.R. Kandel et al., Principles of Neuroscience, 5th Edition, 2013.

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Neural field model

$$\frac{d}{dt}V(\mathbf{x},t) = -V(\mathbf{x},t) + \int_{\Omega} J(\mathbf{x},\mathbf{y})S(\sigma V(\mathbf{y},t))d\mathbf{y}$$

with

 $J(\mathbf{x}, \mathbf{y}) = J_{
m loc}(\|\mathbf{x}-\mathbf{y}\|) + \varepsilon J_{
m LR}(\mathbf{x}, \mathbf{y}) \quad \varepsilon << 1 \quad$ weaker strength of LR connection

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 $J_{\rm loc}$ is "Mexican-hat"-like (local excitation, remote inhibition)

Bifurcation analysis of the homogeneous model ($\varepsilon = 0$)

- Solve in a square Ω with periodic boundary conditions (Ω is a torus)
- Emergence of Turing patterns S. Coombes, P. beim Graben, R. Potthast, Tutorial on neural field theory. In: Coombes S, beim Graben P, Potthast R, Wright J, editors. Neural fields. Berlin: Springer; 2014
- Stripes always unstable
- Spots stable on a very short interval
- Two subcritical secondary branches recover stability



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Symmetries

- Because of the translation invariance, solutions "live" on a torus
- Assume that the spatial period matches that of the PO map T. Kenet et al., Nature, 2003
- Black lines: hypercolumns
- White lines: fundamental domain Ω₀
- Pinwheels are clockwise or counterclockwise



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Long range connections

$$J_{\rm LR}(\mathbf{x}, \mathbf{y}) = G_{\sigma_{\theta}}(\theta(\mathbf{x}) - \theta(\mathbf{y})) J_0\left(\chi, \mathbf{R}^{\mathbf{o}}_{-2\theta(\mathbf{x})}(\mathbf{x} - \mathbf{y})\right)$$

 θ is the PO map. χ controls the anisotropy.



Neuroscience, 5th Edition, 2013.

From P. Bressloff et al., Philos Trans R Soc Lond B, 2001 (→ → + = → + = →) ⊂ へへ

Symmetries of the PO map

Action of the rotations on the PO map:

$$\theta(\mathbf{R}^{\rho}_{\Phi_0}\mathbf{x}) = \theta(\mathbf{x}) + \epsilon \Phi_0/2 \mod \pi/2 \qquad \Phi_0 = \pi/2$$

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 $\epsilon = 1$ if p is counterclockwise, -1 otherwise

Symmetries

- Form Ω with $(2N + 1)^2$ copies of Ω_0
- Symmetries are $D_4 \rtimes (\mathbb{Z}/2N\mathbb{Z})^2$
- The PO map reduces it to one of the 17 wallpaper groups Evgraf Fedorov, 1891. D. Schattschneider, Am Math Mon, 1978
- Further constraints from the PO map reduces it to pmm (or cmm)



Remaining symmetries when $\varepsilon \neq 0$

As shown in R. Veltz, P. Chossat, O.F., 2015 the symmetry group Γ of the NF equations is p4m or p4 meaning:

Write

$$F(V,\lambda) \stackrel{\text{def}}{=} -\frac{d}{dt}V(\mathbf{x},t) - V(\mathbf{x},t) + [\mathbf{J} \cdot S(\lambda V)](\mathbf{x},t)$$

The neural field equation F is equivariant w.r.t Γ

$$\gamma \cdot F(V, \lambda) = F(\gamma \cdot V, \lambda)$$

When the long range connections are turned on the system undergoes spontaneous symmetry breaking

Dynamics with the long-range connections active

- Unperturbed torus $\mathcal{T}_0 \stackrel{\text{def}}{=} \{V_0(\cdot + \mathbf{x})\}, \, \mathbf{x} \in \Omega$
- A torus, flow-invariant manifold T_{LR}, persists as long as the perturbations are small (ε << 1) enough (P. Chossat and R. Lauterbach, Methods in equivariant bifurcations and dynamical systems, 2000)
- Questions:
 - How many steady-states do actually persist?
 - What are the observed phase portraits?
- Topology of the torus is an important constraint
- Analysis of the perturbed torus is done by identifying
 V₀(· + x) and x

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Visual orientation

Proposition (R. Veltz, P. Chossat, O.F., 2015)

Let us assume that there is a finite number of equilibria on the perturbed torus which are all non-degenerate when $\varepsilon \neq 0, \ \chi \geq 0$. There are at least 8 equilibria on the perturbed torus, four of which are saddles and the other four are nodes/foci.

Proof:

Application of the Poincaré–Hopf theorem: $n - 2s = \chi$, n = f + s, $\chi = 0$, hence f = s. Looking at rotations: f = 4

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Example of dynamics





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-Visual orientation

Examples of cortical activations



Final Remarks

 Generalizes B. Ermentrout and J. Cowan, Biol Cybern, 1979 and P. Bressloff et al., Philos Trans R Soc Lond B, 2001

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Group theory is underutilized in neuroscience

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Stochastic Neural Fields



Neurons are intrinsically noisy:

Ion channels and thermal noise

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Synaptic noise

Stochastic Neural Fields

Noisy neural field equation

$$dV(t,x) = \left[-V(t,x) + \int_{\mathbb{R}^n} J(x,y)S(V(t,y)) \, dy\right] dt + \sigma(V(t,x)) \, dW(t,x)$$

- S, σ are globally Lipschitz, S is bounded
- $(W(t,x))_{x \in \mathbb{R}^n, t \ge 0}$ is a stochastic process
- What do we mean by "a solution"?
- Must involve an object of the form

$$\int \sigma(V(t,x))\,dW(t,x)$$

This approach has been used "empirically" by Bressloff and Webber (2012), Bressloff and Wilkerson (2012), Kilpatrick and Ermentrout (2013)

Walsh stochastic integral (1986)

- Cleanly defines a white noise \dot{W} on $\mathbb{R}^+ \times \mathbb{R}^n$,
- ▶ a white noise process $W := (W_t(A))_{t \ge 0, A \in \mathcal{B}(\mathbb{R}^n)}$,
- ▶ and a set of functions, P_W, that can be integrated against the white noise process W

Spatial smoothing

- Informally we may think of the object W(t,x) as a random distribution, hence
- any solution of the stochastic neural field equation is distribution valued in the x-direction
- ▶ For $\varphi \in L^2(\mathbb{R}^n)$ define the (Gaussian) random field

$$W^{\varphi}(t,x) := \int_0^t \int_{\mathbb{R}^n} \varphi(x-y) W(ds \, dy)$$

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• Easy to verify that $\varphi(x - \cdot) \in \mathcal{P}_W$

Time and space regularity of W^{φ}

- The time regularity is the same as that of a Brownian path
- Hölder continuity in space can be enforced by imposing a (very) weak regularity on φ:

Condition (C1):

 $\exists C_{\varphi}$ such that

$$\| arphi - oldsymbol{ au}_z(arphi) \|_{L^2(\mathbb{R}^n)} \leq C_arphi |z|^lpha, \, orall z \in \mathbb{R}^n$$

for some $lpha \in (0,1]$ where $oldsymbol{ au}_z(arphi)(x) = arphi(x+z)$

Then for all t ≥ 0 x → W^φ(t, x) has an η-Hölder continuous modification for all η ∈ (0, α).

The SNF equation driven by spatially smoothed space-time white noise

$$\partial_t V(t,x) = -V(t,x) + \int_{\mathbb{R}^n} J(x,y) S(V(t,y)) \, dy + \sigma(V(t,x)) \, \frac{\partial}{\partial t} W^{\varphi}(t,x)$$

By a solution we mean a real-valued random field (V(t,x))_{t≥0,x∈ℝⁿ}

$$V(t,x) = e^{-t}V_0(x) + \int_0^t e^{-(t-s)} \int_{\mathbb{R}^n} J(x,y)S(V(s,y)) \, dy \, ds + \int_0^t \int_{\mathbb{R}^n} e^{-(t-s)} \sigma(V(s,x))\varphi(x-y) \, W(ds \, dy)$$

Sufficient conditions on J for the existence of a solution Condition (C2):

$$\forall x \in \mathbb{R}^n (y \to J(x, y)) \in L^1(\mathbb{R}^n), \quad \text{and} \quad \sup_{x \in \mathbb{R}^n} \|J(x, \cdot)\|_{L^1(\mathbb{R}^n)} \leq C_J$$

Theorem (Existence, uniqueness: O.F., J. Inglis, Journal of Mathematical Biology, 2015)

If $x \to V_0(x)$ is Borel measurable almost surely and

$$\sup_{x\in\mathbb{R}^n}\mathbb{E}\left[|V_0(x)|^2\right]<\infty$$

If the NF kernel J satisfies the condition (C2), then there exists an almost surely unique predictable random field $(V(t,x))_{t\geq 0, x\in\mathbb{R}^n}$ which is a solution of the SNF equation in the sense of the above definition and such that

$$\sup_{t \in [0,T], x \in \mathbb{R}^n} \mathbb{E}\left[|V(t,x)|^2 \right] < \infty \quad \text{for any } T > 0$$

Stochastic Neural Fields

Spatio-temporal regularity of the solution Condition (C3):

 $\begin{aligned} \exists \mathcal{K}_J \quad \text{s.t.} \quad & \|J(x_1,\cdot) - J(x_2,\cdot)\|_{L^1(\mathbb{R}^n)} \leq \mathcal{K}_J |x_1 - x_2|^{\alpha}, \; \forall x_1, \, x_2 \in \mathbb{R}^n \\ & \alpha \in (0,1]. \end{aligned}$

Theorem (Regularity: O.F., J. Inglis, Journal of Mathematical Biology, 2015)

If $x \to V_0(x)$ is Borel measurable almost surely and $\sup_{x \in \mathbb{R}^n} \mathbb{E} \left[|V_0(x)|^2 \right] < \infty$. Suppose moreover that $\exists \alpha \in (0, 1]$ s.t.

- ► J satisfies (C2) and (C3)
- φ satisfies (C1)

• $x \to V_0(x)$ is α -Hölder continuous

Then $(V(t,x))_{t\geq 0, x\in\mathbb{R}^n}$ has a modification such that $(t,x) \rightarrow V(t,x)$ is (η_1,η_2) -Hölder continuous, for any $\eta_1 \in (0,1/2)$ and $\eta_2 \in (0,\alpha)$.

Stochastic Neural Fields

Final remarks

Probability Theory

- Stochastic calculus
- Large deviations

are important tools in mathematical neuroscience

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- In the late 80s, consciousness was a tolerated hobby for the aging scientist
- "consciousness" is loaded with fuzzy meanings
- Distinguish between at least three meanings, vigilance (wakefulness), attention (focusing of mental resources), conscious access (reportable)

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Three ingredients for making consciousness accessible to experiments

- focusing on conscious access
- manipulating conscious perception
- carefully recording introspection

Thanks to the "new" brain "imaging" modalities, fMRI, EEG, MEG, Optical Imaging



From S. Dehaene, Consciousness and the brain, 2014.

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Unconscious versus conscious

Virtually all the brain's regions can participate in both conscious and unconscious thought

- Unconscious meaning: unseen color word primes the corresponding color Anthony Marcel, 1970
- Unconscious number comparisons: deciding whether a seen digit is larger or less than 5 depends on whether you have been primed with an invisible congruent/non-congruent digit
- Unconscious Attention: an invisible spot of light makes you more accurate to responding to stimuli presented tn the same area

What is the unconscious?

Henri Poincaré Science and Hypothesis, 1902 The subliminal self is in no way inferior to the conscious self; it is not purely automatic; it is capable of discernment; it has tact, delicacy; ...

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What is consciousness good for?

- Unconscious statistics versus conscious sampling: the brain performs Bayesian inference
- Lasting thoughts: once a piece of information is conscious, it stays fresh in our mind for as long as we care to attend to it and remember it
- The human Turing machine: consciousness gives us the power of a sophisticated serial computer

A social sharing device

I: Sudden ignition of parietal and prefrontal circuits

- Perception of masked words (images)
- Making additional sounds conscious



From S. Dehaene, Consciousness and

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the brain, 2014.

- II: Late slow wave: the P3 wave
 - Masked digits
 - Increase of delay between digit and mask
 - Sudden ignition of activity if delay > 50ms, digit visible:
 - Bifurcation in the dynamics of the neural networks



From S. Dehaene, Consciousness and

the brain, 2014. 《 문 · 《 문 · 《 문 · 문 · 문 · 이익이

- III: Late and sudden burst of high-frequency oscillations (deep electrodes)
 - Masked image
 - Increase of delay between mask and face image
 - Lasting burst of high-frequency activity



From S. Dehaene, Consciousness and

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the brain, 2014.

IV: Synchronization of information exchanges across distant brain regions

- Invisible faces/words
- Synchronization of many distinct brain areas



From S. Dehaene, Consciousness and

the brain, 2014. A B R A B R A B R A C

Theorizing consciousness

Awareness of a piece of information ^{def} = it has entered a specific storage area which makes it available to the rest of the brain

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Theorizing consciousness

- Awareness of a piece of information ^{def}= it has entered a specific storage area which makes it available to the rest of the brain
- Hippolite Taine's "theater of consciousness" (1870): "The conscious mind is like a narrow stage that lets us hear only a single actor". But there are lots of invisible actors.

 Raj Reddy et al.'s "blackboard" (1973) of the HEARSAY system

Theorizing consciousness

- Bernard Baars' "global workspace" (1989),
- Changeux and Dehaene's "global neuronal workspace" (1998)



Mathematics and consciousness

- Graph G := (V, E)
- ► Each vertex v ∈ V represents some brain areas (visual, auditory, somatosensory, etc...)
- Its mathematical description is a set of (S)NF equations
- ► Each edge e ∈ E represents the connections between the various brain areas.

- These can be anatomical (connectome) or functional
- They introduce delays (delayed (S)NF equations)

Mathematical challenges

- Provide mathematical models of the four signatures of consciousness
- e.g. bifurcations, metastable states, etc....
- Study rigorously how and when they can appear in the solutions of the (delayed) (S)NF equations
- Provide mathematical models of the global neuronal workspace

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Other challenges

Computational, e.g. HBP, TVB (Jirsa et al) ...

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Clinical (Naccache, Dehaene)

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Conclusion

- The "unreasonable effectiveness of mathematics" in Physics E. Wigner, 1960) is not (yet) verified in neuroscience or in biology in general.
- Conversely, there are few of the mathematical problems raised by these disciplines that are studied by "professional" mathematicians.

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Announcing a new Journal: Mathematical Neuroscience and Applications

- It bifurcates from the Journal of Mathematical Neuroscience (ends end of this year)
- Focuses on using mathematics as the primary tool for elucidating the fundamental mechanisms responsible for experimentally observed behaviours in neuroscience.
- Publishes work that uses advanced mathematical techniques to illuminate these questions.
- Papers that introduce and help develop those new pieces of mathematical theory which are likely to be relevant to future studies of the nervous system are welcome.
- Diamond Open Access model: free Open Access, thanks to the support of episciences.org

Announcing a new Journal: Mathematical Neuroscience and Applications

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	Mathematical Neuroscience and Applications
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Browse all volume Articles	, Mathematical Neuroscience and Applications is an international journal that publishes research articles on the mathematical modeling and analysis of all areas of neuroscience. i.e. the study of the neurous system and its dysfunctions. The focus is on using mathematica as the primary
About the journal	tool for elucidating the fundamental mechanisms responsible for experimentally observed behaviours in neuroscience at all relevant scales, from the molecular world to that of cognition. The aim is to publish work that uses advanced mathematical techniques to illuminate these questions. Pagers that introduce and help develop those new projects of mathematical theory which are lifely to be relevant to future studies of the nervous
Help	system in general and the human brain in particular are also welcome.) It publishes full length original papers, rapid communications and review articles. Papers that combine theoretical results supported by convincing
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	The journal is in part a continuation of the Journal of Mathematical Neuroscience that started publishing articles in 2011 and will cease publication in December 2021. These articles can be freely accessed at the following address.
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Supplementary slides

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Mathematical state of the art: stochastic *partial* differential equations (SPDEs)

- Functional approach: Da Prato and Zabczyk (1992), Prévôt and Röckner (2007)
- Solutions are random processes that take their values in a Hilbert space of functions
- Integration theory w.r.t. a class of random measures: Walsh (1986)
- Solutions are random fields in both t and x

The two approaches are equivalent



Walsh's stochastic integral (1986)

Centered Gaussian random field

$$\dot{W} := (\dot{W}(A))_{A \in \mathcal{B}(\mathbb{R}^+ \times \mathbb{R}^n)},$$

with covariance function

$$\mathbb{E}\left[\dot{W}(A)\dot{W}(B)\right] = |A \cap B|$$

W is a white noise on ℝ⁺ × ℝⁿ
White noise process W := (W_t(A))_{t≥0, A∈B(ℝⁿ)}

$$W_t(A) := \dot{W}([0,t] \times A), \ t \ge 0$$

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Define the norm

$$\|f\|_W^2 = \mathbb{E}\left[\int_0^T \int_{\mathbb{R}^n} |f(t,x)|^2 dt dx\right],$$

for any (random) function that is knowable at time t given $(W_s(A))_{s \le t, A \in \mathcal{B}(\mathbb{R}^n)}$

- ▶ \mathcal{P}_W is the set of all such functions f for which $||f||_W < \infty$
- It is the set of functions that can be integrated against the white noise process W

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Theorem (Walsh 1986) For all $f \in \mathcal{P}_W$, $t \in [0, T]$ and $A \in \mathcal{B}(\mathbb{R}^n)$,

$$\int_0^t \int_A f(s,x) W(ds \, dx)$$

can be well-defined. Moreover for all $t \in (0, T]$ and $A, B, \in \mathcal{B}(\mathbb{R}^n), \mathbb{E}\left[\int_0^t \int_A f(s, x) W(ds dx)\right] = 0$ and

$$\mathbb{E}\left[\int_0^t \int_A f(s, x) W(ds \, dx) \int_0^t \int_B f(s, x) W(ds \, dx)\right] = \mathbb{E}\left[\int_0^t \int_{A \cap B} f(s, x) W(ds \, dx)\right]$$

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Computational challenges

- Already partially addressed
- Bluebrain (Europe), Human Brain Project (Europe), The Virtual Brain Project (Europe), Brain (USA)etc...



V. Jirsa, in S. Coombes et al., Neural Fields, Theory and Applications, 2014 🍙 🧠

Testing in the clinic

Epileptic seizures, e.g. TVB: V. Jirsa et al.

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Vegetative states, S. Dehaene et al.