# **Randomness Increases Order in Biological Evolution, But ... what is biological randomness?**

Giuseppe Longo

CREA, CNRS - Ecole Polytechnique et Cirphles, Ens, Paris

F. Bailly, G. Longo. Mathematics and Natural Sciences. The physical singularity of Life. Imperial College, 2011

G. Longo, MM.aël Montévil. Randomness Increases Order in Biological Evolution. C. Calude's conference on "**Computations, Physics and Beyond**", Auckland, NZ, Feb. 21-24, 2012; LNCS (Dinneen et al. eds), Springer, 2012

# Randomness from Physics to in Biology

## **Physical Determination (Classical)**

Laplace's view:

A) determination *implies* predictibility

and

B) determination =/= randomness

[Laplace, Philosophie des Probabilités, 1786]

### **Physical Determination (Classical)**

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#### A) determination *implies* predictibility (false: Poincaré, 1890) and

B) determination =/= randomness (= determ. unpredictab., 1890)

[Laplace, Philosophie des Probabilités, 1786]

#### Thus, Poincaré broadened determinism

by including classical randomness: a fluctuation/perturbation *below measure*, may yield an observable effect, over time:

"et nous avons un phénomène aléatoire", [Poincaré, 1902]

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**Turing**, 1950:

#### From the LCM to the DSM (*Discrete State Machine*): my DSM is **laplacian!**

**Turing**, 1952:

Morphogenesis as a "continuous dynamics", non-linear ("exponential drift", dynamic unpreditability = classical randomness: **Poincaré**)

## **Consequences from Physical Determination (Classical)**

Laplace's view:

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B) determination =/= randomness (= determ. unpredictab., 1890)

[Laplace, Philosophie des Probabilités, 1786] [J. Monod, Le hasard et la nécessité, 1970]

Consequences of the Laplacian view:

the "DNA is a program" theory, since any *predictable determination* is **programmable** 

## **Consequences from Physical Determination (Classical)**

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[J. Monod, Le hasard et la nécessité, 1970]

#### Schrödinger, What is life? 1944:

« In calling the structure of the chromosomes a code-script, we mean that the all-penetrating mind, **once conceived by Laplace**... could tell from their structure how the egg would develop....»

Schrödinger's **right consequences** of his principles! Today, the code-script has been fully decoded...

# More on randomness as deterministic unpredictability

Classical/Relativistic systems are State Determined Systems: randomness is an **epistemic** issue

*Examples*: dies, coin tossing, a double pendulum ...

the Planetary System (Poincaré, 1890; Laskar, 1992)... finite (short and long) time unpredictability

(*the dies, a SDS, 'know' where they go*: along a geodetics, determined by Hamilton's principle).

#### Recall Laplace:

- infinitary daimon: **OK** (over space-time *classical* continua);
- determination *implies* predictability (except singularities): **Wrong!** 
  - The role of physical measurement

# Quantum unpredictability as intrinsic indeterminism

Quantum Mechanics is *not* deterministic:

intrinsic/objective role of probabilities in constituting the theory:

- measure of conjugated variables;
- entanglement, no hidden variables.

*Schrödinger's* idea: the **equational determination** of a "*law of probability*" (thus back to the indeterministic nature of QM)

Quantum Mechanics: you *can't even think* of an infinitary daimon (key difference: measure of conjugate variables).

Recent survey/reflections: [Bailly, Longo, 2011], [Longo, Paul, 2008]

# **Physical Randomness and Irreversibility**

- **1 Classical randomness = deterministic unpredictability**
- 2 Quantum randomness = intrinsic undetermination (& entanglement)

Different probabilities, different theories of randomness ...

Yet, common points:

- Randomness = unpredictability
- **Randomness** is *correlated to* (co-present with) **irreversiblity** of time (classically: bifurcations ... ; quantum: measurement)

Cf also **Thermodynamics**: II principle; diffusion as random paths.

# Some more philosophy

1 - Laplace (strong, fantastic) program: the (*written*) equational determination allows to deduce/predict completely the properties of the physical World

(Newton: "one has to write and solve equations" ... )

Poincaré: No, it does not work (1892: deterministic unpredictability)

# Some more philosophy

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**Poincaré**: No, it does not work (1892: deterministic unpredictability)

**2 - Hilbert** program: the *finite axiomatic writing* of Mathematics allows to formally deduce/predict completely the properties of Mathematics

Gödel: No, it does not work (1931: undecidability)

Provable correlations between *consequences*, as forms of randomness

# **Towards Biology**

#### 3 - Crick, Monod ....

"the *finite string* of DNA base letters A, C, T, G *completely determine* embryogenesis, ontogenesis ... evolution"

More: « the DNA code ... is the *program* for the *behavioral computer* of this individual » (Mayr 1961)

And the *two ways interactions* 

DNA – proteome/cell/organisms/ecosystem ? None (Crick's central Dogma, 1958), or just « noise », « bad copies »

**Randomness** (= noise) is "laplacian" (extraneous to determination and theory)

### The constitutive role of randomness in Biology

One of the crucial « change of perspective », in Biology:

#### **Randomness** is **not noise** and it *implies* **variability** *implies* **diversity** An essential component of structural stability

*Compare*: Randomness as intrinsic to Quantum Mechanics (change measure and the « structure of determination »)

Kupiec, 1983 .... Buiatti M., Longo G. *Randomness and Multilevel Interactions in Biology*, Ongoing work.

#### **Biological relevance of randomness**

Each mitosis (cell division), a critical phase transition:

Asymmetric partitions of proteomes; differences in DNA copies; changes in membranes ...

*In multicellular organisms*: varying reconstruction of tissues' matrix (collegen structure, cell-to-cell connections)

Not « noise », « mistakes » in polymerase as a Turing's program, but non-specificity and randomness is at the core not only of *variability* and *diversity* (the main biological invariants), but even of **cell differentiation** (in embrogenesis: sensitivity in a critical transition; e.g. variability in Zebrafisch, N. Peyreiras, ongoing).

Randomness enhances robusteness, by diversity : ecosystem, organism ...

# Which form of randomness ?

# **Quantum Randomness in Biology**

**Quantum tunneling:** non-zero probability of passing any physical barrier (cell respiration, Gray, 2003; destabilizing tautomeric enol forms – migration of a proton: Perez, 2010)

**Quantum coherence:** electron transport (in many biogical processes: Winkler, 2005)

**Proton transfer** (quantum probability): RNA mutations (G-C pairs: Ceron-Carrasco, 2009)

Empowered **metabolic random** activities by (water) "QED coherence" (Del Giudice, 2005; Plankar, 2011)

**REFERENCES IN:** 

Buiatti M., Longo G. *Randomness and Multilevel Interactions in Biology*, Downloadable

# **Classical Randomness in Biology**

### Non linear affects (molecular level):

• Molecular enthalpic oscillations

•

• **Turbulence** in the cytoplasm of Eukaryote cells

(see also J.-J. Kupiec, A. Paldi, T. Heams, B. Laforge ... )

# Classical and Quantum Randomness in Biology

Molecular level:

**non linear dynamics** (classical) *and* **quantum processes** *superpose* 

That is:

They happen simultaneously and interfere (not analyzed in Physics)

Morover:

a quantum effect may be amplified by a (classical non-linear) dynamics

# Proper (?) Biological Randomness 1

Randomness *within* other levels of organization *in an organism*:

- [*Besides:* Molecular activities (classical+quantum randomness)]
- Cellular dynamics and interactions in a tissue
- Developmental dynamics (contact inhibition between cells: Soto et al., 1999)
- **Fractal bifurcations** (mammary glands development, *ongoing work*)

# **Proper Biological Randomness 2:**

*Recall*: since Poincaré, randomness as "planetary **resonance**" Extended to general non-linear dynamics:

at **one level** of (mathematical) determination (*far from equilibrium*: Pollicott-Ruelle resonance, dynamical entropy in open

systems (Gaspard, 2007))

# **Proper Biological Randomness 2:**

*Recall*: since Poincaré, randomness as "planetary **resonance**" ... Extended to general non-linear dynamics:

at one level of (mathematical) determination

(*far from equilibrium*: Pollicott-Ruelle resonance, dynamical entropy in open systems (Gaspard, 2007))

#### **Bio-resonance** (Buiatti, Longo, 2011):

Randomness *between* different levels of organization *in an organism*: thus, resonance (as interference) between **different levels** of (mathematical) **determination** 

*The mathematical challenge:* the Mathematics (of Physics) does **not** deal with **heterogeneous structures** (of determination)

# **Bio-resonance**

**Physical resonance** (at equilibrium / far from equilibrium) is related to "destabilization" (growth of entropy or disorder)

# **Bio-resonance** includes "integration and regulation", thus it *stabilizes* and *destabilizes*

Examples:

- The lungs, the drosophila eyes ...
- In embryogenesis ...
- In "colonies" of *Myxococcus Xanthus*, a prokaryote, and *Dictyostelium discoideum*, an eukaryote (Buiatti, Longo, 2011)

# **Randomness in critical transitions**

Life is (not only) a dynamics, a process, but an extended (permanent, ongoing ... in time, space ..) critical transition (Bailly, Longo, Montévil: book and papers)

A critical interval, not just a (mathematical) point, as in Physics.

Key understanding: continual symmetry changes

In Physics, the *determination* of trajectories is *given by symmetries* (the conservation properties)

An biological (ontophylogenetic) trajectory is a *cascade of symmetry changes*.

# The 'double' irreversibility of Biological Time

Increasing complexity (Gould) in evolution is the result of a **random** *asymmetric* **diffusion** 

F. Bailly, G. Longo. *Biological Organization and Anti-Entropy*, in **J. of Biological Systems**, Vol. 17, n.1, 2009.

Evolution, morphogenesis and death are strictly irreversible, but their irreversibility is proper, it *adds on top* of the physical irreversibility of time (thermo-dynamical):

e. g., increasing order induces (also some) disorder.

**Thesis** (the role of randomness):

a random event is (always) correlated to a symmetry breaking.

One more reason for an *intrinsic*, proper Biological Randomness. END

## Some references (more on

http://www.di.ens.fr/users/longo

Buiatti M., Longo G. *Randomness and Multilevel Interactions in Biology,* In progress (downloadable <u>http://www.di.ens.fr/users/longo</u>).

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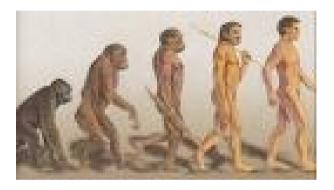
# But ... what is biological randomness? V

## **Randomness Increases Order in Biological Evolution**

G. Longo, M. Montévil. Randomness Increases Order in Biological Evolution. C. Calude's conference on "**Computations, Physics and Beyond**", Auckland, NZ, Feb. 21-24, 2012; LNCS (Dinneen et al. eds), Springer, 2012

### **Evolution and "Complexity"**

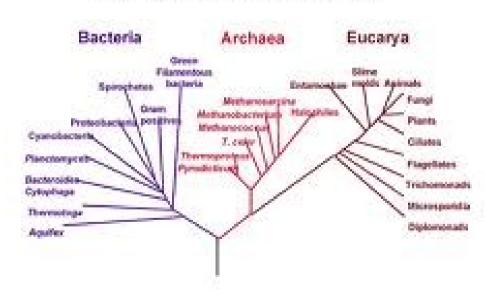
J.-S. Gould's fight against the *wrong* image (progress?):



S.J. Gould. Full house: The spread of excellence from Plato to Darwin . Three Rivers Pr, 1997.

#### **Growing complexity in Evolution?**

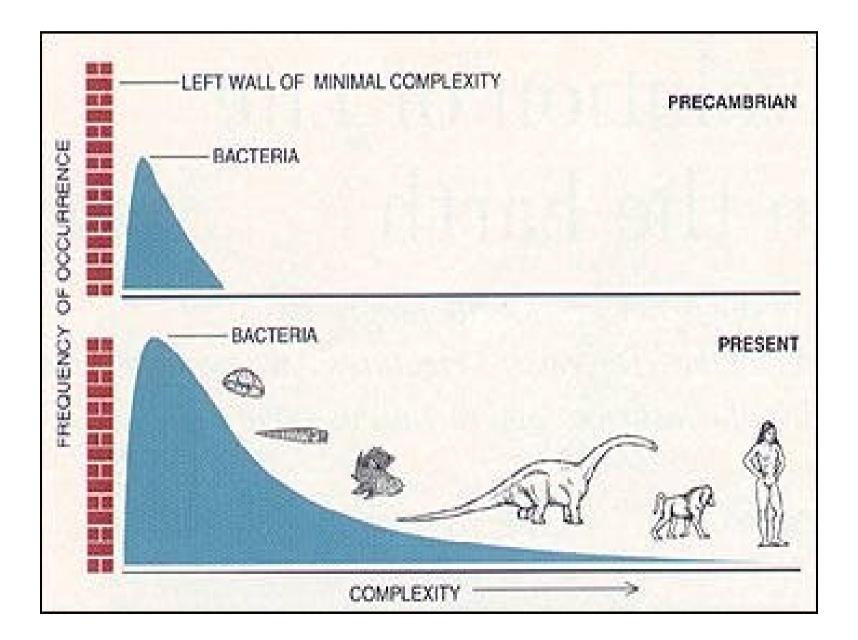
Which "complexity"? Evolutionary complexity?



#### **Phylogenetic Tree of Life**

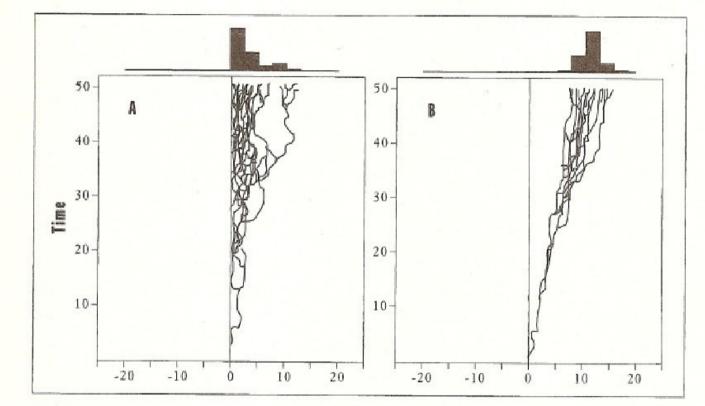
### However: Gould's growth of "morphological" complexity [Full House, 1989]

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#### **Random increase of complexity** [Gould, 1989]

#### Asymmetric Diffusion Biased Increase



#### FIGURE 33

Passive and driven trends in McShea's terminology. A passive trend (A) begins near a left wall, retains a constant mode at this beginning position, and expands in the only open direction toward the right. In a driven trend (B), both minimum and maximum values increase through time.

### How to understand increasing complexity?

No way to explain this in terms of random mutations (only):

- 1. DNA's (genotype) **random mutations** statistically have probability 0 to cause globally increasing complexity of phenotype (examples: mayfly (ephemeral); equus...[Longo, Tendero, 2007])
- 1. Darwin's evolution is **selection of the** *incompatible* ("the best" makes no general sense)
- Greater probabilities of survival and reproduction *do not imply* greater complexity (bacteria, ... lizard...) [Maynard-Smith, 1969]

Gould's idea: symmetry breaking in a diffusion...

### **Mathematical analysis as a distribution of** Biomass (density) over Complexity (bio-organization)

F. Bailly, G. Longo. *Biological organization and anti-entropy*. J. Bio-systems, 17-1, 2009.

### Derive Gould's empirical curb from

- general (mathematical) principles,
- specify the phase space
- explicit (and correct) the time dependence

# Write a suitable **diffusion equation** inspired by Schrödinger operatorial approach

Note: any diffusion is based on random paths!

# Morphological Complexity along phylogenesis and embryogenesis

Specify (quantify) Gould's informal "complexity" as static morphological
 complexity K

 $\mathbf{K} = \alpha K_{c} + \beta K_{m} + \gamma K_{f}$  ( $\alpha + \beta + \gamma = 1$ )

- K<sub>c</sub> (combinatorial complexity) = cellular combinatorics as differentiations between cellular lineages (tissues)
- K<sub>m</sub> (phenotipic complexity) = topological forms and structures (e.g., connexity and fractal structures)
- K<sub>f</sub> (relational complexity) = metabolic relations, neuronal and cellular (interaction) networks

Main idea: formalize **K** as **anti-entropy**,  $-S \neq$  negentropy (not 0-sum, coding dependent) .... in balance equations...

# The theoretical frame: analogies

.... by a *conceptual analogy* with **Quantum Physics**:

In *Quantum Physics* (a "wave diffusion" in Hilbert Spaces): \* The determination is a *dynamics* of a *law of probability*: (Schrödinger Eq.)  $ih\partial \psi/\partial t = h^2 \partial^2 \psi/\partial x^2 + v \psi$ 

In our approach to *Complexity* in *Biological Evolution*:
\* The determination is a *dynamics* of a *potential of variability*:
(PV) ∂f /∂t = D<sub>h</sub>∂<sup>2</sup>f/∂K<sup>2</sup> + α<sub>h</sub>f

What is f? (PV) a diffusion equation, in which spaces?
Random walks ...

# The theoretical frame: dualities

.... by conceptual dualities with Quantum Physics:

In Quantum Physics (Schrödinger equation):

- Energy is an *operator*, *H(f)*, the "main" physical observable.
- Time is a *parameter*,  $f(\underline{x}, t)$ ,

# The theoretical frame: dualities

.... by *conceptual dualities* with **Quantum Physics**:

In Quantum Physics (Schrödinger equation):

- Energy is an *operator*, *H(f)*, the "main" physical observable.
- Time is a *parameter*,  $f(\underline{x}, t)$ ,

In our approach to *Complexity* in *Biological Evolution*:

- Time is an *operator*, identified with entropy production  $\sigma$
- Energy is a *parameter*,  $f(\underline{x}, e)$  (e.g. energy as bio-mass in scalingallometric equations:  $Q = kM^{1/n}$ )

Our f is the density of bio-mass over complexity K (and time t): m(t, K) A diffusion equation:

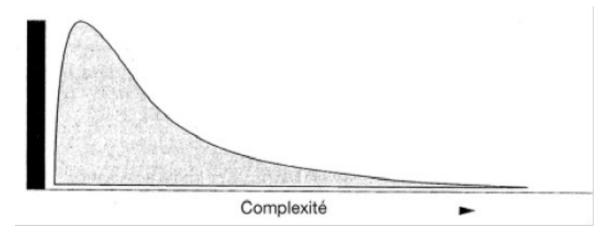
$$\partial \mathbf{m}/\partial \mathbf{t} = \mathbf{D}_{\mathbf{b}}\partial^2 \mathbf{m}/\partial \mathbf{K}^2 + \boldsymbol{\alpha}_{\mathbf{b}}\mathbf{m}(\mathbf{t},\mathbf{K})$$
 (3)

A solution

 $m(t,K) = (A/\sqrt{t}) \exp(at)\exp(-K^2/4Dt)$ 

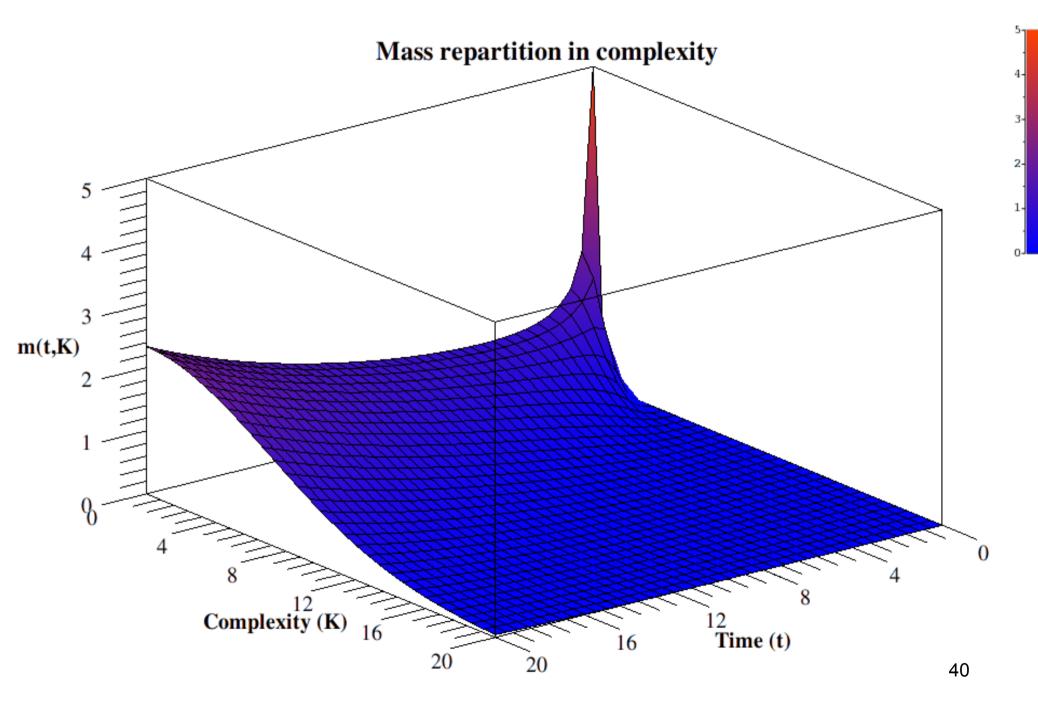
**models** Gould's asymmetric diagram for Complexity in Evolution (a diffusion : *random paths*...), *also along t* : (biomass and the **left wall** for complexity, archeobacteria original formation)

biomass



F. Bailly, G. Longo. Biological Organization and Anti-Entropy...

(Implementation by Maël Montevil; "ponctuated equilibria" smoothed out)



**Some references** (papers downloadable)

http://www.di.ens.fr/users/longo or Google: Giuseppe Longo

- Bailly F., Longo G. Mathematics and Natural Sciences. The physical singularity of Life. Imperial Coll. Press/World Sci., 2011 (*en français* : *Hermann, Paris, 2006*).
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