SYSTEMES COMPLEXES

9^{ème} rencontres de la fédération "Dynamique des Systèmes Complexes" de l'Université Joseph Fourier - Grenoble 1

Heterogeneous collective motion or moving pattern formation?

The exemplary status of multicellular morphogenesis at the border between "informed" physics and "physical" computation

René Doursat

http://www.iscpif.fr/~doursat









Complex systems can be found everywhere around us



- decentralization: the system is made of myriads of "simple" agents (local information, local rules, local interactions)
- emergence: function is a bottom-up collective effect of the agents (asynchrony, balance, combinatorial creativity)
- self-organization: the system operates <u>and changes</u> On its OWN (autonomy, robustness, adaptation)

> Physical, biological, technological, social complex systems



pattern formation O = matter



biological development O = cell



the brain & cognition O = neuron

insect colonies O = ant



Internet & Web O = host/page



social networks O = person





Ex: Pattern formation – Animal colors

animal patterns caused by pigment cells that try to copy their nearest neighbors \checkmark but differentiate from farther cells





(Scott Camazine, http://www.scottcamazine.com)









Ex: <u>Swarm intelligence – Insect colonies</u>

NetLogo Fur simulation

trails form by ants that follow and reinforce each other's pheromone path



archive 2003/EPOW-030811 files/matabele ants.jpg





Harvester ants (Deborah Gordon, Stanford University)



NetLogo Ants simulation



Ex: <u>Collective motion</u> – Flocking, schooling, herding





 Fish school
 Bison herd

 (Eric T. Schultz, University of Connecticut)
 (Montana State University, Bozeman)

- \checkmark thousands of animals that adjust their position,
 - orientation and speed wrt to their nearest neighbors



Separation, alignment and cohesion ("Boids" model, Craig Reynolds)



NetLogo Flocking simulation

Ex: <u>Diffusion and networks</u> – Cities and social links

✓ clusters and cliques of people who aggregate in geographical or social space





NetLogo urban sprawl simulation





NetLogo preferential attachment



\succ All kinds of agents: molecules \rightarrow cells \rightarrow animals / humans \rightarrow technology cells molecules animals living cell physical humans patterns & tech



A vast archipelago of precursor and neighboring disciplines

complexity: measuring the length to describe, time to build, or resources to run, a system

- information theory (Shannon; entropy)
- computational complexity (P, NP)
- Turing machines & cellular automata

→ Toward a unified "complex systems" science and engineering?

dynamics: behavior and activity of a system over time

- nonlinear dynamics & chaos
- stochastic processes
- systems dynamics (macro variables)

PLEX

adaptation: change in typical functional regime of a system

- evolutionary methods
- genetic algorithms
- machine learning

systems sciences: holistic (nonreductionist) view on interacting parts

- systems theory (von Bertalanffy)
- systems engineering (design)
- cybernetics (Wiener; goals & feedback)
- **control theory** (negative feedback)

multitude, statistics: large-scale properties of systems

- graph theory & networks
- statistical physics
- agent-based modeling
- distributed AI systems



French "roadmap" toward a Complex Systems Science

Big questions

- reconstruct multiscale dynam.
- emergence & immergence
- spatiotemp. morphodynamics
- optimal control & steering
- artificial design
- fluctuations out-of-equilib.
- adaptation, learning, evolution



→ Erasmus Mundus MSc/PhD Program in Complex
 Systems Science (Polytechnique, Warwick, Chalmers)
 → Digital University of Complex Systems (Saclay)



French "roadmap" toward a Complex Systems Science



Blue Brain



> The challenges of complex systems (CS) research

Transfersamong systems



CS science: understanding & modeling "natural" CS

(spontaneously emergent, including human-made):

morphogenesis, neural dynamics, cooperative co-evolution, swarm intelligence

Exports

- decentralization
- <u>autonomy</u>, homeostasis
- learning, evolution



- observe, model
- control, harness
- design, use



CS engineering: designing a new generation of "artificial" CS (harnessed & tamed, including nature): collective robotics, synthetic biology, energy networks







Pierre Baudot

Information Theory - Adaptation - Topology - Thermodynamics of perception.

mathematical neuroscience

René Doursat

Artificial development (self-assembly, pattern formation, spatial computing, evolutionary computation) - Mesoscopic neurodynamics (segmentation, schematization, categorization, perception, cognitive linguistics).

artificial life / neural computing

Francesco d'Ovidio

Applied nonlinear dynamics - Transport and mixing in geophysical flows - Interaction of physical and ecological processes in the ocean.





Francesco Ginelli

Nonequilibrium statistical mechanics (Active matter, collective motion, flockng, nonequilibrium wetting, directed percolation, long range interactions) - Dynamical system theory (Lyapunov exponents, Lyapunov vectors, synchronization, stable chaos, spatiotemporal chaos, structural stability, hyperbolicity).

statistical mechanics / collective motion

nonlinear dynamics / oceanography



Ivan Junier

Bio-related: Genetic regulation - Cellular organization - DNA/chromatin modeling --omics (Genomics, Transcriptomics, proteomics,...) - Condensed matter theory -Inference problems in statistical physics - Network analysis (topology, geometry) Dynamical behaviors of complex systems. Statistical physics: Out-of equilibrium syste

structural genomics Thermodynamic description of small syster



Taras Kowaliw

Evolutionary computation, artificial development, computer vision, visualization and electronic art.



computational evolution / development

Telmo Menezes

Complex network analysis and simulation - Social networks - Evolutionary search for multi-agent models, Genetic programming applied to programmable networks -Bio-inspired algorithms.

social networks



Peer-to-Peer networks, Blog networks, Complex networks, Statistical mechanics, Networks modeling, Optical networks, Wireless Internet.

peer-to-peer networks









Romain Reuillon

High performance computing - Grid computing - Scientific workflows - Model exploration - Distributed stochastic simulations - Paralell pseudo-random number generation - Coffee maker.

high performance computing

Jean-Baptiste Rouquier

Complex networks: communities, structure, dynamics. Links between fields. Large datasets.

Cellular automata: model of complex systems, perturbation, asynchronism, robustness.

complex networks / cellular automata

Camilo Melani

Grid Computing, Bioemergences Platform (workflow), Mophodynamics reconstruction, Images processing algorithms.

embryogenesis

David Chavalarias

Web mining and Quantitative Epistemology - Cognitive economics and modelling of cultural dynamics - Collective discovery and scientific discovery.

web mining / social intelligence

Srdjan Ostojic

Neuroscience théoriques - Spiking Neurons - Dynamiques Stochastic-ques.

spiking neural dynamics

Andrea Perna

Morphogenesis, Collective behavior, Spatial patterns, Spatial networks.

spatial networks / swarm intelligence

Fernando Peruani

Biophysique - Active Matter - Complex Networks.

active matter / complex networks

Fabien Tarissan

Informatique théorique, méthodes formelles, théorie des langages de programmation. Systèmes complexes, graphes dynamiques, modélisations des systèmes biologiques. Recherche opérationnelle, programmation mathématique, problèmes inverses. theoretical computer science



Reisdent Researchers





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Minichino, Michele

Andrei - Inst Curi.

SBAlli, Sylvie



The self-made puzzle of embryogenesis

1. Self-organized *and* structured systems

- 2. A two-side challenge: heterogeneous motion / moving patterns
- 3. Artificial Multi-Agent Embryogenesis
- 4. Artificial Evo-Devo & Future Work



Architecture Without Architects

"Simple"/random vs. architectured complex systems



biological patterns

iving cell





biology strikingly demonstrates \geq the possibility of combining pure self-organization and elaborate architecture, i.e.:

flocks

- a non-trivial, sophisticated morphology
 - *hierarchical* (multi-scale): regions, parts, details
 - modular: reuse of parts, quasi-repetition
 - heterogeneous: differentiation, division of labor
- random at agent level, reproducible at system level



Architecture Without Architects

Ex: Morphogenesis – Biological development







Nadine Peyriéras, Paul Bourgine et al. (Embryomics & BioEmergences)

cells build sophisticated organisms by division, genetic differentiation and biomechanical selfassembly

Ex: Swarm intelligence – Termite mounds



Termite mound (J. McLaughlin, Penn State University)



http://cas.bellarmine.edu/tietjen/ TermiteMound%20CS.gif



Termite stigmergy (after Paul Grassé; from Solé and Goodwin, "Signs of Life", Perseus Books)

 termite colonies build sophisticated mounds by
 "stigmergy" = loop between modifying the environment and reacting differently to these modifications

Systems that are self-organized and architectured



free self-organization

the scientific challenge of complex systems: how can they integrate a true *architecture?*

the engineering challenge of complicated systems: how can they integrate **self**organization?



Peugeot Picass

architecture





self-organized architecture / architectured self-organization

Heterogeneous collective motion / moving pattern formation





Statistical (self-similar) systems

Many agents, simple rules, "complex" emergent behavior

 \rightarrow diversity of *patterning* (spots, stripes) and/or *motion* (swarms, clusters, flocks), complex networks, etc., but.....



- often like "textures": repetitive, statistically *uniform*, information-poor \checkmark
- spontaneous order arising from amplification of *random* fluctuations \checkmark
- *unpredictable* number and position of mesoscopic entities (spots, groups) \checkmark \rightarrow "missing" ingredient: heterogeneity of the units 18



Morphological (self-dissimilar) systems compositional systems: pattern formation \neq morphogenesis



"I have the stripes, but where is the zebra?" OR *"The stripes are easy, it's the horse part that troubles me"* —attributed to A. Turing, after his 1952 paper on morphogenesis



Statistical vs. morphological systems

Physical pattern formation is "free" – Biological (multicellular) pattern formation is "guided"





larval axolotl limb condensations Gerd B. Müller



Statistical vs. morphological systems

Multicellular forms = a bit of "free" + a lot of "guided"

✓ domains of free patterning embedded in a guided morphology

unlike Drosophila's stripes, these pattern primitives are <u>not</u> regulated by different sets of genes depending on their position

spots, stripes in skin angelfish, www.sheddaquarium.org





ommatidia in compound eye dragonfly, www.phy.duke.edu/-hsg/54

repeated copies of a guided form, distributed in free patterns

entire structures (flowers, segments) can become modules showing up in random positions and/or numbers

flowers in tree cherry tree, www.phy.duke.edu/~fortney





segments in insect centipede, images.encarta.msn.com

From "statistical" to "morphological" CS in inert matter / insect constructions / multicellular organisms



physical pattern formation





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Overview of embryogenesis

> An abstract computational approach to development

- ✓ as a fundamentally *spatial* phenomenon
- highlighting its *broad principles* and proposing a *computational* model of these principles

Broad principles

- 1. biomechanics \rightarrow collective motion \rightarrow "sculpture" of the embryo
- *2. gene regulation* \rightarrow gene expression patterns \rightarrow "painting" of the embryo
- + *coupling* between shapes and colors

Multi-agent models

- best positioned to integrate both
- account for heterogeneity, modularity, hierarchy
 - each agent carries a combined set of *biomechanical* and *regulatory* rules



Embryogenesis couples assembly and patterning

> Sculpture \rightarrow forms







"shape from patterning"

 the forms are
 "sculpted" by the selfassembly of the
 elements, whose
 behavior is triggered
 by the colors

\succ Painting \rightarrow colors



"patterns from shaping

 new color regions appear (domains of genetic expression) triggered by deformations Niki de Saint Phall

Embryogenesis couples mechanics and regulation

> Cellular mechanics

- \checkmark adhesion
- ✓ deformation / reformation
- ✓ migration (motility)
- division / death

cellular Potts model (Graner, Glazier, Hogeweg)









(Doursat)



Gene regulatory pattern formation

Segmentation & identity domains in Drosophila

 ✓ periodic A/P band patterns are controlled by a 5-tier gene regulatory hierarchy



 intersection with other axes creates organ primordia and imaginal discs (identity domains of future legs, wings, antennae, etc.)



From DNA to Diversity, p63

Embryogenesis couples mechanics and regulation







Embryogenesis couples motion and patterns

Collective motion regionalized into patterns

Nadine Peyriéras, Paul Bourgine, Thierry Savy, BioEmergences Benoît Lombardot, Emmanuel Faure et al. http://bingweb.binghamton.edu/~sayam **Hiroki Sayama** (Swarm Chemistry) SwarmChemistry, zebrafish Ż Embryomics

Pattern formation that triggers motion



http://zool33.uni-graz.at/schmickl

Doursat



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Why multi-agent modeling?

Equations and laws can be hard or impossible to find...

- "The study of non-linear physics is like the study of nonelephant biology." —Stanislaw Ulam
 - the physical world is a fundamentally *non-linear* and *out-of-equilibrium* process
 - focusing on linear approximations and stable points is missing the big picture in most cases
- ✓ let's push this quip: "The study of nonanalytical complex systems is like the study of non-elephant biology." —??
 - complex systems have their own "elephant" species, too: dynamical systems that can be described by diff. eqs or statistical laws
 - many real-world complex systems do not obey neat macroscopic laws





Why multi-agent modeling?

Equations and laws can be hard or impossible to find in...

- ✓ systems that *no macroscopic quantity* suffices to explain (DEE)
 - no law of "concentration", "pressure", etc.
 - even if global metrics can be found, they rarely obey a given equation or law
- ✓ systems that require a *non-Cartesian* decomposition of space (Pp€)
 - network of irregularly placed or mobile *agents*
- ✓ systems that contain *heterogeneity*
 - segmentation into different *types of agents*
 - at a fine grain, this would require a "patchwork" of regional equations
 - systems that are dynamically *adaptive*
 - the topology and strength of the interactions depend on the short-term activity of the agents and long-term "fitness" of the system in its environment



Different approaches and families of models

Biological, bio-inspired or artificial models

- focused on spatial differentiation patterns (little or no motion)
 - reaction-diffusion (PDEs, cellular automata)
 - gene networks (Boolean or concentrations) on a fixed lattice
 - "amorphous computing"
- ✓ focused on motion (little or no patterning)
 - Cellular Potts Model (on predefined cell types)
 - aggregation, self-assembly
 - collective motion, flocking, cellular sorting
- \checkmark at different scales
 - macroscopic models (densities, differential geometry) \rightarrow no individual information
 - mesoscopic models (cellular centers, Potts) \rightarrow no membrane geometry or nuclei
 - microscopic models (elastic polyedra, drop models) \rightarrow cellular deformations



a combination that is still rare ; but see Hogeweg / Salazar-Ciudad / Mjolsness..



Capturing the essence of embryogenesis in an Artificial Life agent model









W













Programmed patterning (patt): the hidden embryo atlas

- a) same swarm in different colormaps to visualize the agents' internal patterning variables *X*, *Y*, *B*_i and *I*_k (virtual *in situ hybridization*)
- b) consolidated view of all identity regions I_k for k = 1...9
- c) gene regulatory network used by each agent to calculate its expression levels, here: $B_1 = \sigma(1/3 X)$, $B_3 = \sigma(2/3 Y)$, $I_4 = B_1B_3(1 B_4)$, etc.



From feedforward to recurrent gene regulation

Summary: simple feedforward hypothesis

- developmental genes are broadly organized in tiers, or "generations": earlier genes map the way for later genes
- ✓ gene expression propagates in a directed fashion: first, positional morphogens create domains, then domains intersect



From feedforward to recurrent gene regulation

Toolkit genes are often multivalent

- exception to the feedforward paradigm: "toolkit" genes that are reused at different stages and different places in the organism
- ✓ however, a toolkit gene is triggered by different switch combos, which can be represented by duplicate nodes in different tiers



Endless Forms Most Beautiful, p125

From feedforward to recurrent gene regulation

More realistic variants of GRNs

- ✓ add recurrent links within tiers → domains are not established independently but influence and sharpen each other
- ✓ subdivide tiers into subnetworks → this creates modules that can be reused and starts a hierarchical architecture





Morphological refinement by iterative growth

✓ details are not created in one shot, but gradually added. . .





 \checkmark . . . while, at the same time, the canvas grows





from Coen, E. (2000) The Art of Genes, pp131-135









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Quantitative mutations: limb thickness

Quantitative mutations: body size and limb length

Qualitative mutations: limb position and differentiation

Qualitative mutations: 3rd-level digits

Toward More Realistic Biomodeling

top-down, abstract, heuristic approach

. . . .

bottom-up, data-driven, induction approach

Toward More Realistic Biomodeling

More accurate mechanics

- ✓ 3-D
- ✓ individual cell shapes
- ✓ collective motion, migration
- ✓ adhesion

Better gene regulation

- ✓ recurrent links
- ✓ gene reuse
- ✓ kinetic reaction ODEs
- ✓ attractor dynamics

Toward More Realistic Biomodeling

Latest progress

- ✓ 3D particle-based mechanics ≟
- ✓ kinetic-based gene regulation

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