Brain Science at the Interface of Biological, Physical, and Mathematical Sciences, Computer Science, and Engineering: Analysis of New Opportunities

- 1. How does the brain encode, represent, and transform information?: What are the essential computational properties of the brain?
- 2. Is it possible to develop "biomimetic" neural systems that capture the essential computational properties of the brain?
- 3. Can biomimetic computational systems interact with the brain through real-time, bi-directional communication?

- 1. How does the brain encode, represent, and transform information?: What are the essential computational properties of the brain?
	- advent of multi-neuron recording / fMRI in intact, behaving animals and humans
	- opportunity to explore population dynamics of neural coding: spatiotemporal coding
	- what information can be coded in the spatio-temporal capacity of populations of neurons? – given variability among neurons
	- how does that capacity depend on underlying cellular/molecular mechanisms?
	- how is information transformed as it propagates through the nervous system?
	- how do these transformations map onto "cognitive functions" higher thought processes?
	- advantages to biological coding yet to be discovered?

Multi-Site Recording Array Technologies

- microwire arrays; multiple arrays
- silicon-based micro-fabricated electrodes
- ceramic-based micro-fabricated electrodes; chemical sensors
- need to push the envelope of multi-site recording array technology

Advanced Algorithms for Decoding **Brain Computations**

- population vector
- optimal linear est.
- linear dynamic models
- nonlinear dynamic models
- adaptive models

Song, Berger – USC

5000
Time (msec

9000

Georgopolous – Univ. of Minnesota

Schwartz – Univ. of Pittsburgh

Electrophysiological Indices of Cognitive States and Operations

Delayed Non-Match-to-Sample Memory Task

Deadwyler – Wake Forest Univ. Eichenbaum – Boston Univ.

- 2. Is it possible to develop "biomimetic" neural systems that capture the essential computational properties of the brain?
	- increased fundamental anatomical, physiological, etc., knowledge of neural systems; increased computing power and computational hardware
	- should be able to start building systems that really act like biological systems – computational, robotic, "cognitive" systems, etc.
	- neuronal processes are strongly nonlinear: what types and orders of nonlinearity need to be considered?; nonstationarities (learn./develop.)
	- what does it take to put together "mini-systems" of a cortical column?, a hippocampal slice?, a cerebellar folia? a retina?
	- what types of hardware representations of such neural systems can be realized?
	- can we develop hardware platforms that are both large-scale and flexible to use as investigational tools?

Experimentally-Based, Biomimetic Models of Neural Systems

- "Blue Brain" Project (H. Markram, EPFL, Switzerland) – experimentallybased, functional, structural, biochemical model of a neocortical column – simulation of a 10,000 neuron system using an IBM multiprocessor "Blue-Gene" computing system
- "Neurogrid" Project (K. Boahen, Stanford Univ.): emulating millions of visual cortical neurons using multiprocessor arrays; programmable, hybrid, analog-digital circuitry

Markram – EPFL, Switzerland

Cellular Neural Network (CNN) implementation of a biologically-based, multi-layer model of the retina – Werblin, UC Berkeley; Roska, Hungarian Academy of Sciences

Biomimetic Neural Systems that Capture the Essential Computational Properties of the Brain

DRAM

Multi-Level, Hierarchical Models of the Nervous System: Relating Molecules to Synapses to Neurons to Networks

$$
(a) \frac{\partial \psi}{\partial t} (r_0, t_0) = \nabla_x [D' \nabla_x \psi(r_0, t_0)] + \int_{D_r(r_0)} \rho_p(r) \psi(r, t)
$$

$$
\int_{D_r(r, t_0)} P_r(r_0, t_0, r, t) \pi_g(s, r; r_0) \sigma(s, t) ds dr + \Gamma_x [\psi(r_0, t_0); \psi_{r_0 f_0}]
$$

$$
(b) \frac{\partial \mu}{\partial t} (s, t) = \nabla_x [D' \nabla_x \psi(s, t)] + \int_{D_r(r_0)} \rho_p(r') \int_{D_r(r, t_0)} P_r(s, s') \pi_g(s', r'; r_0) \mu_0(s', t') ds' dr' + \Gamma_x [\mu(s, t)] \quad (4)
$$

Chauvet – Univ. of Angers, France

- need for a comprehensive theoretical framework that can account for molecular, cellular, system, and multi-system function
- capable of taking into account multiple time and space scales of the nervous system
- capable of relating changes at one level of the neuronal hierarchy to changes at other levels
- e.g., n-level field theory of Chauvet; also see J. Cowan

- 3. Can biomimetic computational systems interact with the brain through real-time, bi-directional communication?
	- advanced tissue culture "designer circuit" methods: neuron-silicon "co-interrogation" of reduced neural systems
	- biomimetic system-neural system "co-processing of information" in the brain: can we detect fundamental brain operations such as object recognition or memory formation?
	- can we develop biomimetic systems that assist or enhance fundamental brain operations
	- can we develop biomimetic systems that replace damaged brain regions – implantable neural prostheses
	- can we develop biomimetic systems that actively "query" the brain as to its "state" or its current "computational operation" – hybrid systems that "learn how the brain works"

Surface Chemical Patterning to Create "Designer" Neural Circuits that Interface Predictably with Silicon-Based Multi-Site Electrode Arrays

- surface chemical patterning to attract and adhere cultured neurons to "nodes" in a sensor grid
- physical and molecular constraints to shape "directionality" of circuit structure and function
- combined recordingstimulation protocols to understand computational properties of reduced neural circuits
- possibility of controlling progressively greater complexity of neural circuits and systems

Wheeler – Univ. of Illinois, Urbana-Champaign J. Hickman – Univ. Central Florida

Restoring Hippocampal Nonlinear Dynamics After Replacing CA3 with a Biomimetic, Hardware Model

- *nonlinear dynamic model of the CA3 hippocampal region*
- *experimental removal of the CA3 region in the slice*
- *substitution of CA3 with a hardware model restores circuit nonlinearities*

