# From Neuron to Cognition: What is needed to understand brain mechanisms of higher brain function?

Understanding brain function must be done at multiple levels. Starting from molecules ending at the behavior of a whole society.

EMBO (~1995): How can we understand the Brain using molecular techniques.

V. Braitenberg: For every problem there is the right scale, you do not want to read the newspaper with a microscope.

Hamula.... 100m

Family..... 10m

Body.....m

Brain.....10 cm

Region.....cm

Network...mm

Neuron.... 10 µm

Synapse...µm

Channel... nm

Atom.....Å

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Where is the largest gap in knowledge when brain mechanisms of cognition are concerned?

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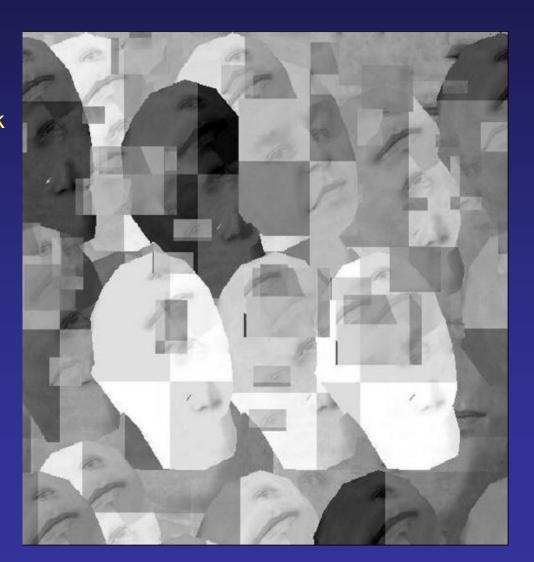
Atom.....Å

Where is the largest gap in knowledge when brain mechanisms of cognition are concerned?

Interactions among neurons in networks and networks among regions. Binding.

#### **Binding**

Courtesy of E. Bienenstock



#### **Neural networks**

Perceptron
Multi-layer Perceptron
Attractor Neural Networks
Syn-Fire Chains
Cannonical Cortical Circuit

#### The most needed technology:

Measure activity of each of many(~1000) neighboring neurons with high time resolution (~1ms).

#### **Expected results:**

What is the code?

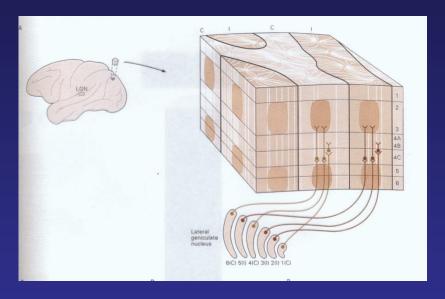
What neural network produces it?

How is it "read" by other networks?

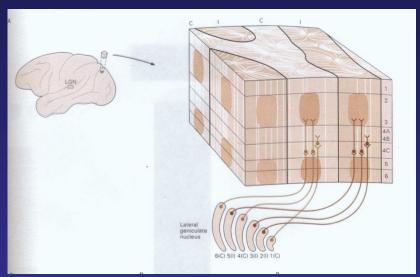
How are complex mental constructs built from 'simpler ones (the binding problem)?

What is the "most important" discovery?

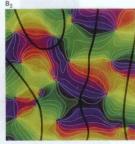
#### What is the most devastating discovery?



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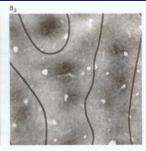


Figure 27-17 Organization of orientation columns, ocular dominance columns, and blobs in primary visual cortex.

A An array of functional columns of cells in the visual cortex contains the neural machinery necessary to analyze a discrete region of the visual field and can be thought of as a functional module. Each module contains one complete set of orientation columns, one set of ocular dominance columns (right and left eye), and several blobs (regions of the cortex associated with color processing). The entire visual field can be represented in the visual cortex by a regular array of such modules.

B. Images depicting ocular dominance columns, orientation columns, and blobs from the same region of primary visual cortex (Courtesy of Gary Blasdel.) 1, Images of ocular dominance

columns were obtained using optical imaging and independently stimulating the left and right ocular dominance columns in a particular region. Because neural activity decreases cortical reflectance, the subtraction of one left eye image from one right eye image produces the characteristic pattern of dark and light bands, representing the right and left eyes respectively. 2. In this image the borders of the ocular dominance columns shown in 1 appear as black lines superimposed on the pattern of orientation-specific columns depicted in Figure 27-14. 3. The borders of the ocular dominance columns shown in 1 are superimposed on tissue reacted for cytochrome oxidase, which visualizes the blobs. The blobs are thus seen localized in the centers of the ocular dominance columns.

#### Example auditory cortex

#### Response of neighboring neurons in A1 to tone sweeps.

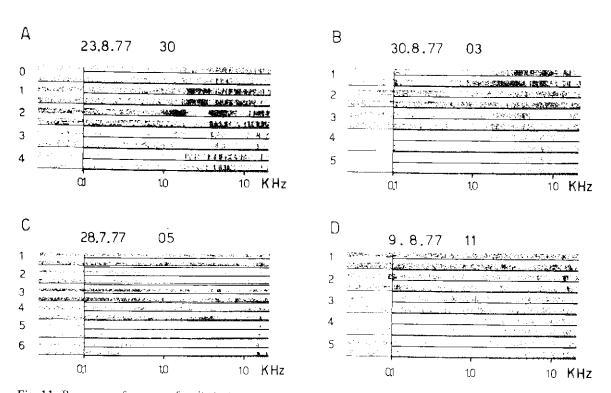


Fig. 11. Response of groups of units to tone sweeps.

Responses of few groups of cells from different experiments are shown. Same representation as in Fig. 10.

(By courtesy of R. Frostig)

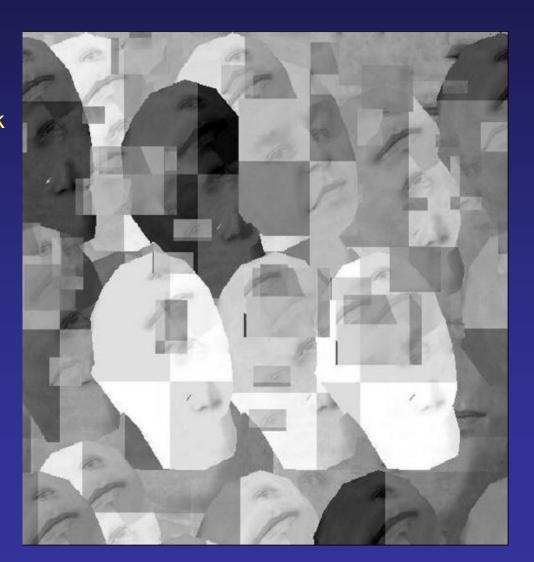
Abeles, 1982 Local Cortical Circuits

What is the "most important" discovery? What is the most devastating discovery? What is the most devastating idea?

It is sufficient to record one neuron at a time to understand how neurons interact while processing information.

#### **Binding**

Courtesy of E. Bienenstock



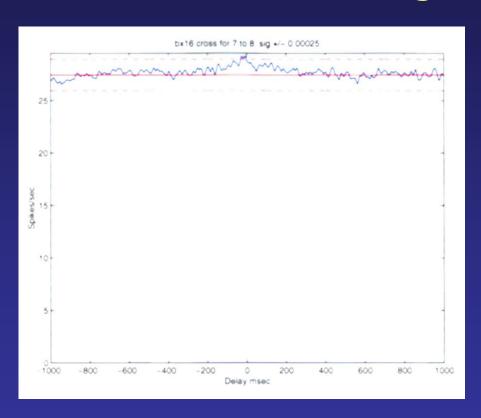
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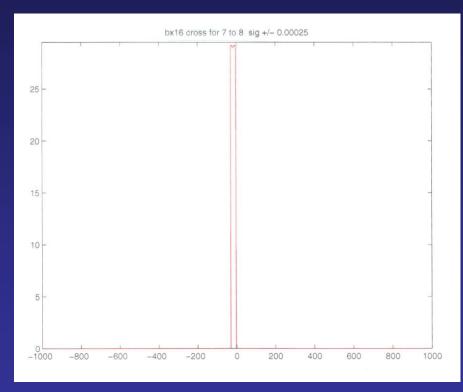
It is sufficient to record one neuron at a time to understand how neurons interact while processing information.

One can understand how the brain processes information by in a bottom up fashion alone. (Blue Brain, Digital Anatomy)

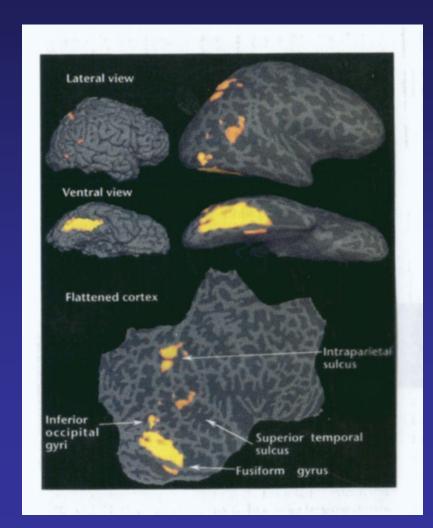
What is the "most important" discovery?
What is the most devastating discovery?
What is the most devastating idea?
What is the most devastating technology?

#### The most devastating technology:

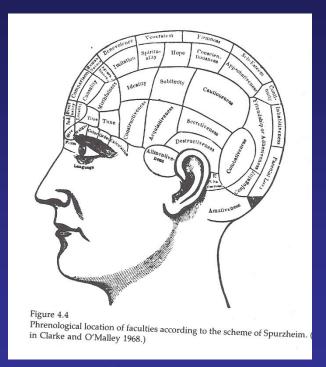




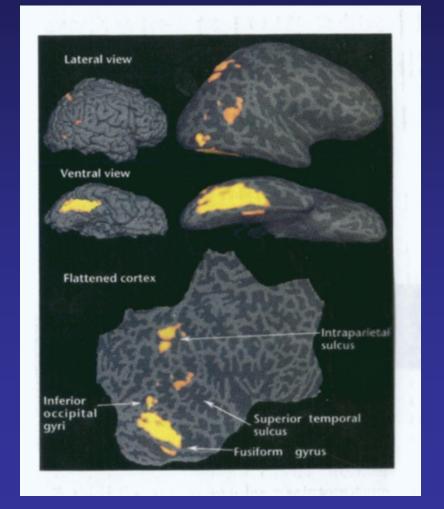
#### The most devastating technology:



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#### Neophrenology

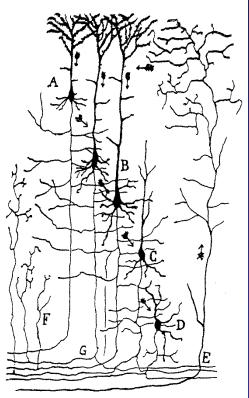


What is the "most important" discovery?

**Early** 

Cajal – The Neuron doctrine

Direction of processing



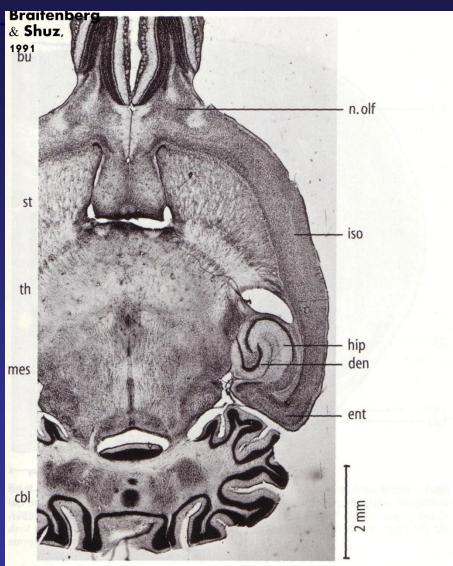
[FIG. 37] Fig. 13. Scheme designed to show the probable course of the [impulses] and the axodendritic connections in the cells of the cortex. A, small pyramid; B, giant pyramid; C and D, polymorphic cells; E, terminal fiber arriving from other centers; F, [axonal] collaterals of the white matter; G, axon bifurcating in the white matter.

Calel 1892

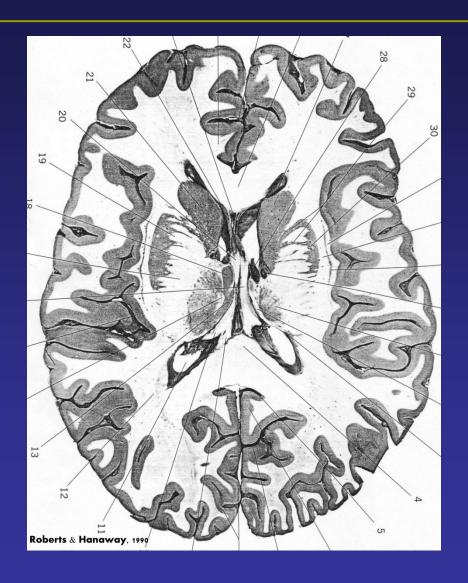
Anda de Cienças Medicas de Barcellara 18: 457-445

## Macroscopic View

## Horizontal section through the mouse brain



**Fig. 2.** Horizontal Nissl-stained section through the mouse brain. Demarcation of the olfactory bulb from the cortex. Transition of the cortex into the hippocampus. *bu* olfactory bulb; *st* striatum; *th* thalamus; *mes* mesencephalon; *cbl* cerebellum; *n.olf* olfactory nucleus; *iso* isocortex; *hip* hippocampus; *den* dentate gyrus; *ent* entorhinal cortex



What is the "most important" discovery?

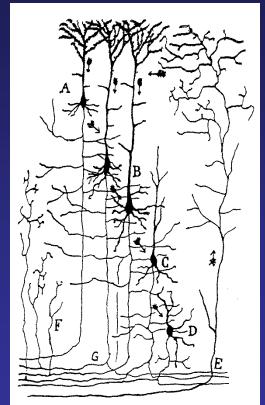
#### Early

Cajal – The Neuron doctrine

Direction of processing

#### Late

Braitenberg – Quantitative anatomy



[FIG. 37] Fig. 13. Scheme designed to show the probable course of the [impulses] and the axodendritic connections in the cells of the cortex. A, small pyramid; B, giant pyramid; C and D, polymorphic cells; E, terminal fiber arriving from other centers; F, [axonal] collaterals of the white matter; G, axon bifurcating in the white matter.

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Amola de Ciencias Medicas de Barcellana 18: 457-446

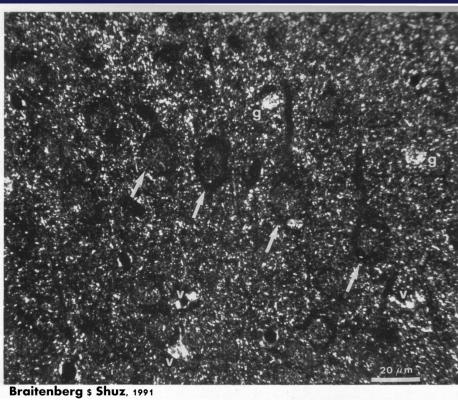
#### Quantitative anatomy

Table 1.5.4. Typical compositions of cortical tissues

Variable	Value
Neuronal density	40,000/mm <sup>3</sup>
Neuronal composition:	
Pyramidal	75%
Smooth stellate	15%
Spiny stellate	10%
Synaptic density	$8 \cdot 10^8 / \text{mm}^3$
Axonal length density	$3,200 \text{ m/mm}^3$
Dendritic length density	$400 \text{ m/mm}^3$
Synapses per neuron	20,000
Inhibitory synapses per neuron	2,000
Excitatory synapses from remote sources per neuron	9,000
Excitatory synapses from local sources per neuron	9,000
Dendritic length per neuron	10 mm

From: Corticonics Abeles 1991





Is this because "nature" cannot build orderly nervous system?

Braitenberg: On the texture of brains.

Abeles 2003

These numbers are crucial because they tell us that there is plenty of hardware to carry very complex computations.

The idea that all the neurons in a small piece of cortex do essentially the same thing seem ridiculous and does not fit the electrophysiology of single units in awake, behaving, animals.

What then is needed?

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Atom.....Å

Where is the largest gap in knowledge when brain mechanisms of cognition are concerned?

Interactions among neurons in networks and networks among regions. Binding.

Theories about the network structure in the cortex.

Measurements to try and refute the theories.

#### The most needed technology:

Measure activity of each of many(~1000) neighboring neurons with high time resolution (~1ms).

## Thank you