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Harvard Medical School

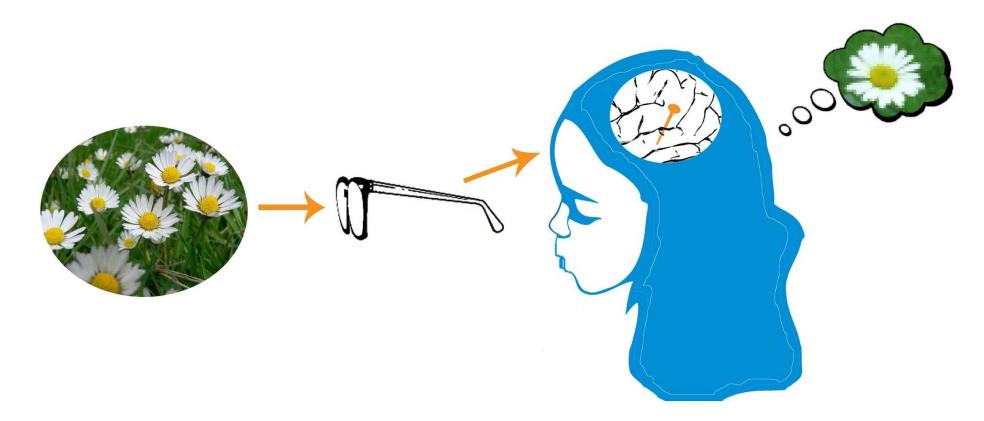
#### **Collaborators and Disclosures**

- HMSClay Reid
- MGH
   Emad Eskandar
   Nathan Killian
   Milena Vurro
   Bechir Bourkiza
   Margee Kyada
   Sarah Keith
   Ailsa Jeffries
   Anne Marie Crowell
- Disclosures
   No financial conflicts
   Speculative use of therapies not approved by FDA

#### The Problem: Blindness

- Blindness in Developed Countries
  - millions affected
  - eyes no longer sensitive to light
    - glaucoma
    - macular degeneration
    - retinitis pigmentosa
    - trauma (military personnel, civilians, children)
    - surgical removal
    - (congenital?)
  - rest of visual brain intact
  - is system repairable?

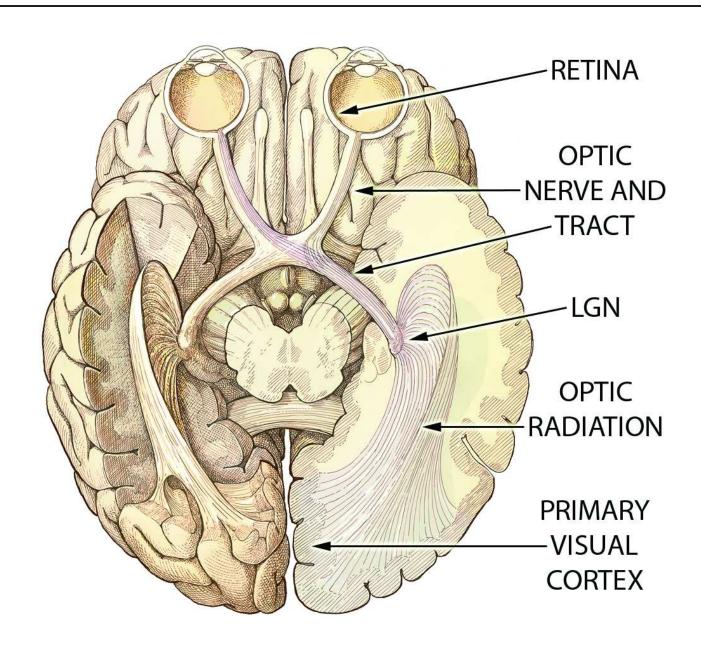
#### **A Solution: Visual Prosthesis**



Digital Camera + Wireless Electrode = **Sight for** Eyeglasses + Implant in Brain **the Blind** 

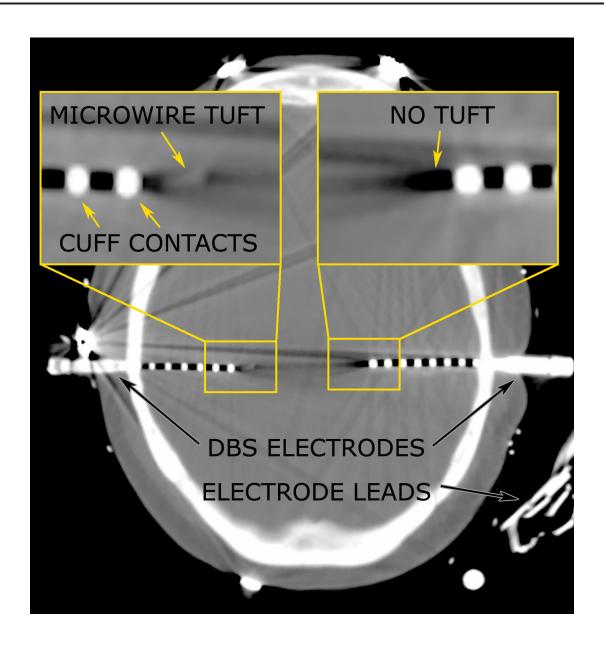
www.sight2blind.org

## **Early Visual Pathway**

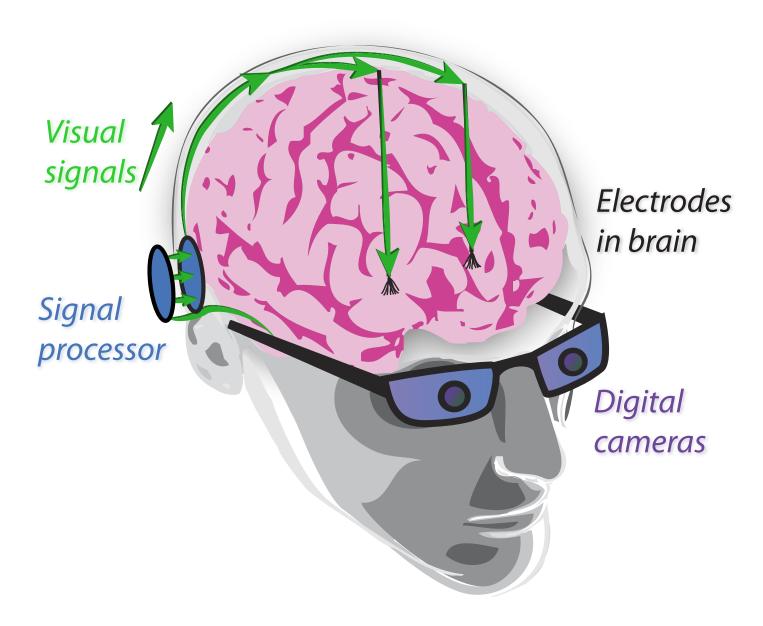


Pezaris, Eskandar, 2009, Neurosurgical Focus

## **DBS** / Depth Electrodes in the Clinic



## Prosthesis through Microstimulation in LGN



### Thalamic Visual Prosthesis Project

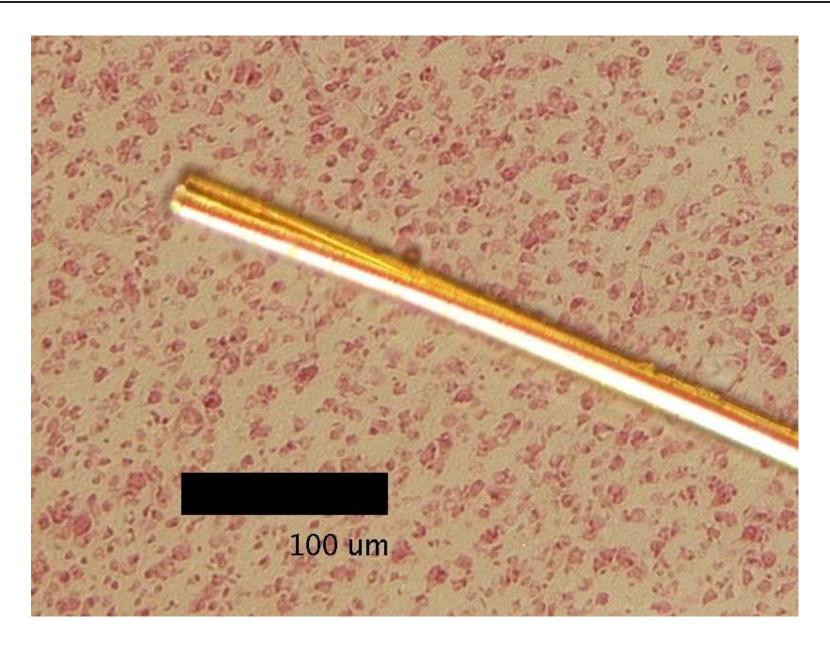
- Expt. 1: Proof of Concept
- Expt. 2: Engineering Validation
- Expt. 3: Human Acuity
- Expt. 4: Human Reading
- Expt. 5: Monkey Acuity
- Expt. 6: Monkey Prosthetic Vision
- Expt. 7: Human Prosthetic Vision

### **Experiment 1: Proof of Concept**

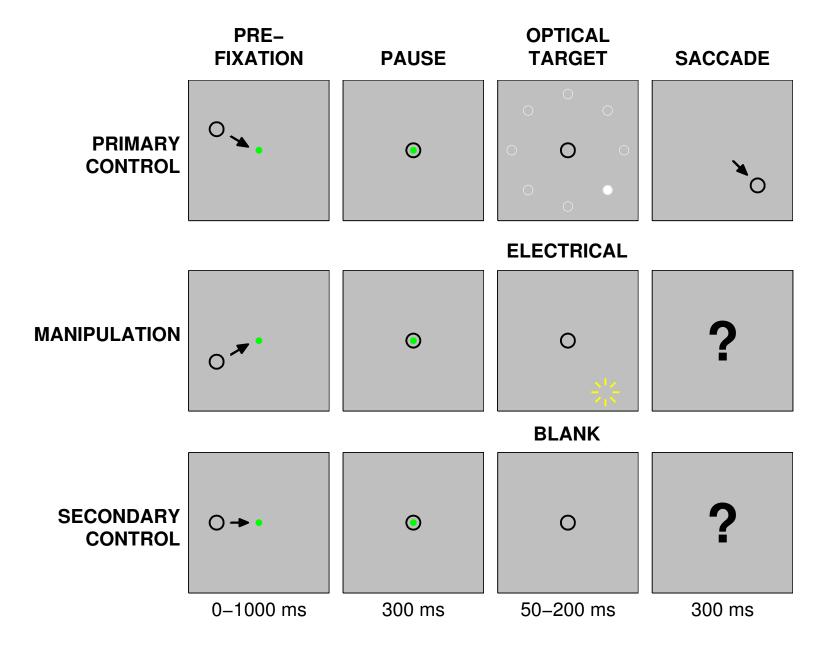
Q: Does LGN microstimulation elicit predictable visual percepts?

(Electrical stimulation in monkeys with behavioral report.)

# **Experiment 1: Methods**



## **Experiment 1: Methods**



## **Experiment 1: Results**

vid/pezaris-reid-2007-pnas-movie-sh

Movie of monkey saccading from central optically presented point to peripheral points presented both optically and electrically.

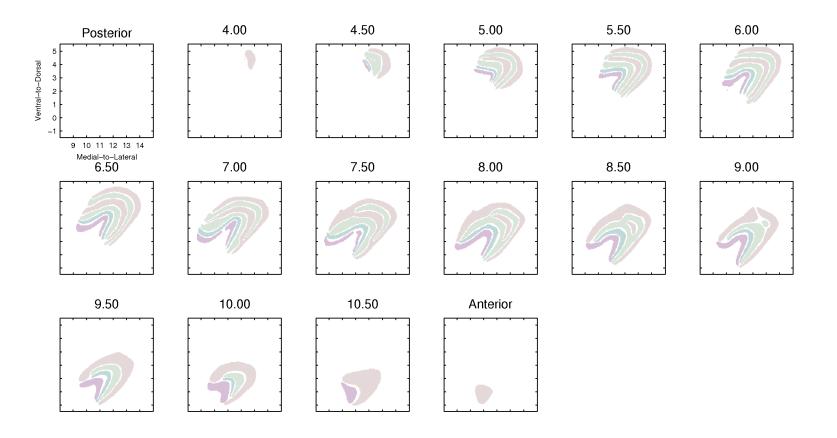
# Expt 2: How Many Electrodes Fit in LGN?

Q: How many microwires fit comfortably in LGN?

(Computer stimulation.)

### **Experiment 2: Methods**

- electrode contacts evenly spaced in 3D
- how many fit in LGN?
- what does it look like?



### **Experiment 2: Results**

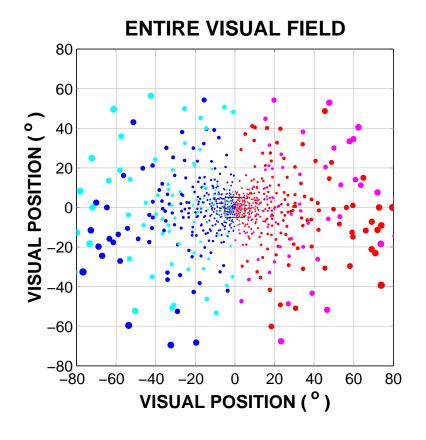
$\lambda$ ( $\mu m$ )	Total Points	Within 10°	Magno	Parvo
1000	110 $\pm$ 4	<b>24</b> ± <b>2</b>	<b>12</b> ± <b>3</b>	<b>43</b> ± <b>3</b>
800	217 $\pm$ 4	<b>48</b> ± <b>3</b>	<b>23</b> ± <b>3</b>	<b>87</b> ± <b>4</b>
<b>⇒ 600</b>	<b>522</b> ± <b>8</b>	<b>124</b> ± <b>6</b>	<b>52</b> ± <b>5</b>	$211 \pm  7$
400	$\boldsymbol{1707}{\pm}\boldsymbol{13}$	$\textbf{372} \! \pm  \textbf{7}$	$\textbf{180} \! \pm  \textbf{8}$	$686 {\pm} 11$

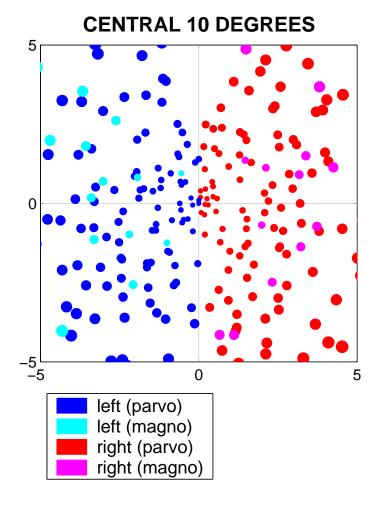
TABLE 1. LGN Electrode Tip Count for Isotropic Placement Matrix. The number of electrode tips landing in LGN were counted for varying mean densities based on Cartesian electrode spacing  $(\lambda)$ . The total count of electrode tips falling bilaterally in LGN is given, along with the number of these which fell within the central 10 degrees of visual space, and the breakdown for those in magnocellular and parvocellular layers.

Adapted from Pezaris, Reid, 2009, IEEE-TR-BME

### **Experiment 2: Results**

- 500 total electrodes / 250 per hemisphere
- 110 total within 10 degrees

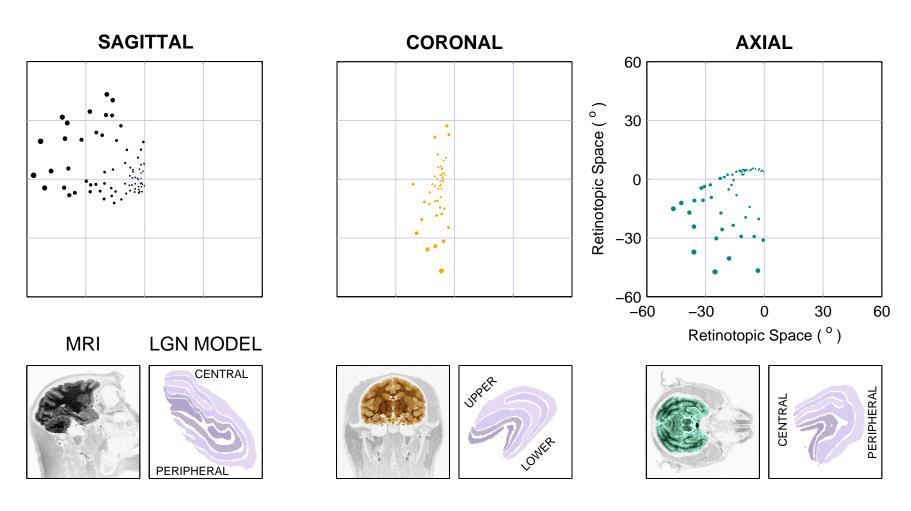




Pezaris, Reid, 2009, IEEE-TR-BME

### **Experiment 2: Results**

Assessment of Surgical Approach (50 contacts)



Pezaris, Reid, 2009, IEEE-TR-BME

### **Experiment 2: Simulated Artificial Vision**

vid/pezaris-simulated-prosthetic-vision.flv

Movie of simulated artificial vision.

# **Experiment 3: Acuity vs Electrode Count**

Q: What acuity with reasonable electrode counts?

(Virtual Reality simulation in sighted humans.)

# **Experiment 3: Background**



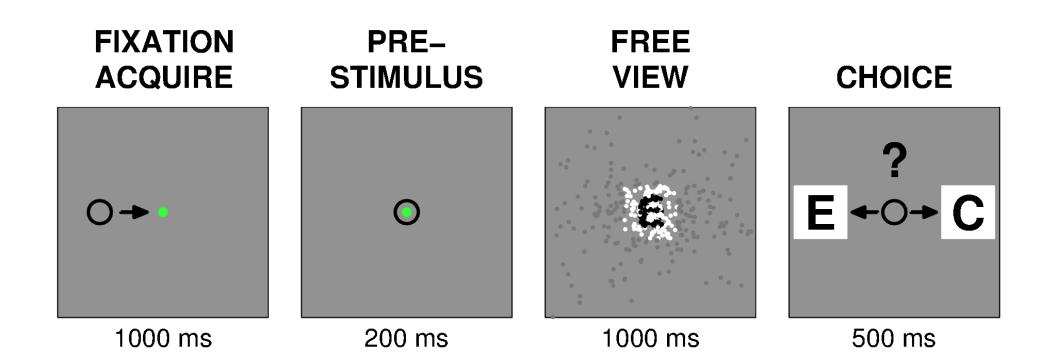
**Snellen Chart for Visual Acuity** 

## **Experiment 3: Methods**



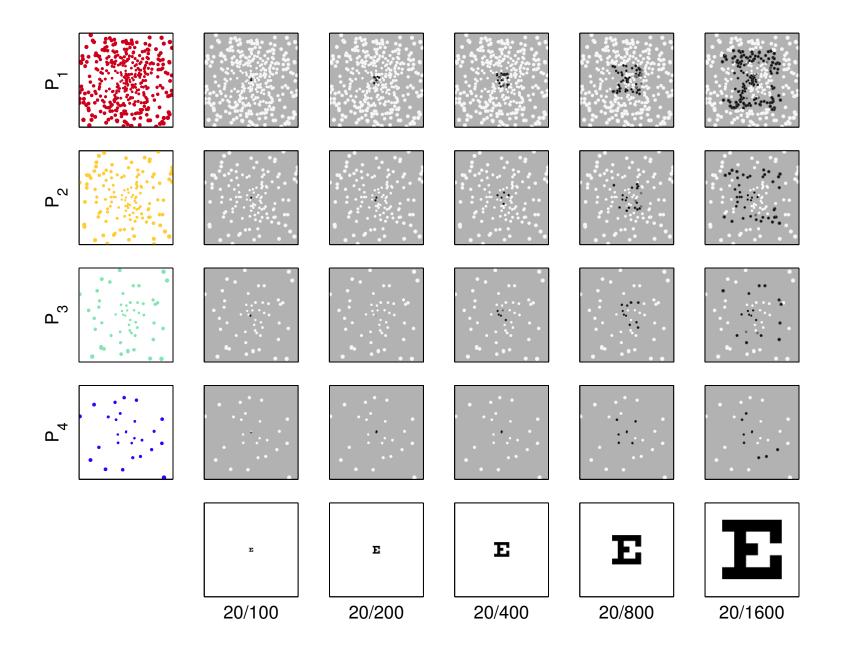
Bourkiza, Vurro, Jeffries, Pezaris, 2013, PLoS ONE

## **Experiment 3: Methods**



**2AFC** Letter Recognition Task

# **Experiment 3: Methods**

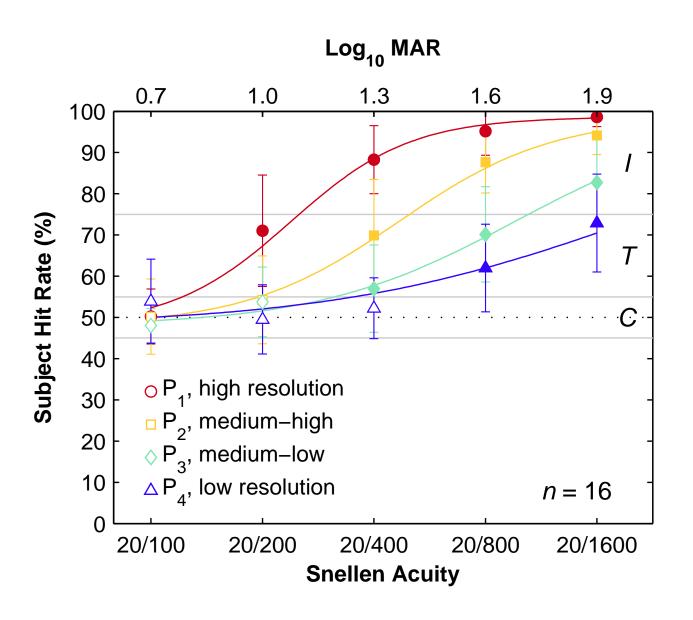


## **Experiment 3: Movie**

vid/pezaris-letter-recognition-movi

Movie of human letter recognition task.

### **Experiment 3: Results**



# **Experiment 4: Human Simulations, Reading**

Q: Can LGN phosphene patterns be used to read?

(Virtual Reality simulation in sighted humans.)

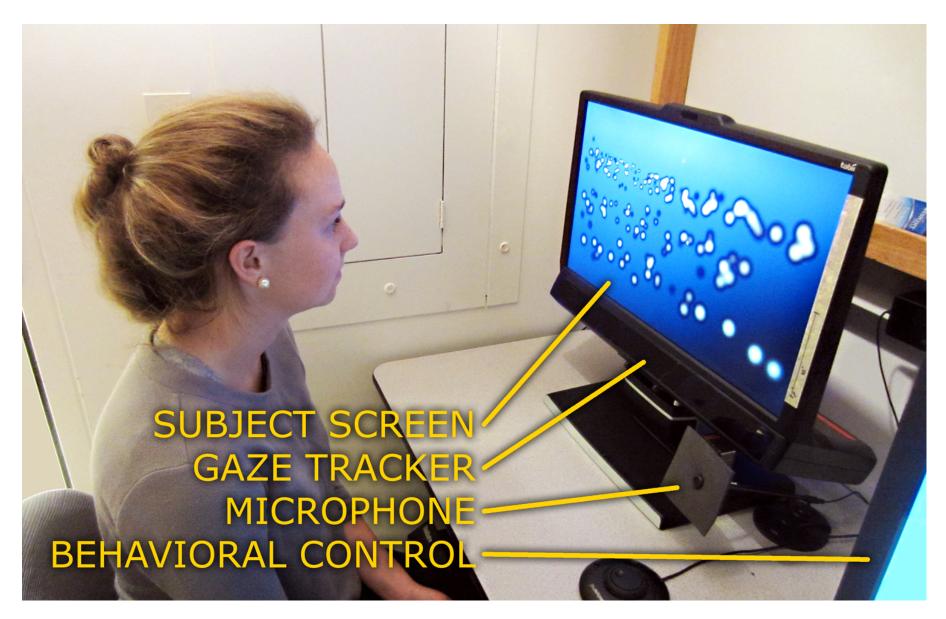
# **Experiment 4: Background**



	MNREAD* LOW-VISION READING ACUITY CHA	NRT 1			
M size	Mr. fathan asked ma	Snellen Iar 40cm (1	Snellen logMAR for 45cm (15 inches)		
	My father asked me				
4.0	to help the two men	20/200	1.0		
	carry the box inside				
	Three of my friends				
3.2	had never been to a	20/160	0.9		
circus before today					
	My grandfather has				
2.5	a large garden with fruit and vegetables		0.8		
2.0	He told a long story about ducks before	20/100	0.7		
_	his son went to bed		•••		
1.6	My mother loves to hear the young girls	20/80	0.6		
_	sing in the morning	200.50			
1.3	The young boy held his hand high to ask quantities in school	20/63	0.5		
1.0	this, breathers resemble against of mallit with his online offers breach	20/50	0.4		
0.8	1 de set minorene de mindig fair to play	20/40	0.3		
0.6	Selection description of the selection o	20/32	0.2		
0.5 0.4	BATTANIA BATTANIA	20/25 20/20	0.1		

Minnesota Reading Task (MNREAD)

### **Experiment 4: Methods**



Vurro, Crowell, Pezaris, 2014, Front. Human Neurosci.

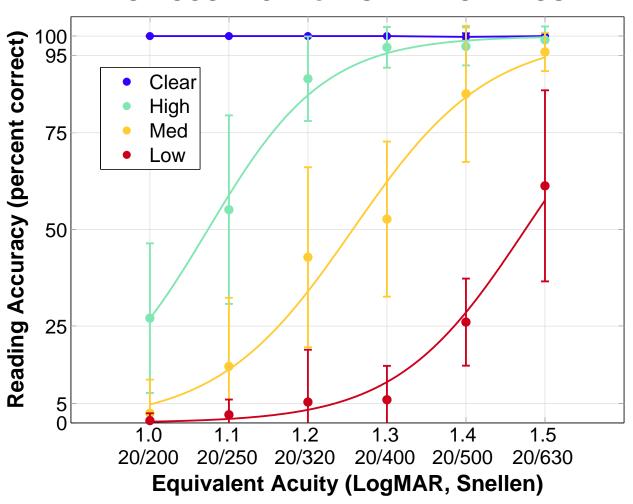
# Experiment 4: Movie (1800 phosphenes)

vid/reading-h-flowers.flv

Ten different kinds / of flowers grow by / the side of the road.

#### **Experiment 4: Results**

#### READING ACCURACY vs. NUMBER OF PHOSPHENES



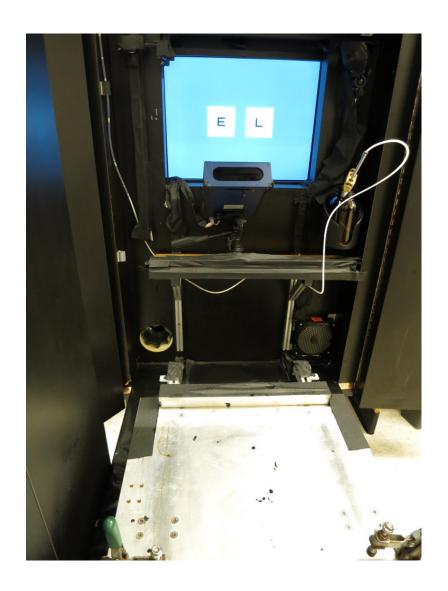
H,M,L: 1800(370), 1000(230), 500(124) Vurro, Crowell, Pezaris, 2014, Front Hum Neurosci

## **Experiment 5: Monkey Acuity**

Q: What is acuity available for reasonable electrode counts?

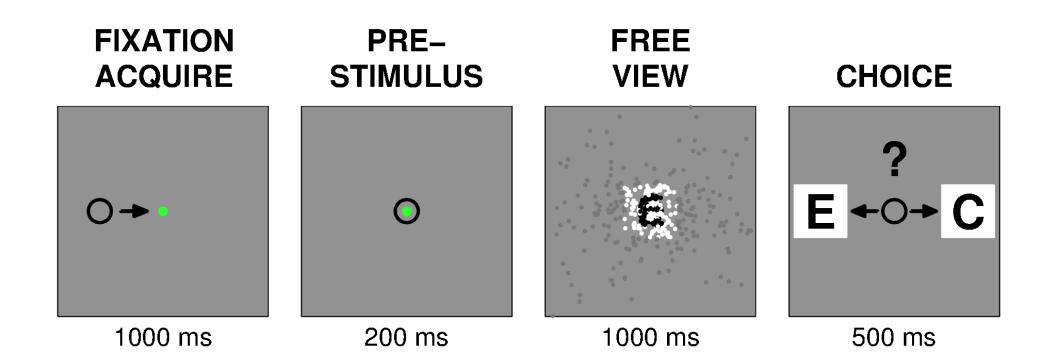
(Virtual reality simulation in sighted monkeys.)

# **Experiment 5: Methods**



Monkey version of human psychophysics setup

## **Experiment 5: Task**



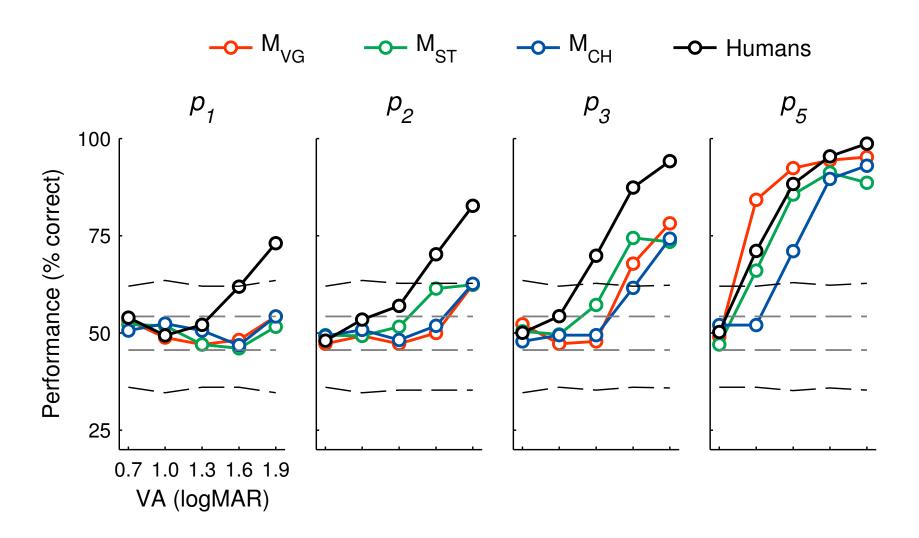
Same as human task, except . . .

#### **Experiment 5: Methods**

ABCDEFGHIJKLMNOPQRSTUVWXX

Stelio Font, inspired by Sloan optotypes

# **Experiment 5: Results, Acuity**



p1-p5: 250(50), 500(120), 1000(230), 2000(450) phosphenes

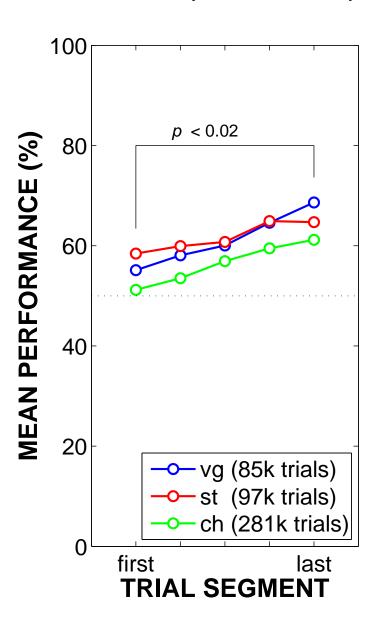
Killian, et al., under revision

## **Experiment 5: Results, Learning**

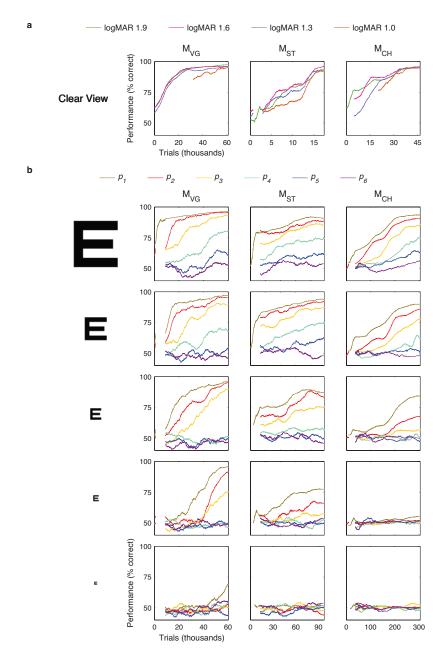
#### **Human** (population)

# 100 p < 0.02**MEAN PERFORMANCE (%)** 80 65% 60 40 20 200 trials/segment n = 16first last TRIAL SEGMENT

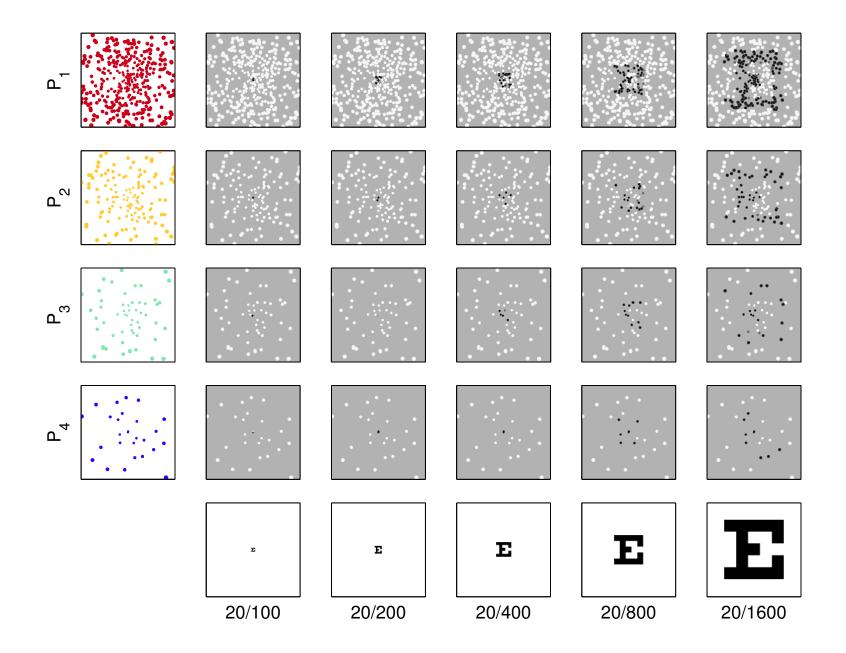
#### Monkey (individuals)



# **Experiment 5: Results, Learning**



# **Experiment 5: Results, Learning**



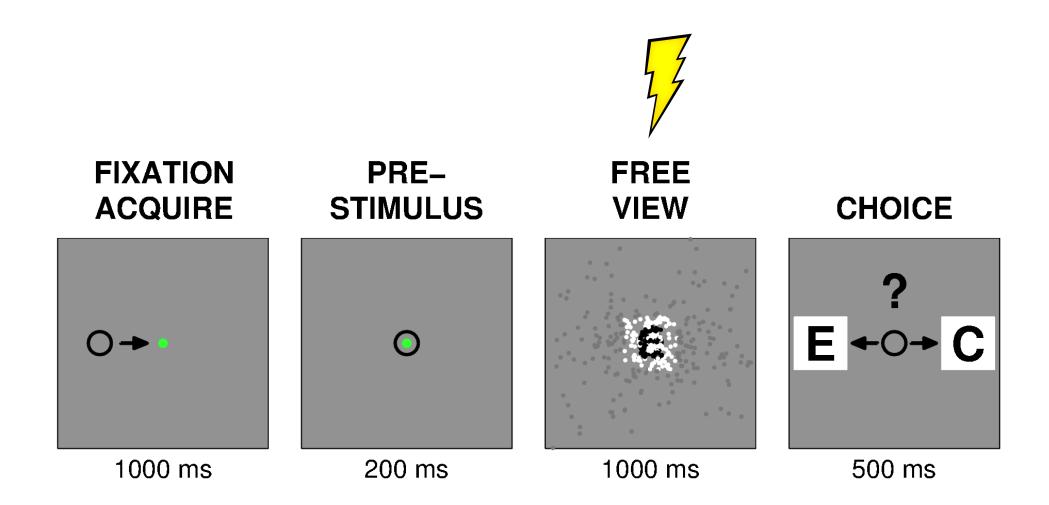
## **Experiment 6: Monkey Prosthetic Vision**

Q: Can monkeys use prosthetic vision in a cognitive task?

(Real visual prosthesis in sighted monkeys.)

— ongoing —

## **Experiment 6: Task**



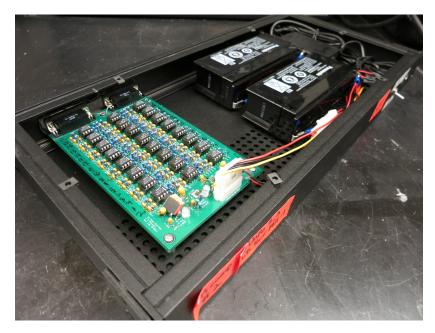
Task as before, except cues presented electrically.

# **Experiment 6: Methods**

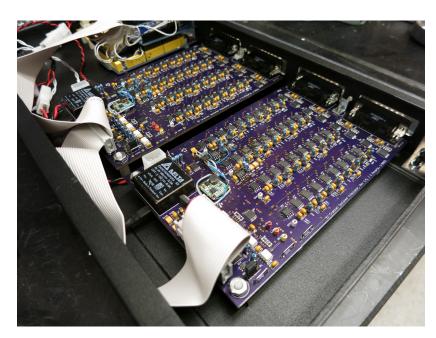


64 Channel Microwire Bundle Electrode

# **Experiment 6: Methods**



16 Channel Precision Amp



**64 Channel Stimulator** 

# **Experiment 6: Results**

— Results Shortly —

### **Experiment 7: Human Prosthetic Vision**

Q: Can humans use prosthetic vision in a cognitive task?

(Real visual prosthesis in blind humans.)

— Near-future work —

# **Summary**

Expt.	1:	Proof of Concept	•	•	•	•	•	•	•	done
Expt.	2:	<b>Engineering Validation</b> .					ple	enty	of	room
Expt.	3:	Human Acuity				go	od	perf	f <mark>or</mark> n	nance
Expt.	4:	Human Reading				go	od	perf	f <mark>or</mark> n	nance
Expt.	<b>5</b> :	Monkey Acuity					ł	nard	to	learn
Expt.	6:	Monkey Prosthetic Vision							on	going
Expt.	<b>7</b> :	Human Prosthetic Vision						ne	ar f	future

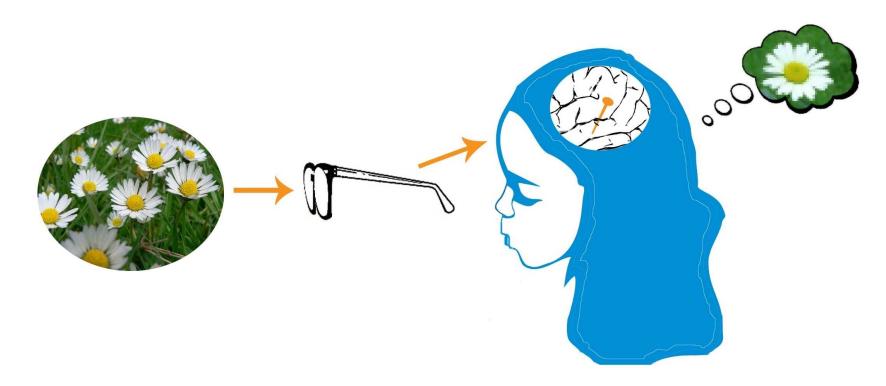
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Christopher Lewis
Michael Gersh
Teddy and Sumi Cho

### Sight2Blind



Digital Camera + Wireless Electrode = **Sight for** Eyeglasses + Implant in Brain = **the Blind** 

http://www.sight2blind.org