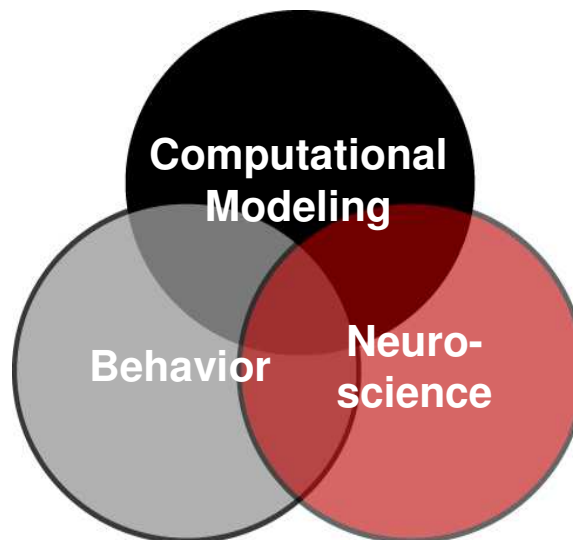




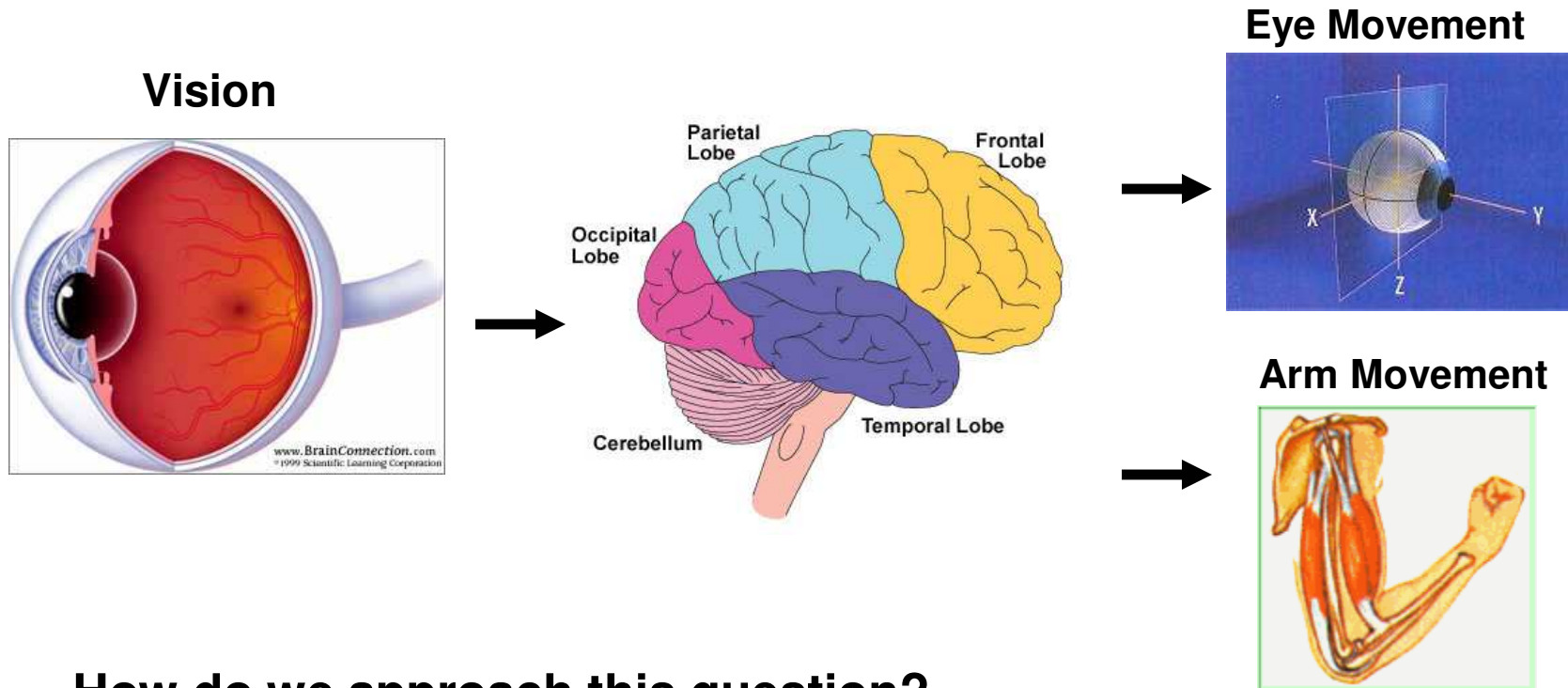
## Workshop on Perspectives for Future Directions in Computational and Mathematical Neuroscience

*Levels of theory in Sensorimotor Neuroscience*



## **General Workshop Goals:**

- (i) Definition of the field**
- (ii) Research highlights in the field**
- (iii) Critical considerations for someone wanting to enter the field today**
- (iv) Ideal type(s) of training**
- (v) Suggested changes and directions for the field.**



**How do we approach this question?**

**The Scientific Method: Experimental tests of hypotheses**

**Problem:** Intuitive hypotheses often do not capture the complexity of the brain and behavior

**Solution:** Formalize hypotheses as computational models that can be tested

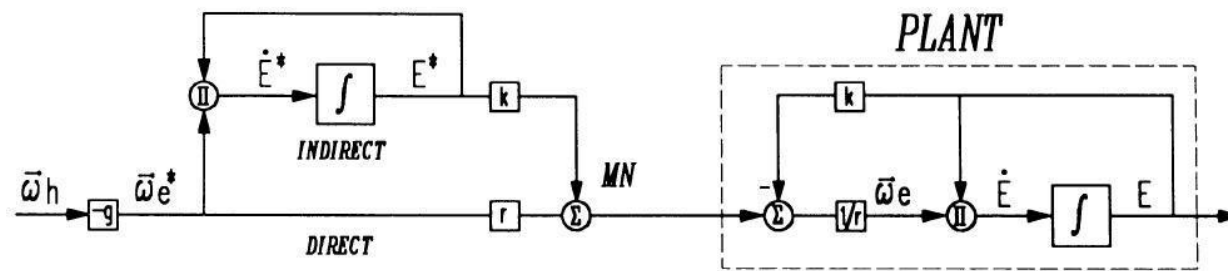
## Survey of Examples From My Lab:



- Role that Computational Modeling has played in our work
  - Vestibulo-ocular Reflex
  - Rapid Gaze Movements
  - Spatial Updating
  - Visually Guided Reach
- Key points:
  - Synergy between theory and experiment
  - Predictive power of models as formal Hypotheses
  - Moving between different levels of theory



### 3-D VOR Model (1988-1991)



Crawford & Vilis (1991) *J. Neurophysiol.*

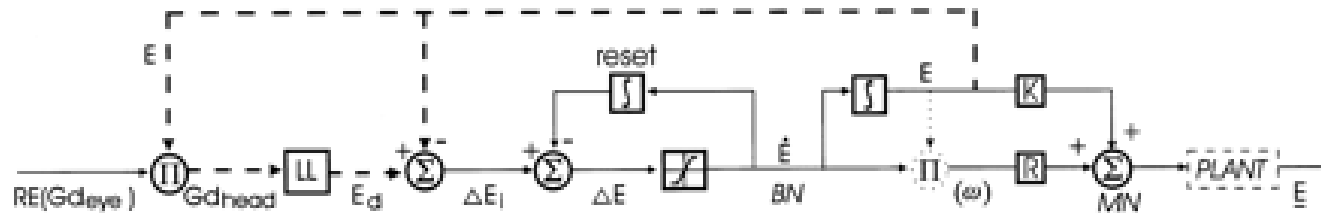
### Predictions

- 1) eye rotates about same axis as head during VOR
- 2) VOR violates Listing's law of eye

Crawford JD, Vilis T. (1991) Axes of eye rotation and Listing's law during rotations of the head. *J Neurophysiol* 65:407-23.

- 3) need a three-dimensional 'neural integrator'

Crawford JD, Cadera W, Vilis T. (1991) Generation of torsional and vertical eye position signals by the interstitial nucleus of Cajal. *Science* 252:1551-3.



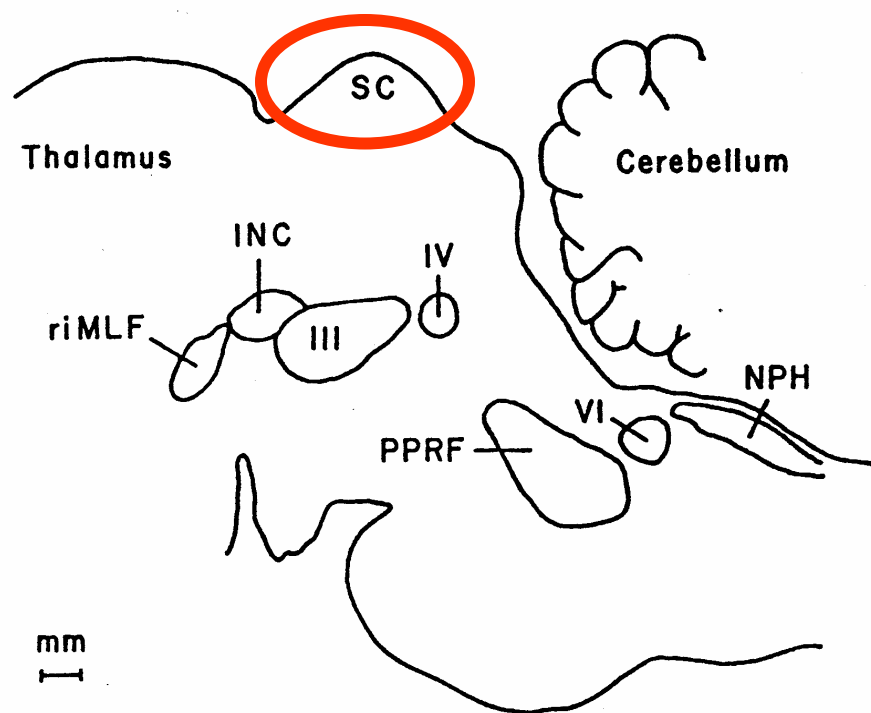
## Predictions

**Crawford JD. (1994) The oculomotor neural integrator uses a behavior-related coordinate system. *J Neurosci.* 14:6911-23.**

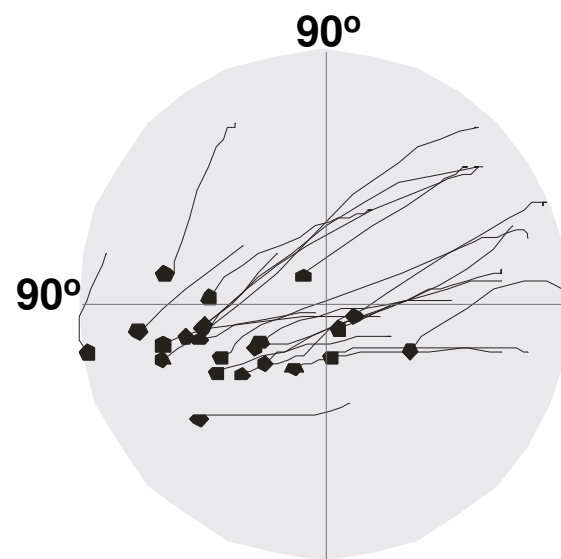
**Klier EM, Crawford JD. (1998) Human oculomotor system accounts for 3-D eye orientation in the visual-motor transformation for saccades. *J Neurophysiol.* 80:2274-94.**

**Klier EM, Wang H, Crawford JD. (2001) The superior colliculus encodes gaze commands in retinal coordinates. *Nat Neurosci.* 4:627-32.**

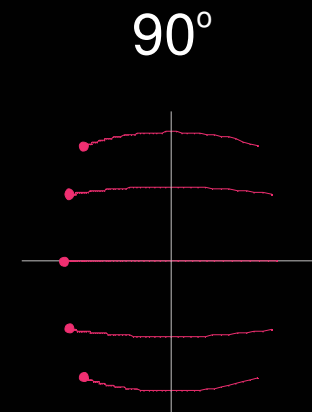
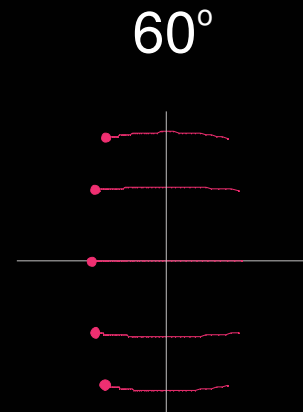
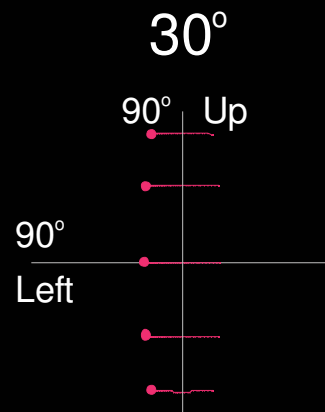
## Gaze Coding in Superior Colliculus (SC)



Gaze Shifts  
During SC Stimulation

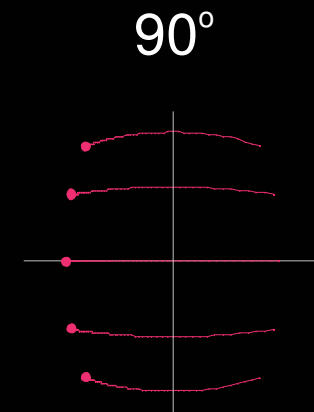
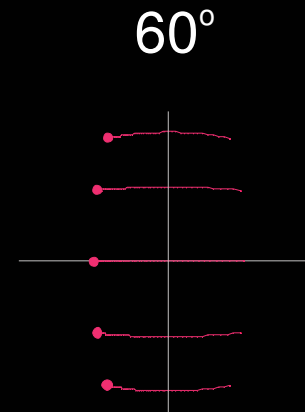
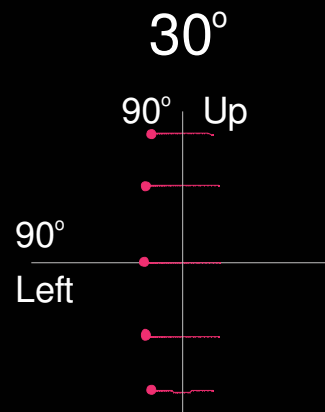


# Fixed-vector Model

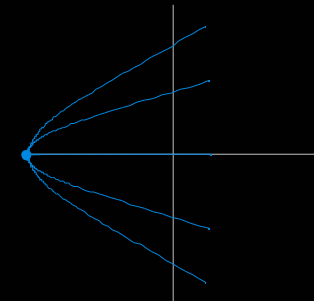
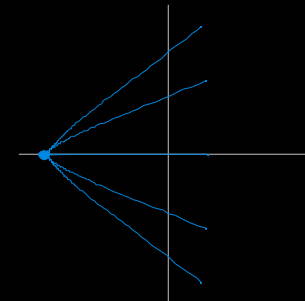
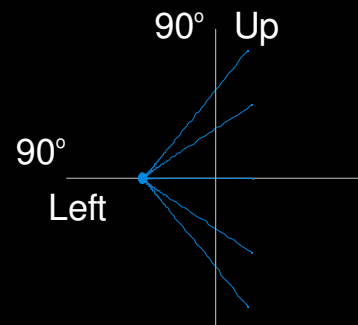




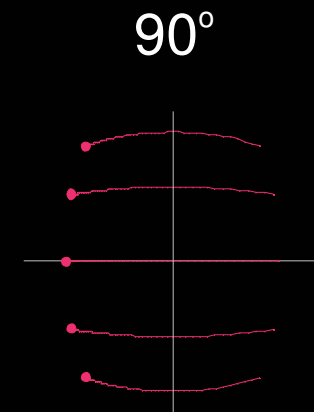
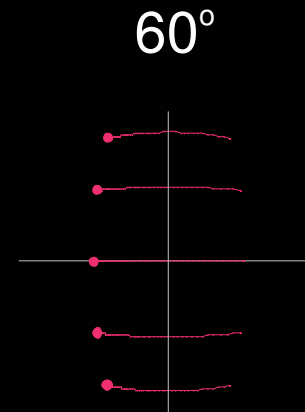
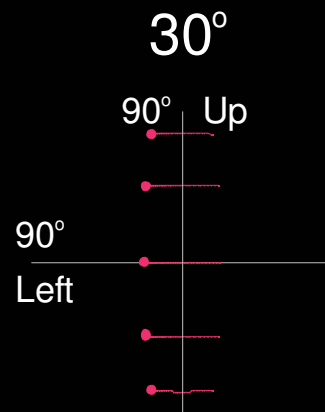
## Fixed-vector Model



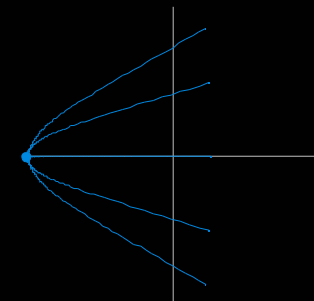
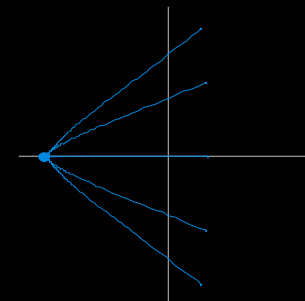
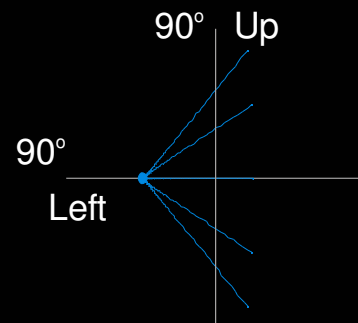
## Space-Centred Model



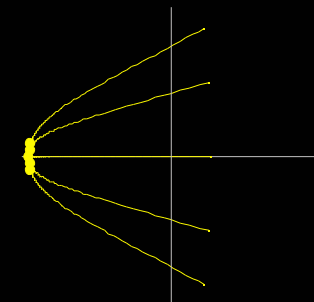
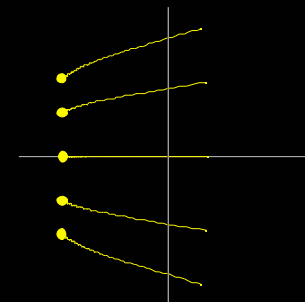
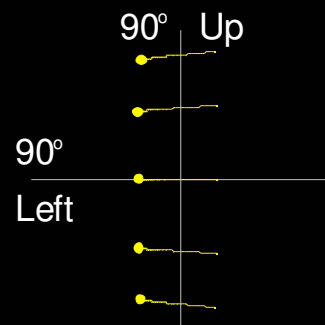
## Fixed-vector Model



## Space-Centred Model

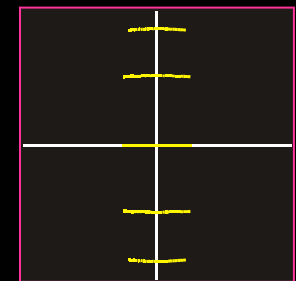
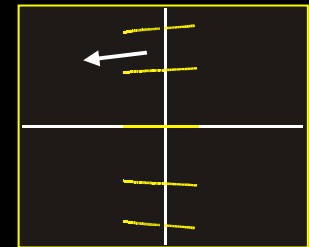
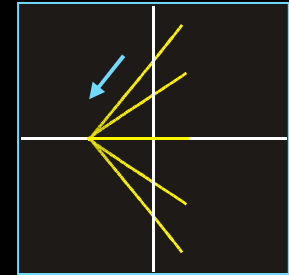
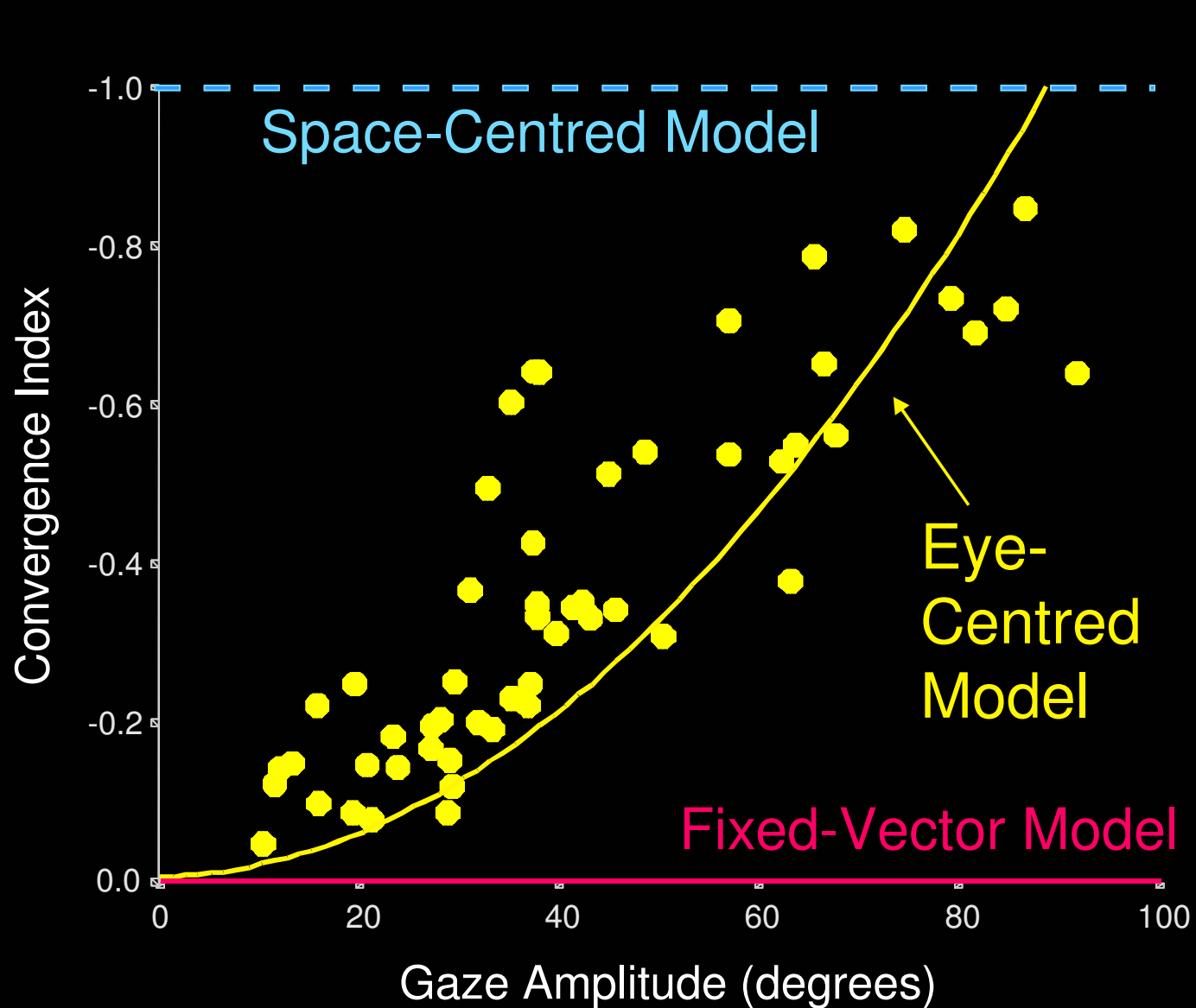


## Eye-Centred Model



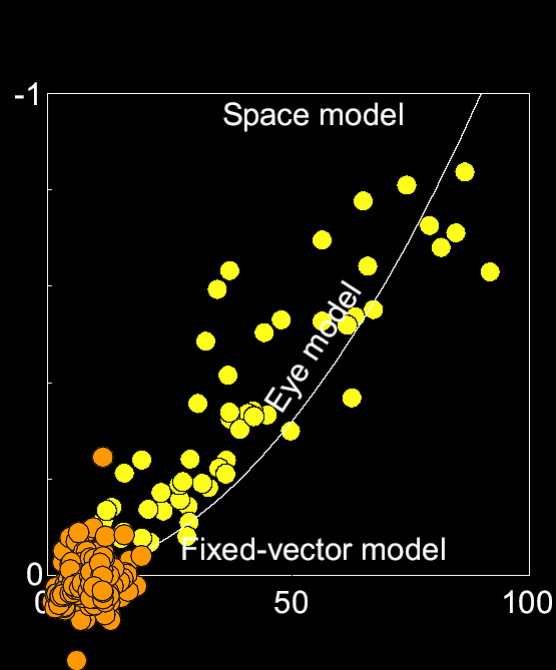
(Crawford & Guitton *J. Neurophysiol.* 1997)

(Stimulation Data: Klier et al. *Nature Neuroscience*, 2001)



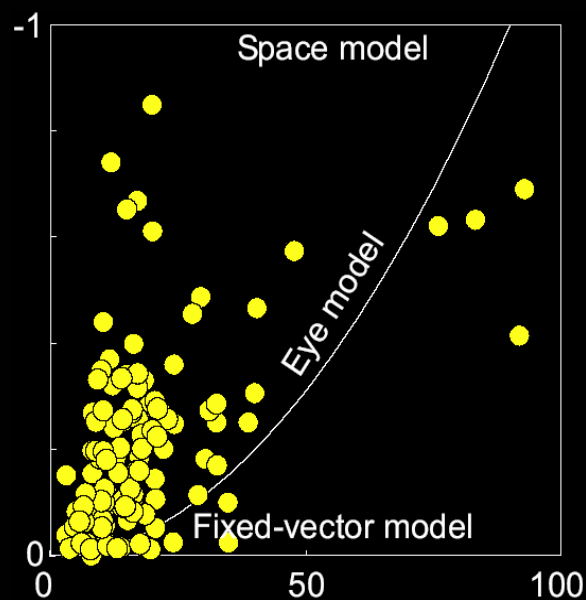
# Comparison across brain sites

SC & LIP



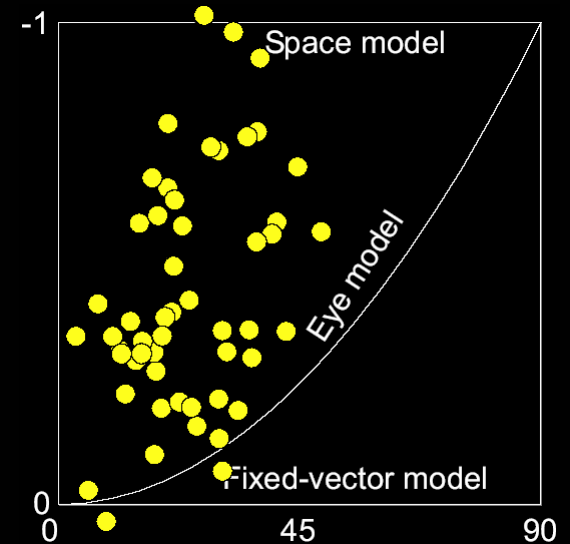
Eye-centered RF  
(Klier et al. 2001  
*Nature Neurosci.*)

FEF



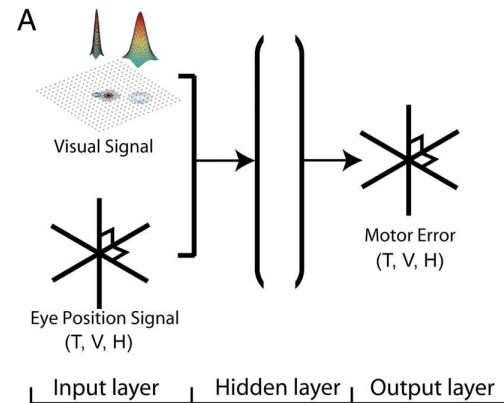
Transition between SC  
and SEF

SEF

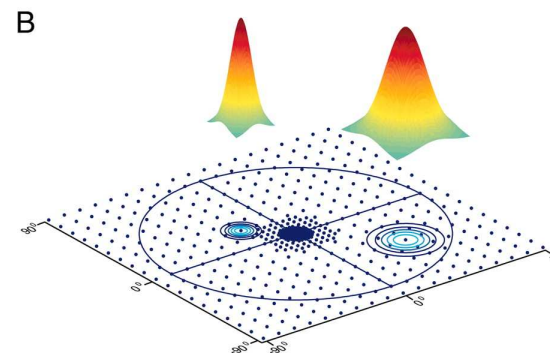


Multiple RFs  
(Martinez-Trujillo  
et al. *Neuron* 2004)

## Neural Net Version (2001-2005)



Smith & Crawford (2005)  
*J. Neurophysiol.*

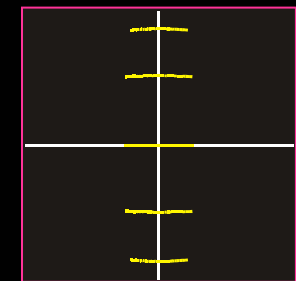
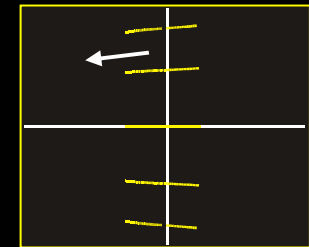
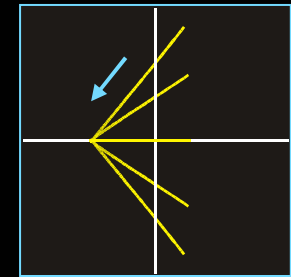
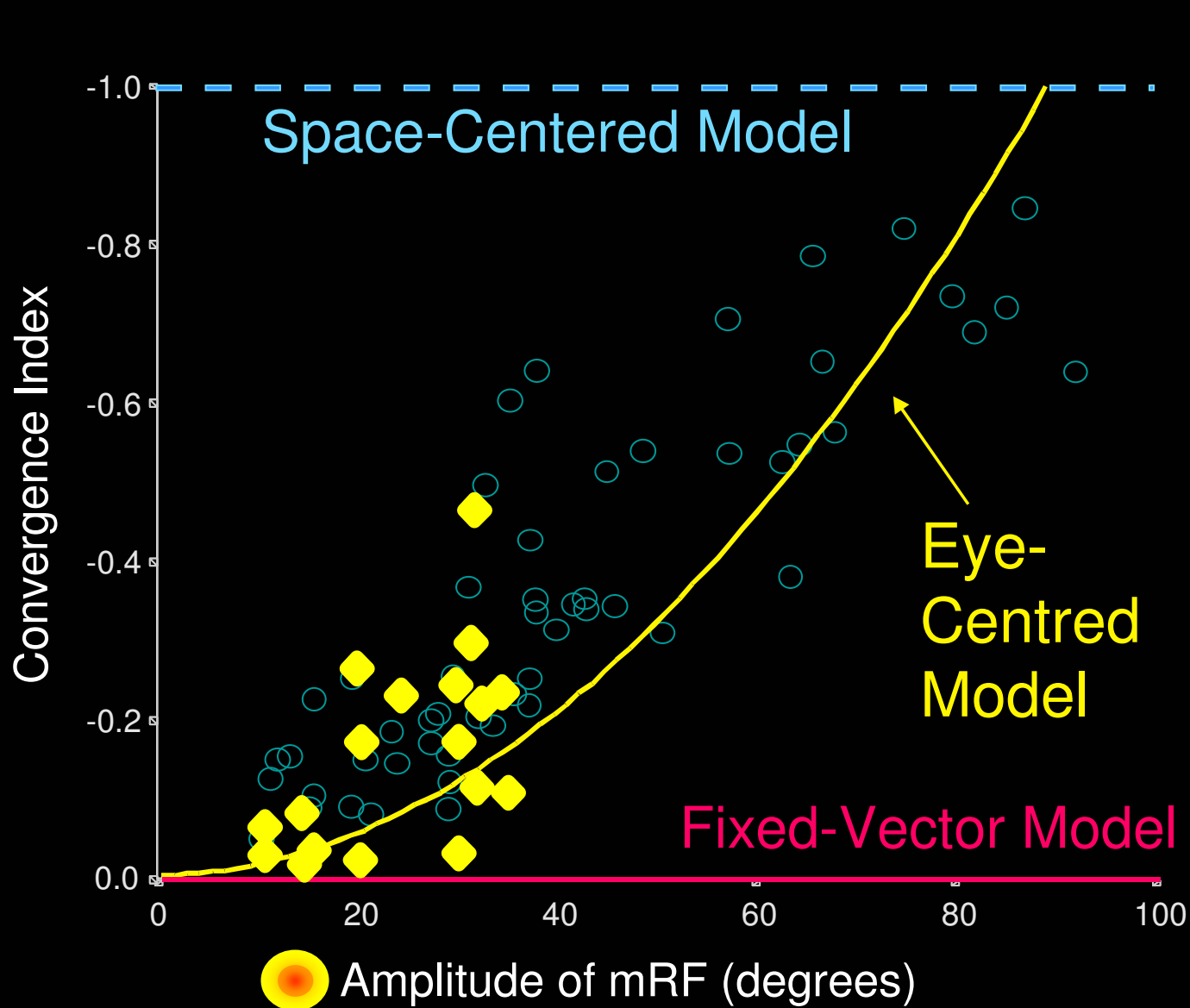


### Predictions

**1) superior colliculus units have eye-centered receptive fields with non-linear eye position modulation**

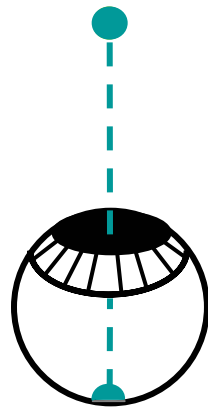
DeSouza, J.F.X., Yan, X., Blohm, G., Wang, H., & Crawford, J.D. (2006). Gaze position effects and position-dependent motor tuning in primate superior colliculus (SC) neurons during head-unrestrained visually guided movements. *Society for Neuroscience Abstracts*.

(DeSouza et al. *CPS*, 2007)

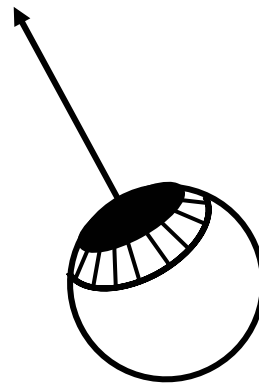


○ Stimulation of SC (Klier et al. *Nature Neuroscience*, 2001)

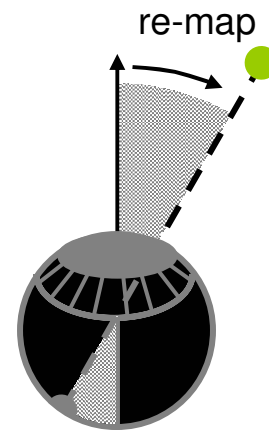
# Spatial Updating & 'Remapping'



before

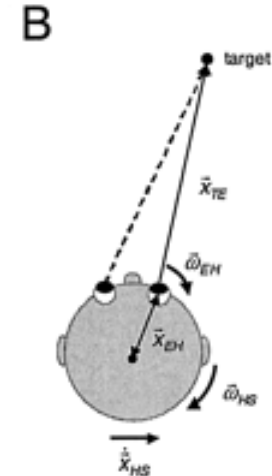


movement



after

**Medendorp et al.**  
(2002, 2003)  
*J. Neurosci.*



## 1) parietal cortex activity 'updated' during eye movements

**Medendorp WP, Goltz HC, Vilis T, Crawford JD. (2003) Gaze-centered updating of visual space in human parietal cortex. *J Neurosci.* 23:6209-14.**



# Pointing Topography



Rightward Pointing



Leftward Pointing



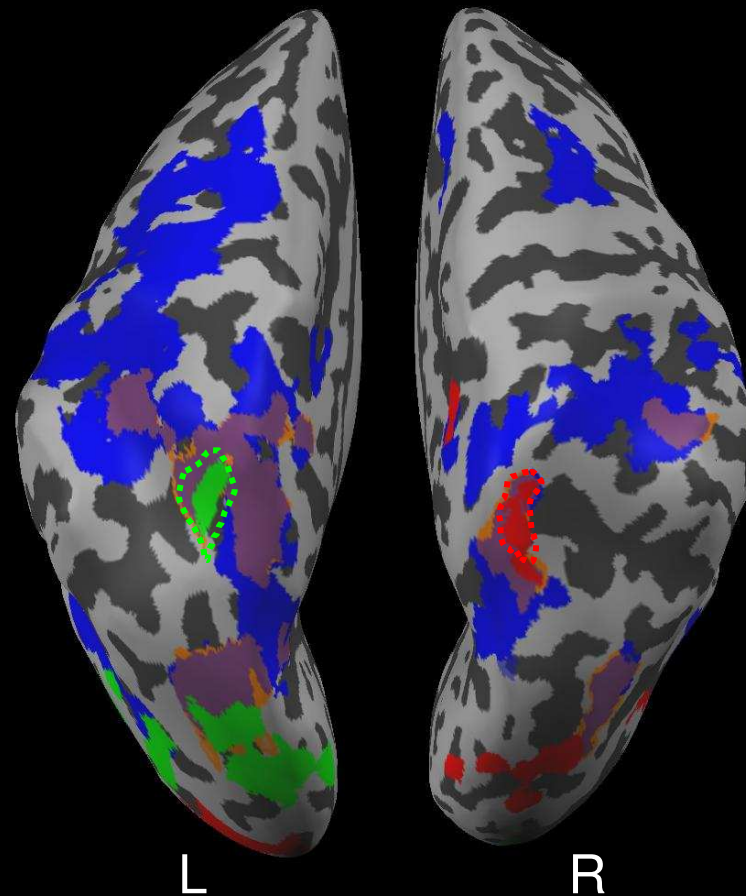
Saccades



Pointing



Both



Medendorp et al.  
*J. Neuroscience*  
2003

Medendorp  
et al.  
*J. Neuroscience*  
2003

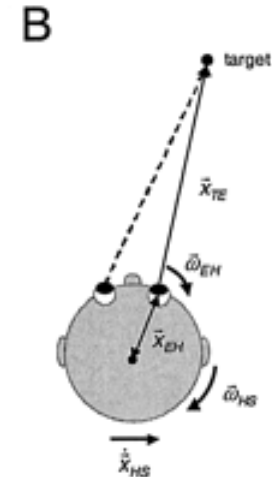
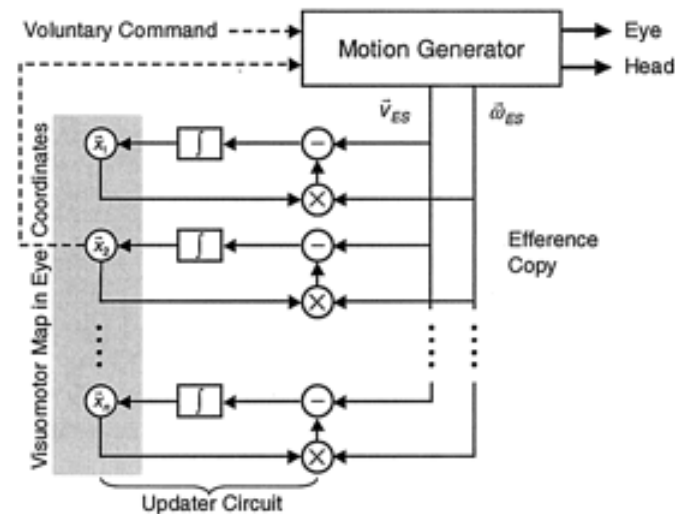
Does this spatially-selective activity remap during saccades?

## Spatial Updating Models (1999-2003)



Smith et al. (2001)  
*J. Neurophysiol.*

Medendorp et al.  
(2002, 2003)  
*J. Neurosci.*



## Predictions

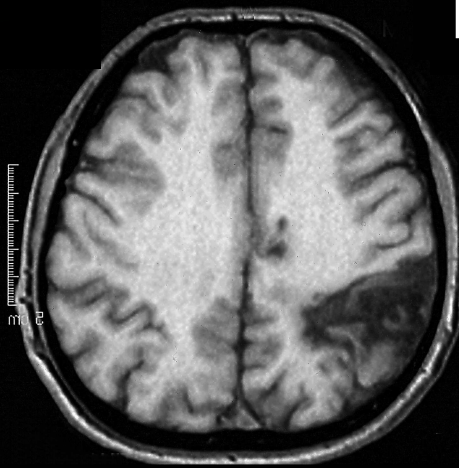
### 1) parietal cortex activity 'updated' during eye movements

Medendorp WP, Goltz HC, Vilis T, Crawford JD. (2003) Gaze-centered updating of visual space in human parietal cortex. *J Neurosci.* 23:6209-14.

### 2) parietal damage leads to eye-centered deficits in reaching

Khan AZ, Pisella L, Vighetto A, Cotton F, Luaute J, Boisson D, Salemme R, Crawford JD, Rossetti Y. (2005) Optic ataxia errors depend on remapped, not viewed, target location. *Nat Neurosci.* 8:418-20.

O.K.

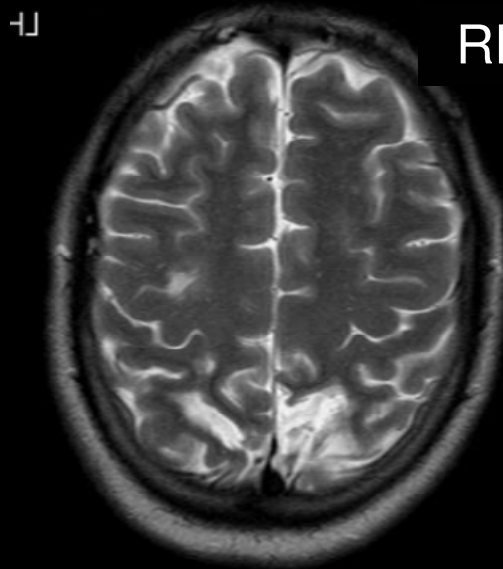


RH

- 39 year-old right-handed male
- Right Parietal Cortex Lesion

A. Khan et al.  
*Nature Neuroscience*  
2005

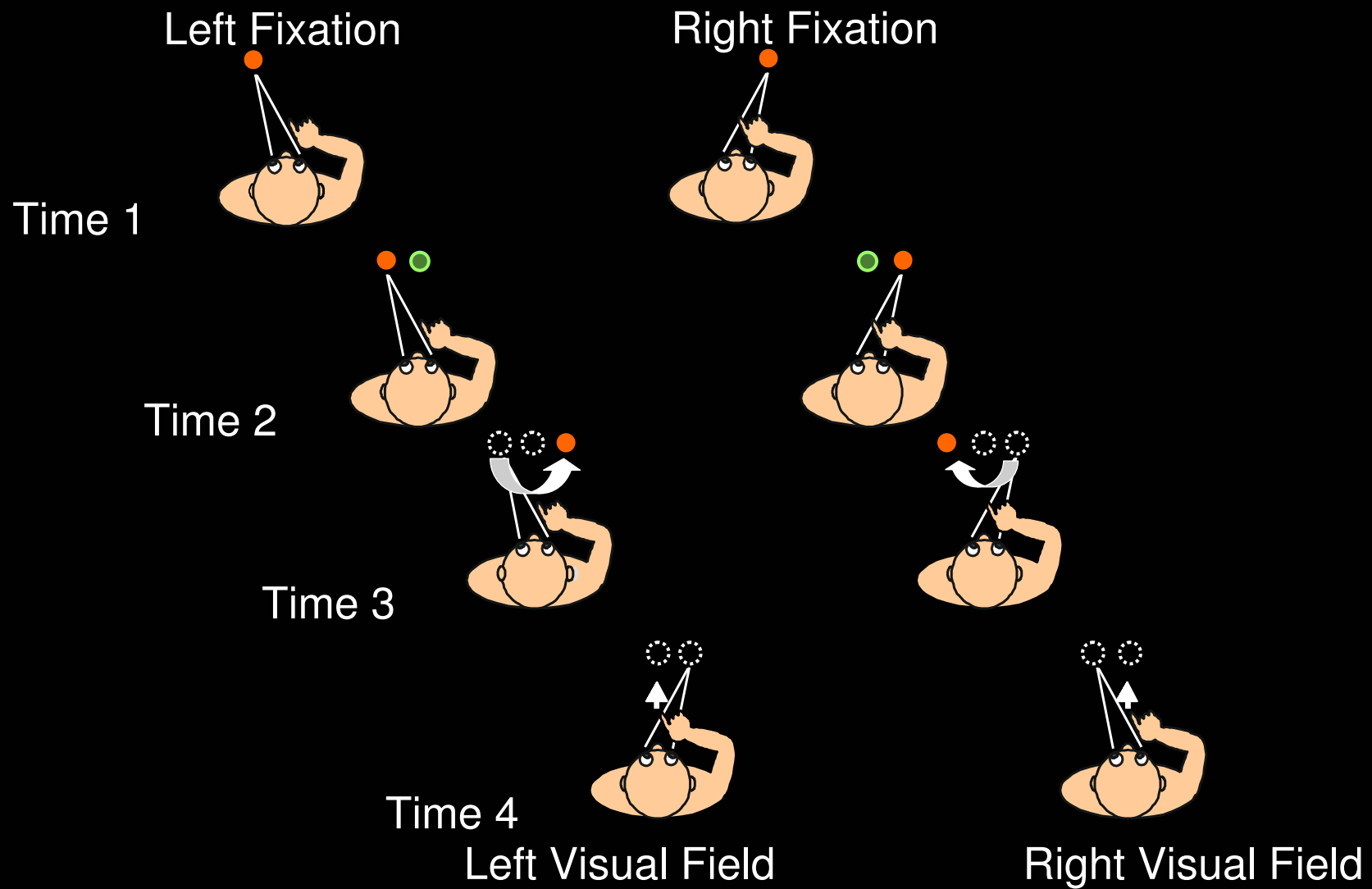
C.F.

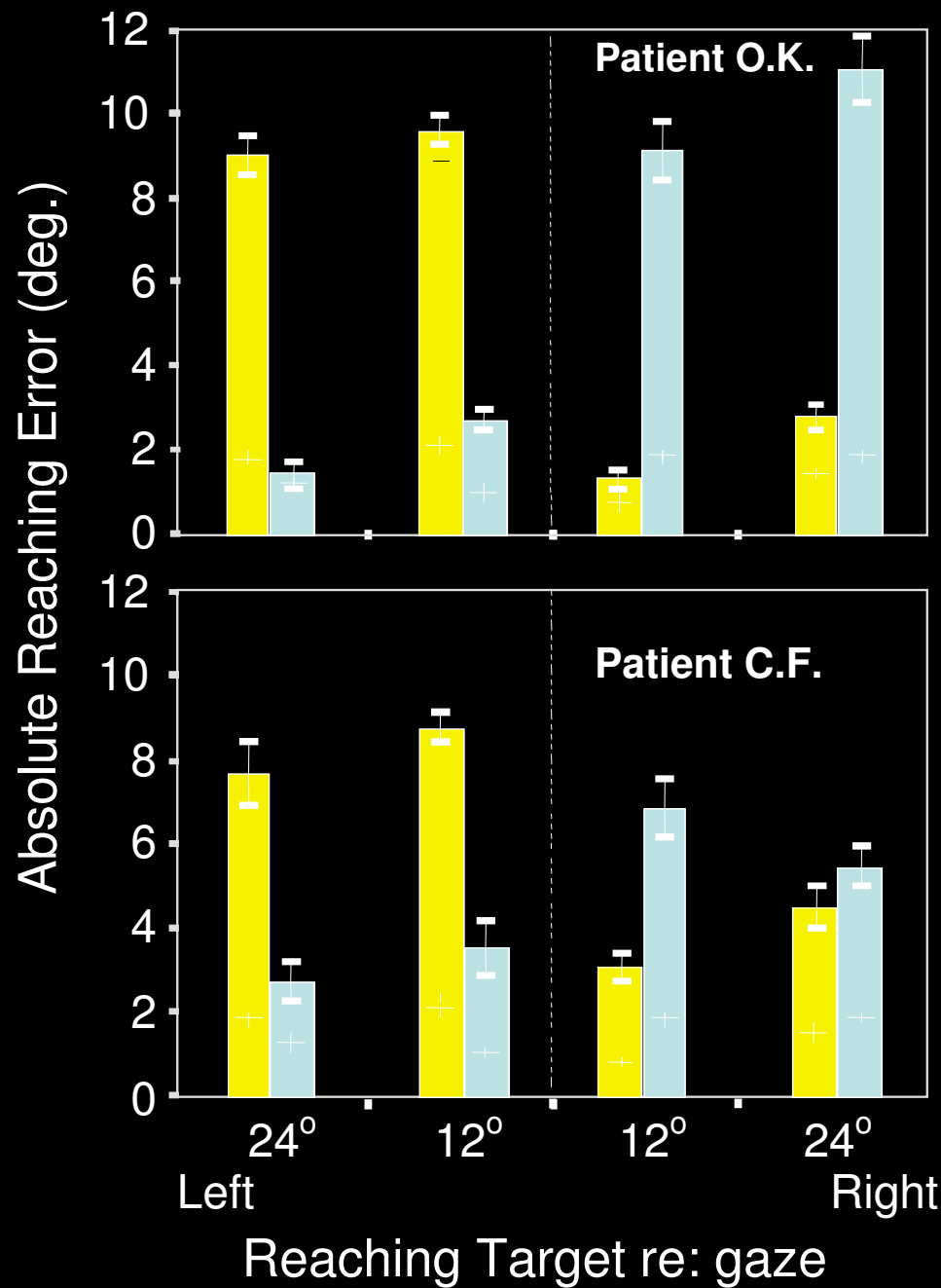


RH

- 27 year-old right-handed male
- Bilateral Damage
- Unilateral (Right Parietal) OA

# Opposite-Field Saccade Task





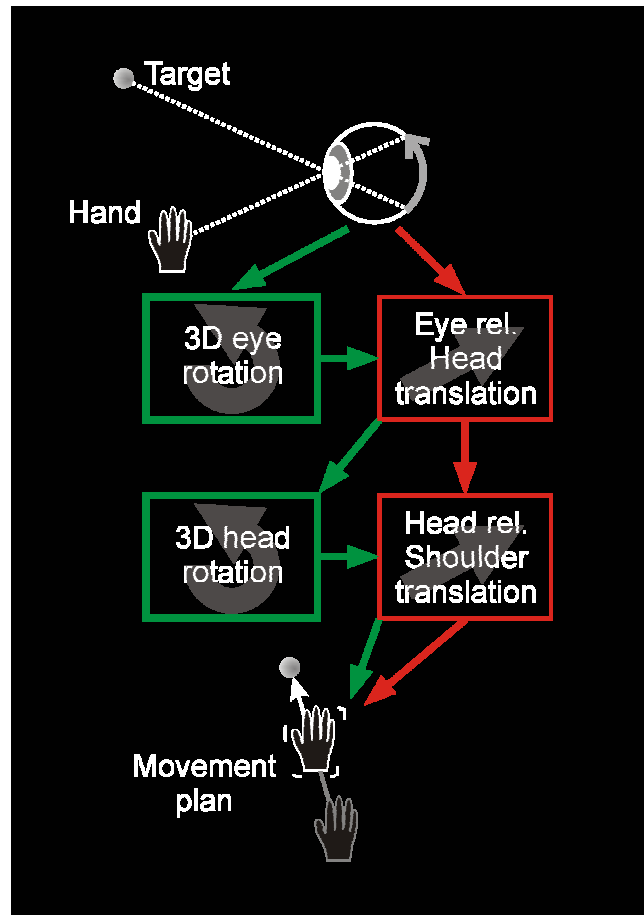
A. Khan et al.  
*Nature Neuroscience*  
2005

Fixation Task

Opposite Field Saccade Task

*(Target in initial gaze coordinates, before Saccade)*

# Visually-Guided Reach (2005-2007)



## Predictions

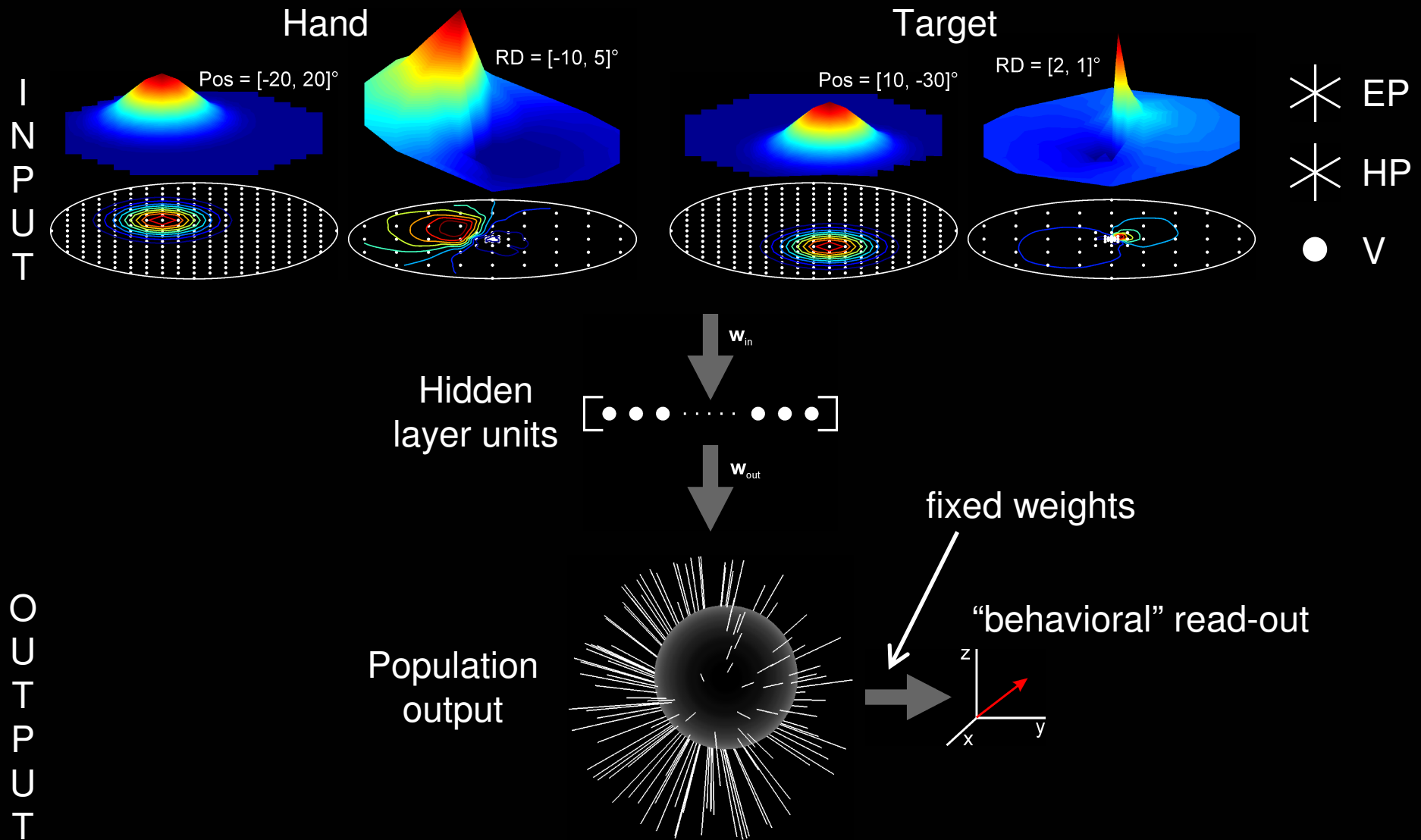
### 1) Optimal Reaching in Normal Subjects

Blohm, G. and Crawford, J.D. (2007) Geometric Computations for Visually- guided reaching in 3-D Space *The Journal of Vision*.

### 2) Specific parametric deficits in stroke patients with frontal / parietal damage

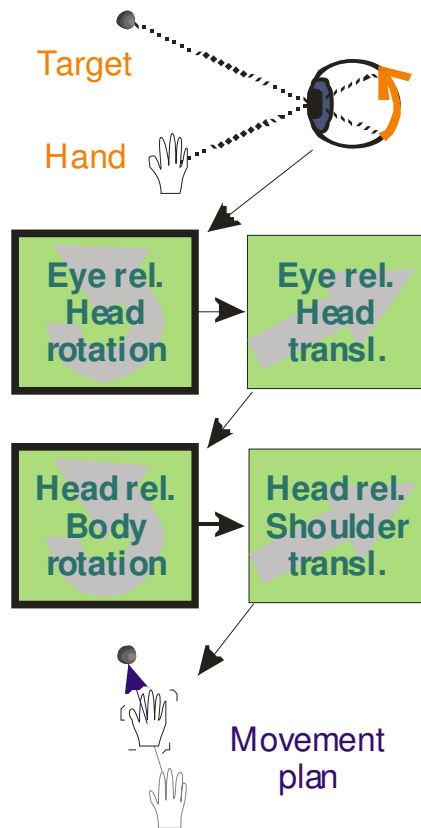
Currently Being Investigated.

## Neural Network Version (2006-2007)

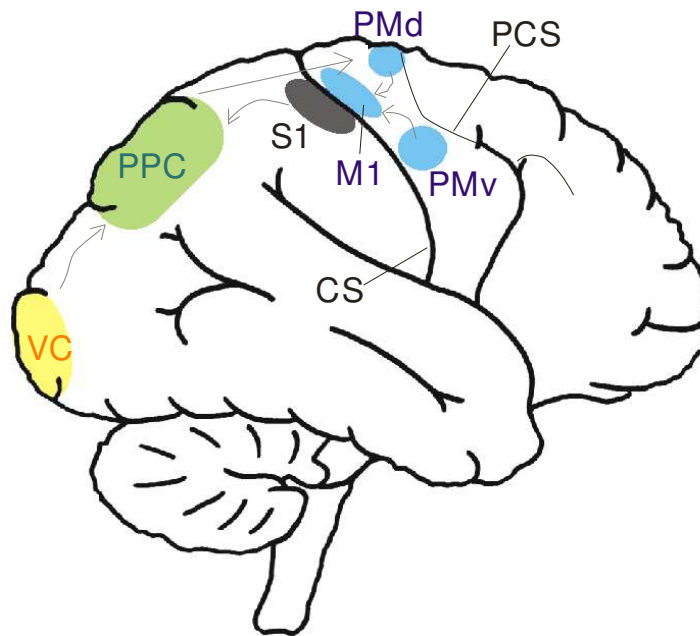


# Moving Between Different Levels of Theory of Brain Function

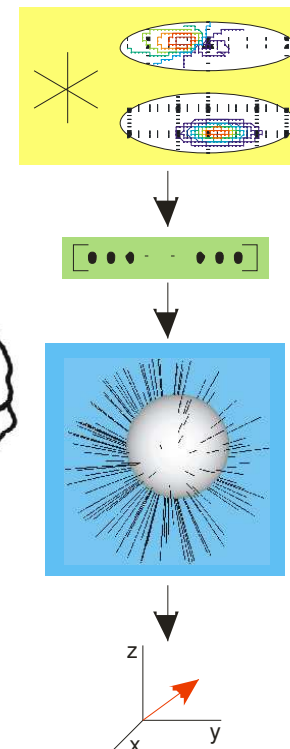
## 1) Algorithm



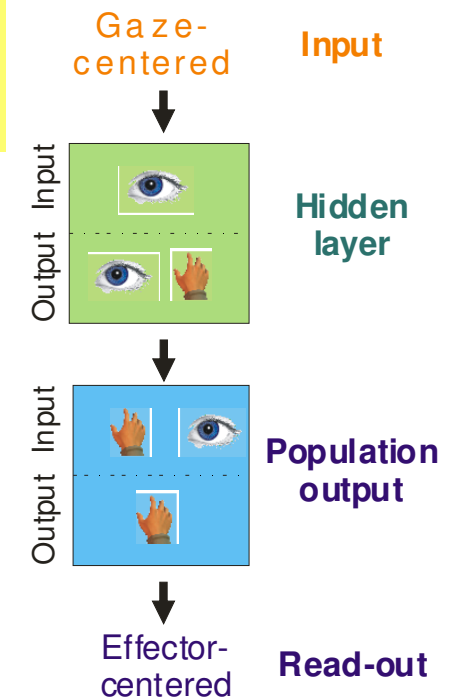
## 2) Known Physiology



## 3) Networks



## 4) Implementation





## **General Workshop Goals:**

**(i) Definition of the field**

## **'Systems Level' Sensorimotor Research:**

- **'Black Box' models of sensorimotor algorithms**
- **'Neural Network' Models**
- **Quantification of data...motion analysis, spikes etc.**

## **General Workshop Goals:**

**(ii) Research highlights in the field**

**(just a few examples):**

**1) Models have provides us with testable framework to understand Mechanisms of**

- 3-D VOR**
- Spatial Frames for Gaze Coding**
- Spatial Updating for saccades & pointing**
- Visually-Guided Reach**

**2) Related mathematical tools for data analysis**

## **General Workshop Goals:**

**(iii) Critical considerations for someone wanting to enter the field today**

**1) What do we already know?**

**2) What do we not know?**

**3) Can I design models that will have an impact on**

- Basic experimental work**
- Clinical applications**

**4) How will I realize those applications?**

## **General Workshop Goals:**

### **(iv) Ideal type(s) of training**

- 1) Training environments that provide more than one approach (theoretical & experimental)**
- 2) Training environments that provide a synergy between theory and experimentation**
  - Within projects**
  - As part of collaborative ventures**

## **General Workshop Goals:**

**(v) Suggested changes and directions for the field.**

- 1) More synergy between theory and experiment!**
- 2) Better communication (and mutual appreciation)  
between theorists, experimentalists, and clinicians**
- 3) Better understanding of the relationships between  
different 'levels' of theory & experiment**  
  
**e.g., a formal theory of how sensorimotor  
algorithms are distributed within networks**



# ACKNOWLEDGEMENTS



Mike  
Smith



Pieter  
Medendorp



Gunnar  
Blohm



Gerry  
Keith



Canada Research  
Chairs

