## Come presentare una proposta di successo in ambito ERC: Panel LS - Life Sciences

# Aldo Genovesio

Dipartimento di Fisiologia e Farmacologia Vittorio Erspamer Sapienza Università di Roma

## ERC 2014: HUMO Project

What is everybody doing? Prediction, categorization, and monitoring in the prefrontal cortex.

## Aldo Genovesio

### Machiavelli intelligent hypothesis

Nicholas Humphrey's "The Social Function of Intellect" (1976)

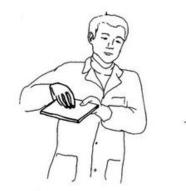
Machiavellian Intelligence (Byrne and Whiten 1988)



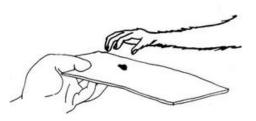


### **Mirror neurons**

### Sharing the representation of action



_	II II AN III III I	
_111		
1		

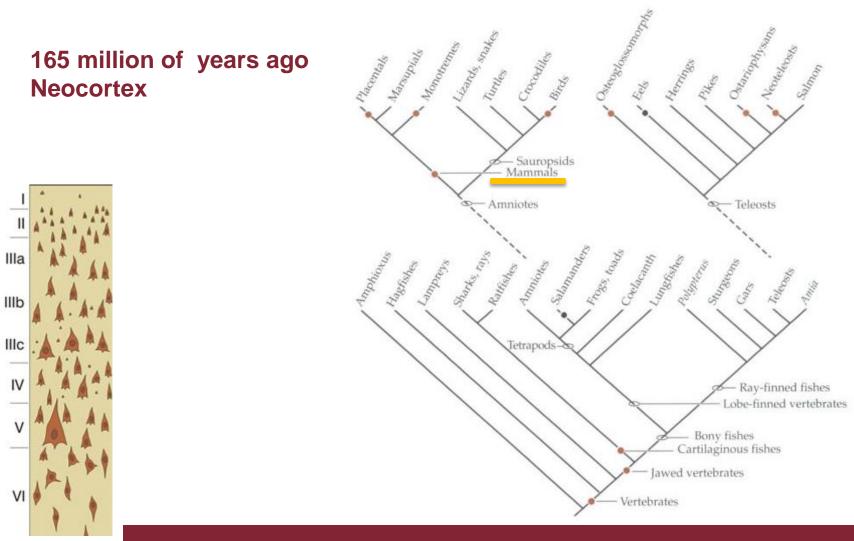


♥ 1001 0000	
UL BUU <b>UL</b> I I	

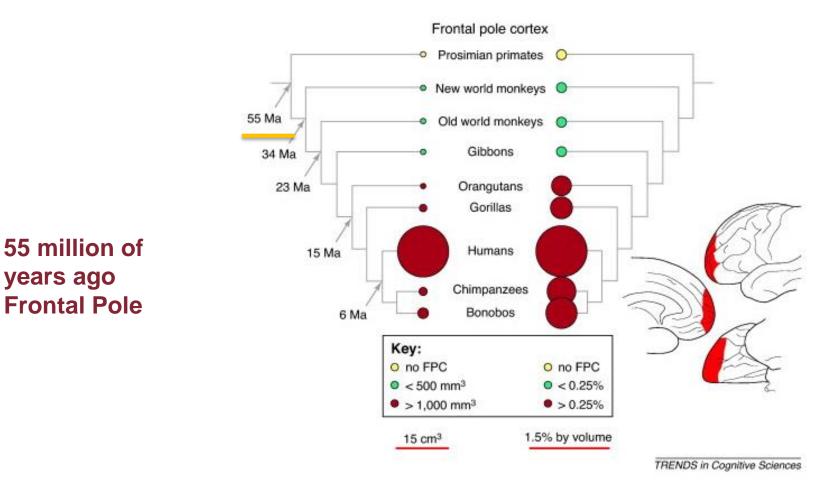
500 msec



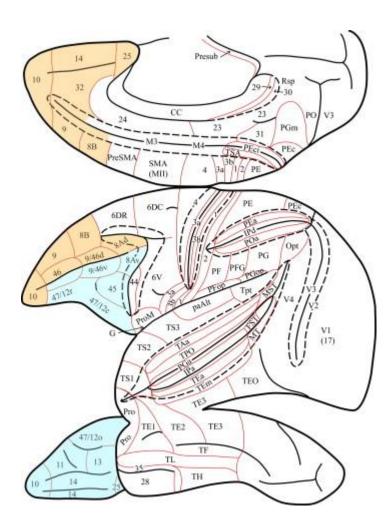
### **Mammals Innovation**



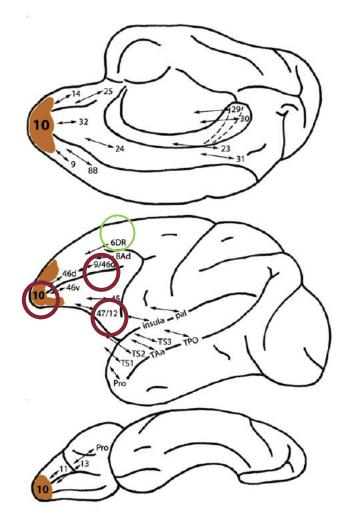
### **Primates Innovation**



### **Frontal Pole Connectivity**



HUMO



Special issue: Review

CORTEX 48 (2012) 58-81

The cortical connectivity of the prefrontal cortex in the monkey brain

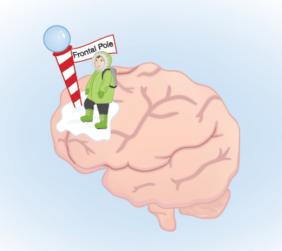
Edward H. Yeterian  ${}^{a,b,c,*},$  Deepak N. Pandy<br/>a ${}^{b,c,d},$  Francesco Tomaiuolo ${}^e$  and Michael Petrides<br/>  ${}^f$ 

### Monitoring of goal achievement in the Frontal Pole cortex

Jonathan D Wallis

The frontal pole cortex is thought to be the most complex of all frontal cortex areas. Overcoming technical obstacles to direct recordings, a study in this issue finds that neurons in this area have unexpectedly simple response properties.

The polar explorer Ernest Shackleton said "Difficulties are just things to overcome, after all." The last decade has seen rapid progress in our understanding of the functional organization of the frontal lobe. Much of our knowledge of this organization has come from studies recording the electrical activity of single neurons in awake, behaving monkeys. One area, however, has resisted exploration. The frontal pole (Brodmann area 10) is the most anterior region of the frontal cortex (Fig. 1). Until now, no one has been able to record neuronal activity from this area because it lies underneath a bony air sinus. In this issue, Tsujimoto et al.1 treated this difficulty as just an obstacle to overcome and their intrepid exploring has produced some surprising results.



#### ARTICLES

nature neuroscience

# Evaluating self-generated decisions in frontal pole cortex of monkeys

Satoshi Tsujimoto<sup>1,2</sup>, Aldo Genovesio<sup>1,3</sup> & Steven P Wise<sup>1</sup>





### H-M paradigm



Rossella Falcone, Emiliano Brunamonti, Stefano Ferraina, Aldo Genovesio\*

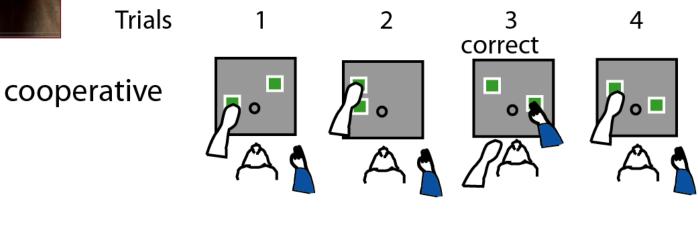
Department of Physiology and Pharmacology, Sapienza University of Rome, Rome, Italy

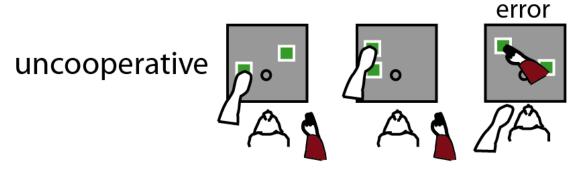


PLos one



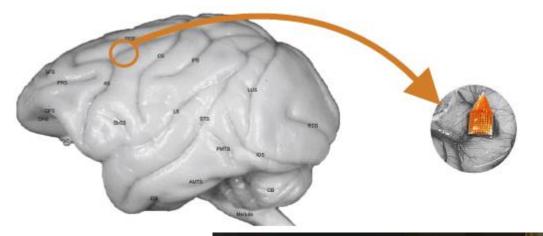
### H-M paradigm







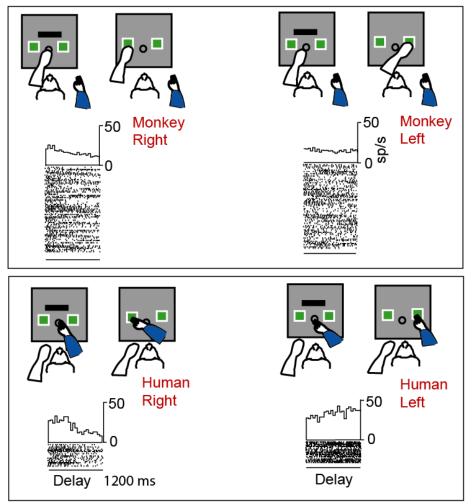
### Multi-array recordings, array in premotor cortex

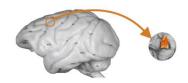






# Preliminary data with H-M paradigm in Premotor cortex

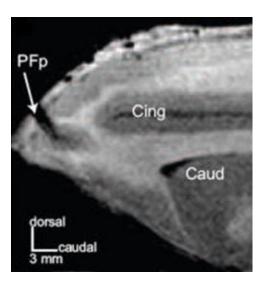


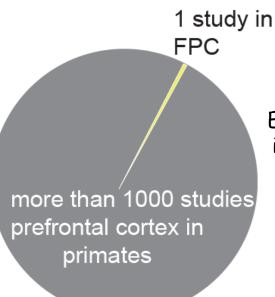






## Only one study





Evaluating self-generated decisions in frontal pole cortex of monkeys.

Why do we study the frontal pole cortex in Monkeys?

Why do we study social behavior in Monkeys?

Why should be my lab to address this question?



### Several years to answer many questions

### **Frontal Pole**

- Does it monitor the choices of other agents?
- Does it represent the confidence with which a goal is accomplished?
- Is it involved in learning from single events (object-in-scene learning task)?
- Does it have a function in observation learning?

### **Dorsolateral Prefrontal cortex**

- Does it predict what the others will do next?
- Does it predict the behaviour of others based on their category (cooperative and
- uncoperative agents)
- Does it mantein in memory the previous choices of others?
- Does it distinguish self from others?
- Does it track the history of who acted before?
- Does it show a neural bias in the condition of free choice?

### Orbitofrontal

• Does it monitor the past action of other agents and the outcome?

## **People in the Department of Physiology**

### Faculty



Alexandra Battaglia-Mayer





Roberto Caminiti



C Paul Verschure Robotic

Collaborators re Paolo Mazzone C Neurosurgery

Giovanni Pezzullo Modelling











# **Additional Slides**

Social group size and Frontal Pole

Frontal Pole and Theory of Mind

Frontal Pole and Autism

Oxytocin

Object-in-place task

Insula, TPO, orbitofrontal

**Observation learning** 

Spatial and Object tasks



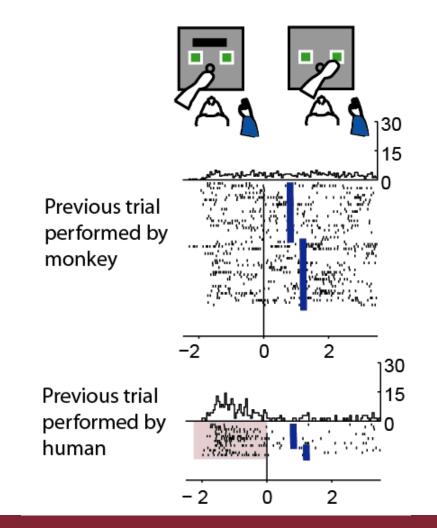
### **Time table**

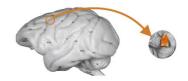
A Monkeys 1,2,3); NMTG task B (Monkeys 3, 4): Object-in-place scene learning task

Months	Month	Month	Month	Month	Total
	1-18	19-36	37-54	55-60	(M1-60)
TRAINING	A-B				
RECORDING	A-B	А			
DATA ANALYISIS	А	A1-B	A-B	А	А
PUBLICATION			А	A-B	A-B



# Preliminary data on H-M paradigm in premotor cortex





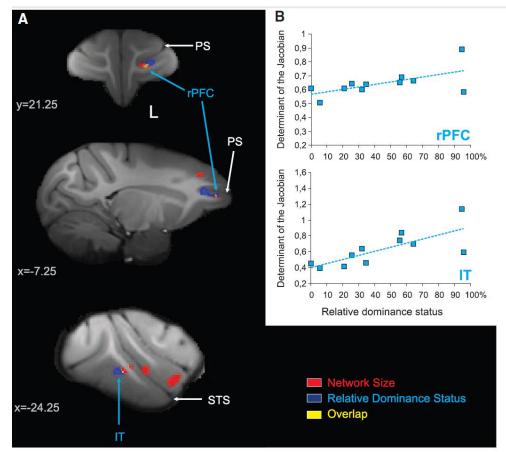


### Social group dimensions affects Frontal pole

# Social Network Size Affects Neural Circuits in Macaques

J. Sallet,<sup>1,2\*</sup>† R. B. Mars,<sup>1,2\*</sup> M. P. Noonan,<sup>1,2\*</sup> J. L. Andersson,<sup>2</sup> J. X. O'Reilly,<sup>2</sup> S. Jbabdi,<sup>2</sup> P. L. Croxson,<sup>1,3</sup> M. Jenkinson,<sup>2</sup> K. L. Miller,<sup>2</sup> M. F. S. Rushworth<sup>1,2</sup>

HUMO



rupt emotional responsiveness in both humans and macaques (13, 14). Bickart and colleagues (2) have reported a similar correlation in human subjects but, unlike here, they were unable to determine whether differences in amygdala size were the consequence of differences in experienced social network size.

Our initial analysis also identified clusters in rPFC, in the rostral principal sulcus, at the border between dorsolateral prefrontal cortex (area 46) and the frontal pole (area 10) (Fig. 1, A and B, and table S3). In human subjects, an rPFC area rostral to the paracingulate sulcus, together with the STS, is active when predictions are made and updated about the intentions of others (15, 16). Such predictions will have to be made about more individuals, and combinations of individuals, as social network size increases. It is possible that the rPFC region that we have identified in the macaques in the present study is similar to the human rPFC area (17, 18). In humans, rPFC, together with STS and TPJ, is associated with "theory of mind" and prediction of another individual's overt behavior as well as their intentions (15, 16). It is not clear if macaques have a theory of mind and are able to represent others' intentions (15, 16), but they can make decisions based on inferences about what others can see (19), which might be a precursor of such an ability. There is also evidence that the activ-

### **Theory of mind and Frontal pole**

## Neuronal correlates of theory of mind and empathy: A functional magnetic resonance imaging study in a nonverbal task

Birgit A. Völlm<sup>a,</sup> ▲· ➡, Alexander N.W. Taylor<sup>a</sup>, Paul Richardson<sup>a</sup>, Rhiannon Corcoran<sup>b</sup>, John Stirling<sup>c</sup>, Shane McKie<sup>a</sup>, John F.W. Deakin<sup>a</sup>, Rebecca Elliott<sup>a</sup>

+ Show more

doi:10.1016/j.neuroimage.2005.07.022

Get rights and content

#### Abstract

Theory of Mind (ToM), the ability to attribute mental states to others, and empathy, the ability to infer emotional experiences, are important processes in social cognition. Brain imaging studies in healthy subjects have described a brain system involving medial prefrontal cortex, superior temporal sulcus and temporal pole in ToM processing. Studies investigating networks associated with empathic responding also suggest involvement of temporal and frontal lobe regions. In this fMRI study, we used a cartoon task derived from Sarfati et al. (1997) [Sarfati, Y., Hardy-Bayle, M.C., Besche, C., Widlocher, D. 1997. Attribution of intentions to others in people with schizophrenia: a non-verbal exploration with comic strips. Schizophrenia Research 25, 199-209.] with both ToM and empathy stimuli in order to allow comparison of brain activations in these two processes. Results of 13 right-handed, healthy, male volunteers were included. Functional images were acquired using a 1.5 T Phillips Gyroscan. Our results confirmed that ToM and empathy stimuli are associated with overlapping but distinct neuronal networks. Common areas of activation included the medial prefrontal cortex, temporoparietal junction and temporal poles. Compared to the empathy condition, ToM stimuli revealed increased activations in lateral orbitofrontal cortex, middle frontal gyrus, cuneus and superior temporal gyrus. Empathy, on the other hand, was associated with enhanced activations of paracingulate, anterior and posterior cingulate and amygdala. We therefore suggest that ToM and empathy both rely on networks associated with making inferences about mental states of others. However, empathic responding requires the additional recruitment of networks involved in emotional processing. These results have implications for our understanding of disorders characterized by impairments of social cognition, such as autism and psychopathy.



# Patients with area 10 lesions have theory of mind deficits

The role of Area 10 (BA10) in human multitasking and in social cognition: A lesion study

María Roca<sup>a, b, c,</sup> **a**, **a**, **b**, c, **a**, Teresa Torralva<sup>a, b</sup>, Ezequiel Gleichgerrcht<sup>a, b</sup>, Alexandra Woolgar<sup>d</sup>, Russell Thompson<sup>d</sup>, John Duncan<sup>d</sup>, Facundo Manes<sup>a, b,</sup> **a**, **a** 

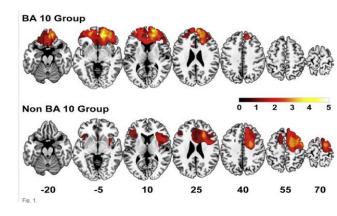
+ Show more

doi:10.1016/j.neuropsychologia.2011.09.003

Get rights and content

#### Abstract

A role for rostral prefrontal cortex (BA10) has been proposed in multitasking, in particular, the selection and maintenance of higher order internal goals while other sub-goals are being performed. BA10 has also been implicated in the ability to infer someone else's feelings and thoughts, often referred to as theory of mind. While most of the data to support these views come from functional neuroimaging studies, lesion studies are scant. In the present study, we compared the performance of a group of frontal patients whose lesions involved BA10, a group of frontal patients whose lesions did not affect this area (nonBA10), and a group of healthy controls on tests requiring multitasking and complex theory of mind judgments. Only the group with lesions involving BA10 showed deficits on multitasking and theory of mind tasks when compared with control subjects. NonBA10 patients performed more poorly than controls on an executive function screening tool, particularly on measures of response inhibition and abstract reasoning, suggesting that theory of mind and multitasking deficits following lesions to BA10 cannot be explained by a general worsening of executive function. In addition, we searched for correlations between performance and volume of damage within different subregions of BA10. Significant correlations were found between multitasking performance and volume of damage in right lateral BA10, and between theory of mind and total BA10 lesion volume. These findings stress the potential pivotal role of BA10 in higher order cognitive functions.



### Mentalizing

### Functional Specialization within Rostral Prefrontal Cortex (Area 10): A Meta-analysis

Sam J. Gilbert<sup>1</sup>, Stephanie Spengler<sup>1</sup>, Jon S. Simons<sup>1</sup>, J. Douglas Steele<sup>2</sup>, Stephen M. Lawrie<sup>3</sup>, Christopher D. Frith<sup>1</sup>, and Paul W. Burgess<sup>1</sup>

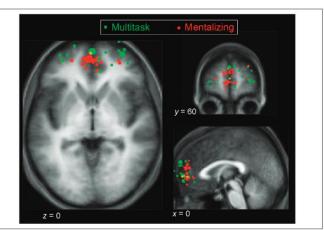
#### Abstract

■ One of the least well understood regions of the human brain is rostral prefrontal cortex, approximating Brodmann's area 10. Here, we investigate the possibility that there are functional subdivisions within this region by conducting a meta-analysis of 104 functional neuroimaging studies (using positron emission tomography/functional magnetic resonance imaging). Studies involving working memory and episodic memory retrieval were disproportionately associated with lateral activations, whereas studies involving mentalizing (i.e., attending to one's own emotions and mental states or those of other agents) were disproportionately associated with me-

<u>HUMO</u>

dial activations. Functional variation was also observed along a rostral–caudal axis, with studies involving mentalizing yielding relatively caudal activations and studies involving multiple-task coordination yielding relatively rostral activations. A classification algorithm was trained to predict the task, given the coordinates of each activation peak. Performance was well above chance levels (74% for the three most common tasks; 45% across all eight tasks investigated) and generalized to data not included in the training set. These results point to considerable functional segregation within rostral prefrontal cortex. ■

Figure 4. Location of activations in the Mentalizing and Multitask categorics, plotted separately on axial (projected onto z = 0), coronal (projected onto y =60), and sagittal (projected onto x = 0) slices of a structural scan (mean of 14 normalized T1-weighted images). Where more than one activation occupies the same location, the area of each blob is increased proportionately.



### **Autism and Frontal Pole**

### A voxel-based investigation of brain structure in male adolescents with autistic spectrum disorder

Gordon D Waiter ▲ · ª· ➡, Justin H.G Williams<sup>b</sup>, Alison D Murray<sup>a</sup>, Anne Gilchrist<sup>c</sup>, David I Perrett<sup>d</sup>, Andrew Whiten<sup>d</sup>

+ Show more

doi:10.1016/j.neuroimage.2004.02.029

Get rights and content

#### Abstract

Autistic spectrum disorder (ASD) has been associated with abnormal neuroanatomy in many imaging and neuropathological studies. Both global brain volume differences and differences in the size of specific neural structures have been reported. Here, we report a voxel-based morphometric whole brain analysis, using a group specific template, on 16 individuals of normal intelligence with autistic spectrum disorder (ASD), and a group of 16 age-, sex- and IQ-matched controls. Total grey matter volume was increased in the ASD group relative to the control group, with local volume increases in the right fusiform gyrus, the right temporo-occipital region and the left frontal pole extending to the medial frontal cortex. A local decrease in grey matter volume was found in the right thalamus. A decrease in global white matter volume in the ASD group did not reach significance. We found the increase in grey matter volume in ASD subjects was greatest in those areas recognised for their role in social cognition, particularly face recognition (right fusiform gyrus), mental state attribution: 'theory of mind' (anterior cingulate and superior temporal sulcus) and perception of eye gaze (superior temporal gyrus). The picture as a whole may reflect an abnormally functioning social cognitive neural network. We suggest that increased grey matter volume may play a pivotal role in the aetiology of the autistic syndrome.



### Area 10 minicolums in autism

### The Neuropathology of Autism

Manuel F. Casanova, MD

Department of Psychiatry and Behavioral Sciences, University of Louisville, Louisville, Ky.

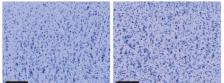
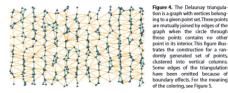


Figure 3. The Boolean germ spain model supposes that a spatial pattern, in this case the Neiss-tained main in a segmented micrograph, is the union of random closed sets (grain advams from some distribution and positioned at random points in space (the germ process). This model is a simplification in that if does not account for clustering of the grains equite minicipal columnar structure. Linder these assumptions, the mean grain area ( $\beta$ ) and perimeter ( $\beta$ ), along with the intensity (2) of the germ process, completely determine the spatial pattern, providing for a node-based estimate of neuronal cross-section and density. These quantities are found to be reduced and increased, respectively, in autism. Left-cortical area 9, right hemisphere, liminal III from a 2-Syearch dim ans Wittou autism. A = 123 µm<sup>2</sup>,  $\lambda$  = 0.0059 µm<sup>-2</sup>. Scale bars measure 200 µm.



ing a posteroanterior trend across neocortex toward decreased mini-column width. Significant diagnosis by area interactions were found in frontopolar (area 10) and the anterior cingulate gyrus (area 24) of paralimbic cortex (21).

What are the functional implications of increased numbers of narrower minicolumns containing smaller projection neurons? These modular microcircuit assemblies are interconnected by thousands of collateral projections within larger networks. Each mini-column is linked to local networks through myelinated bundles in superficial, or radiate, white matter, and to more distant cortical areas via deeper white matter tracts. The neocortex is presumed to have expanded in evolution through addition of mini-columns by proliferation of their clonal progenitors across an embryonic germinal layer (101, 102). Additive increase in mini-column numbers would entail a geometric increase in short- and many loci. Combinations of such polymorphisms in turn could support a bias toward a developmental extreme of this evolution ary trend. In primates, more complex prefrontal connections require an extended period of maturation. Developmental insults or epigenetic interactions during this period would unmask underlying genetic vulnerabilities. Autistic syndromes may represent various manifestations of this phenomenon. A putative increase in local interconnectivity and reduced prefrontal transcortical connections in areas subserving cognitive flexibility and prioritizing (area 10) and emotional and social cognition (area 25) is consistent with the clinical picture of stereotypy, rigidity and interpersonal deficits characterizing autism.

#### NEUROINFLAMMATION

Classical descriptions of inflammatory processes involve a vascular component leading to the accumulation of cells and ing to a given point set. Three points fluid within the extravascular space. In this are mutually joined by edges of the regard autism lacks a vascular component graph when the circle through those points contains no other and a classical inflammatory response. point in its interior. This figure illus-Levels of quinolinic acid and neopterin in trates the construction for a rancerebrospinal fluid are either normal or domly generated set of points, paradoxically reduced in comparison to clustered into vertical columns. controls (139). Samples of cerebrospinal have been omitted because of fluid obtained from live patients show boundary effects. For the meaning normal cell counts and protein electrophoresis. There is no evidence of recruitment of leukocytes into the central nervous long-distance projection fibers in order to system. Reported cellular changes do not maintain a constant degree of transcortical support their participation in tissue repair connectivity among modules (61). Longer and recovery. Furthermore, the clinical white matter fibers occupy more space, symptoms in autism denote lack of conrequire disproportionately larger soma to comitant fever, myalgia, arthralgia, anorsupport increased metabolic costs and result exia or somnolescence that would sustain a in signal processing delays. Selection pressystemic effect for any given inflammatory sure would therefore be expected to have agent. Rather, the tissue response and their given rise to modules internally linked by effects appear confined to the brain and indicate an intact blood-brain barrier.

radially oriented processes and integrated into local networks by short collaterals. Pro-There is no evidence of an adaptive of portionately less white matter would be specific immune response in the brains of autistic patients (89, 125). Immunocydevoted to longer-range connections, encouraging regional functional specializatochemical studies have failed to identify tion. Neuropathological descriptions of T- and B-lymphocyte infiltration and depodecreased cell size and narrow minisition of immunoglobulin/complement in columns (20), studies revealing increased brain tissue (125). Perivascular CD3 and superficial white matter (58), and func-CD20 lymphocyte cell subsets have been tional imaging studies revealing decreases in rare and reported in both patients and conactivity linking prefrontal and posterior trols (125). The immune system appears areas (64) support this view. This reorganiactivated exclusively in terms of its innate zation might require an increase in frecomponents, astrocytes and microglial cells quency of multiple rare polymorphisms at and the influx of macrophages (7, 52, 125).

ing a posteroanterior trend across neocortex toward decreased mini-column width. Significant diagnosis by area interactions were found in frontopolar (area 10) and the anterior cingulate gyrus (area 24) of paralimbic cortex (21).

phenomenon. A putative increase in local interconnectivity and reduced prefrontal transcortical connections in areas subserving cognitive flexibility and prioritizing (area 10) and emotional and social cognition (area 25) is consistent with the clinical picture of stereotypy, rigidity and interpersonal deficits characterizing autism.



### Autism and frontal pole

frontiers in HUMAN NEUROSCIENCE REVIEW ARTICLE published: 21 June 2012 doi: 10.3389/fnhum.2012.00184

### A framework for interpreting functional networks in schizophrenia

HUMQ

#### Peter C. Williamson<sup>1</sup>\* and John M. Allman<sup>2</sup>

<sup>1</sup> Tanna Schulich Chair in Neuroscience and Mental Health, University of Western Ontario, London, ON, Canada
<sup>2</sup> Frank P. Hixon Professor of Neurobiology, Division of Biology, California Institute of Technology, Pasadena, CA, USA

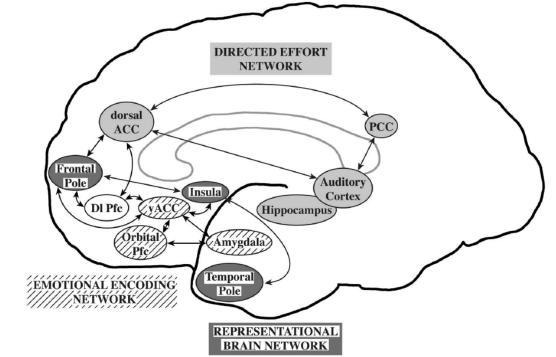


FIGURE 1 | The representational brain. The representational brain network (dark gray), proposed to underlie autism, includes the frontal pole, insula, and temporal pole. The directed effort network (light gray), proposed to underlie schizophrenia, includes the dorsal anterior cingulate cortex (ACC), posterior cingulate cortex (PCC), auditory cortex, and hippocampus. The emotional encoding network (lined), proposed to underlie mood disorders, includes the

orbital prefrontal cortex (Pfc), ventral anterior ACC, and amygdala. The directed effort and emotional encoding networks interact with the representational network and the dorsolateral prefrontal cortex (DI Pfc, not shaded). Reprinted with permission from Williamson and Allman. *The Human Illnesses: Neuropsychiatric Disorders and the Nature of the Human Brain.* New York, NY: Oxford University Press, Copyright 2011.

### Autism and abnormal development of area 10

doi:10.1093/brain/awu083

Brain 2014: 137; 1799–1812 | 1799

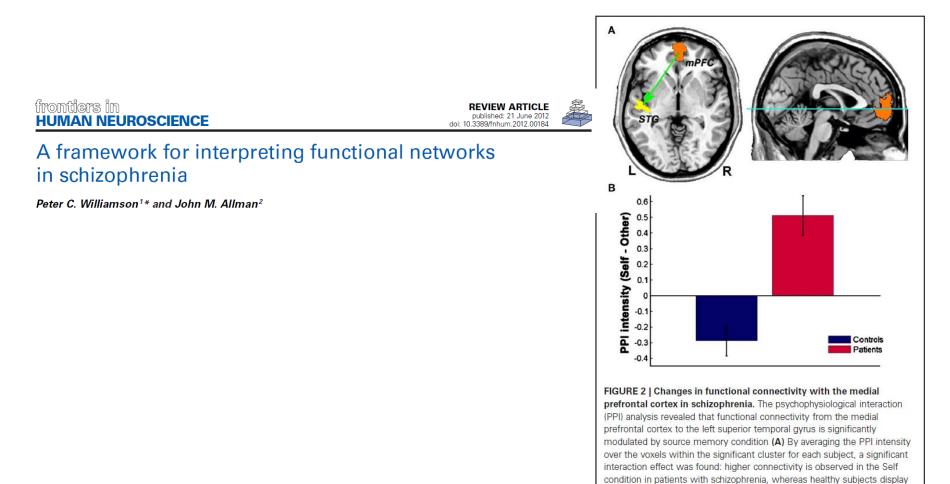


# Longitudinal changes in cortical thickness in autism and typical development

Brandon A. Zielinski,<sup>1,2,3</sup> Molly B. D. Prigge,<sup>1,4</sup> Jared A. Nielsen,<sup>4</sup> Alyson L. Froehlich,<sup>5</sup> Tracy J. Abildskov,<sup>6</sup> Jeffrey S. Anderson,<sup>4,7</sup> P. Thomas Fletcher,<sup>8,9</sup> Kristen M. Zygmunt,<sup>8</sup> Brittany G. Travers,<sup>10</sup> Nicholas Lange,<sup>11,12,13</sup> Andrew L. Alexander,<sup>10,14,15</sup> Erin D. Bigler<sup>6,16</sup> and Janet E. Lainhart<sup>10,15</sup>

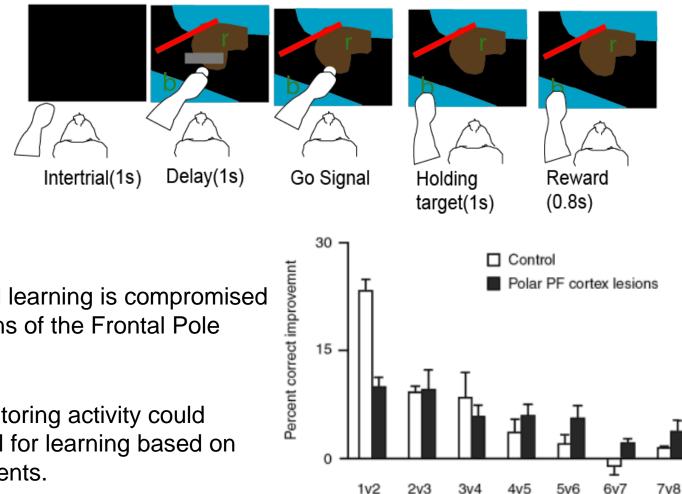
The natural history of brain growth in autism spectrum disorders remains unclear. Cross-sectional studies have identified regional abnormalities in brain volume and cortical thickness in autism, although substantial discrepancies have been reported. Preliminary longitudinal studies using two time points and small samples have identified specific regional differences in cortical thickness in the disorder. To clarify age-related trajectories of cortical development, we examined longitudinal changes in cortical thickness within a large mixed cross-sectional and longitudinal sample of autistic subjects and age- and gender-matched typically developing controls. Three hundred and forty-five magnetic resonance imaging scans were examined from 97 males with autism (mean age = 16.8 years; range 3–36 years) and 60 males with typical development (mean age = 18 years; range 4–39 years), with an average interscan interval of 2.6 years. FreeSurfer image analysis software was used to parcellate the cortex into 34 regions of interest per hemisphere and to calculate mean cortical thickness for each region. Longitudinal linear mixed effects models were used to further characterize these findings and identify regions with between-group differences in longitudinal age-related trajectories. Using mean age at time of first scan as a reference (15 years), differences were observed in bilateral inferior frontal gyrus, pars opercularis and pars triangularis, right caudal middle frontal and left rostral middle frontal regions, and left frontal pole. However, group differences in cortical thickness varied by developmental stage, and were influenced by IQ. Differences in age-related trajectories emerged in bilateral parietal and occipital regions (postcentral gyrus, cuneus, lingual gyrus, pericalcarine cortex), left frontal regions (pars opercularis, rostral middle frontal and frontal pole), left

### Change in functional connectivity in Schizofrenia



higher connectivity in the Other condition (B) Reprinted from Wang et al. (2011). Aberrant connectivity during self-other source monitoring in schizophrenia. *Schizophr. Res.* 125, 136–142 with permission from Elsevier.

### **Object-in-place scene learning task**



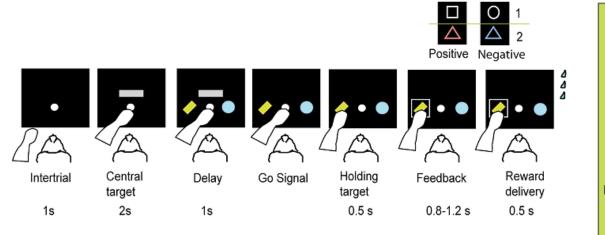
Block v block

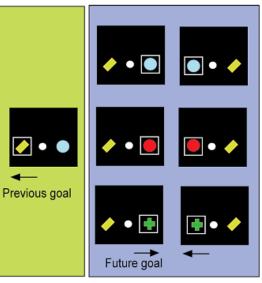
One trial learning is compromised by lesions of the Frontal Pole

The monitoring activity could be usefull for learning based on single events.



### **Object NMTG**



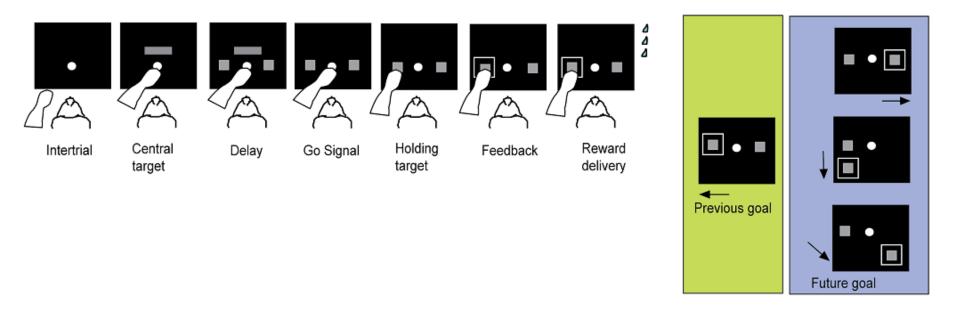


Feedbacks



-

### **Spatial NMTG**





### **Insula shows social functions**



Volume 21, Issue 3, 8 February 2011, Pages 195-199

Report

### Emotional and Social Behaviors Elicited by Electrical Stimulation of the Insula in the Macaque Monkey

Fausto Caruana<sup>1, 2, 3,</sup> 📥 🖾, Ahmad Jezzini<sup>1, 3</sup>, Beatrice Sbriscia-Fioretti<sup>1</sup>, Giacomo Rizzolatti<sup>1, 2,</sup> 📥 🖾, Vittorio Gallese<sup>1, 2</sup>

+ Show more

doi:10.1016/j.cub.2010.12.042

Under an Elsevier user license

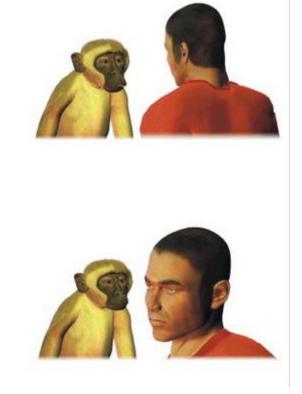




Open Archive

#### Summary

Evidence from a large number of brain imaging studies has shown that, in humans, the insula, and especially its anterior part, is involved in emotions and emotion recognition [1, 2, 3, 4, 5, 6, 7 and 8]. Typically, however, these studies revealed that, besides the insula, a variety of other cortical and subcortical areas are also active. Brain imaging studies are correlative in nature, and, as such, they cannot give indications about the necessary contribution of the different centers involved in emotions. In the present study, we aimed to define more clearly the role of the insula in emotional and social behavior of the monkey by stimulating it electrically. Using this technique, one may determine whether direct activation of the insula can produce specific emotional or social behaviors and exactly which parts of this structure are responsible for these behaviors. The results showed that two emotional behaviors, a basic one (disgust) and a social one (affiliative state), were easily elicited by electrical stimulation of specific parts of the insula. Both behaviors were characterized by specific motor and vegetative responses and by a dramatic change in the monkey's responsiveness to external stimuli.



Stimulation evokats lips smaking behavior only under condition of human eye contact.



### **TPO is sensitive to gaze direction**

Proc R Soc Lond B Biol Sci. 1985 Jan 22;223(1232):293-317.

### Visual cells in the temporal cortex sensitive to face view and gaze direction.

Perrett DI, Smith PA, Potter DD, Mistlin AJ, Head AS, Milner AD, Jeeves MA.

#### Abstract

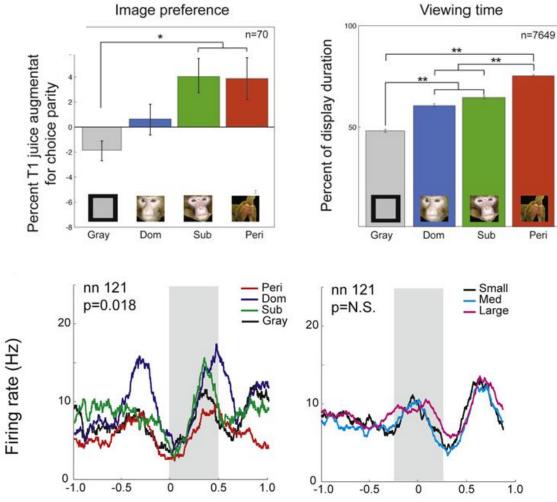
The direction of eve gaze and orientation of the face towards or away from another are important social signals for man and for macaque monkey. We have studied the effects of these signals in a region of the macaque temporal cortex where cells have been found to be responsive to the sight of faces. Of cells selectively responsive to the sight of the face or head but not to other objects (182 cells) 63% were sensitive to the orientation of the head. Different views of the head (full face, profile, back or top of the head, face rotated by 45 degrees up to the ceiling or down to the floor) maximally activated different classes of cell. All classes of cell, however, remained active as the preferred view was rotated isomorphically or was changed in size or distance. Isomorphic rotation by 90-180 degrees increased cell response latencies by 10-60 ms. Sensitivity to gaze direction was found for 64% of the cells tested that were tuned to head orientation. Eighteen cells most responsive to the full face preferred eye contact, while 18 cells tuned to the profile face preferred averted gaze. Sensitivity to gaze was thus compatible with, but could be independent of, sensitivity to head orientation. Results suggest that the recognition of one type of object may proceed via the independent high level analysis of several restricted views of the object (viewer-centred descriptions).



### **Orbitofrontal cortex**

### Social Signals in Primate Orbitofrontal Cortex

Karli K. Watson<sup>1,2,\*</sup> and Michael L. Platt<sup>1,2</sup> Current Biology 2012



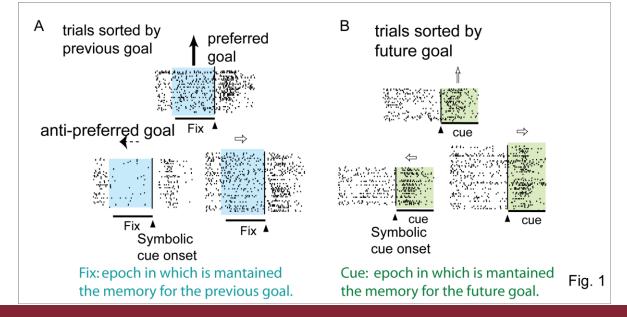
### **Dorsolateral prefrontal cortex**

Can neurons that encode previous and future goals be agent-specific?

Behavioral/Systems/Cognitive

### Representation of Future and Previous Spatial Goals by Separate Neural Populations in Prefrontal Cortex

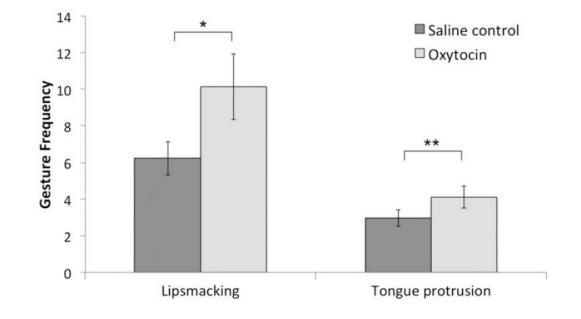
Aldo Genovesio, Peter J. Brasted, and Steven P. Wise



### Oxytocin

# Inhaled oxytocin increases positive social behaviors in newborn macaques

Elizabeth A. Simpson<sup>a,b,1</sup>, Valentina Sclafani<sup>b</sup>, Annika Paukner<sup>a</sup>, Amanda F. Hamel<sup>c</sup>, Melinda A. Novak<sup>c</sup>, Jerrold S. Meyer<sup>c</sup>, Stephen J. Suomi<sup>a</sup>, and Pier Francesco Ferrari<sup>b</sup>



### **Observation learning is possible in H-M**

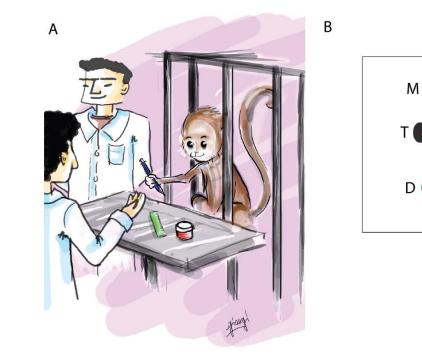
OPEN O ACCESS Freely available online

PLOS ONE

HN

### Macaque Monkeys Can Learn Token Values from Human Models through Vicarious Reward

Sara Bevacqua, Erika Cerasti, Rossella Falcone, Milena Cervelloni, Emiliano Brunamonti, Stefano Ferraina, Aldo Genovesio\*

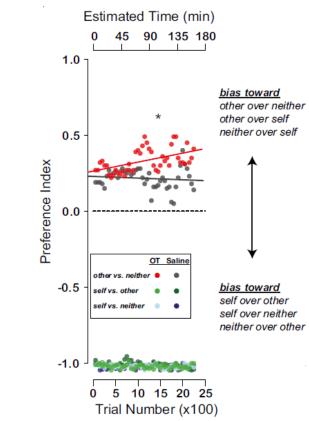


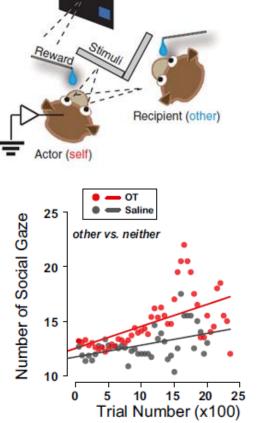


### **Oxtocin increases prosocial behaviour**

# Neuronal reference frames for social decisions in primate frontal cortex

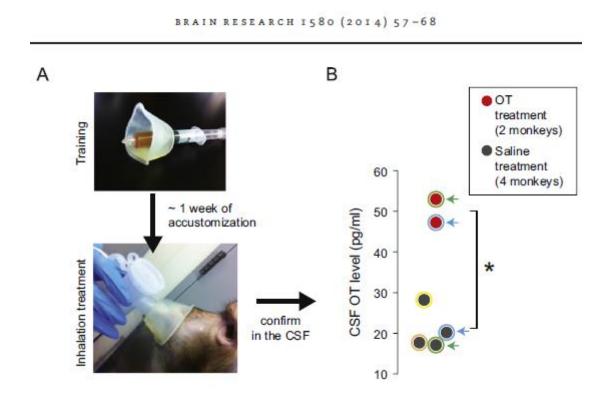
Steve W C Chang<sup>1,2</sup>, Jean-François Gariépy<sup>2</sup> & Michael L Platt<sup>1-3</sup>





## **Oxytocin inhalation**

Inhalation of oxytocin increeases its concentration in the cerebrospinal fluid





### Monkeys represents others' knowledge

DOI: 10.1111/j.1467-7687.2011.01085.x

### **Developmental Science**

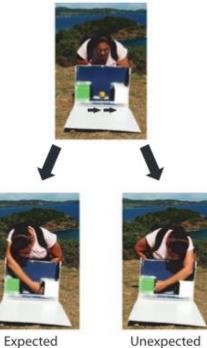
Developmental Science 14:6 (2011), pp 1406-1416

### PAPER

Monkeys represent others' knowledge but not their beliefs

Drew C.W. Marticorena, April M. Ruiz, Cora Mukerji, Anna Goddu and Laurie R. Santos

**Different Side** 





**HUMO** 









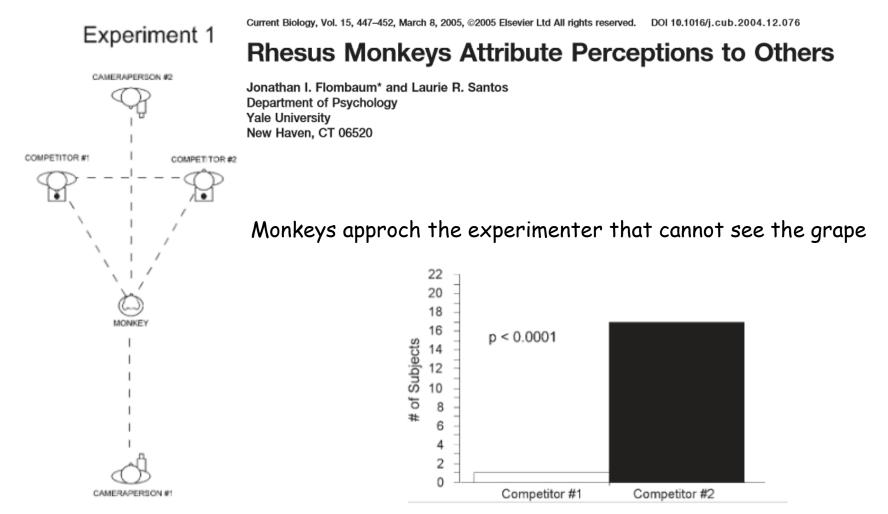






Unexpected

### **Attribution of perceptions**



Subjects were allowed to approach the grape in front of one of the two competitors while two camerapersons filmed them.

## **Capuchins understand intentions**

### **Developmental Science**

Developmental Science 12:6 (2009), pp 938-945

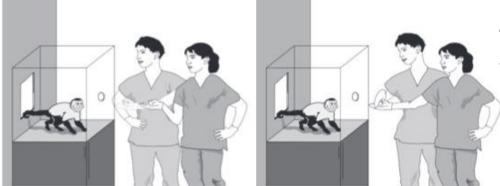
DOI: 10.1111/j.1467-7687.2009.00840.x

### PAPER

'Unwilling' versus 'unable': capuchin monkeys' (*Cebus apella*) understanding of human intentional action

Webb Phillips,<sup>1</sup> Jennifer L. Barnes,<sup>1</sup> Neha Mahajan,<sup>1</sup> Mariko Yamaguchi<sup>2</sup> and Laurie R. Santos<sup>1</sup>

Unwilling vs. unable in capuchin monkeys

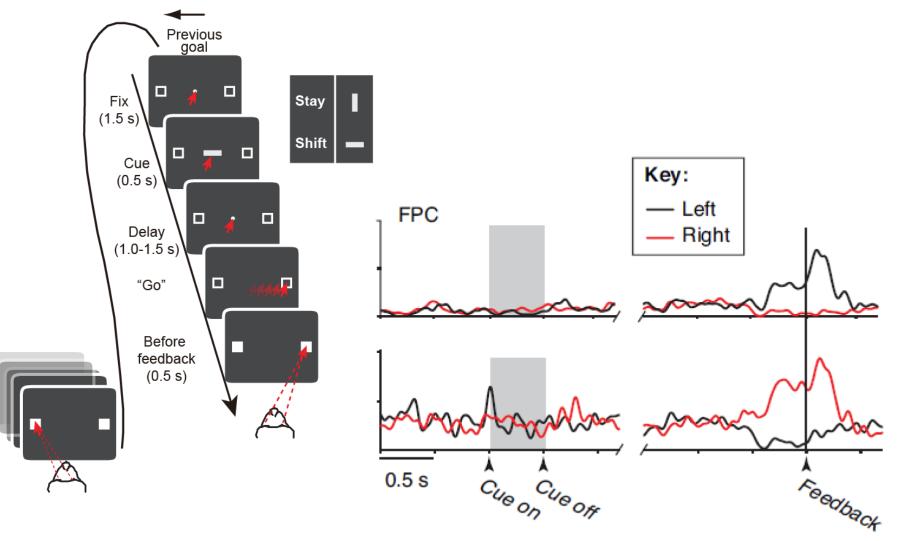


The monkey stays longer to wait in the unable condition.

A depiction of unwilling (left) versus unable (right) test events in Experiment 2.



### **Frontal Pole previously used task**



### **Differences between prefrontal areas**

