ICON Session S06:

Cross-modal integration and plasticity of sensory systems in the normal and peripherally deprived brain

- **Speakers:**
- **Franco Lepore** Université de Montréal
- (Canada)
- Krish Sathian- Emory University (USA)
- **Stephen G. Lomber** University of Western Ontario (Canada)
- **Amir Amedi** Hebrew University of Jerusalem (Israël)

Cross-modal compensation and plasticity in the blind and deaf: These two modalities do not always show similar outcomes

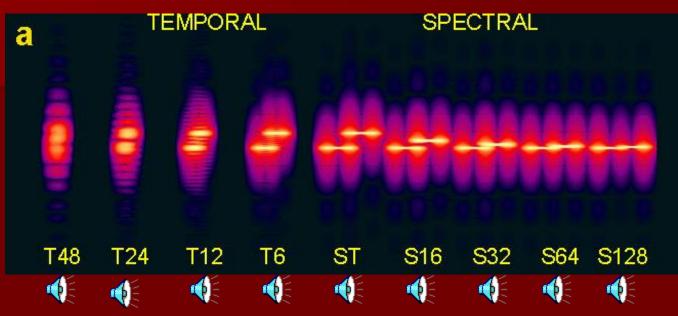
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With respect to the blind we have examined two hypotheses: 1- are they better than the sighted in various cross-modal tasks? 2- does visual cortex participate in task resolution in a functionally relevant manner?

Simple task: Pitch discrimination

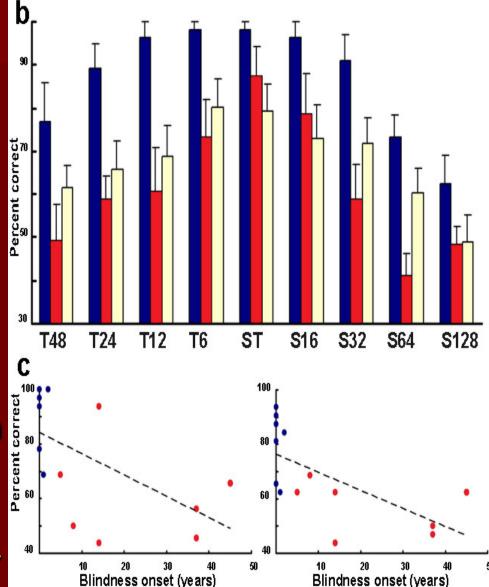


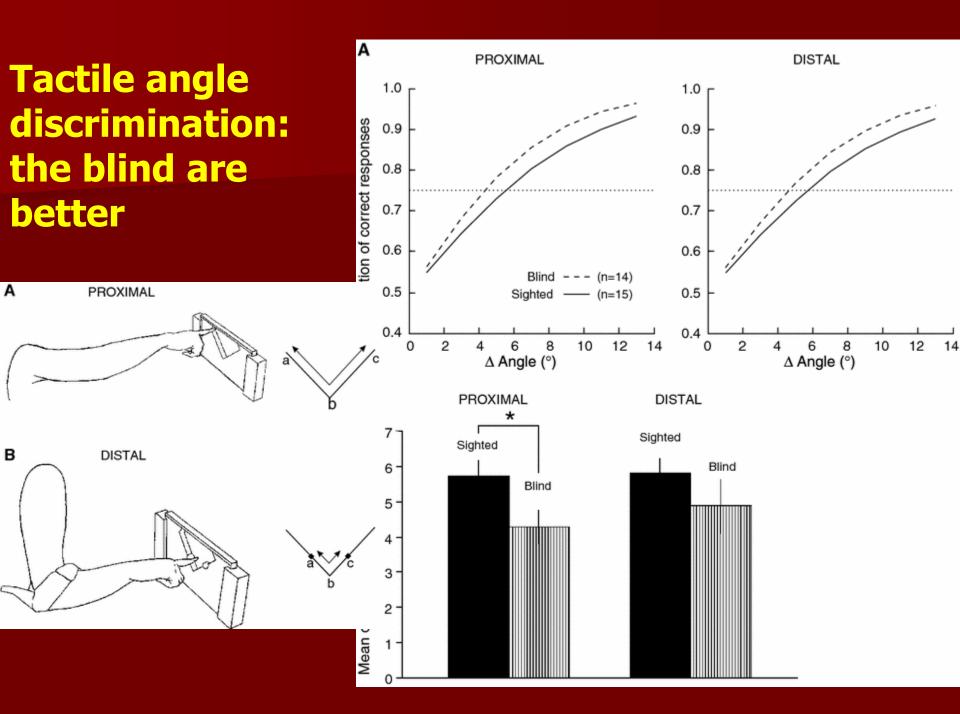
- Two tones of different frequencies or time intervals
- Is frequency decreasing or increasing?
- Standard stimulus (ST): 1.24 kHz, difference 1/8 octave, duration 333ms
- Increasing difficulty-symmetrical

Gougoux, Lepore, Lassonde, Voss, Zatorre, Belin (2004) Nature

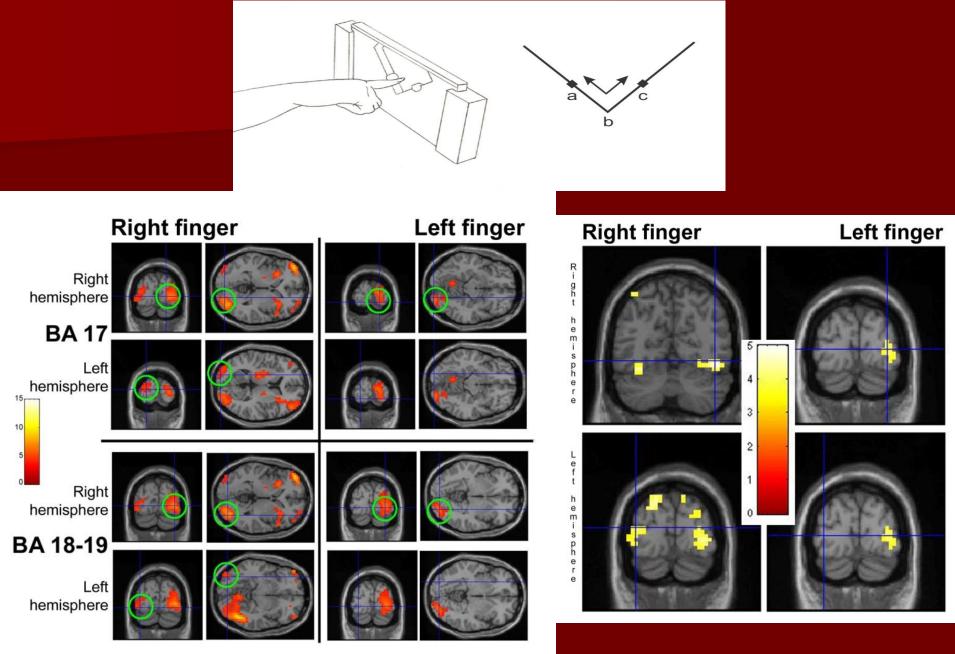
Pitch discrimination in the blind

- Judgement of pitch direction and inter-tone interval
- The early-blind are superior to the sighted but, with respect to the critical period, they are also superior to the lateblind subjects
- Performance was negatively correlated with age of blindness onset
- Gougoux, Lepore, Lassonde, Voss, Zatorre, Belin (2004) Nature

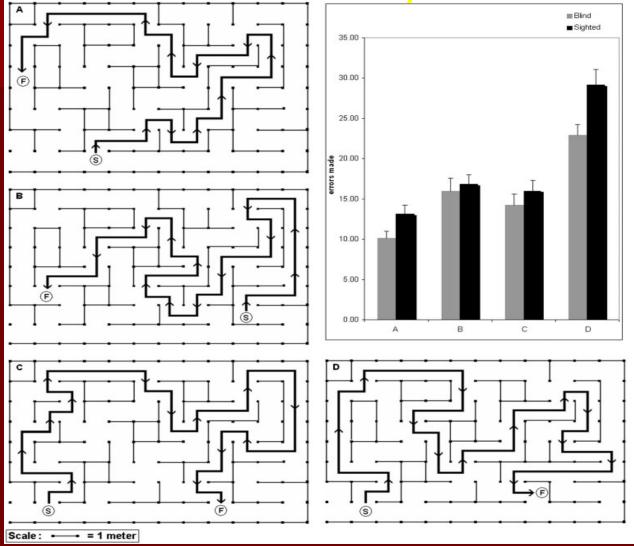




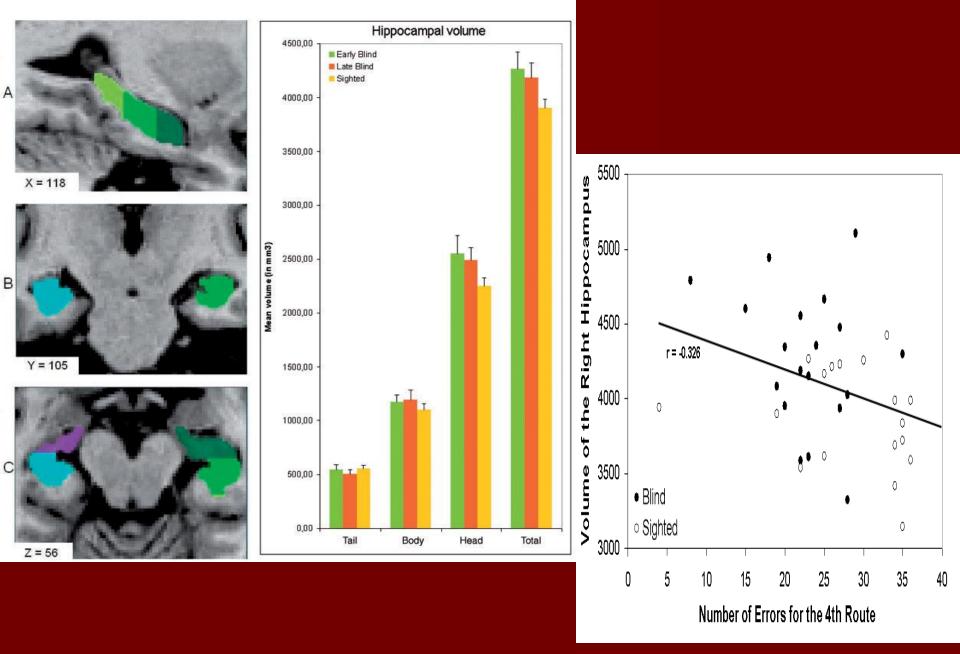
Tactile : angle discrimination in the blind



The blind are better at **navigating** in a human size labyrinth

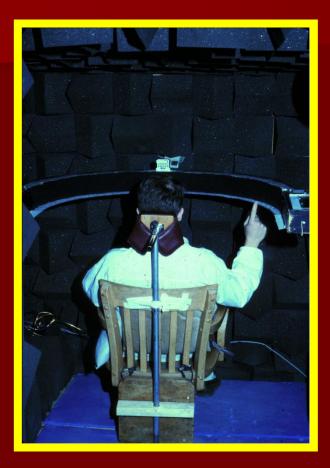


Fortin, Voss, Lassonde, Belin, Zato rre, Lepore, 2008, Brain



Fortin, Voss, Lassonde, Belin, Zatorre, Lepore, 2008, Brain

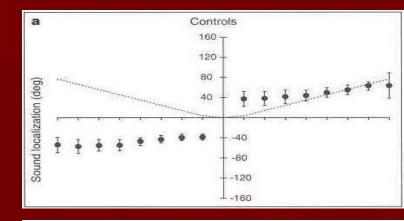
Monaural sound localization

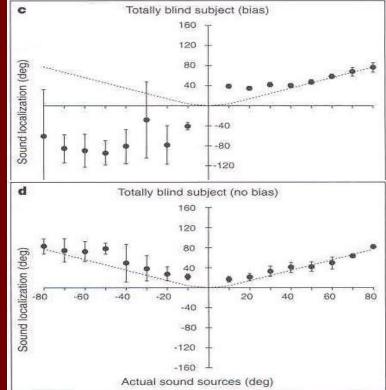


Lessard, Paré, Lepore, Lassonde (1998) Nature





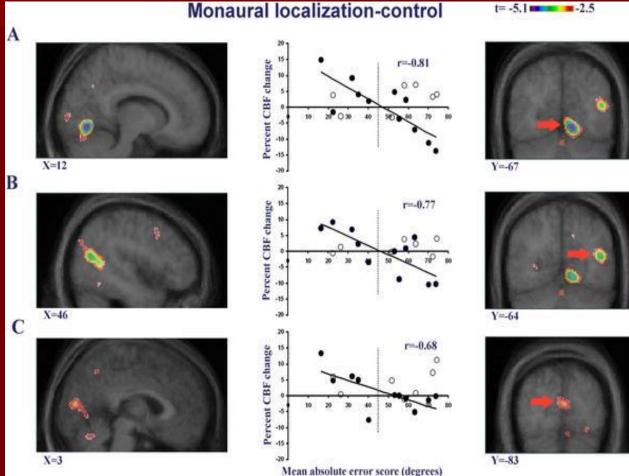




Monaural localization and PET

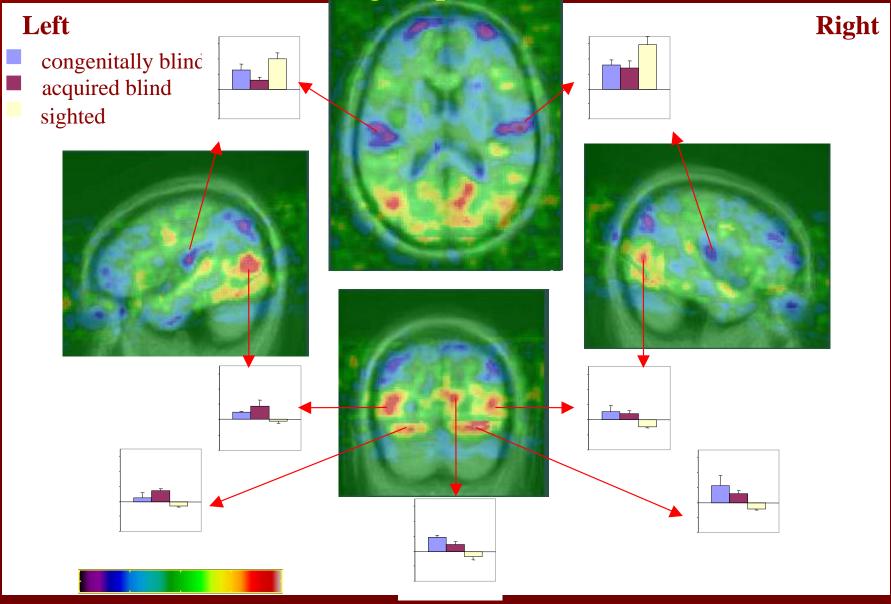
A strong activation was observed in different regions of visual cortex in the blind individuals

But more importantly, there was a correlation between degree of activation and localization performance



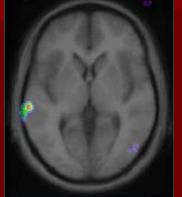
Gougoux, Zatorre, Lassonde, Voss, Lepore, (2005) PLoS Biology

Voice discrimination: All stimuli - silence Inter-group contrasts



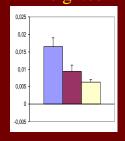
Vocal versus non-vocal Inter-group contrasts

congenital vs acquired



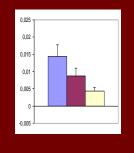


congenitally blind
 acquired blind
 sighted

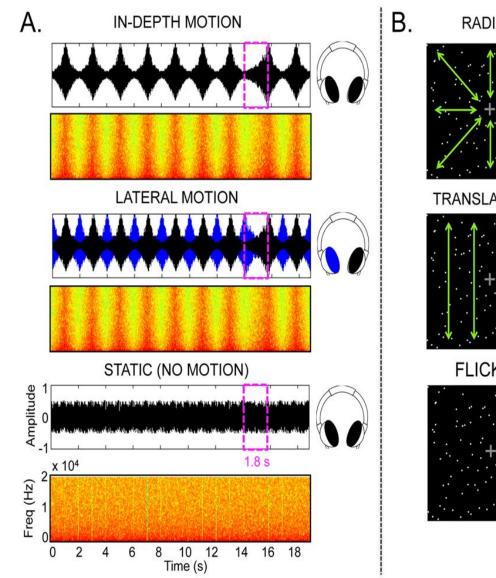


congenital vs sighted

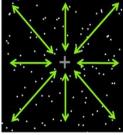




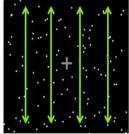
Gougoux, Lassonde, Zatorre, Voss, Belin, Lepore (Neuropsychologia, 2009)



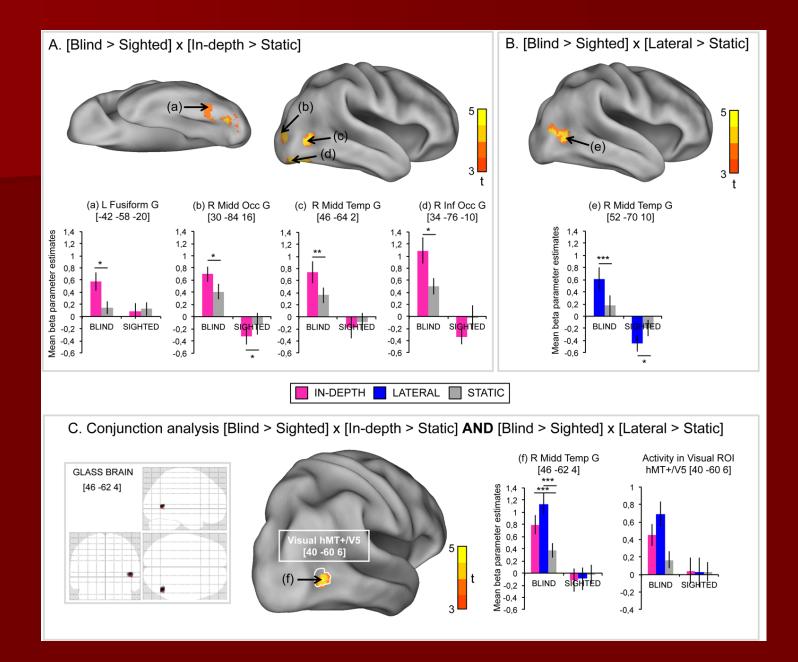
RADIAL



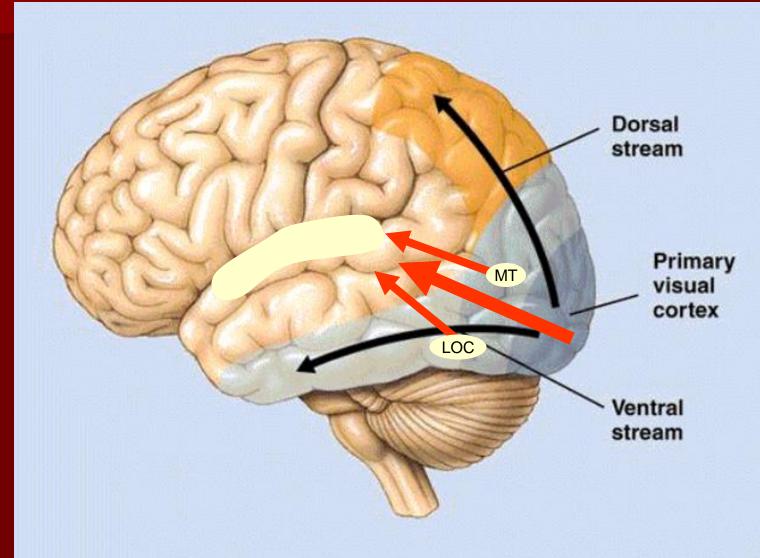
TRANSLATIONAL



FLICKER

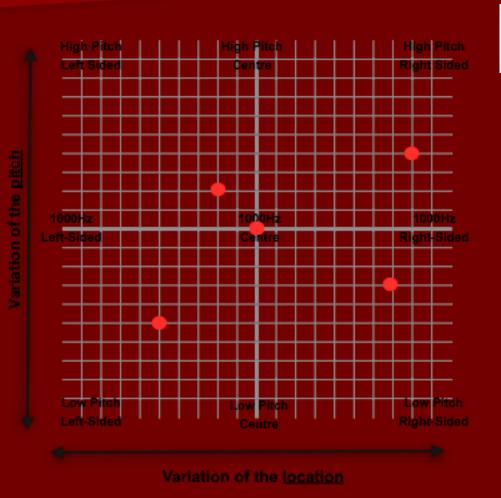


Does reorganisation in blind individuals also respect another critical function : ventral and dorsal streams

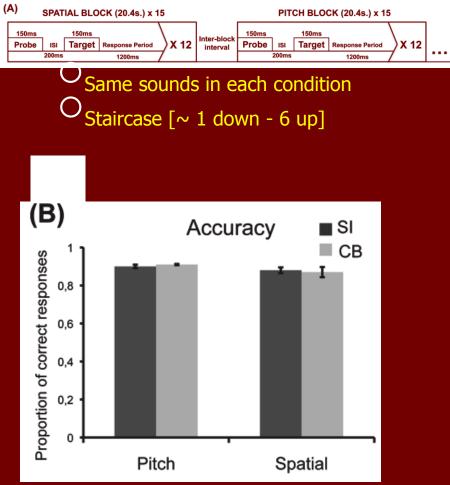




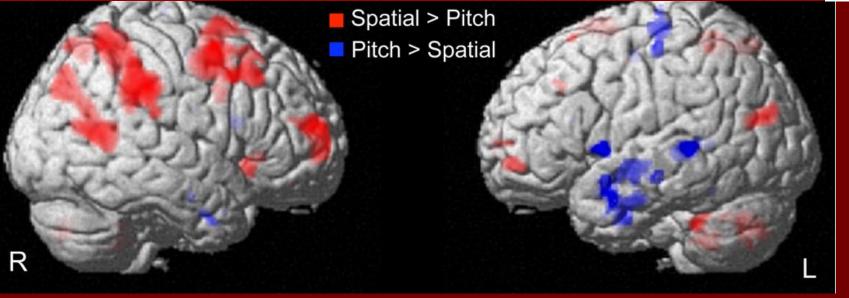
Design:

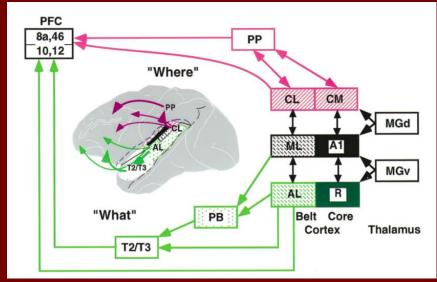


6400 sounds matrix







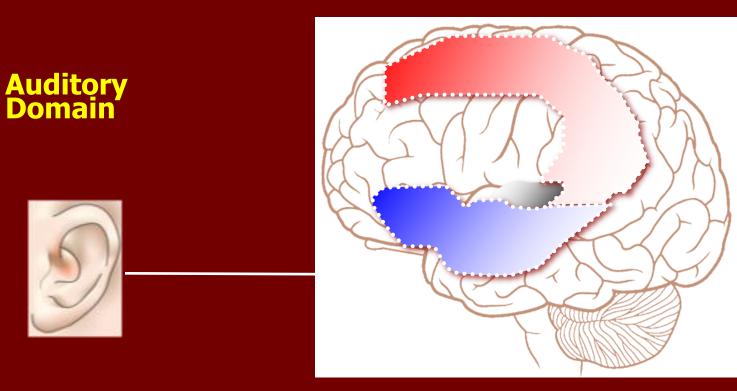


As seen with respect to the blind two comments can be made: 1- They are better than the sighted in various cross-modal tasks 2-The visual cortex participates in task resolution in a functionally significant manner

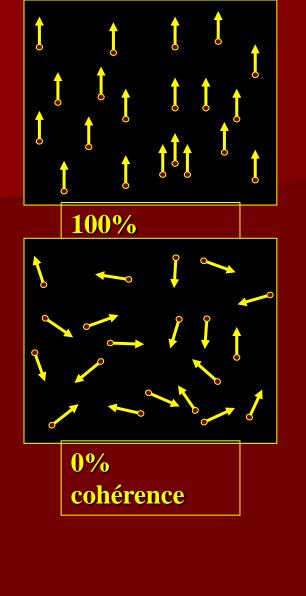
Does this hold for the **deaf** with respect to other functions?

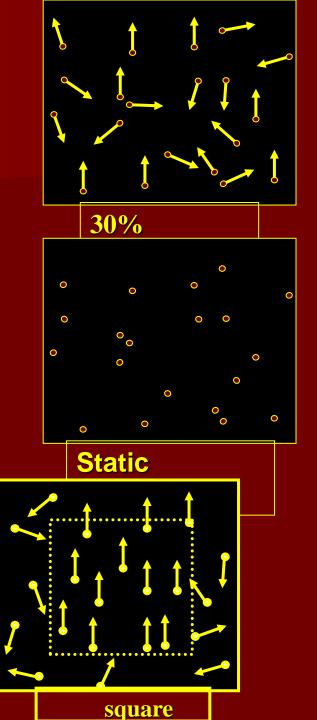


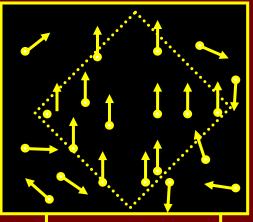
Dorsal Stream - Localization



Ventral Stream - Identification

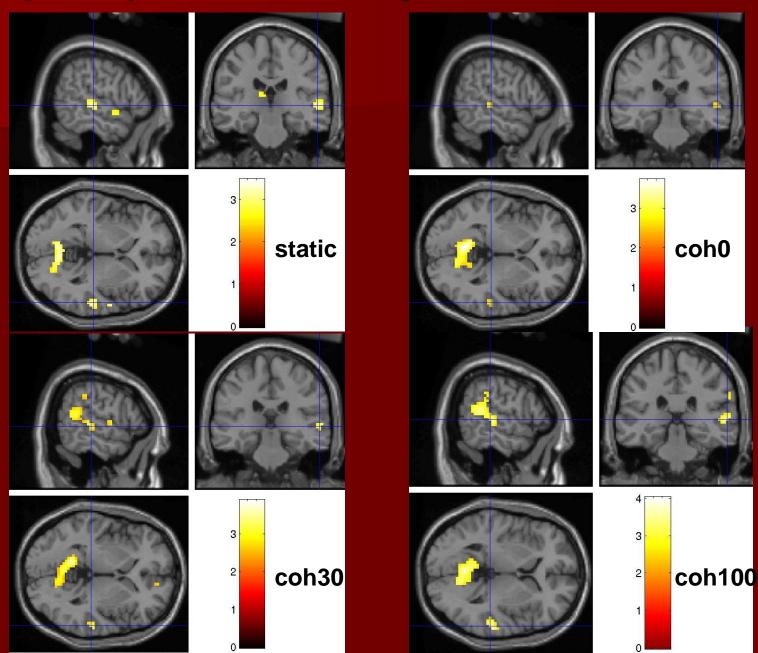




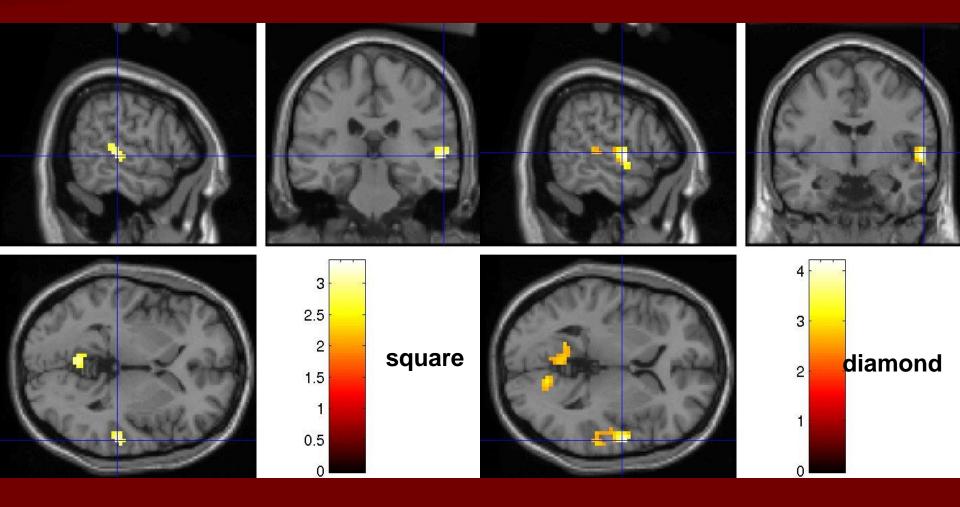


diamond

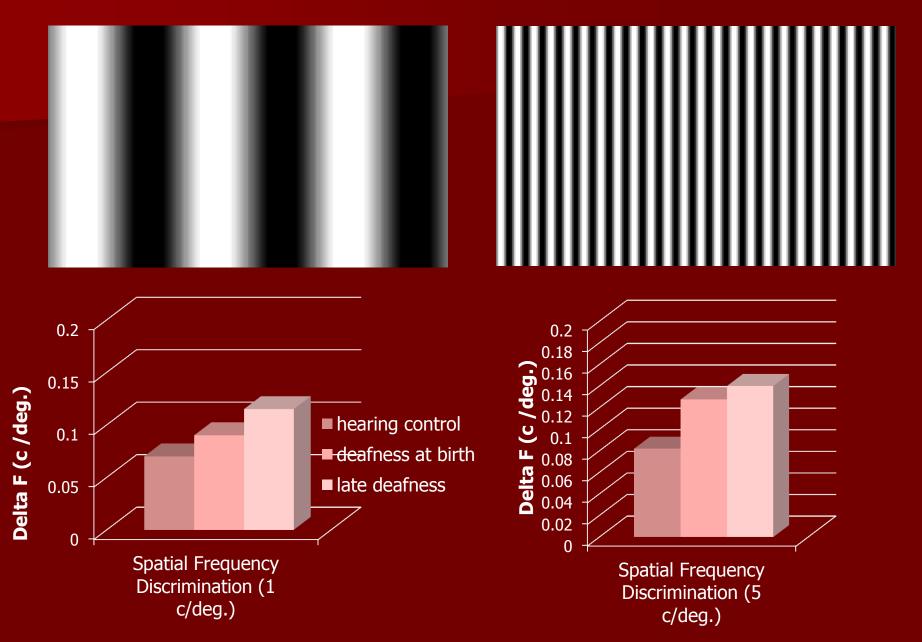
Activations under différent conditions between deaf participants and hearing controls: mouvement



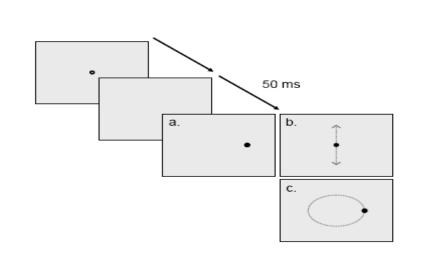
Activations under différent conditions between deaf participants and hearing controls: form

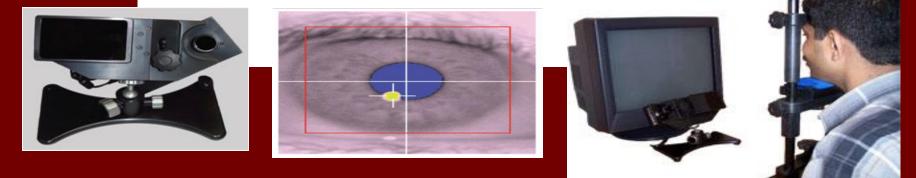


Visual Discrimination



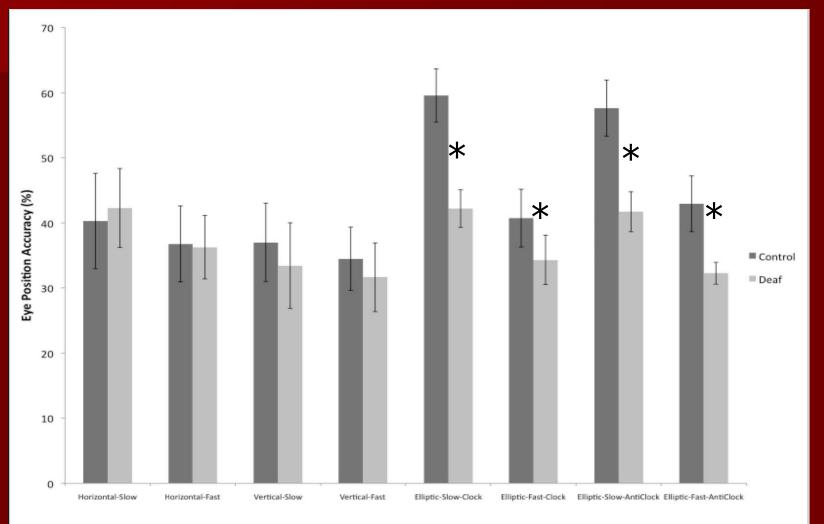
Auditory deprivation during infancy affects eye movement scanning function





Turgeon, Johnson, Lepore, & Ellemberg (CAA 2009)

Pursuit task



fearful

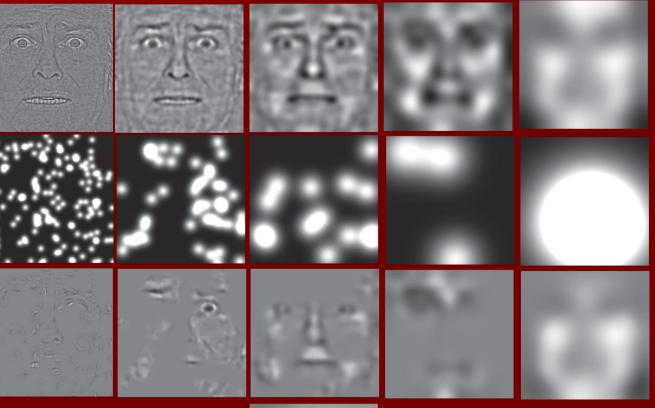




Doucet, Gosselin, Lepore, et al

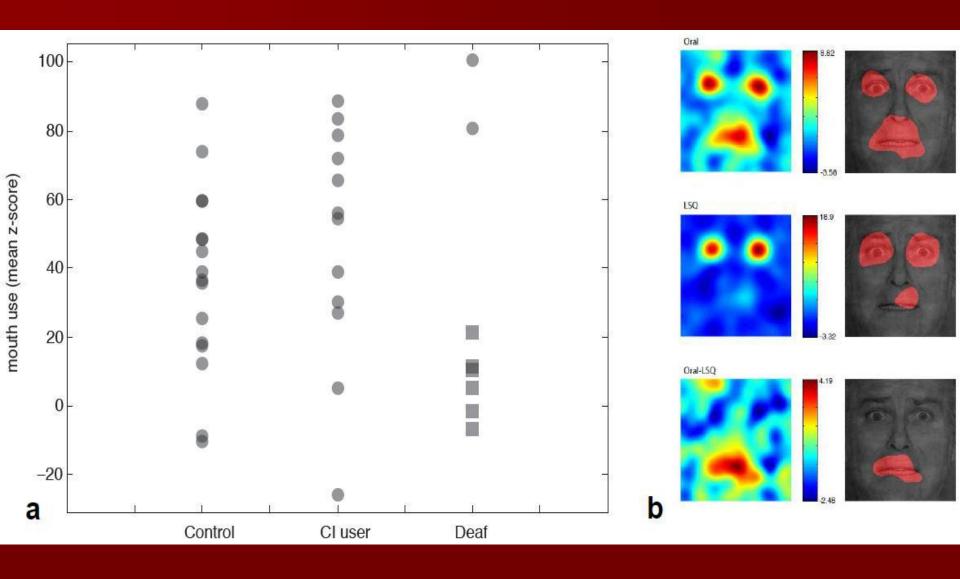
The "bubbles" technique i.e. gaussian apertures







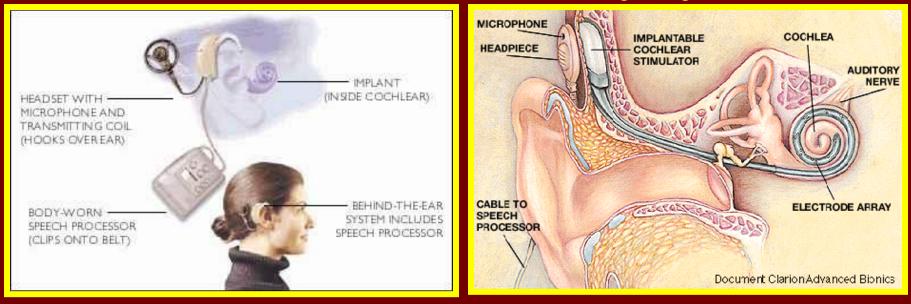
Gosselin et al, 2005; Schyns et al, 2004



The conclusions with respect to the **deaf** are: -Yes vision recruits in a cross-modal fashion auditory areas

-No, in both simple and higher level tasks we found no supra-performance and in fact there is under-performance

Reactivation of a sensory system



Collaborators

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<u>University of California at</u> <u>Los Angeles and University</u> <u>of Southern California</u> Paul Thompson Natasha Lepore

> McGill University Robert Zatorre

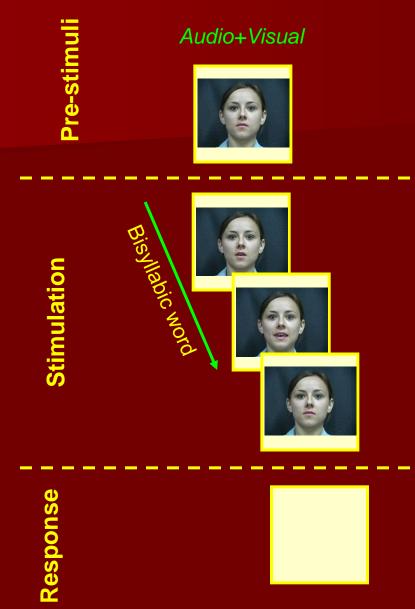
Funded by: NSERC, FRQS, CRC, CIHR, CEDAR

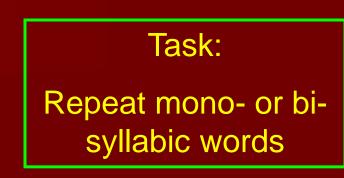
Thank you

« And the Blind shall hear »

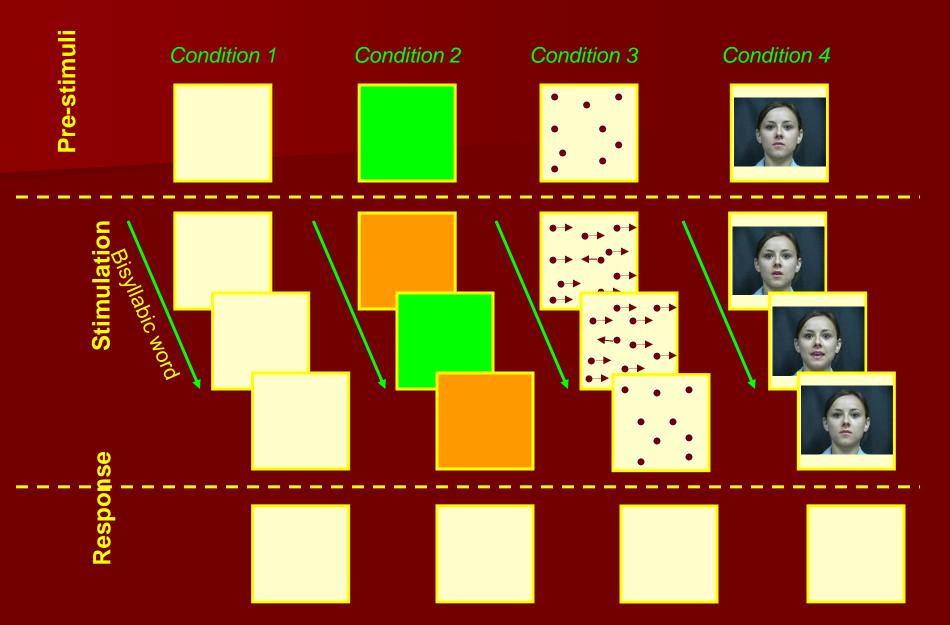
- Results show that cochlear implants are efficient in re-establishing oral language comprehension
- However, they are not always equally efficient for all deaf subjects to recognize language. Why?:
- We examined the conditions that affect performance using four modes of presentations of **40 bi-syllabic words**:
- -presented alone
- -presented simultaneously with a color
- -presented with coherently moving dots
- -presented with facial expressions saying a different bi-syllabic word

What happens in language comprehension?

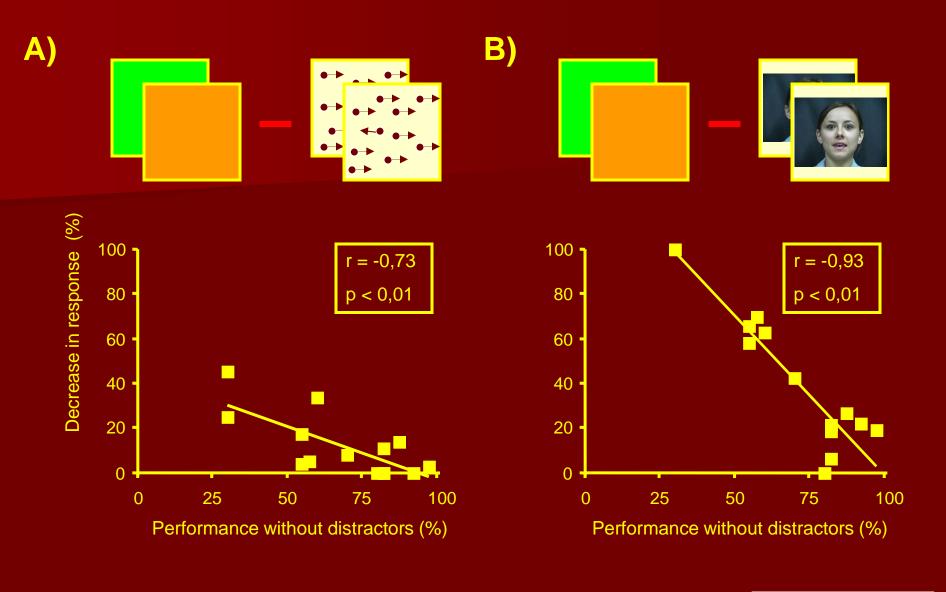




Some deaf subjects who have received a cochlear implant perform well on this task while others do not



Champoux, F, Tremblay, C, Lepore, F, Theoret, H, Neuropsychologia, 2008

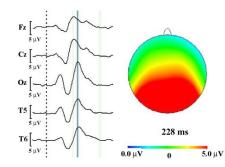


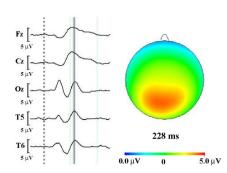
Champoux, F, Tremblay, C, Lepore, F, Theoret, H, Neuropsychologia, 2008

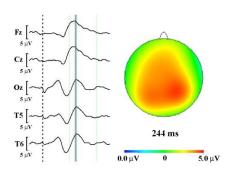
Good performers

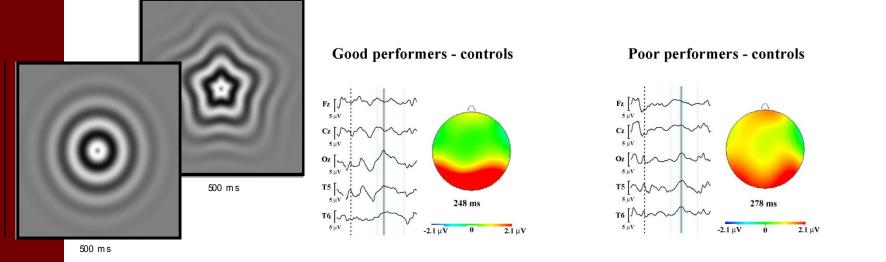


Poor performers









Visual evoked potentials to the presentation of the transformational apparent motion stimulus for good performing and badly performing subjects Doucet, Lassonde, Lepore et al, Brain, 2006 Measure used to study dancing in cochlear implanted subjects: Perception and time reproduction of music using a synchronization and music capture apparatus

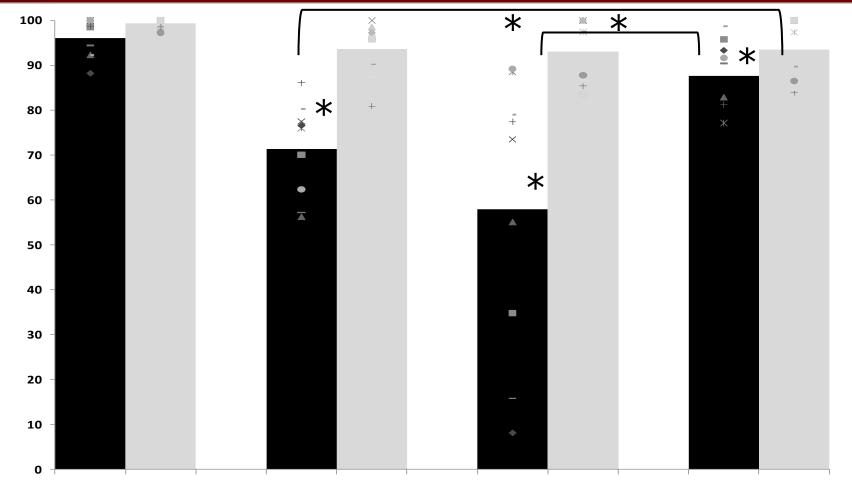
It gives: 1.Periodicity of body mouvements 2.The constancy of this mouvement 3.The ability to compare performance with norms

-)

Wii

Music perception and dance with cochlear implant

Hearing
 Cochlear implants



Métronome

Merengue

Piano

Batterie

These results with hearing restauration using cochlear implants with the deaf raise an interesting question: what happens with sight restauration in the blind- the **Boston Keratoprosthesis (Kpro)**



We have seen that there appears to be a « sensitive period » for plasticity to manifest itself. But is this an absolute rule? The Boston Keratoprosthesis study seems to indicate that this is not true!



Adlave, 2009; Dagher & Dohlman, 2008; Dohlman, 2007; Khan et al., 2008

Sight restoration following lens implant

Patient: 41 years, Rieger Syndrome
 Keratoprosthesis of the right eye

C		OD	OS	OU
	PRE			20/400
	POST +J3	20/100		20/100 - 20/125

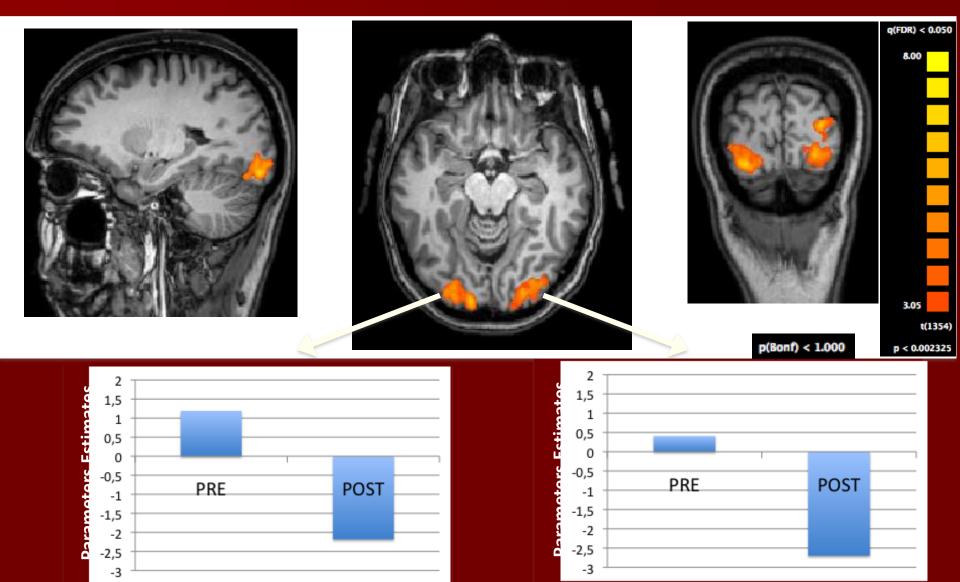
Examined with fMRI 1) Just before implant 2) 3 days after

Auditory and visual tasks



Cross-modal plasticity before Kpro

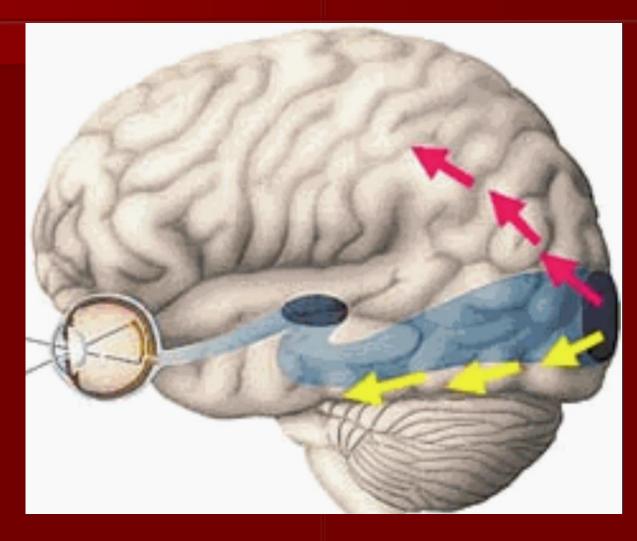
AUDITION Before KPro > After KPro



Face specific ventral visual areas







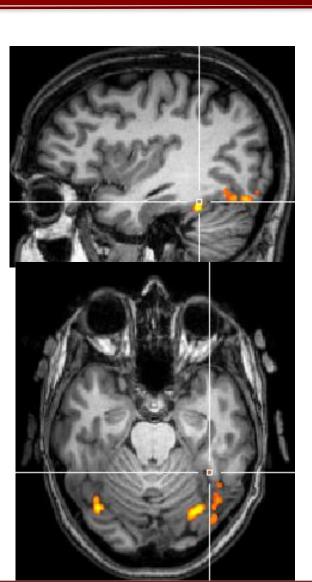
Haxby et al., PNAS, 1991; Kanwisher et al., The Journal of Neuroscience, 1997

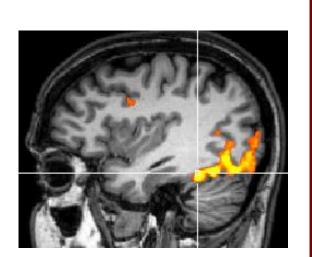
Before KPro

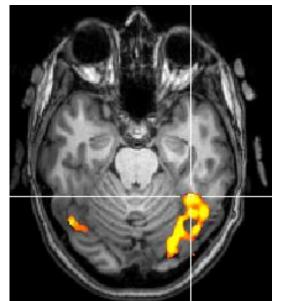
3 days after Kpro

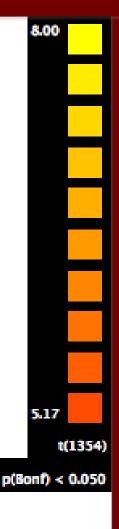




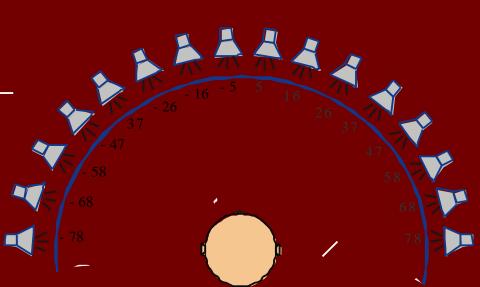




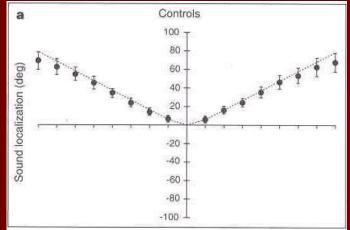




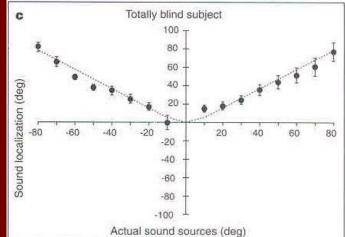
Binaural sound localization



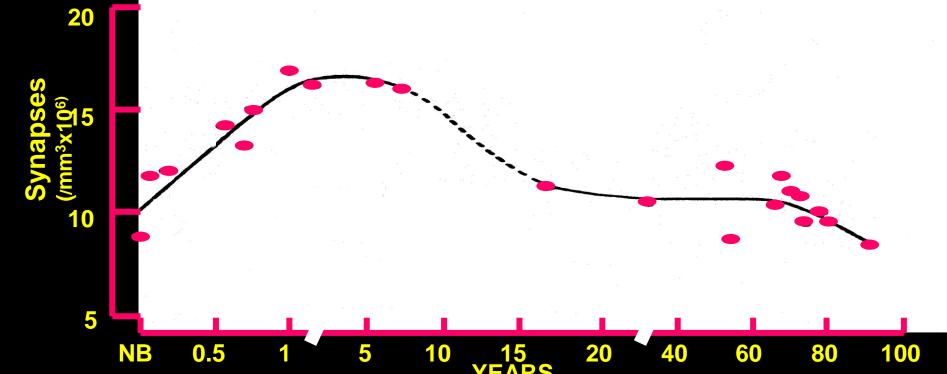
Sighted



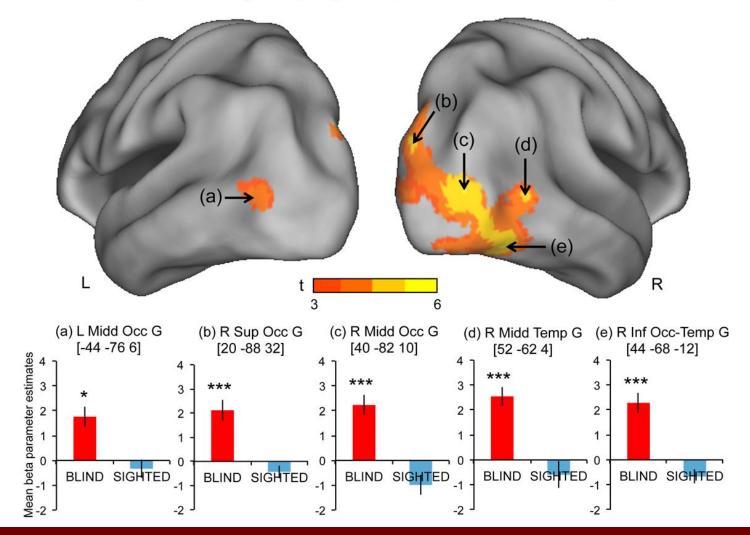
Blind



Lessard, Paré, Lepore, Lassonde (1998) Nature The human brain is hard-wired at birth. However, following development and learning, a number of phenomena take place, such as **synaptic pruning**, which modify substantially its structure. These modifications are normal and allow us to act, feel and adapt to the environment as well as to others. These are time dependent, well known as the **critical period**.



[Blind > Sighted] x [In-depth + Lateral + Static]

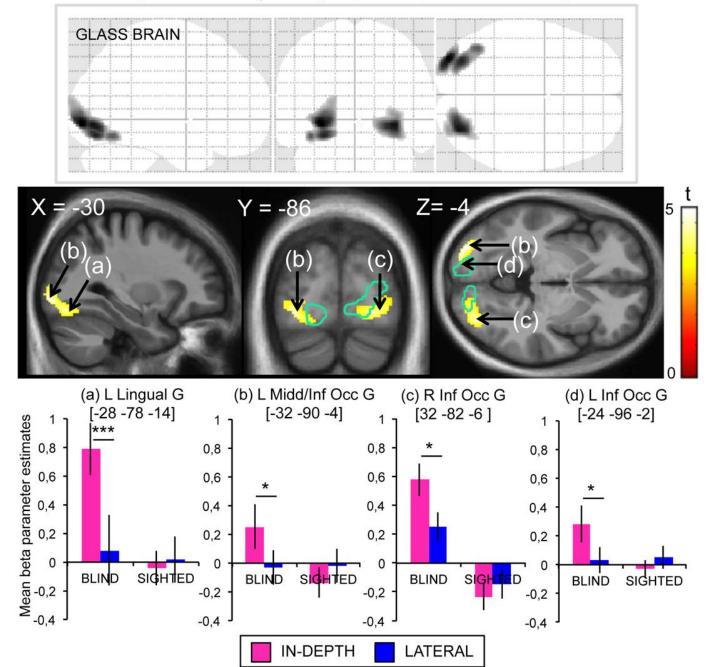


We have in the context of the studies to be presented today two subject populations who had significant sensory impairements: visual loss or deafness

- -with respect to vision loss, we shall examine two subpopulations:
- -blind subjects and individuals with lens restauration

-with respect to the deaf, we shall also look at two sub-populations: deaf subjects and individuals whose auditory functions were re-established with a cochlear implant

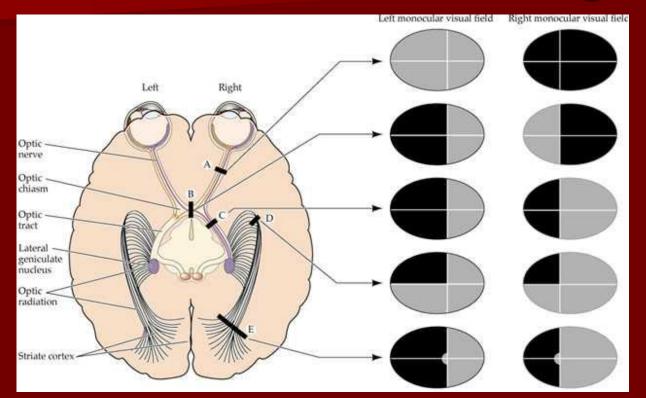
[Blind > Sighted] x [In-depth > Lateral]



We have shown important plasticity and reorganization in individuals who loose sight through peripheral receptor loss. How do persons adapt to vision loss due to cerebral cortical loss: the case of hemianopic subjects

I would like to specify that this is a new project and we only have preliminary data. The reason for presenting it is to show another field in which we are working and where biomedical imaging can furnish an excellent tool to study spared function and the effect of rehabilitation

Hemianopia and blindsight: why is it interesting?



Unilateral loss of posterior visual cortex due to surgery or stroke leads to blindness in contralateral field.

- Sign of a residual cortical or subcortical visual pathway remaining.
 - Residual visual capacities in the blind field.
 - Develop a rehabilitation technique to improve the visual abilities in the blind field.

Our plan...

- Compare the results obtained in two populations.
 Cortical infarct
 Delimited lesion of the visual cortex
- POSTERIOR PARIETAL LOBE WHERE Orsal stream Or of the stream WHAT INFERIOR TEMPORAL LOBE
- Evaluate the **residual visual capacities** of our participants.
- fMRI and Diffusion Tensor Imaging (DTI) to try to define the functional activations and connectivity between areas.
- Training using coupled visual/auditory stimulations and stimuli which are treated by subcortical structures, such as moving dots (to come in the future!)

Our hypothesis ...

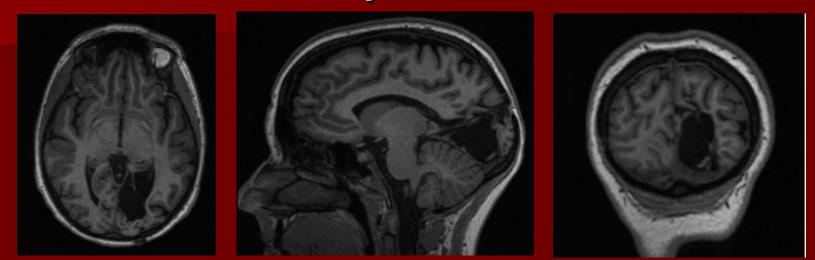
Implication of the **subcortical pathways** passing by the superior colliculus and pulvinar and by the "where" or **parietal dorsal stream pathway**.

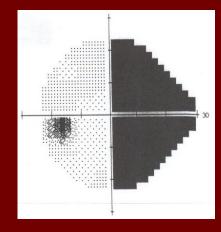
Preliminary behavioural study on two hemianopic participants

- Two participants :
 - Participant 1 : right homonymous hemianopia.
 - Participant 2 : left homonymous hemianopia
- Use the functional specificity of the superior colliculus and of the parietal pathway to evaluate the quality of the residual visual capacities and to determine the implication of these regions.
 - (low frequency, motion, multisensory stimulation and integration).

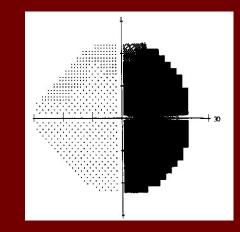
All potential saccadic movements were controlled with an EOG system.

Participant 1





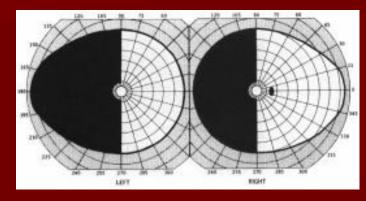
Retinotopy left eye

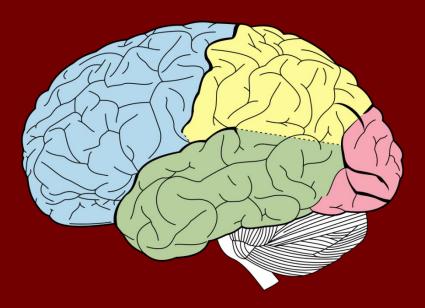


Retinotopy right eye

Participant 2

- Had epilepsy in her right hemisphere.
- Had the removal of the visual cortex the 24 of September 2013.
- Doesn't have any sensation of visual stimulation, including motion in her natural environment.
- Has developed a sweeping oculomotor technique, when concentrated.





Patch of moving dots paradigm

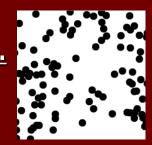
(All the tasks where carried out in the blind field)

First part : Presentation of moving dots versus static dots.

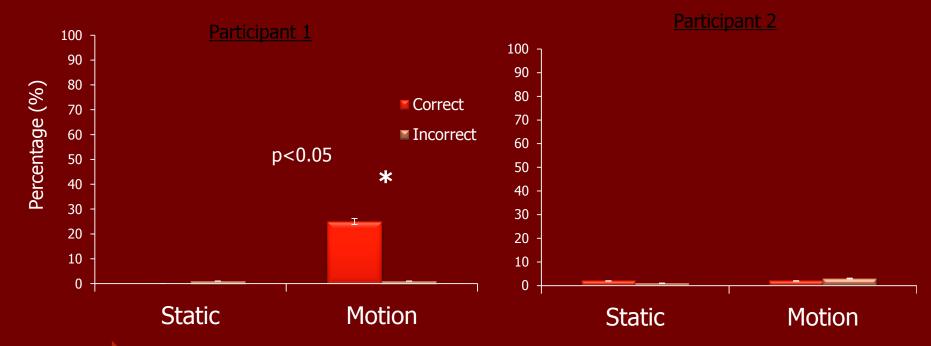
- 12 degree patch
- White background
- 150 dots of 0.5 degree each with a 12°/sec for the vertically moving dots.
- Presented at 12 degrees.

Second part : Discrimination of the direction of the motion.

- Same parameters as in the first part.
- Upward and downward discrimination.
- White noise auditory stimulus to initiate the answer presented at the same position.

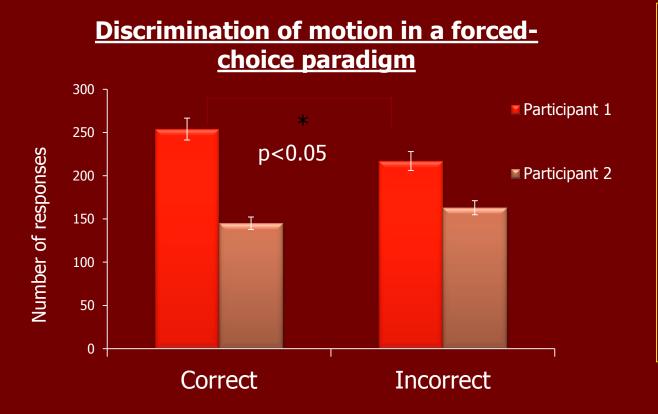


Is there any reported perception in the blind field when presented with moving dots or static dots?



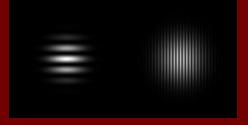
* Participant 1 reported correctly a sensation of motion when moving dots were presented compared to the presentation of static dots. **No false positives.**

Is this awareness reflected in the detection of the motion direction?



* Significant results only in participant 1 who had better performance when discriminating between downward and upward motion. Gabor patch (12°) of low and high spatial frequency

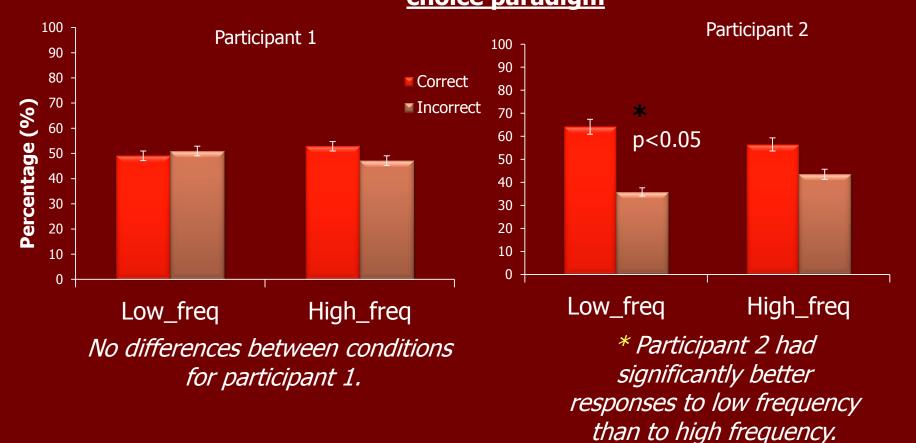
- Low frequency = 1 Hz. High frequency = 5 Hz.
- Two directions = vertical and horizontal.
- Two positions : 20° on the y axis in the upper field and in the lower field, with a 24° on the abscissa.
- Motion detection of a moving single gray bar
 - 4 directions of movement : up, down, left, right.
 - Moving from 10° to 36° on the x axis and from 20° upper field and 20° lower field.
- Pointing task
 - Gray flash presented in random positions.



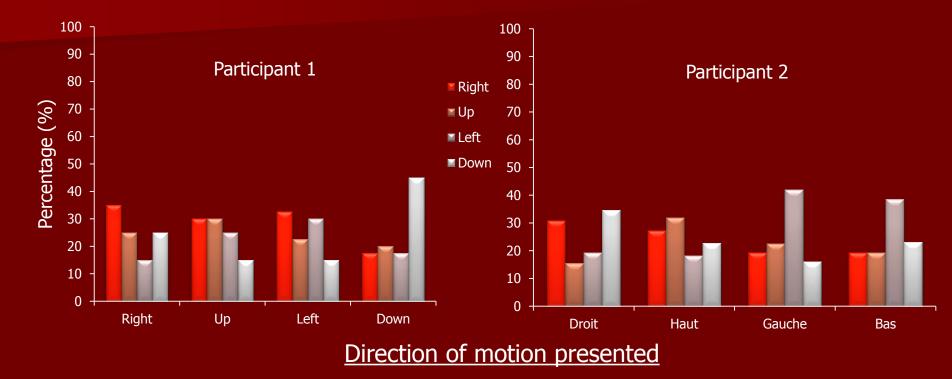


Low and high frequency Gabor patch presented vertically or horizontally Orientation discrimination in a forced-

<u>choice paradigm</u>



Motion detection of a single bar



The participants could not detect the direction of the moving bar when presented in their blind field.

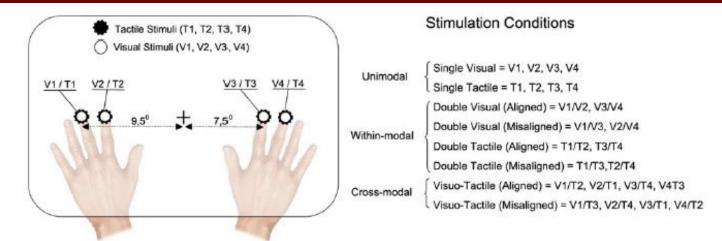
Pointing task: No relation was found between the position of the target and the position aimed.

Multisensory integration

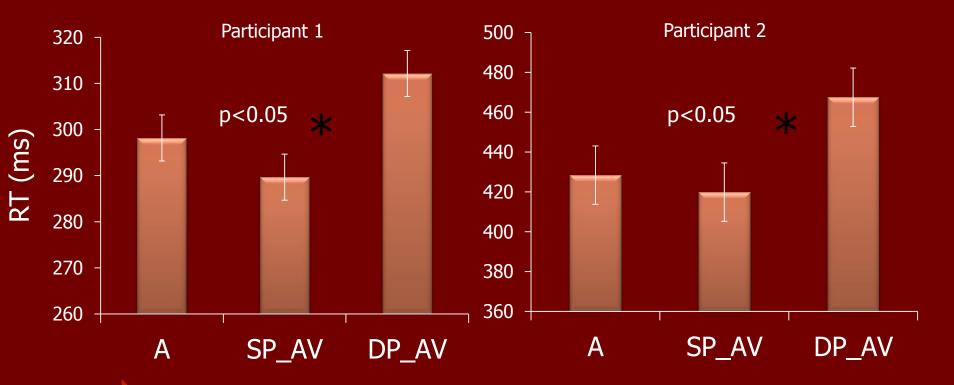
Audio-visual paradigm (gray flash of 2 °)

- 4 positions (x/y): 8°/20°; 8°/-20°; 36°/20°; 36°/-20°.
- Soft white noise presented alone (A), at the same position (SP) or at a different position (DP) of the visual stimulus.
- Localisation of the sound at 8 ° or 36°.
- Reaction time was measured.

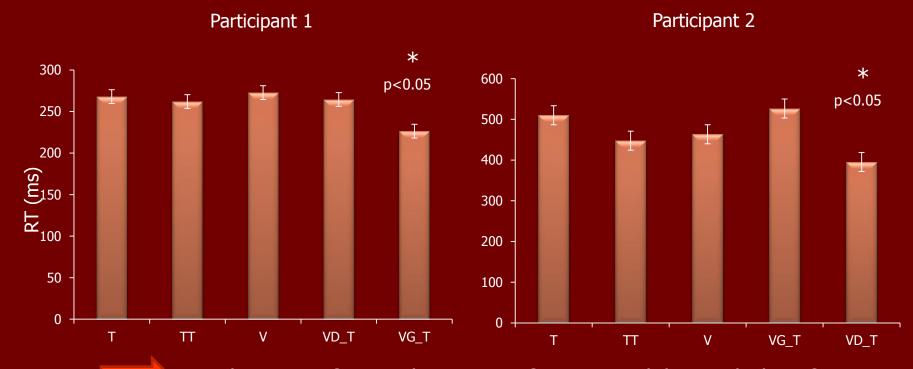
Visuo-tactile paradigm



Audio-visual integration in a localisation task presented in the blind field



* Faster RT in same position (SP_AV) condition compared to different position (DP_AV), reflecting in integration with unseen visual stimuli. Impact of the spatial congruence of redundant targets on within-modal and cross-modal integration (tactile and visual)



Slower RT for combinations of cross-modal stimuli than for combinations of within stimuli shown only when the visual was presented in the normal field.

Magnetic Resonance Imaging Hypothesis & Scanning session

- Hypotheses : Hemianopic participants present different pattern of activation when seeing stimulus movement depending on their blindsight performances
- -they could also present different white matter tracks supporting their behavioural performances.

MRI Scanner Siemens Trio 3T :

- Anatomical scan
- Diffusion Tensor Imaging scan
- BOLD scan :
 - Whole brain
 - 17 slices Thalamic focused scan
- Resting State scan

Statistical approaches

Whole brain analyses :

GLM analyses of the visual mouvement induced activations

ROI based Superior Colliculus (SC) imaging :

 Noise-Reduction ROI based analysis (ROI : SC ; Anterior Cerebellum ; V1 ; V5 ; LGN)

(De Zwart et al. 2008)

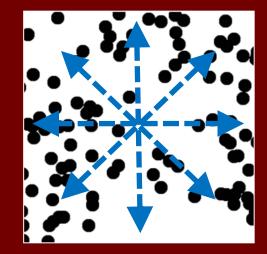
Custom HRF with max amplitude pick at 4s (Wall et al. 2010)

ROI based DTI :

 Using the same ROI : creation of tracks linking vision and movement perception for cortical sub-cortical areas (Leh et al. 2006)

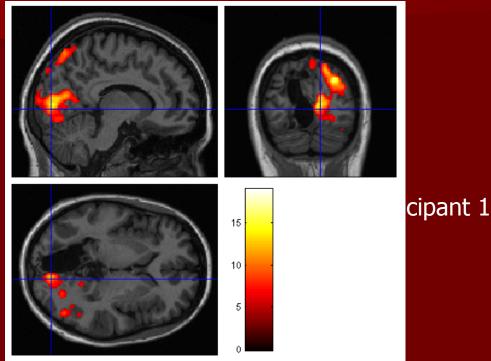
Preliminary results In-scan performances

- Stimuli : Presentation of moving dots patches in the left or the right visual hemifield
- Task : Report any feeling or sensation of presence of the stimulus in the blind field



- Results : Detection in the blind hemifield 89,23 % (P<0,05)
- Control : MR-Video Eye-tracker, the excentricity has been tested in order to avoid any stimulation that could occur in both hemifields.

Preliminary results WB BOLD activation



Left cortical activations (stimuli presented in the right hemifield) (p< 0,05(FWE))

Right cortical activations (stimuli presented in the left hemifield) (p< 0,05(FWE))

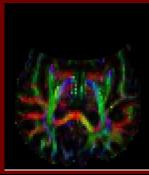
10

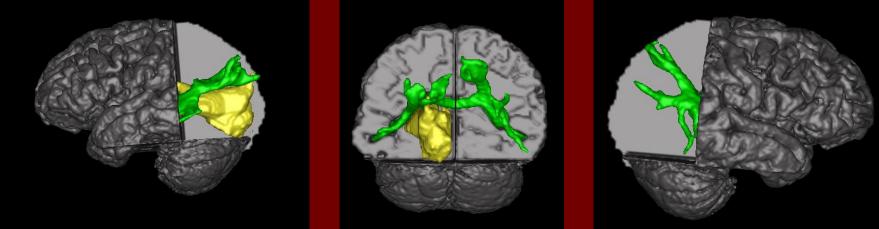


There are contra-lateral BOLD activations during visual presentation in the blind hemifield

Preliminary Results Diffusion Tensor Imaging

Plotted regions : Striate Cortex (V1) ; Extra-Striate Cortex (V5) ; Sub-Cortial areas (Pulvinar ; Superior Colliculus ; Lateral Geniculate Nucleus) Other Regions (Cerebellum; Corpus Callosum)





Corpus Callosum posterior tracts (green) and surgery caused lesion (yellow) (participant 1)

The lesion induces a structural asymetry between left and right tracks As seen with respect to the **blind** the conclusions that can be made are:

-They are better than the sighted in tasks involving other modalities

-Visual cortex participates in task resolution in a functionally significant manner so that more cortex means better performance

Next step...

Test 10 other hemianopic participants (cortical stroke and delimited surgical lesion groups).

Begin the audio-visual and the moving dots training.

Training of 2 hours /day for 10 consecutive days.

Audio-visual training :
 Visual stimulation presented with a white noise to improve the oculomotor search.

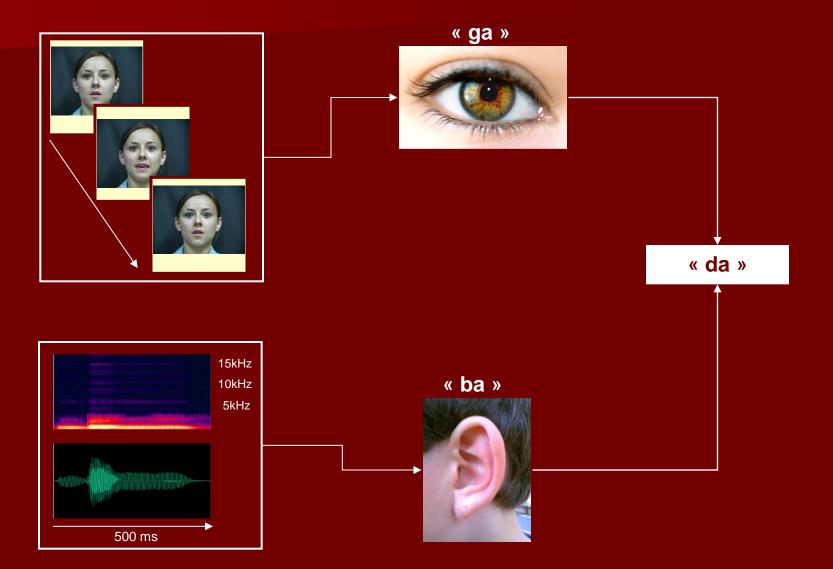
Moving dots training :
 Forced choice paradigm of the discrimination of the direction of moving dots (same pardigm seen before).

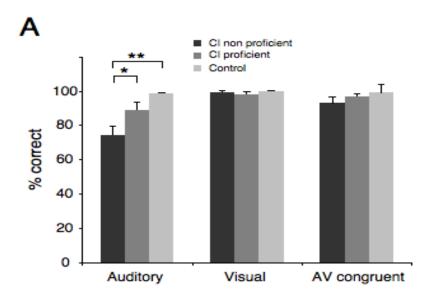
Compare the results obtained in the behaviour and fMRI tasks **before and after training**, while also comparing the **lesion** of each participant.

Develop a readaptive technique to restore partially the vision in the blind field.

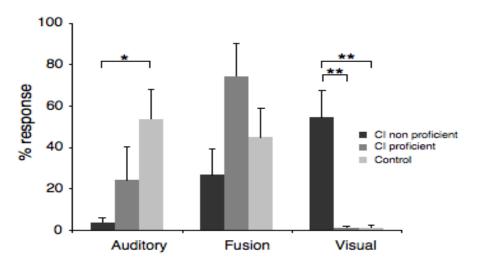
But what about specialized function, which in humans are generally treated in specialized areas or structures? Are they also transferred to these structures? -The spatial navigation and the Hyppocampus (involved in route learning i.e., the London taxi drivers -Human vocalizations, that are generally treated in structures within the superior temporal sulcus (STS)

Visual-Auditory interactions: The McGurk effect



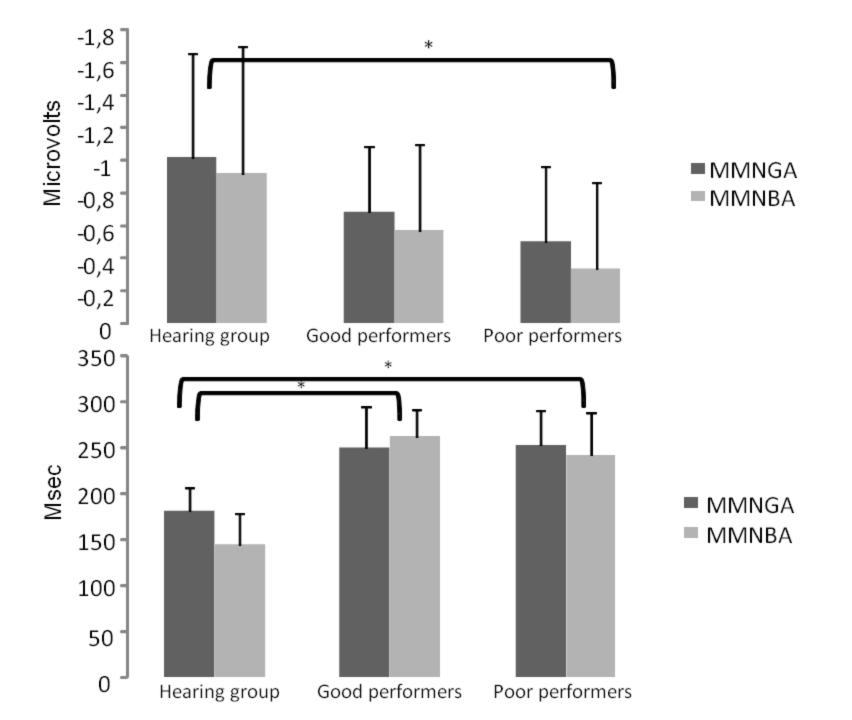


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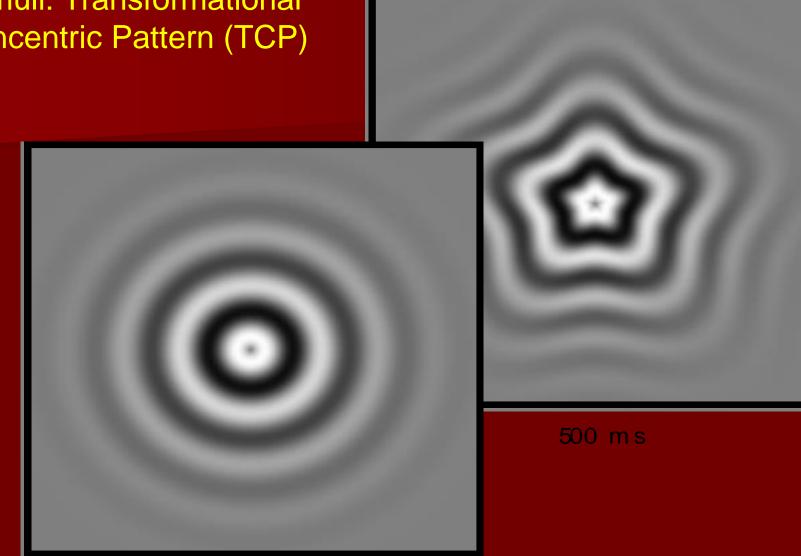


Performance in McGurk audio-visual trials for normal, pCI and npCI groups

Tremblay, Champoux, Lepore et al Rest Neurol Neurosci, 2010



Stimuli: Transformational Concentric Pattern (TCP)



500 m s

Doucet, Lassonde, Lepore et al, Brain, 2006

Control

Deat





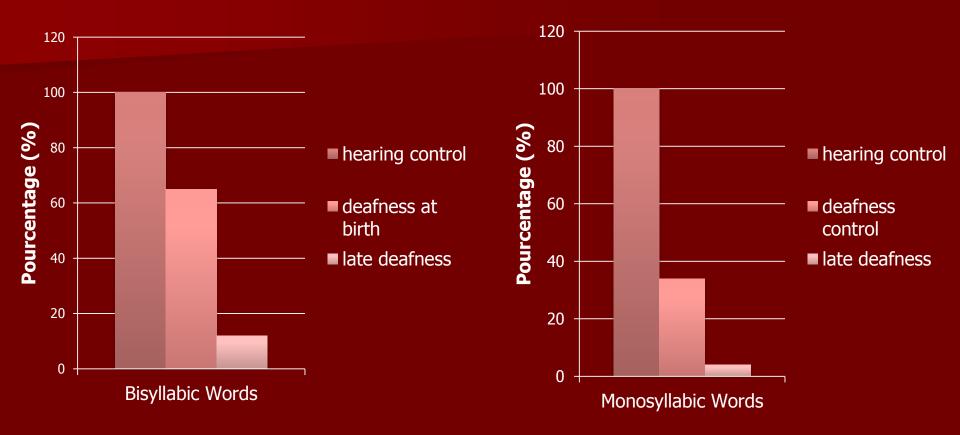


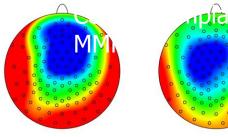


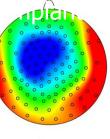
Нарру

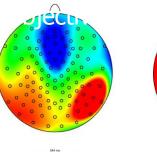
Doucet, Gosselin, Lepore, et al ECVP 2008

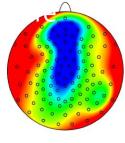
Speech discrimination in silence







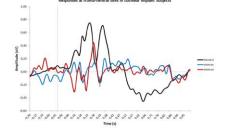


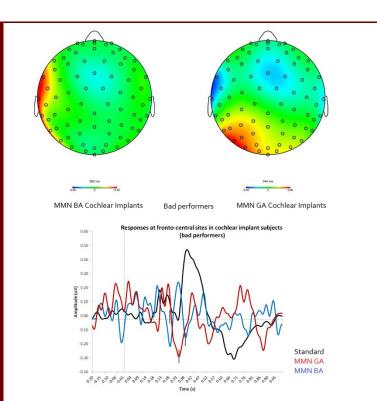


MMN BA Cochlear Implants

0.50 pl MMN GA Cochlear Implants



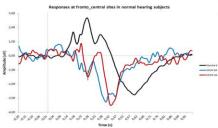




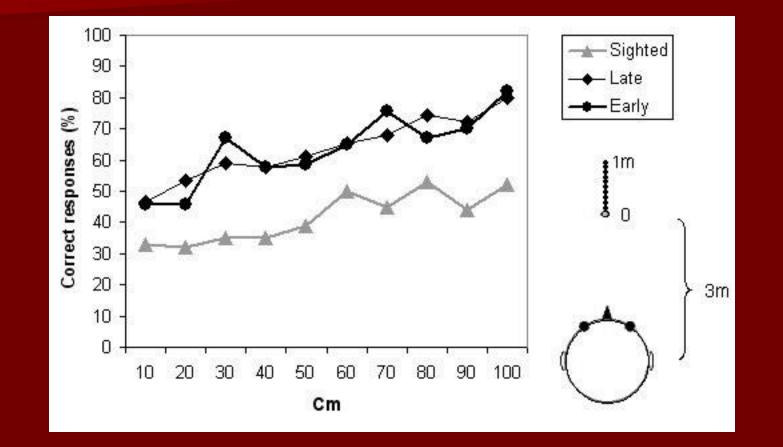
MMN GA Normal Hearing

.....

MMN BA Normal Hearing

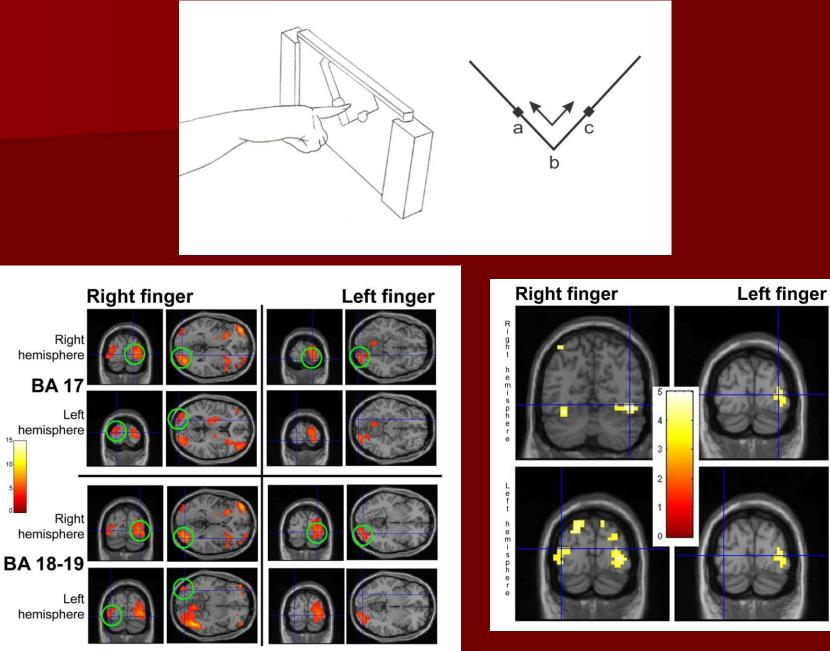


Sound discrimination in far space



Voss, Lassonde, Gougoux, Fortin, Guillemot, Lepore (2004) Current Biology

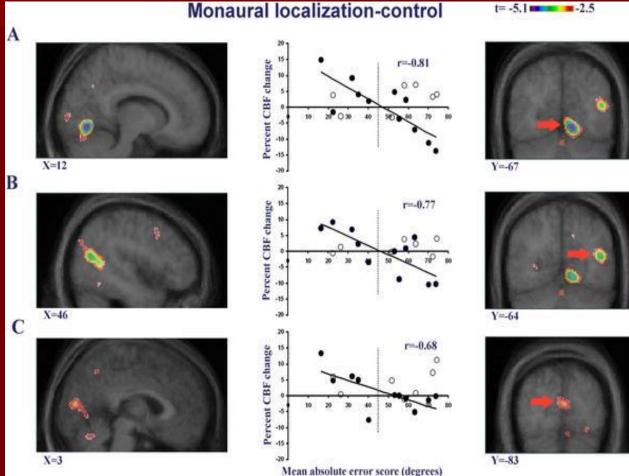
Tactile : angle discrimination in the blind



Monaural localization and PET

A strong activation was observed in different regions of visual cortex in the blind individuals

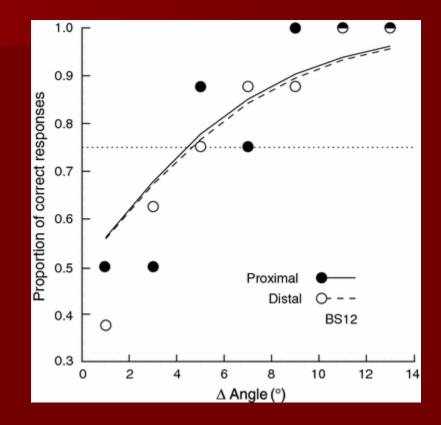
But more importantly, there was a correlation between degree of activation and localization performance

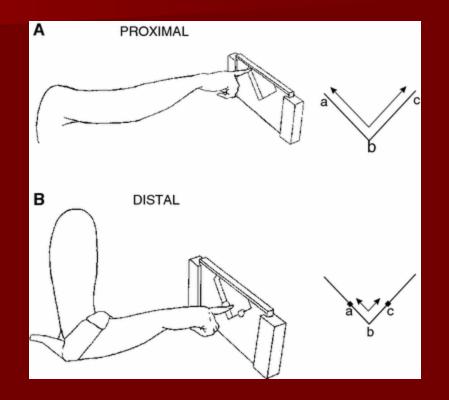


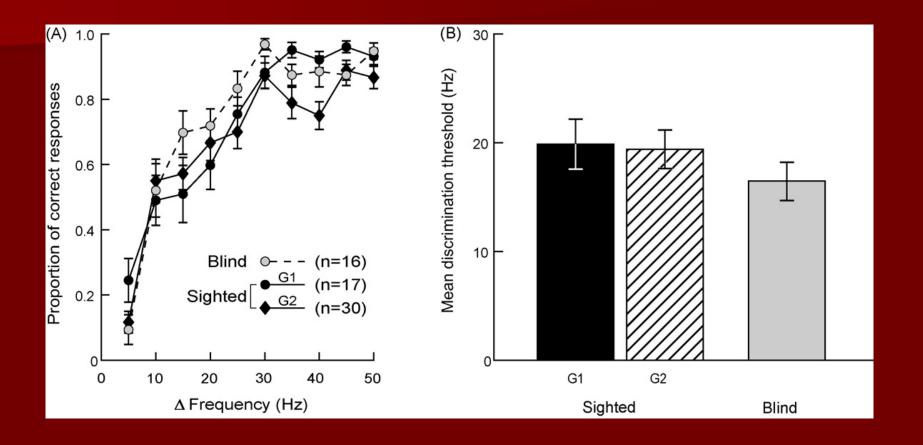
Gougoux, Zatorre, Lassonde, Voss, Lepore, (2005) PLoS Biology

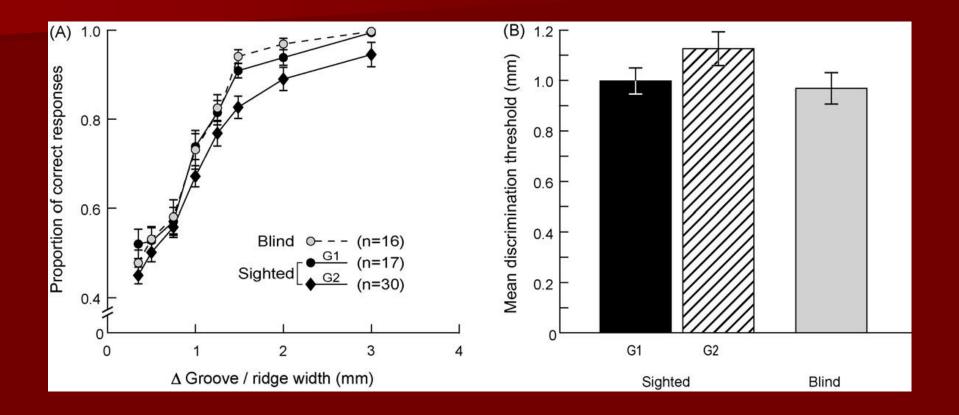
1800 1600 1400 1200 1000 800 600 400 200 0

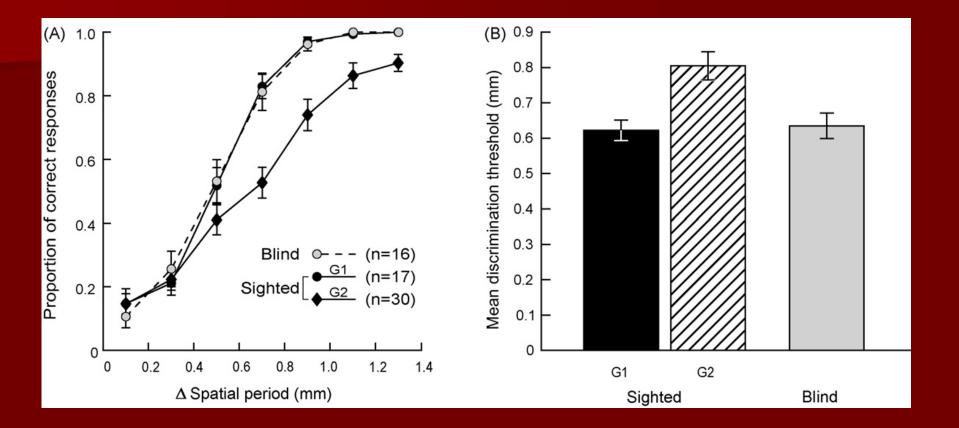
*
 Temps de réactionCongénitaux Tardifs
 Temps de réaction Congénitaux Tardifs Contrôles
 Congénitaux Tardifs Contrôles
 Détection
 Identification
 Localisation









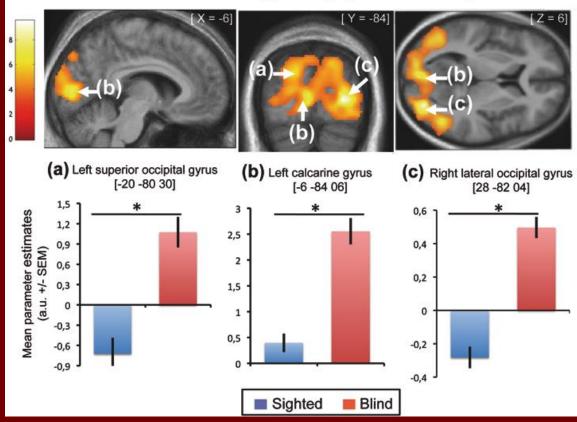


Introduction Behavioral reorganizations **Cerebral reorganizations** Discussion

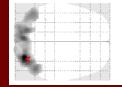


Blind > Sighted [Spatial+Pitch] :

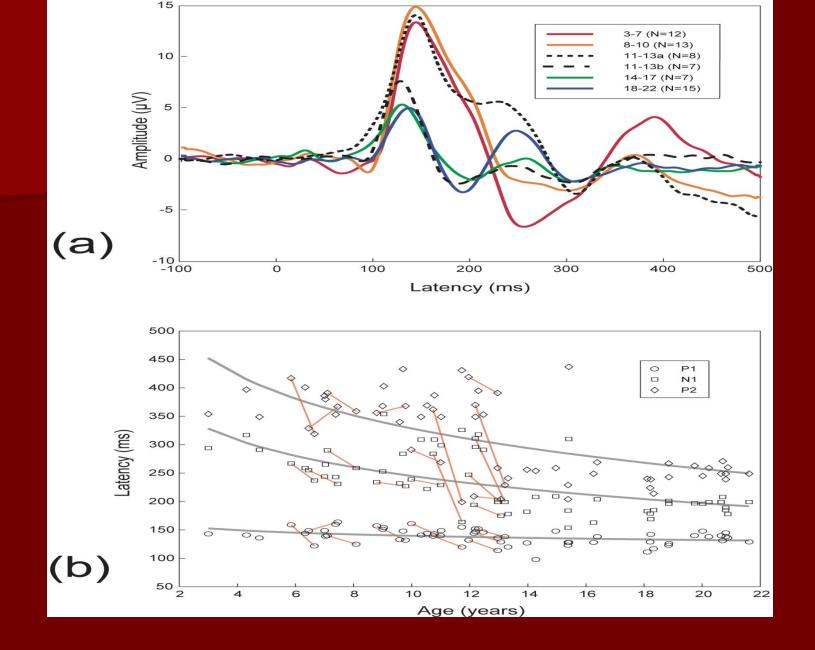
Blind > Sighted [Spatial+Pitch]







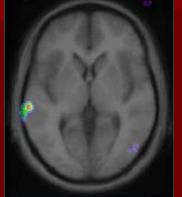
SPM{T₂₀}



Doucet, Lassond, Lepore et al 2005, NeuroReport

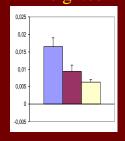
Vocal versus non-vocal Inter-group contrasts

congenital vs acquired



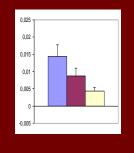


congenitally blind
 acquired blind
 sighted



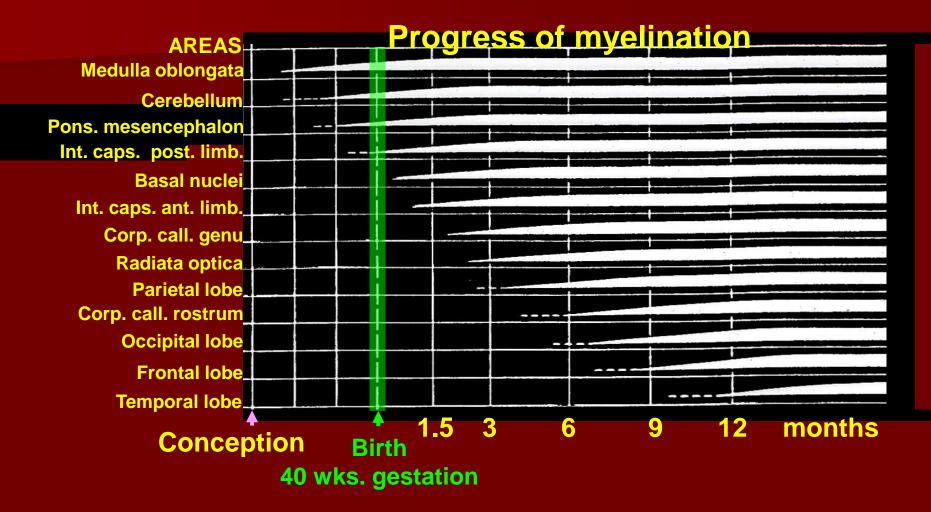
congenital vs sighted





Gougoux, Lassonde, Zatorre, Voss, Belin, Lepore (Neuropsychologia, 2009)

Myelinisation



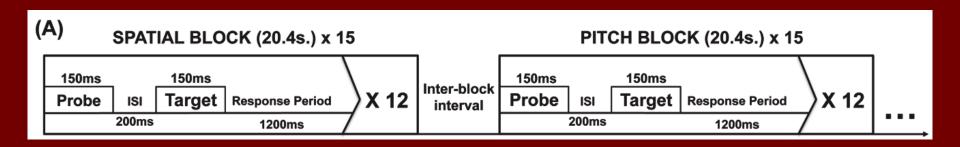


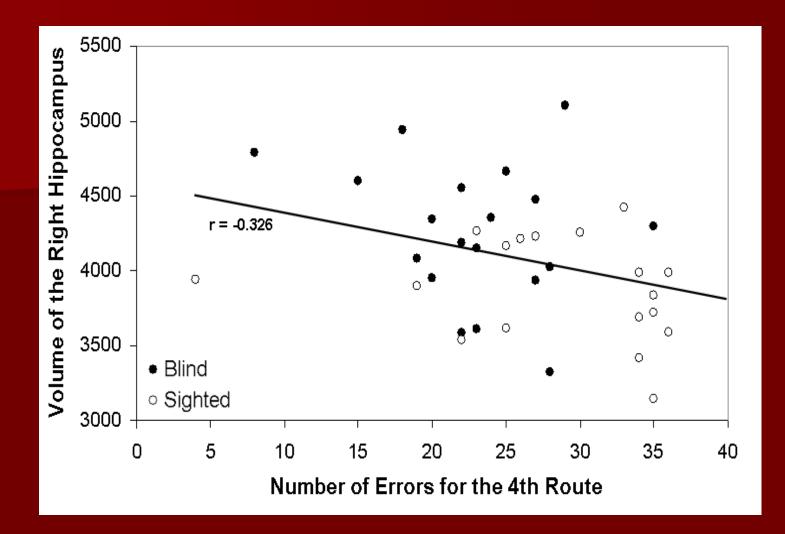


Sample :

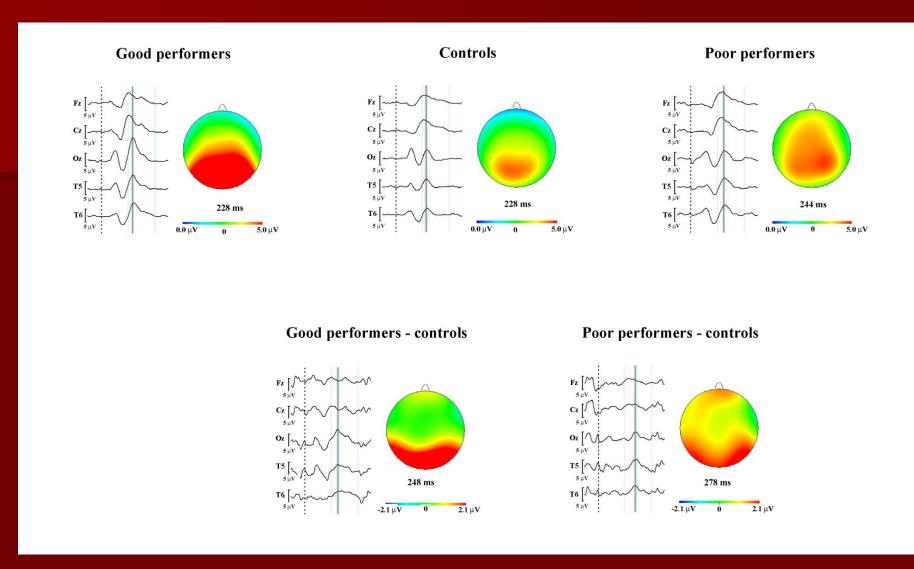
- 11 congenitally blind participants
 11 matched blindfolded sighted controls

functional Magnetic Resonance Imaging (fMRI) [3T Trio-TIM (Siemens)]



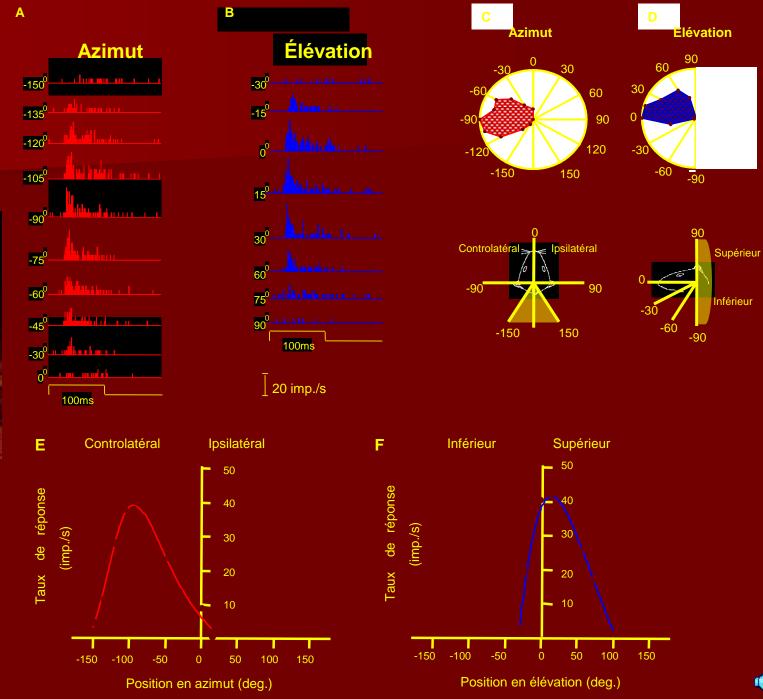


Fortin, Voss, Lassonde, Belin, Zatorre, Lepore, 2008, Brain



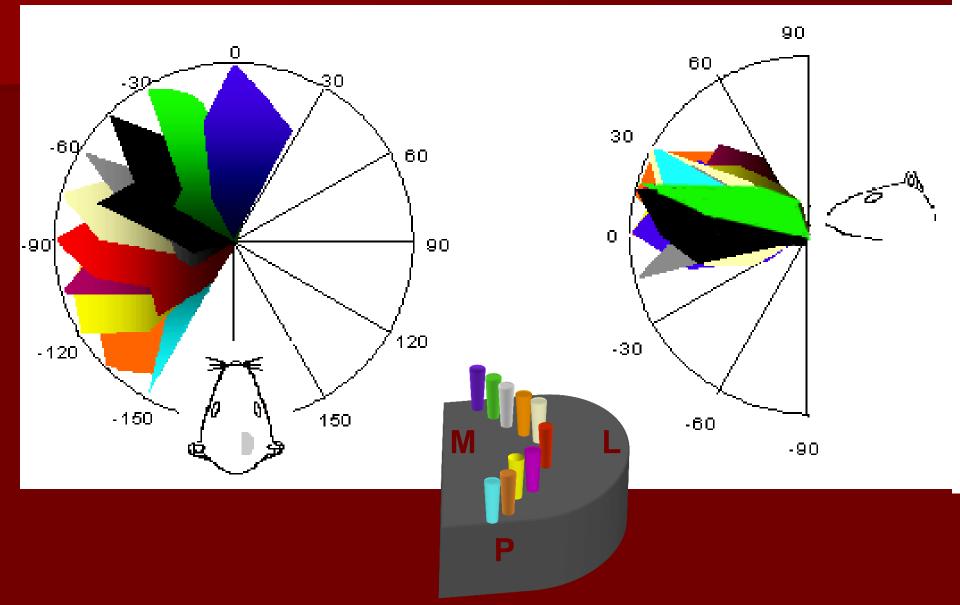
Visual evoked potentials to the presentation of the transformational apparent motion stimulus for good performing and badly performing subjects Doucet, Lassonde, Lepore et al, Brain, 2006

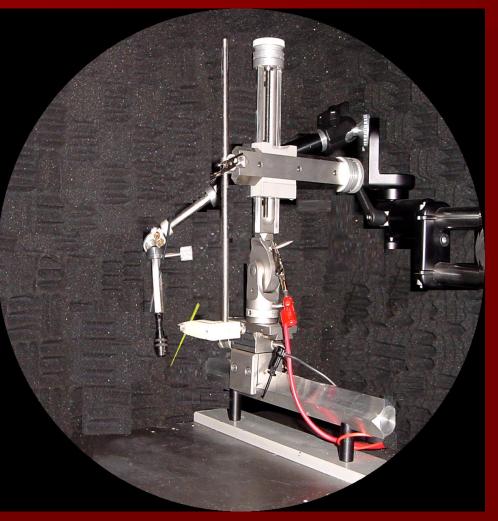


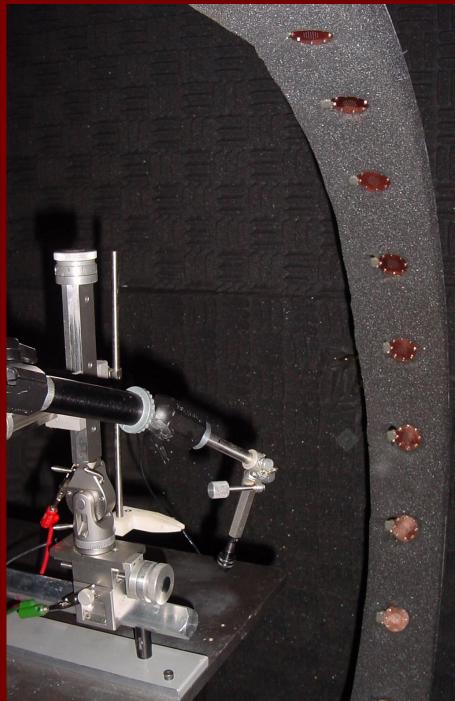


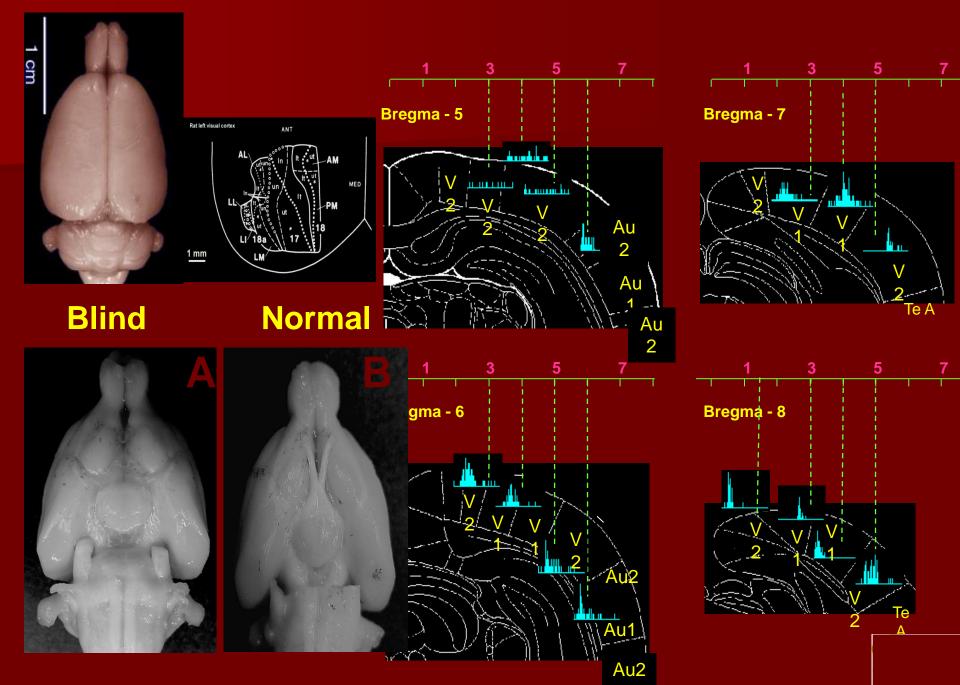
Azimut



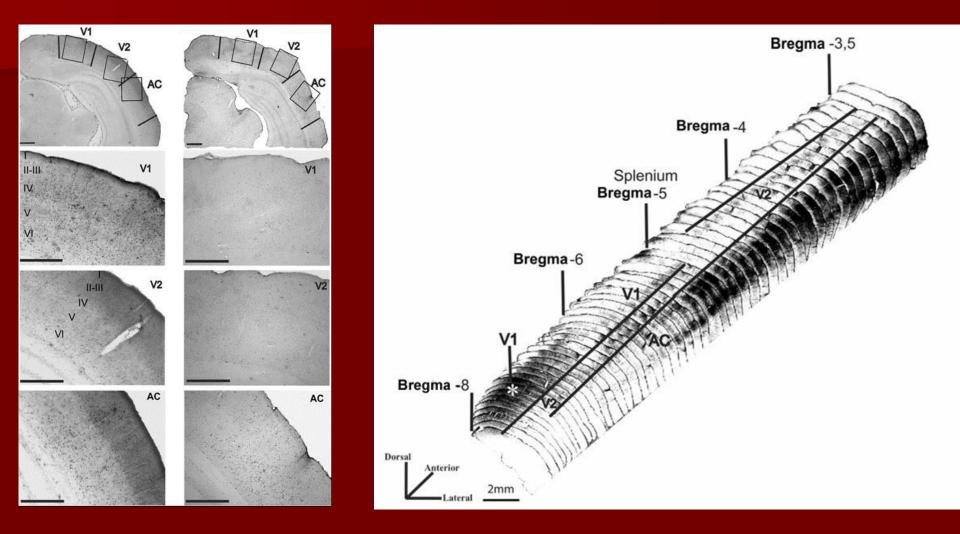


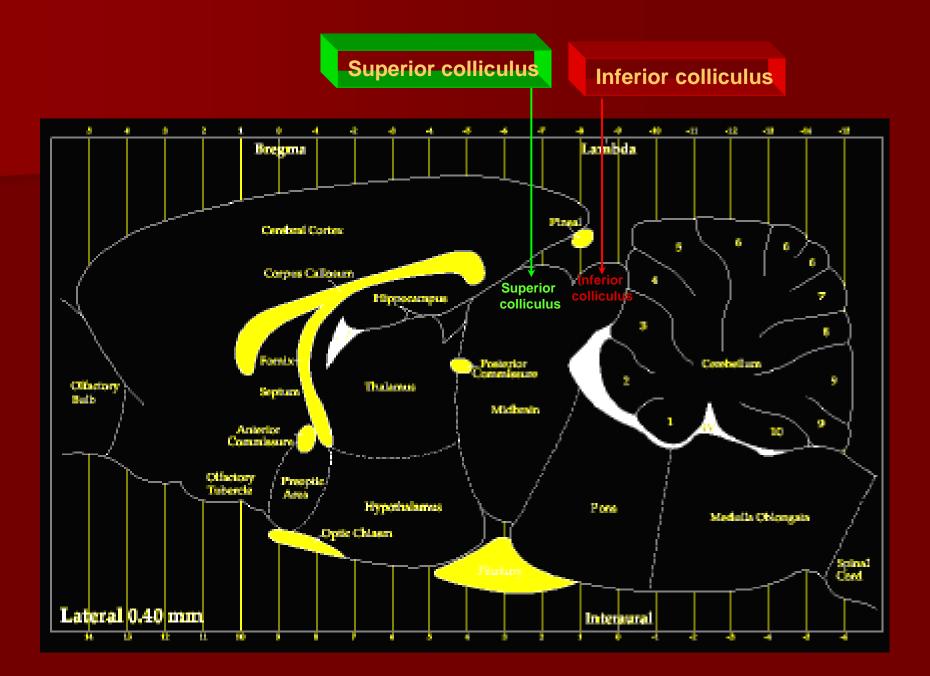




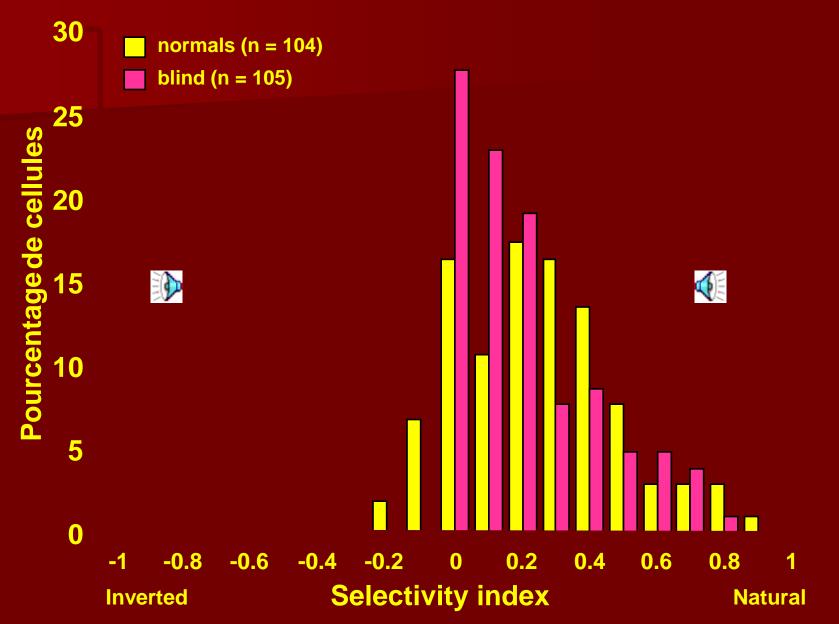


C-Fos Immunohistochemistry





Species specific vocalizations



The first question: is there cortical reorganisation in deaf individuals?

