

**U13 Bedside-to-Bench Conference Series  
Sensory Impairment and Cognitive Decline**

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**October 2 – 3, 2017**

**Bethesda North Marriott Hotel and Conference Center  
White Oak A & B**

**Animal models of hearing loss and brain changes**

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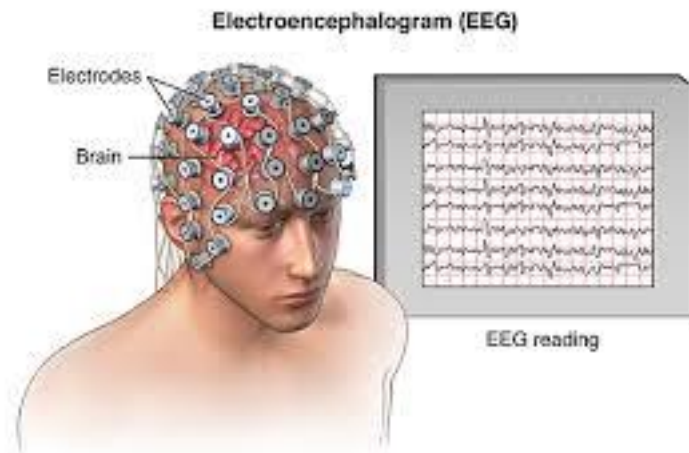
Disclosure:

Current funding: R21AG024372

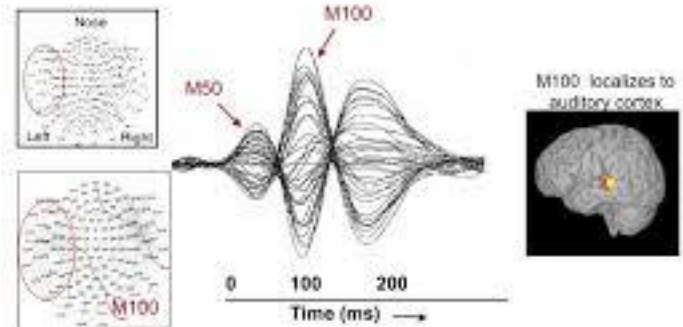
R01DC015232

Other financial relationships: None

Conflicts of Interest: None



### Neuromagnetic Auditory Evoked Field



A prototype auditory evoked neuromagnetic field detected by MEG. 37 channels with y-scale representing evoked response magnitude in units of femtoTesla (fT) are shown collapsed on the same horizontal time axis.

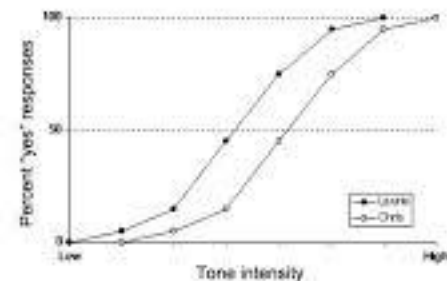
### A. Auditory task response relative to rest



### B. Visual task response relative to rest

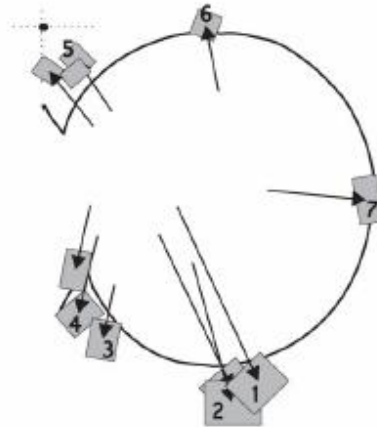
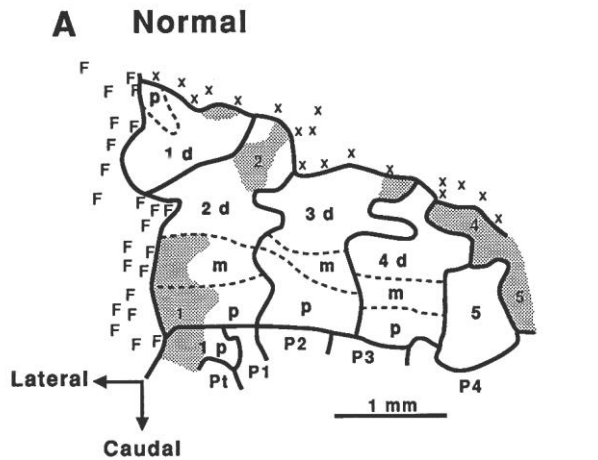


### Yes/no method of constant stimuli

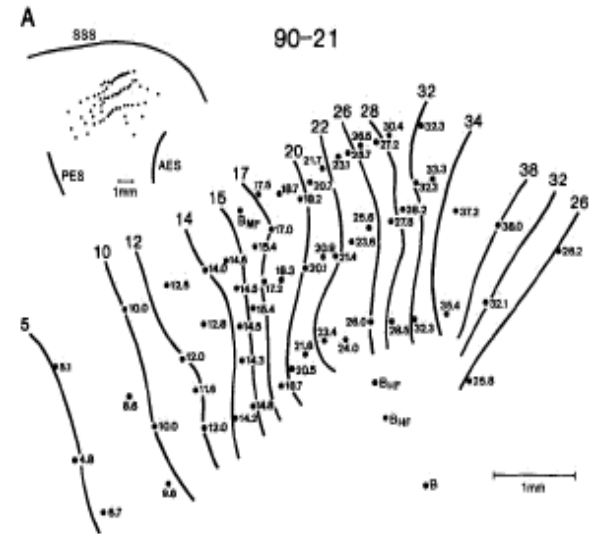


Do these data indicate that Laurie's threshold is lower than Chris's threshold?

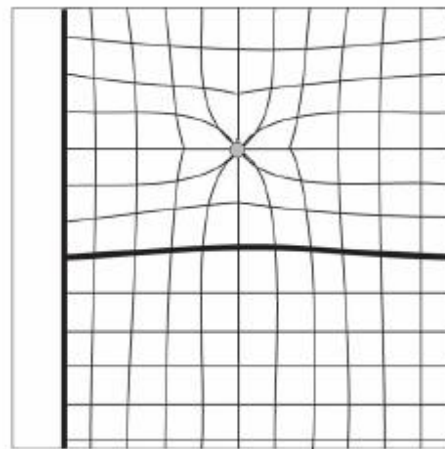
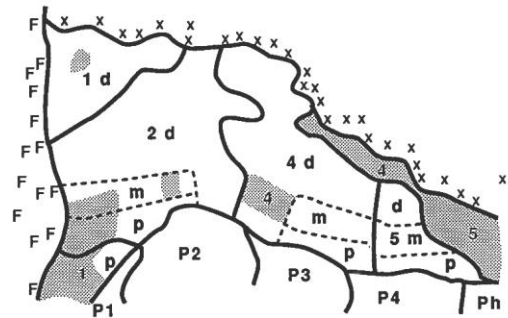
# Sensory Deprivation in Adults leads to Cortical Map Changes



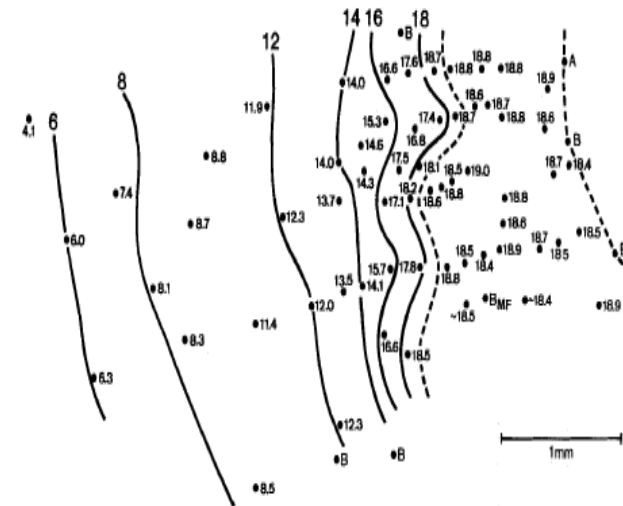
26 weeks post-lesion



**B** 62 days after digit 3 amputation



Cortex -- 2 months



Merzenich et al. (1984) *J. Comp. Neurol.* 224:591-605

Gilbert and Li (2012) *Neuron* 75:250-264

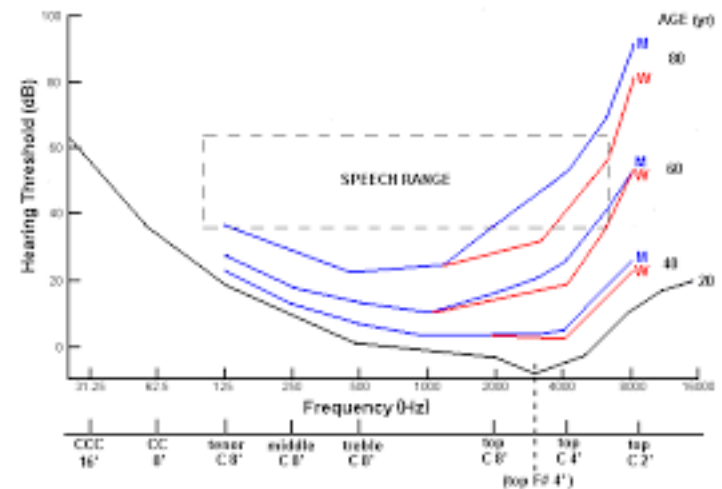

Rajan et al. (1993) *J Comp Neurol* 338:17-49

Normal aging often leads to sensory deficits from the periphery

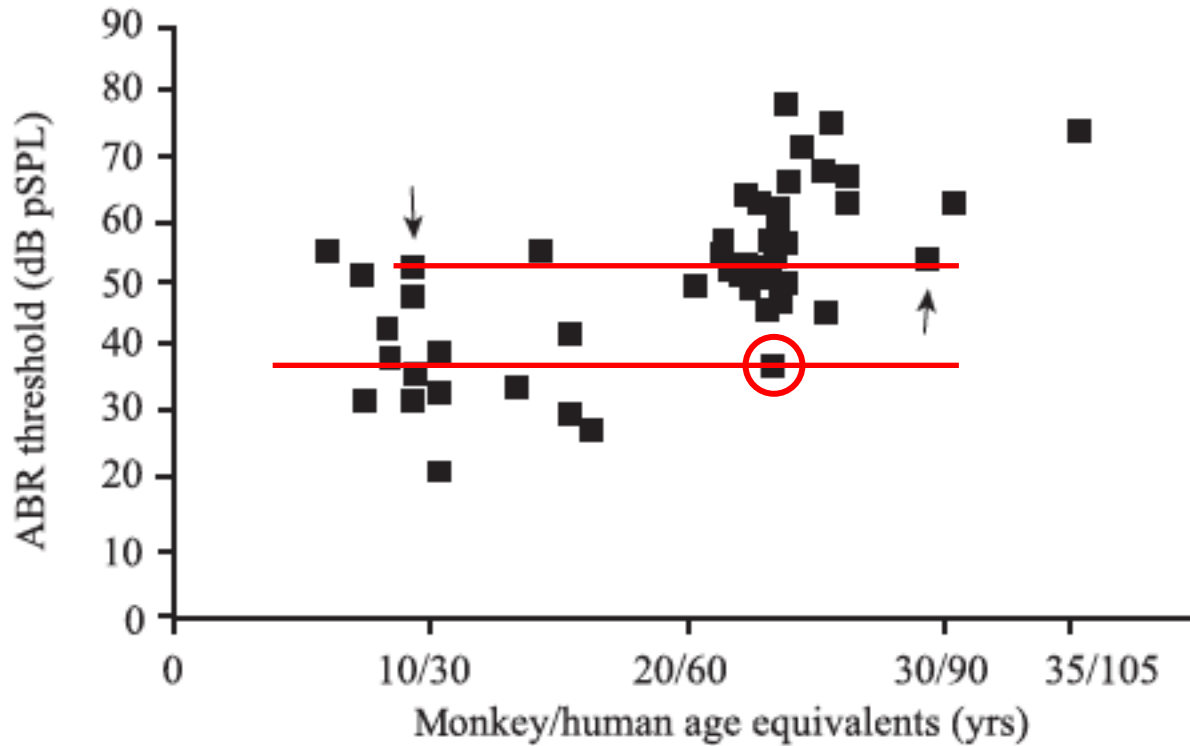


Typical Peripheral Neuropathy Symptoms

- ✓ Loss of Feeling
- ✓ Freezing
- ✓ Tingling
- ✓ Hyper Sensitivity
- ✓ Sharp Jabbing Pain
- ✓ Burning Sensation
- ✓ Numbness

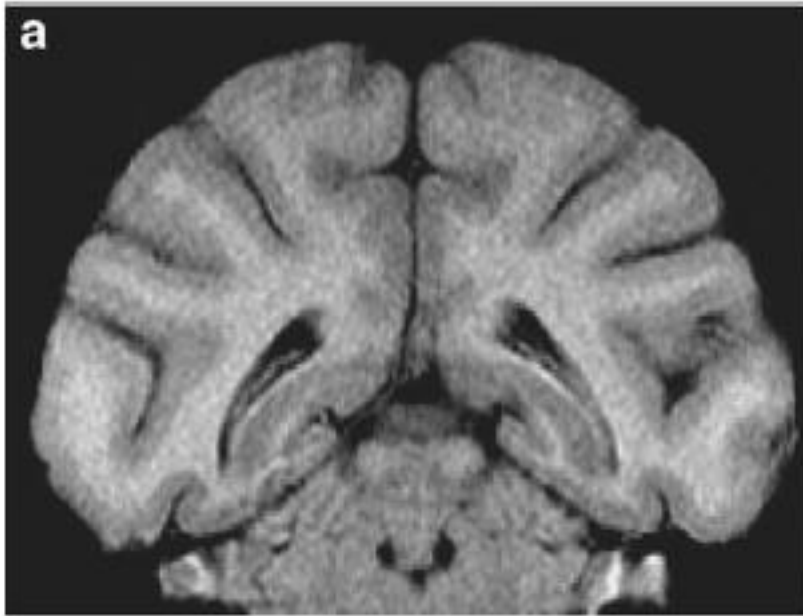


Normal aging often leads to sensory deficits from the periphery  
BUT NOT ALWAYS!

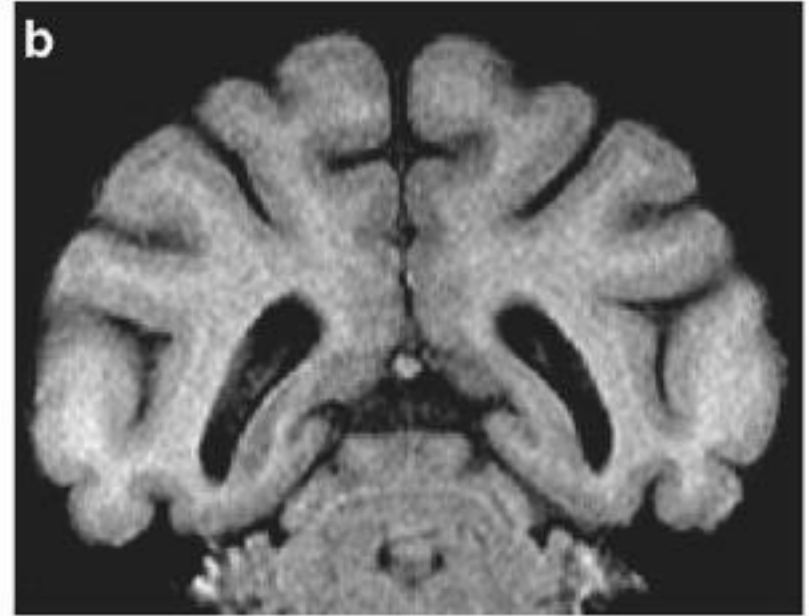


Normal aging does lead to changes in the central nervous system even in the absence of peripheral deficits

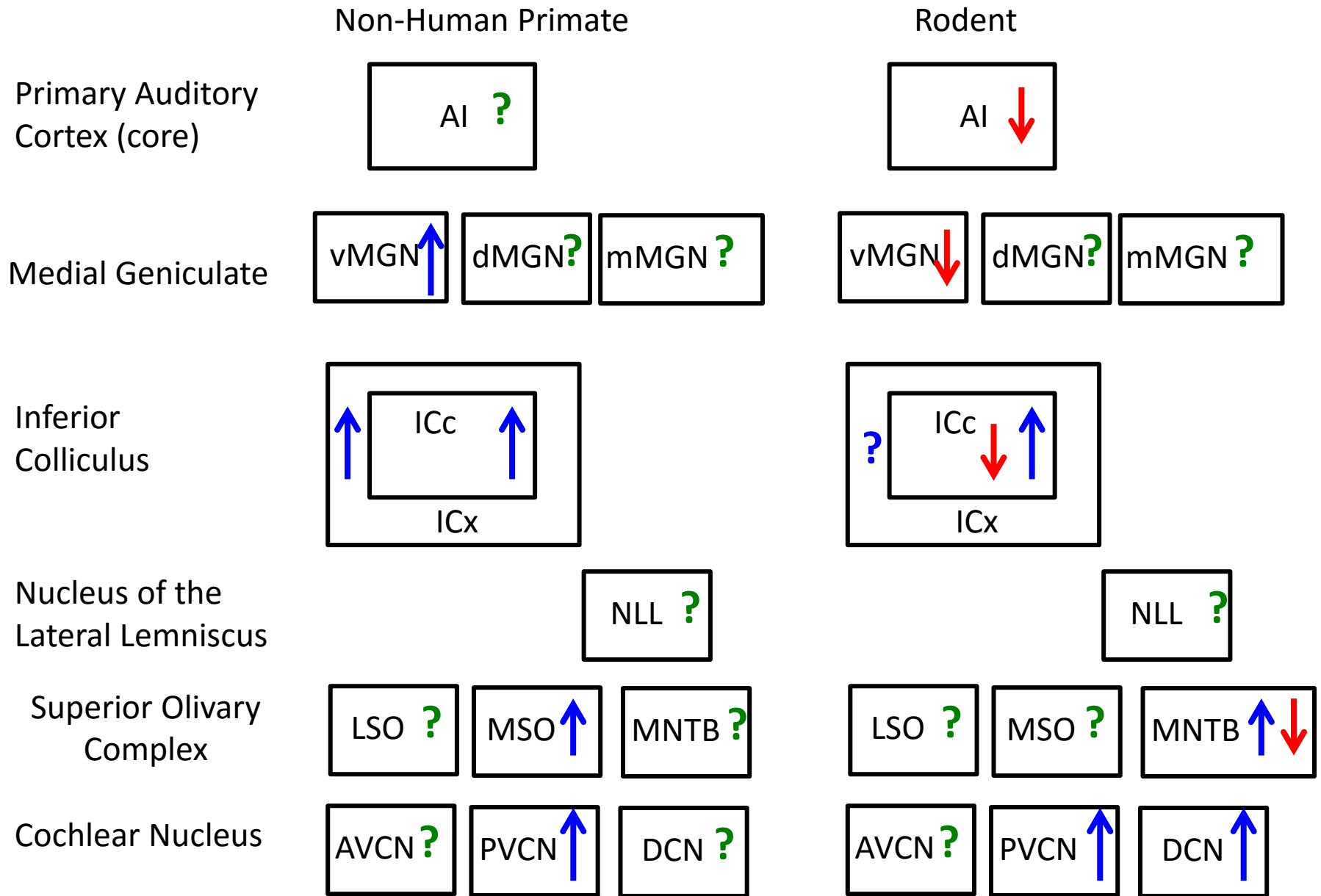
6 year old (~18 human)



24 year old (~72 human)



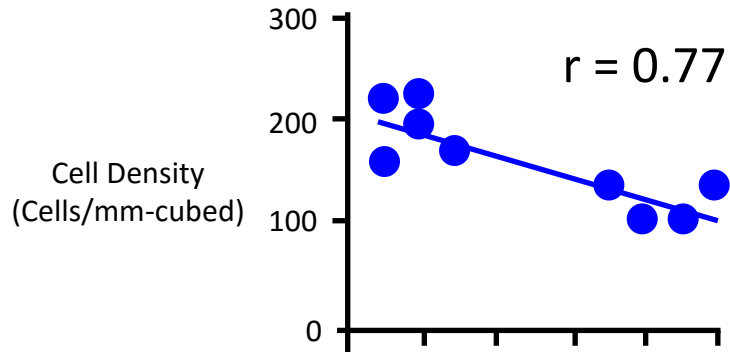
# Neurochemical changes occur throughout the ascending auditory system with age





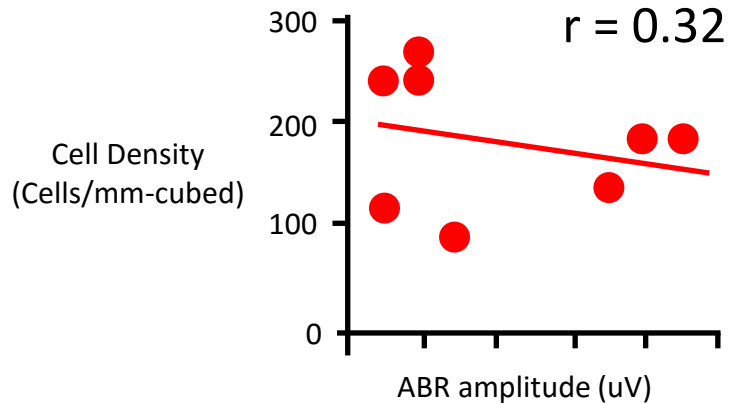
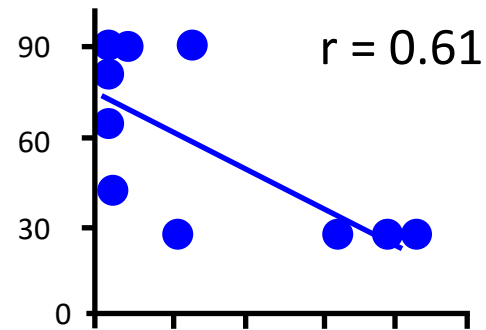
# Neurochemical changes are not always tightly correlated with ABR thresholds

## Cochlear Nucleus

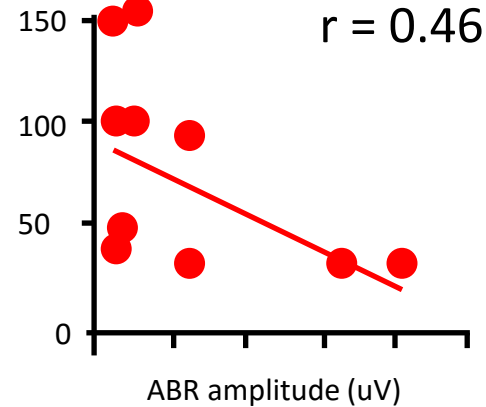


## Superior Olivary Nucleus

NADPHd



Parvalbumin



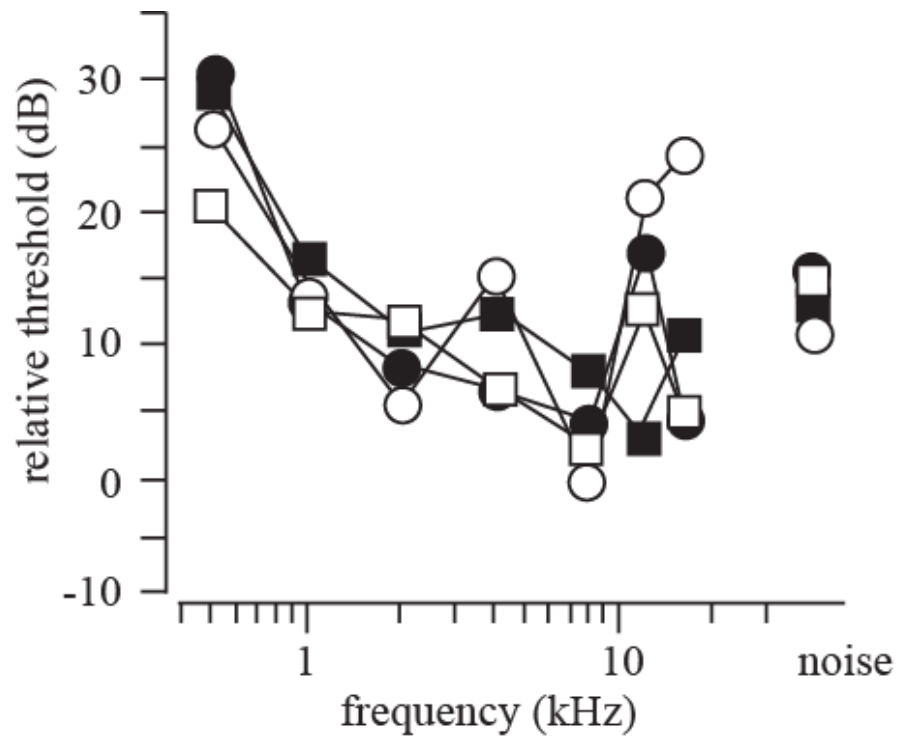
There are varying degrees of sensory loss between individuals with aging

The aging brain shows both morphological and neurochemical changes at least partially independently of sensory loss

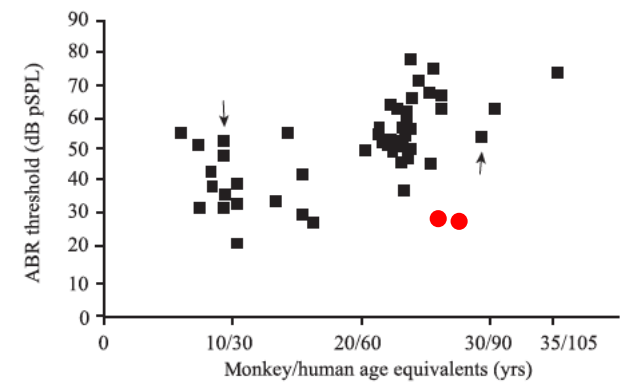
Both sensory loss and aging need to be taken into account when studying central changes

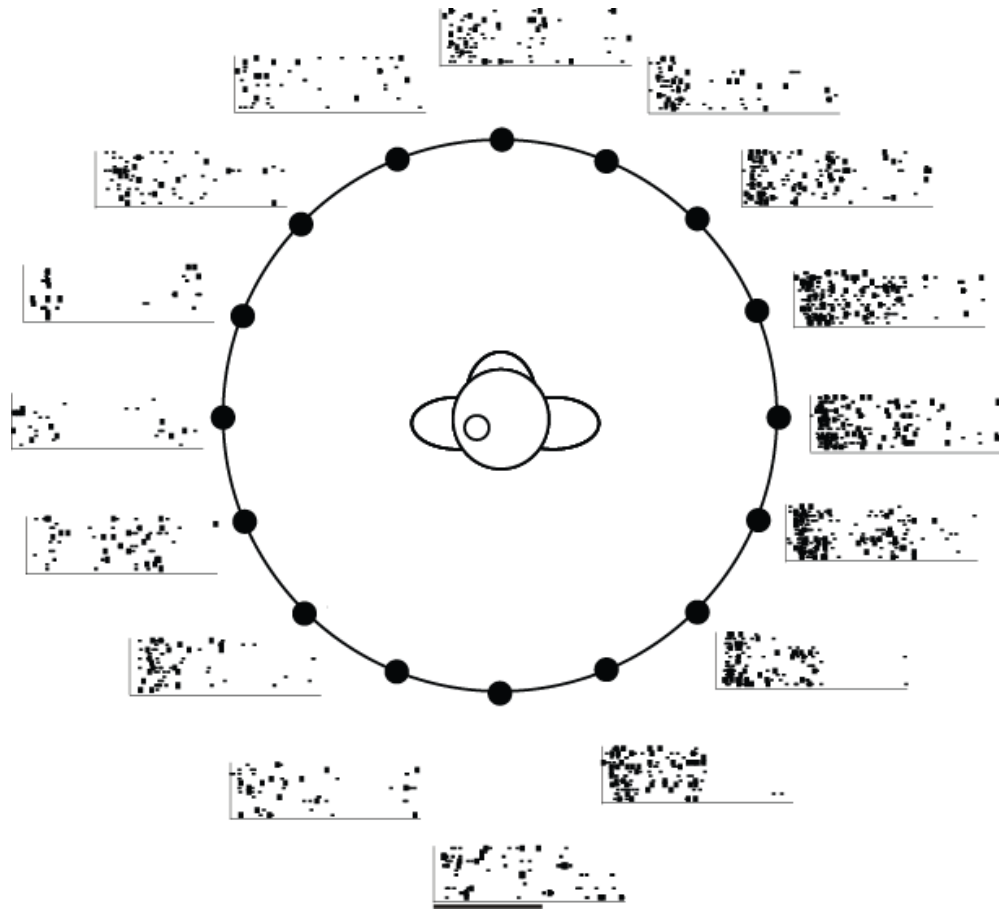
***How much does the brain change just with aging?***

# Comparisons Between Young and Geriatric Monkeys

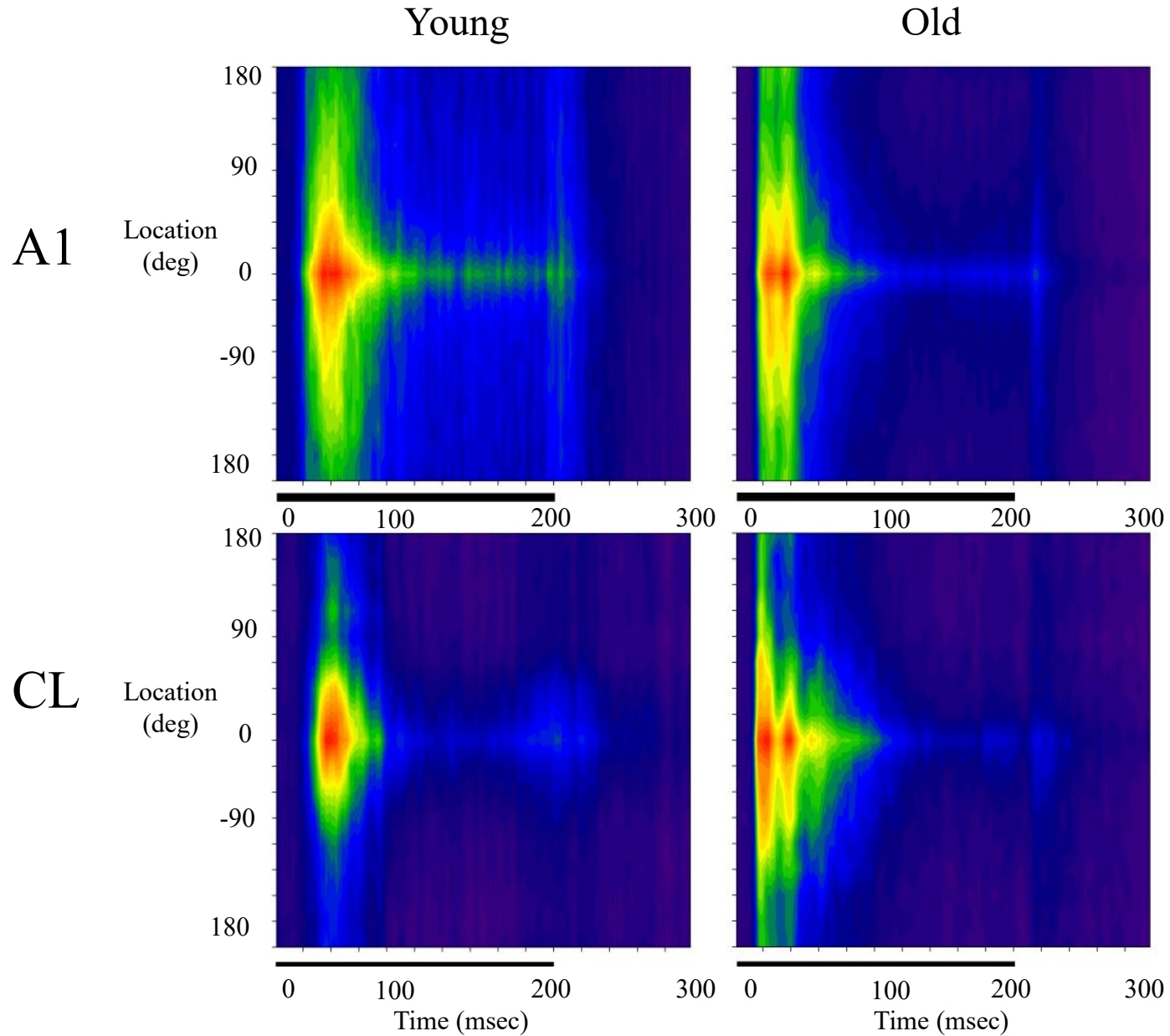


- monkey A: 26 years (77 human)
- monkey B: 27 years (79 human)
- monkey H: 10 years (31 human)
- monkey N: 14 years (43 human)

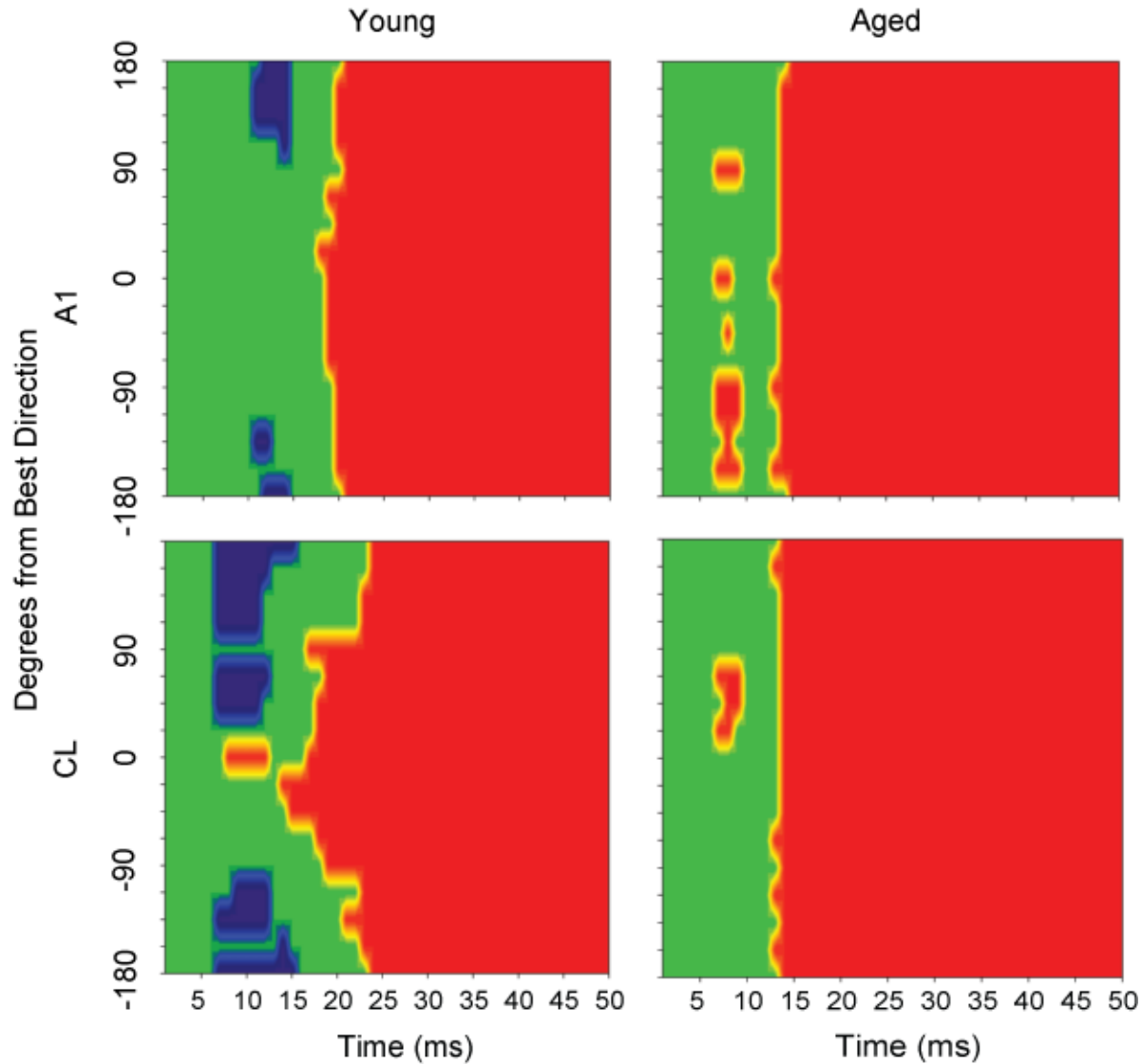




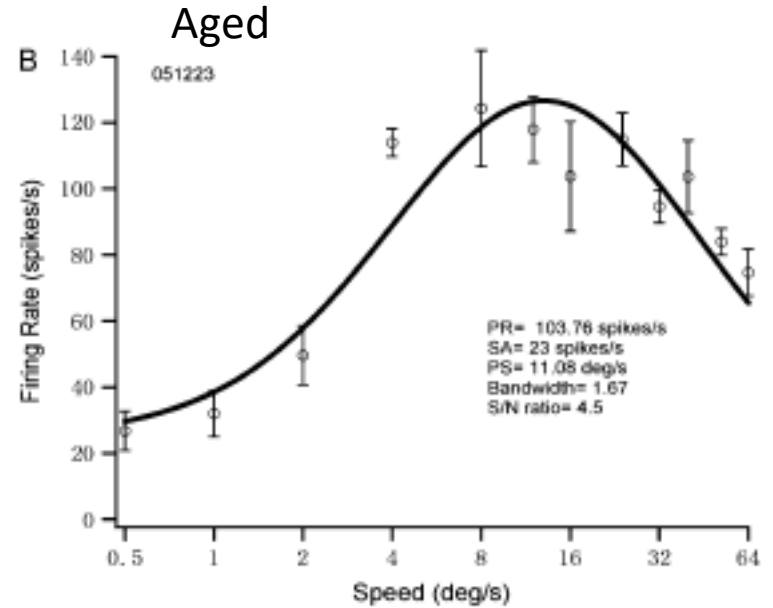
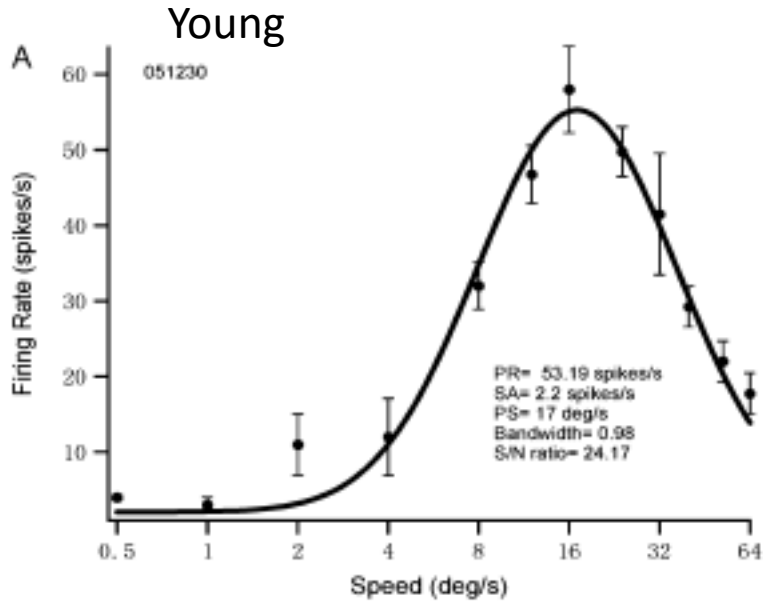
# Spatial tuning of the population of neurons



There is also less spatial inhibition in aged vs. young cortical neurons



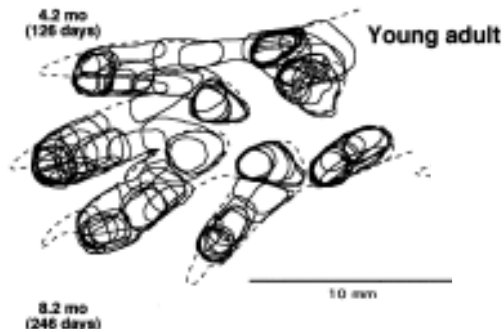
# Speed tuning in MT is broader in aged monkeys



Yang et al (2009) *Cereb Cortex* 19:1957-1967

# Receptive fields in S1 are larger in aged rats

Young



Aged



Coz and Xerri (2001) *Neuroscience* 104:705-715

- Caveats: 1) why are neurons in some animals older than others?  
2) Are the sensory/cognitive functions similar and/or subserved by the same neural machinery?



2 years old  
mouse = 90+ years



3 years old  
Rat = 90+ years



30ish years old  
Monkey = 90+  
years





Studying perception in aged animals or people needs to take into account that the brain is aging with or without sensory loss

Animal studies using invasive techniques is a compliment to non-invasive human techniques and vice versa. Both are necessary to understand the aging brain

Gaps:

- 1) How much is sensory loss and how much is aging?
- 2) How can we ameliorate one, the other, or (preferably) both?

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