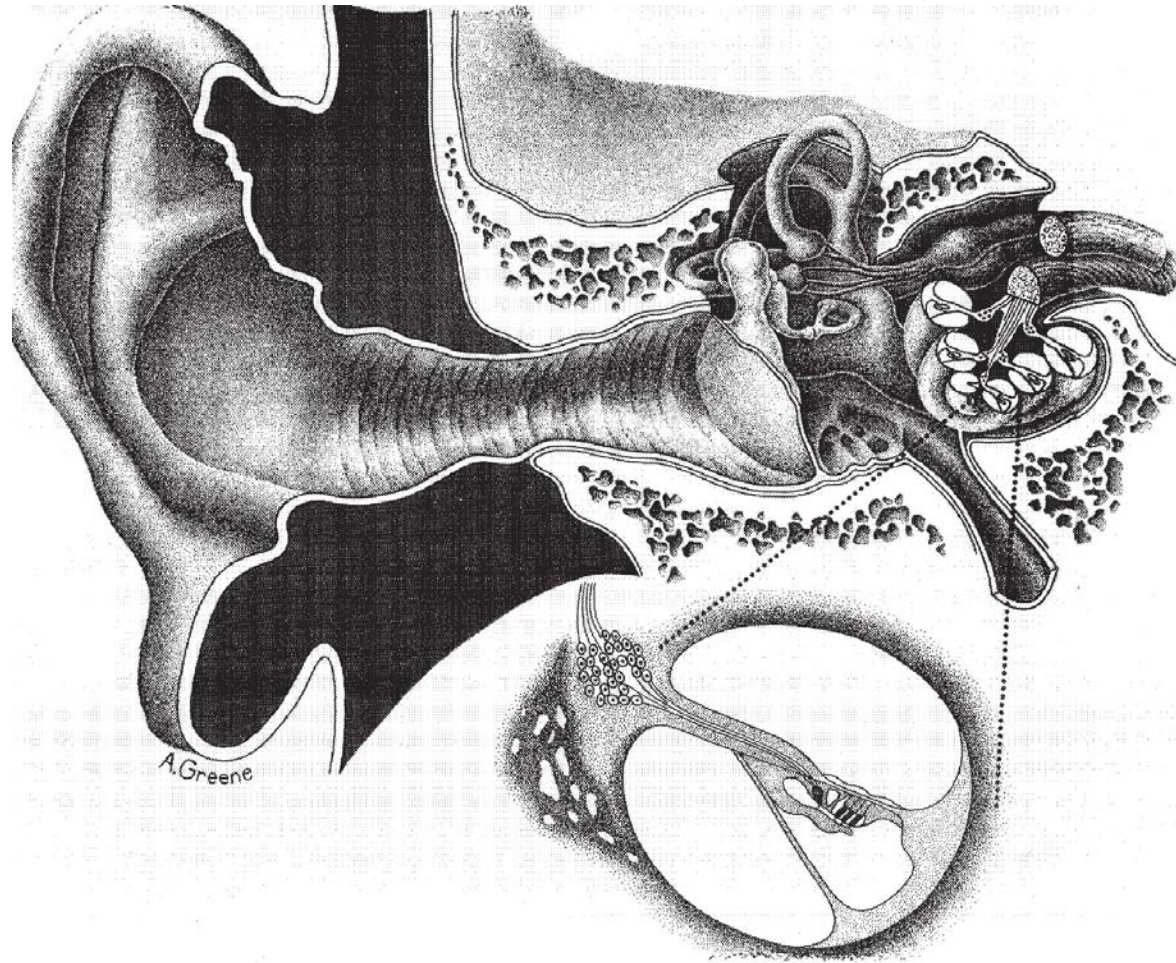
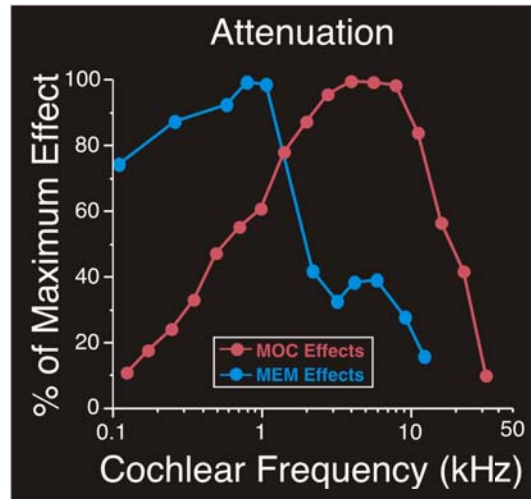
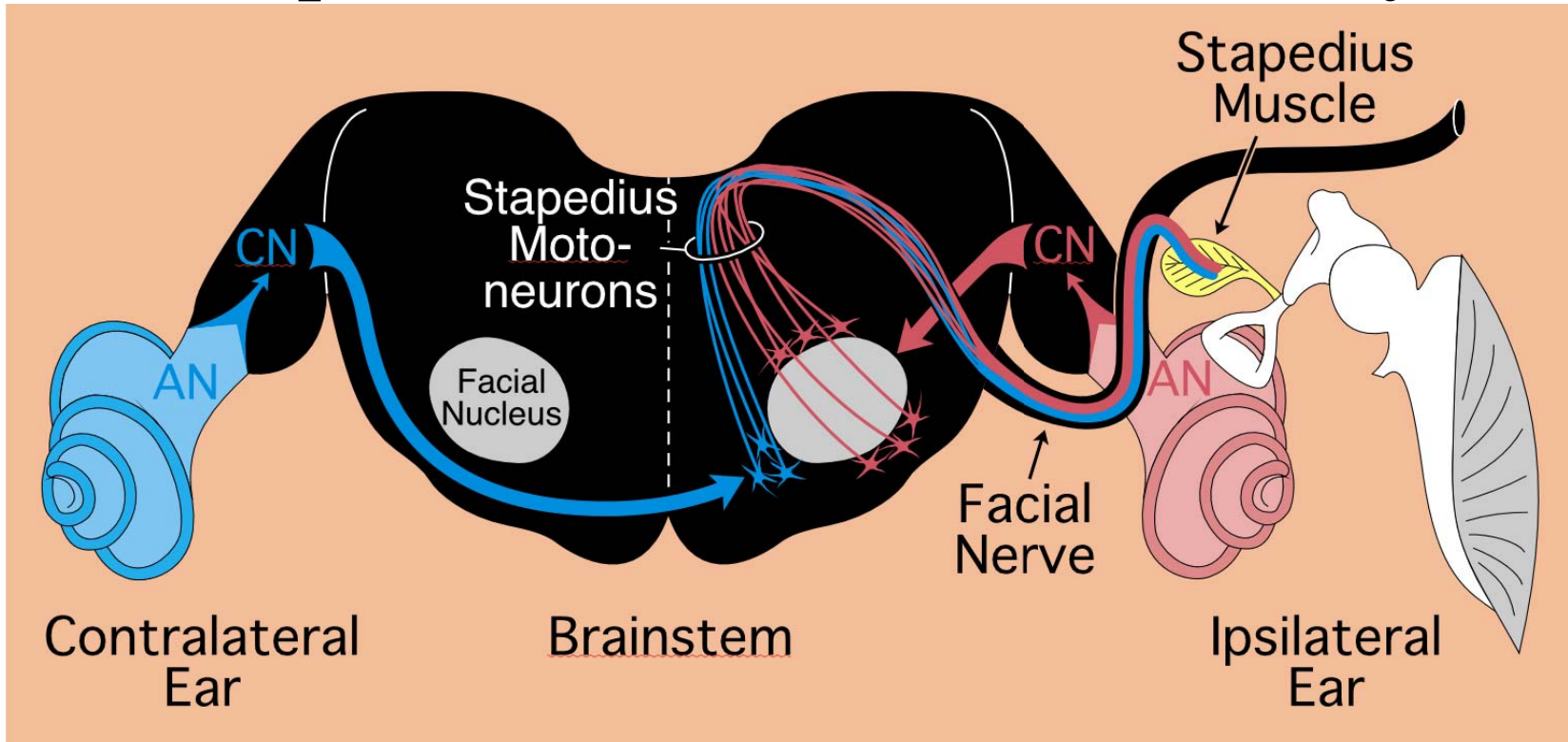


HST 721 Efferent Control Lecture

October 2004



Stapedius Muscle Central Circuitry



Hypotheses for MEM Function

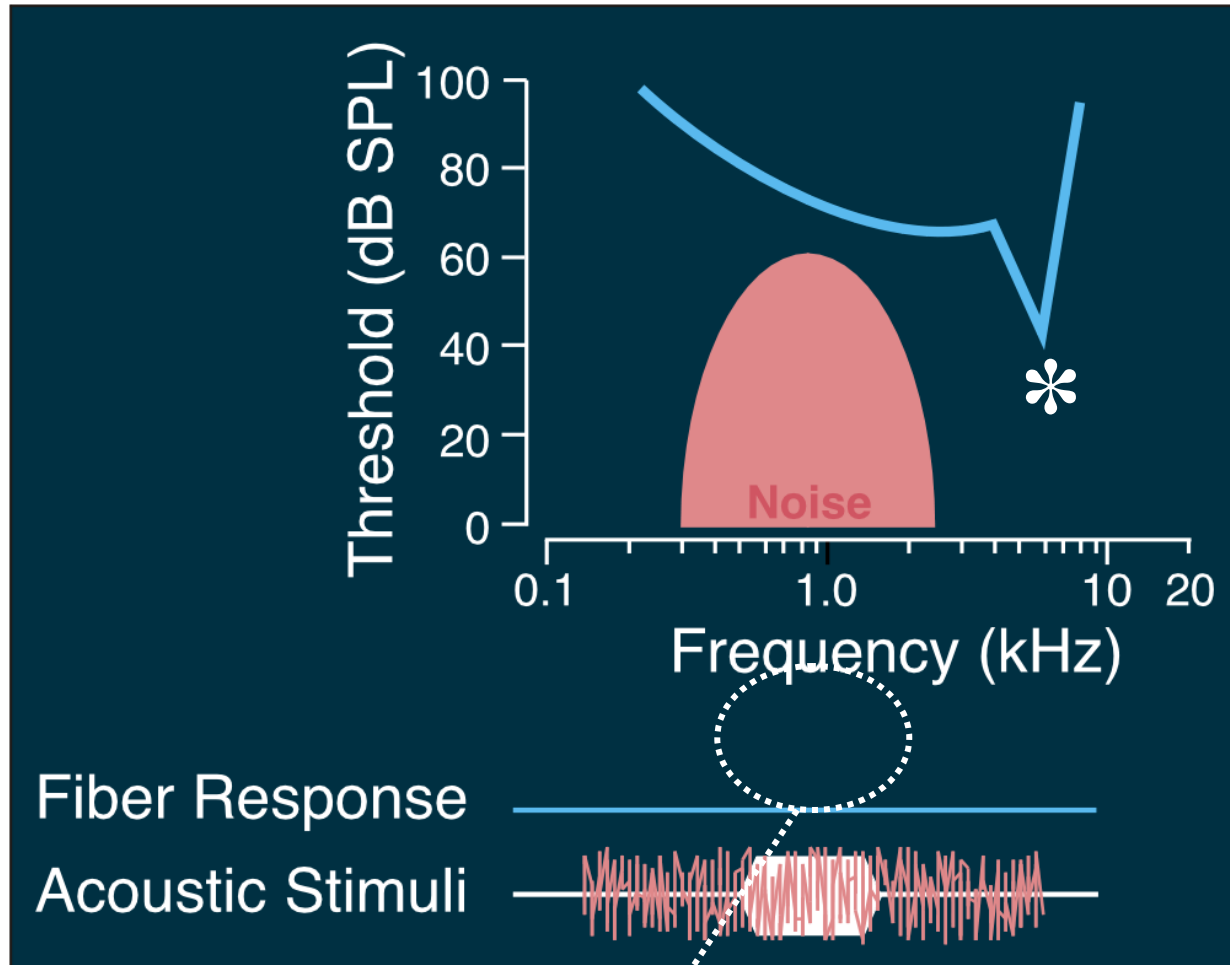
A. Stapedius

- 1. Extend Dynamic Range - a gain control system**
- 2. Protect the Inner Ear from Acoustic Injury**
- 3. Control Masking from Continuous Background Noise**

B. Tensor Tympani

- 1. Aid in Middle-Ear Aeration**

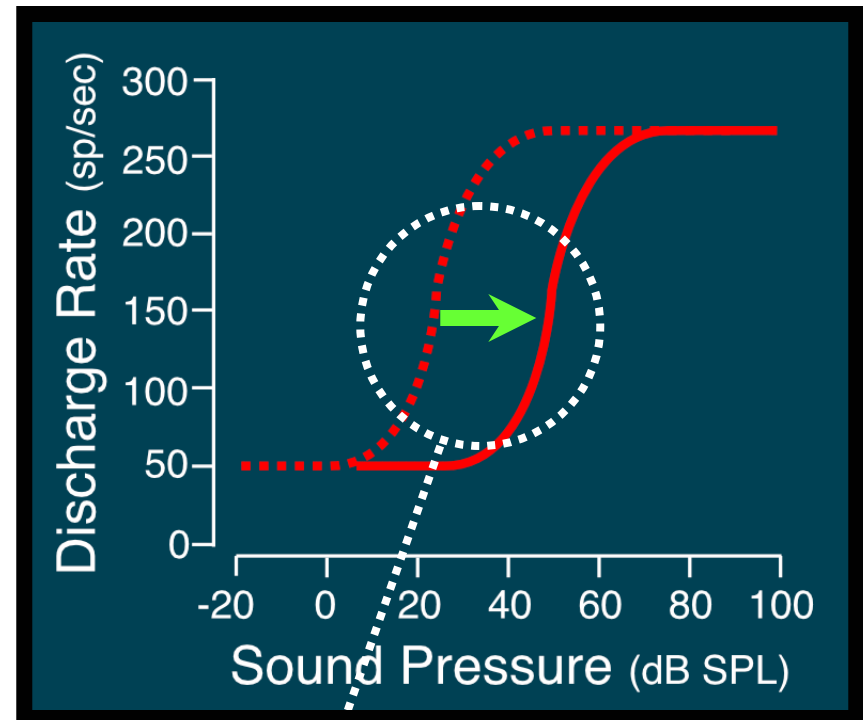
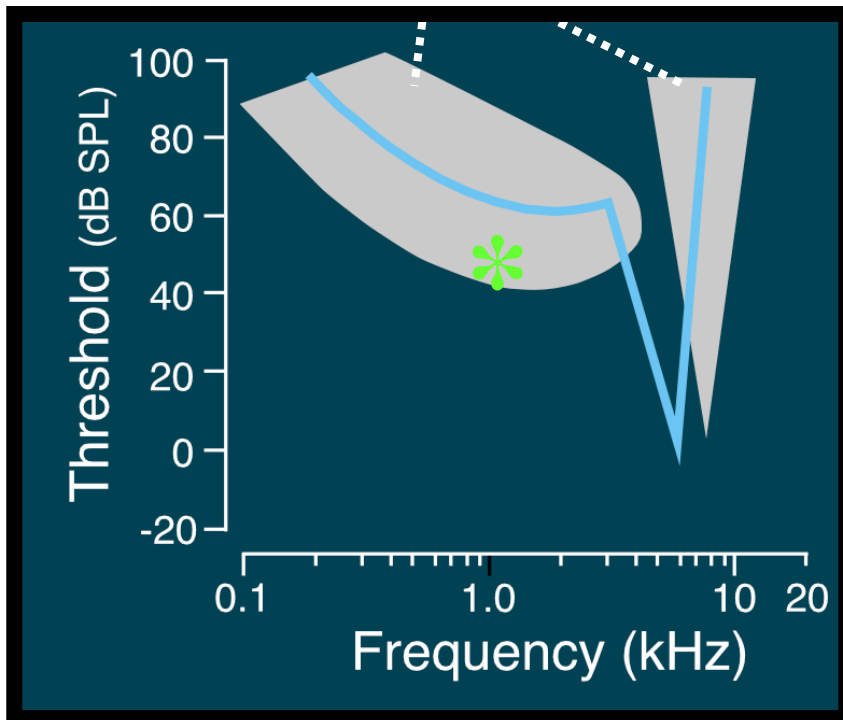
Suppressive Masking



Response to transient suppressed; yet masker elicits no response

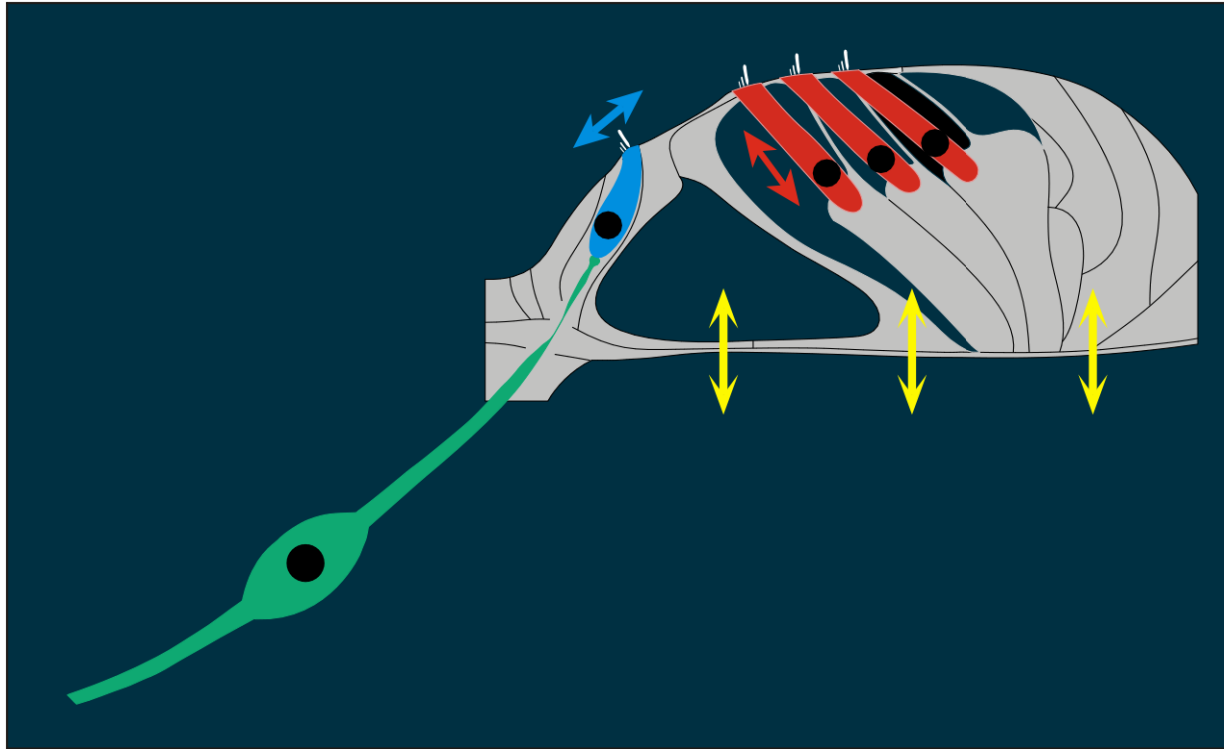
Suppressive masking ~ Two-Tone suppression

Two-Tone Suppression Contours



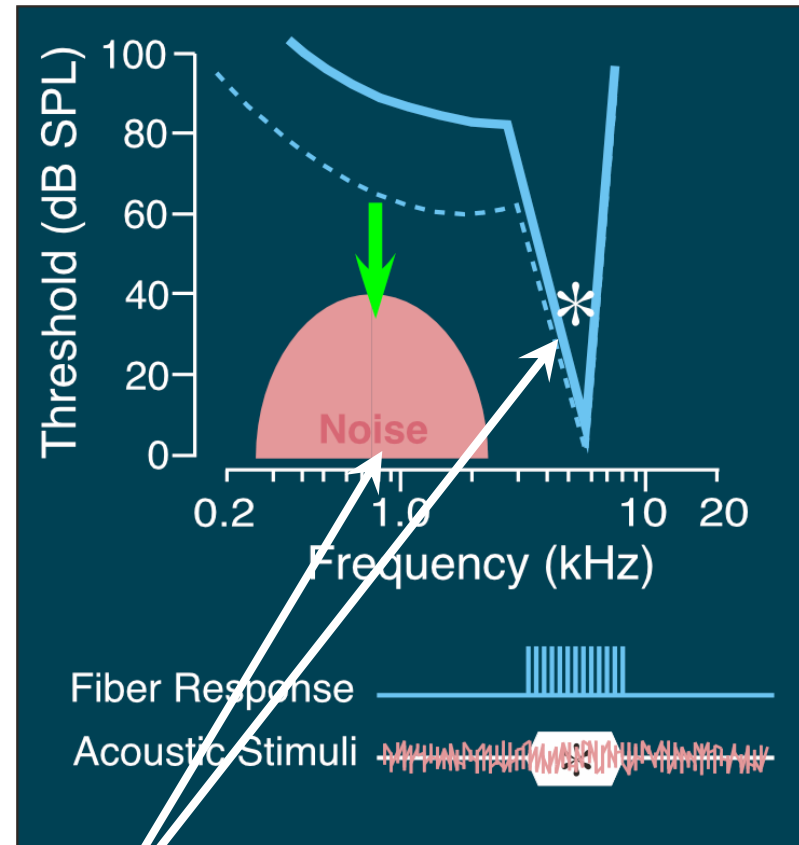
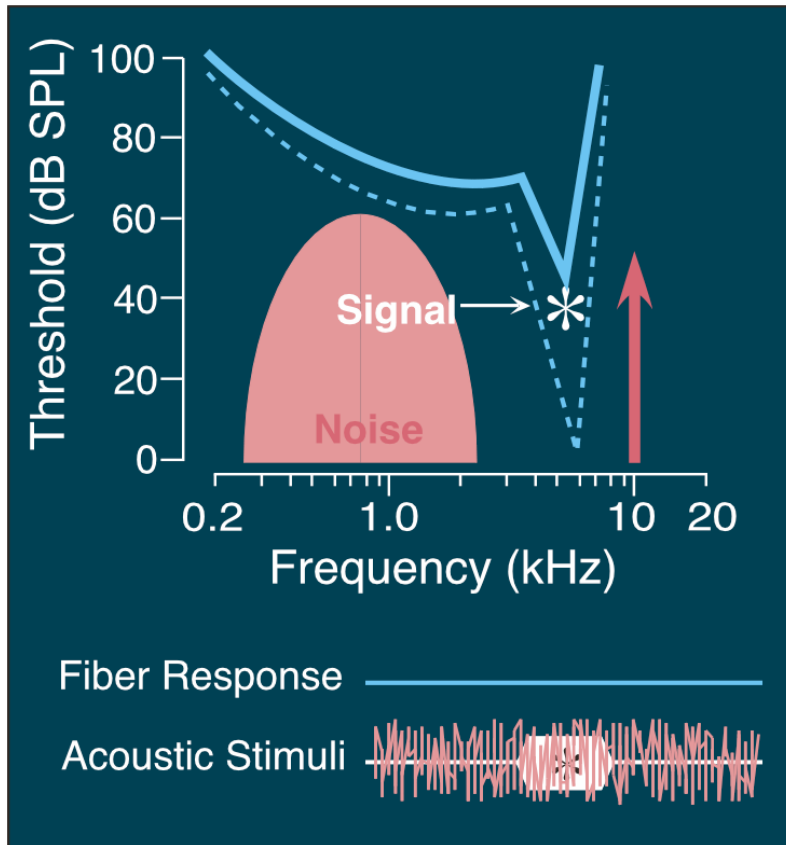
Dynamic Range Shift:
3 dB/dB

Two-Tone Suppression



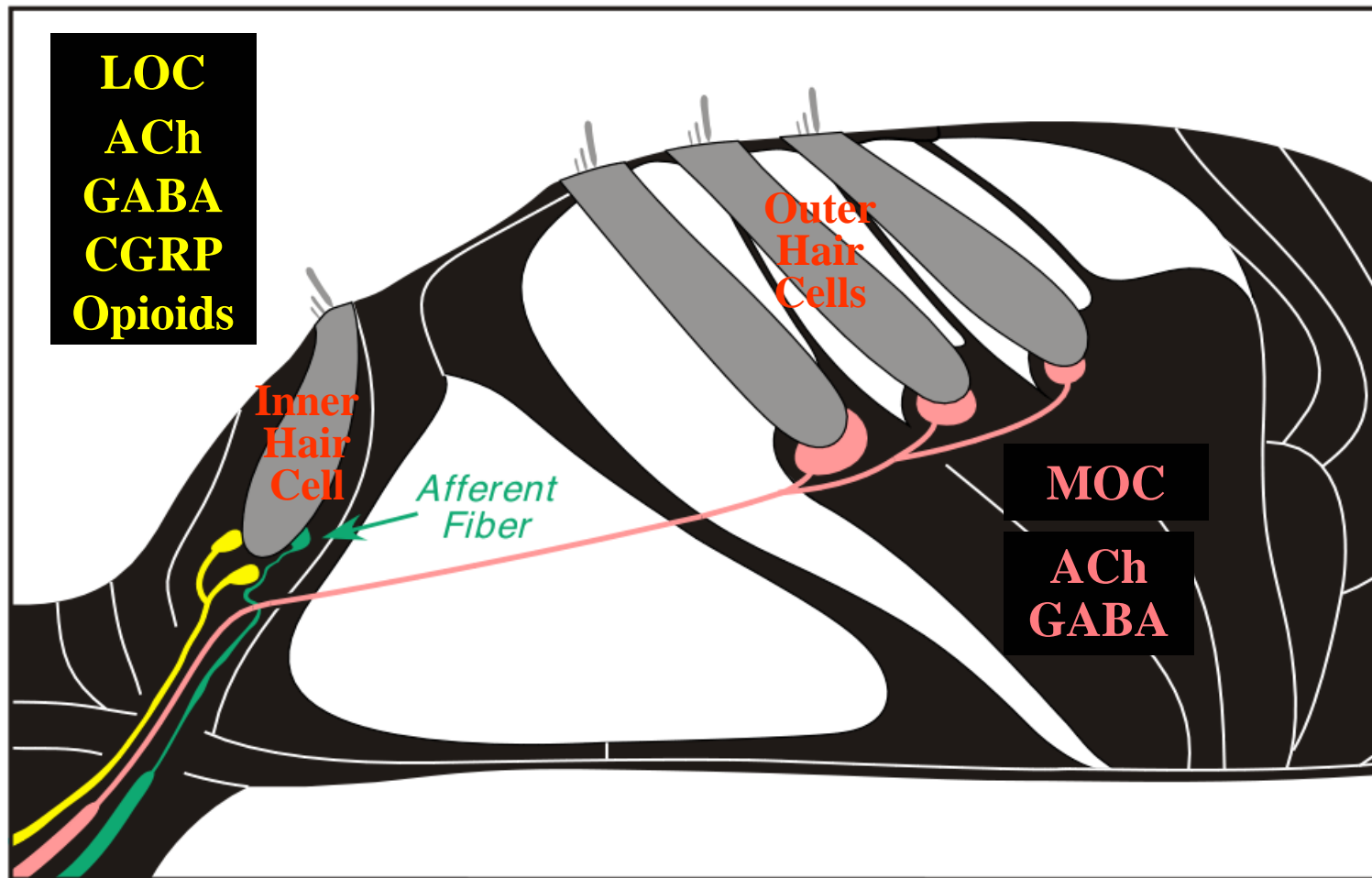
- Measured in basilar membrane motion
- Arises from OHC-based non-linearity
- Requires simultaneous stimuli

MEMs counteract suppressive masking



MEM reflex (Stapedius) useful for high-frequency signals embedded in low frequency noise

Olivocochlear Peripheral Circuitry



Unmyelinated

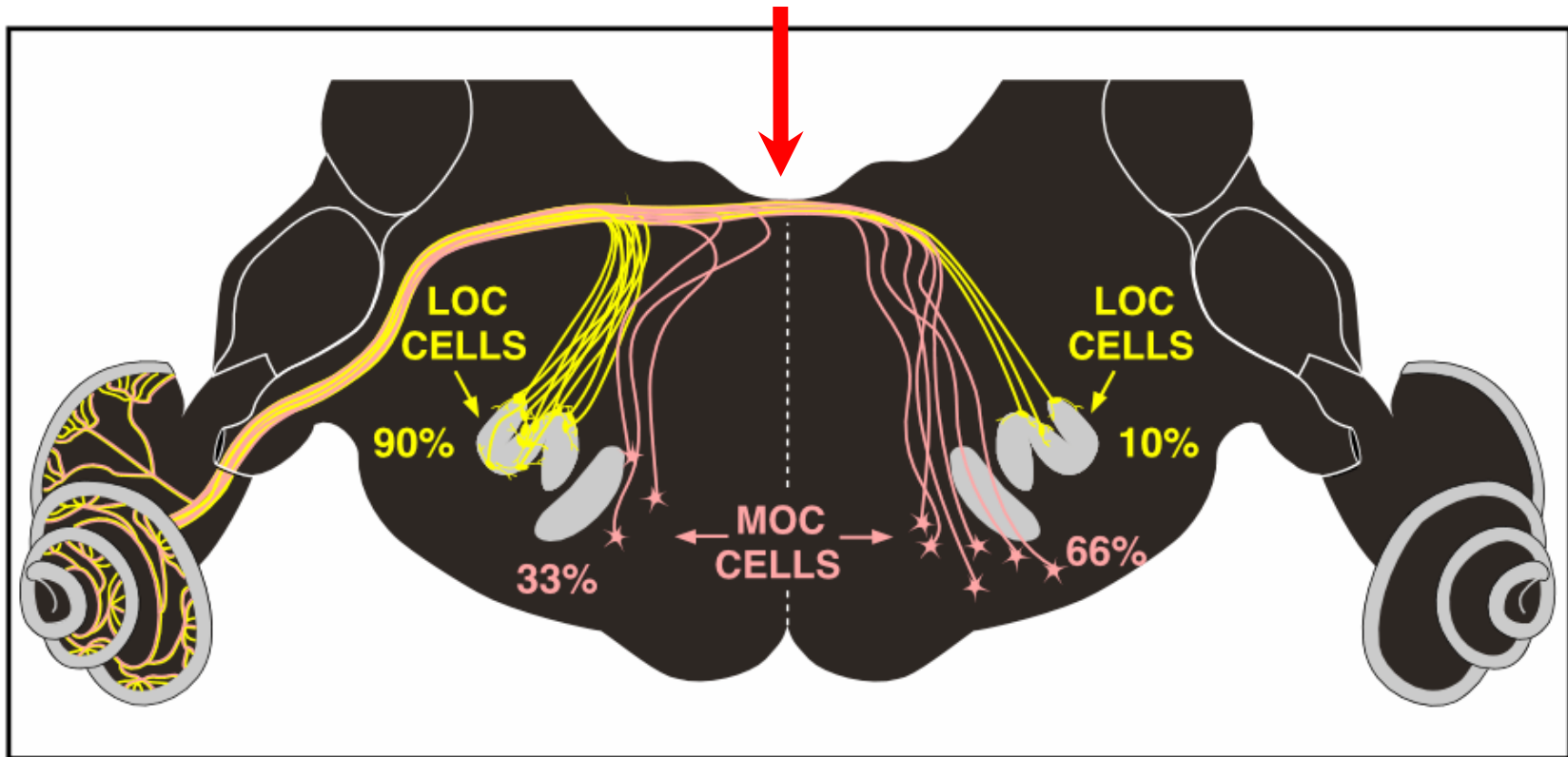
Myelinated

Afferent Excitation/Suppression

Cochlear Suppression

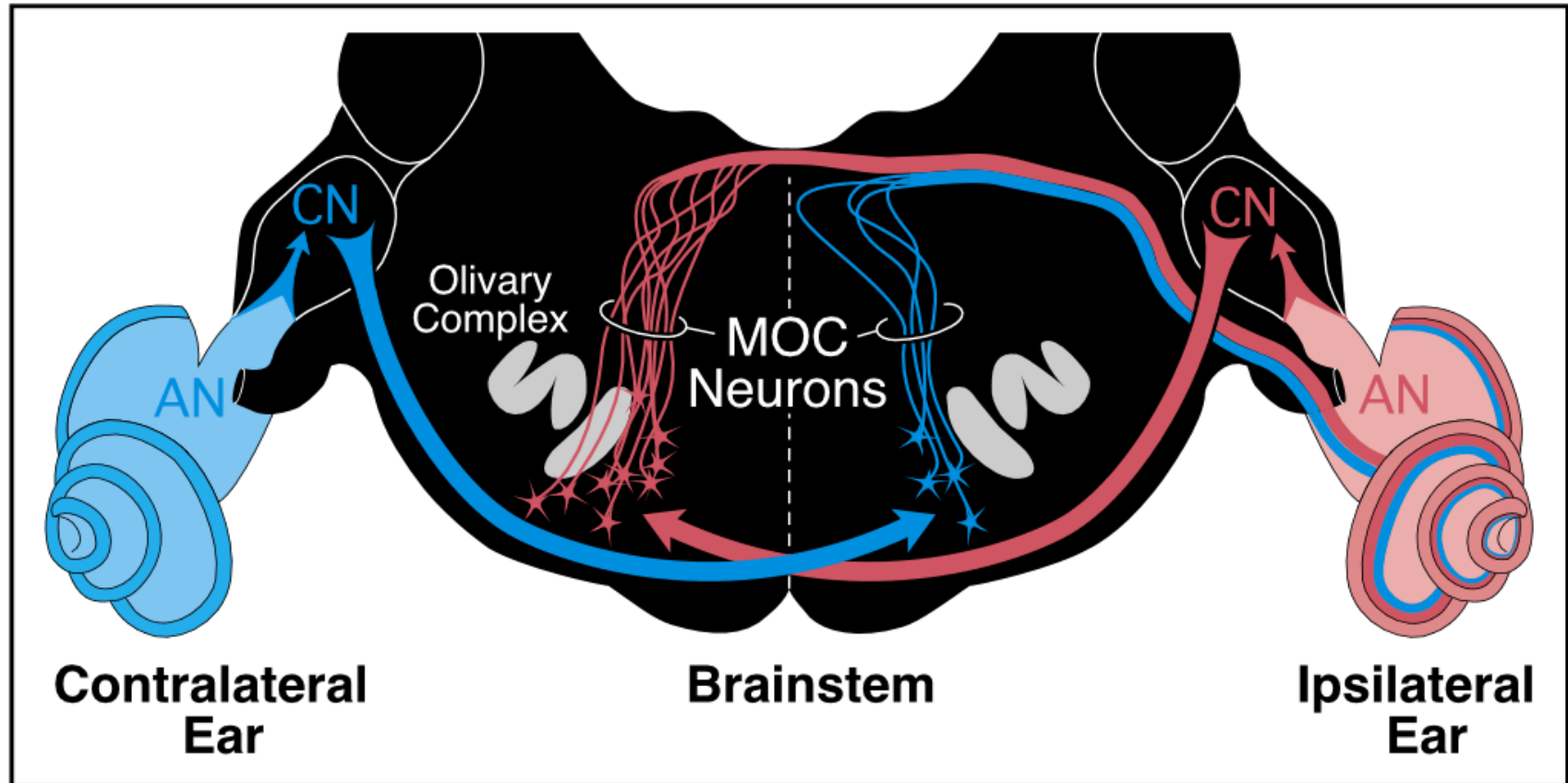
Olivocochlear Central Circuitry

Olivocochlear Bundle:
Superficial at floor of IVth Ventricle



LOC 90% Ipsilateral vs. MOC 66% Contralateral

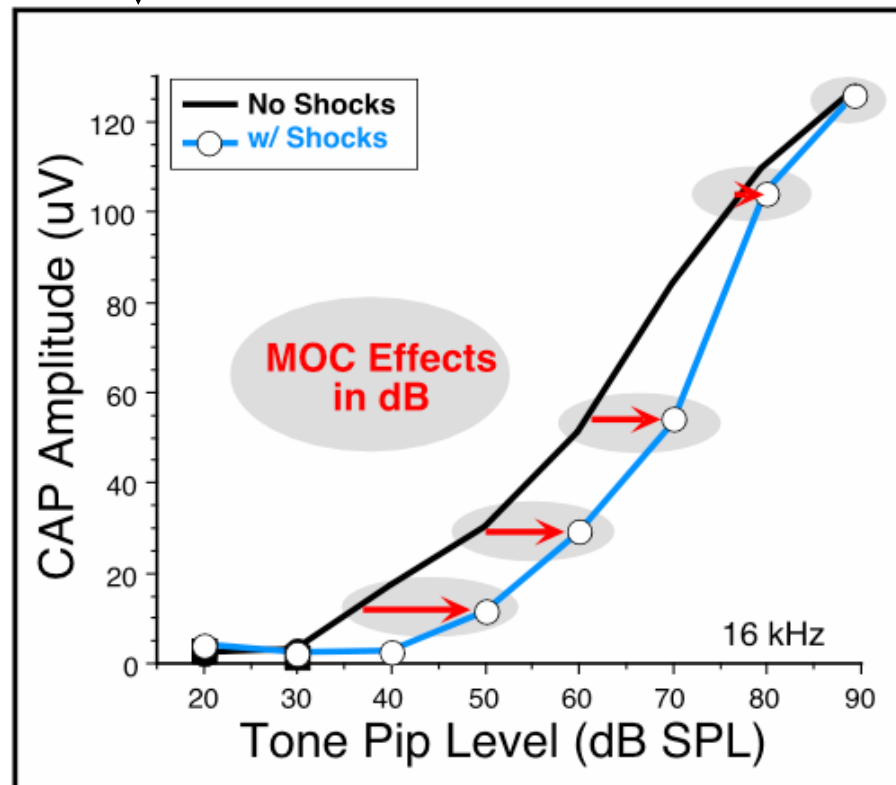
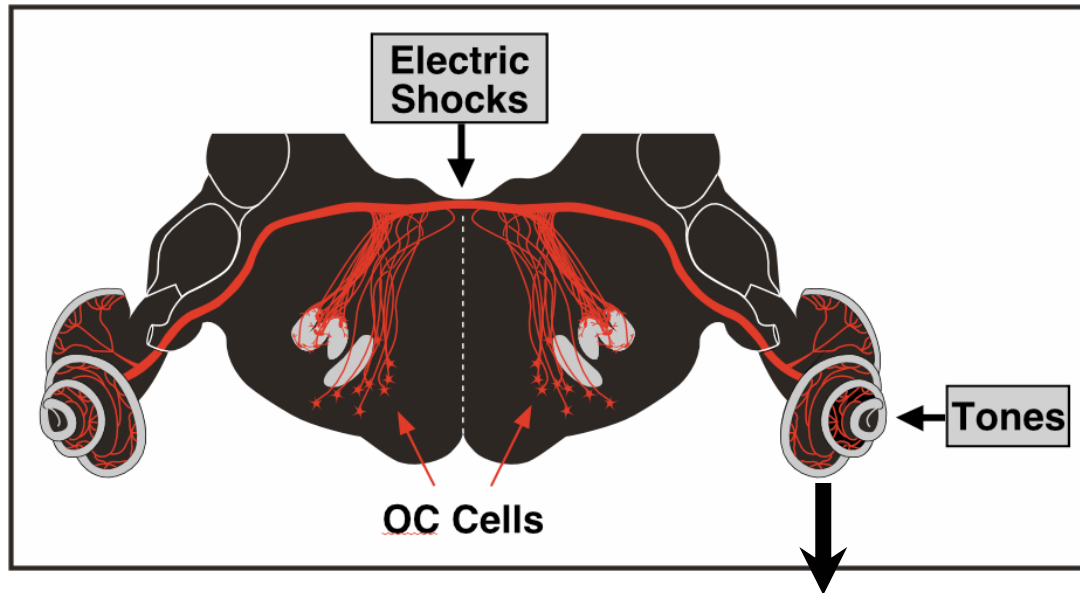
MOC Sound-Evoked Reflex



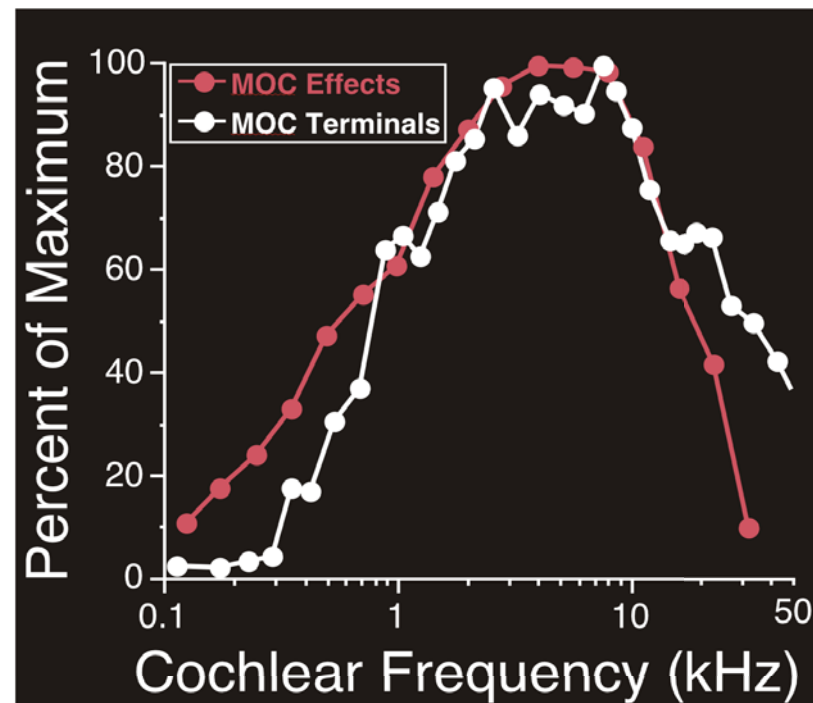
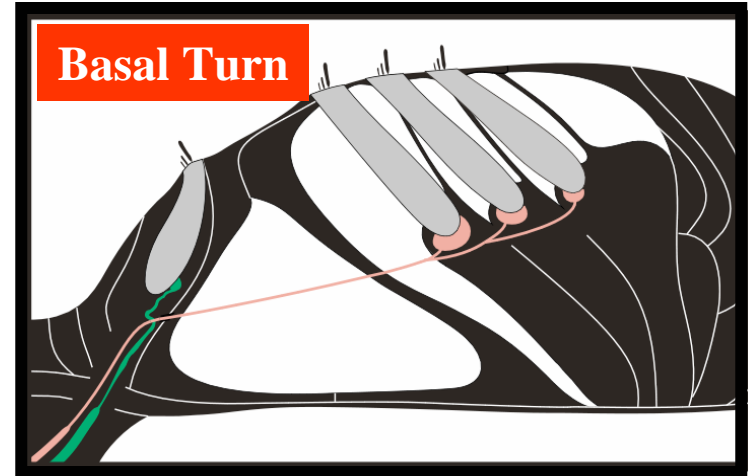
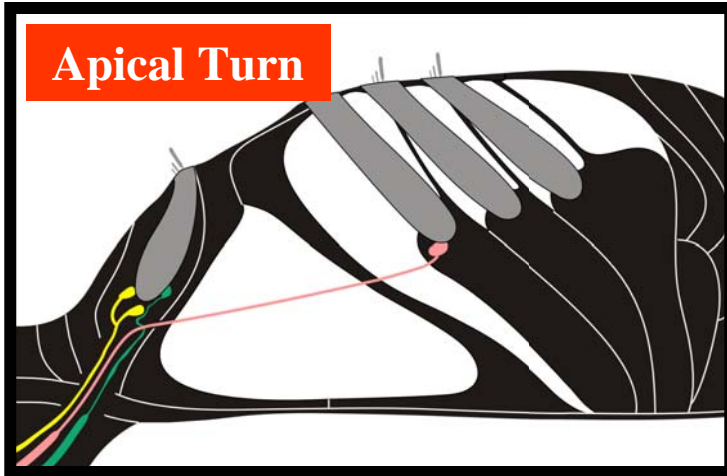
Ipsilaterally Evoked Reflex

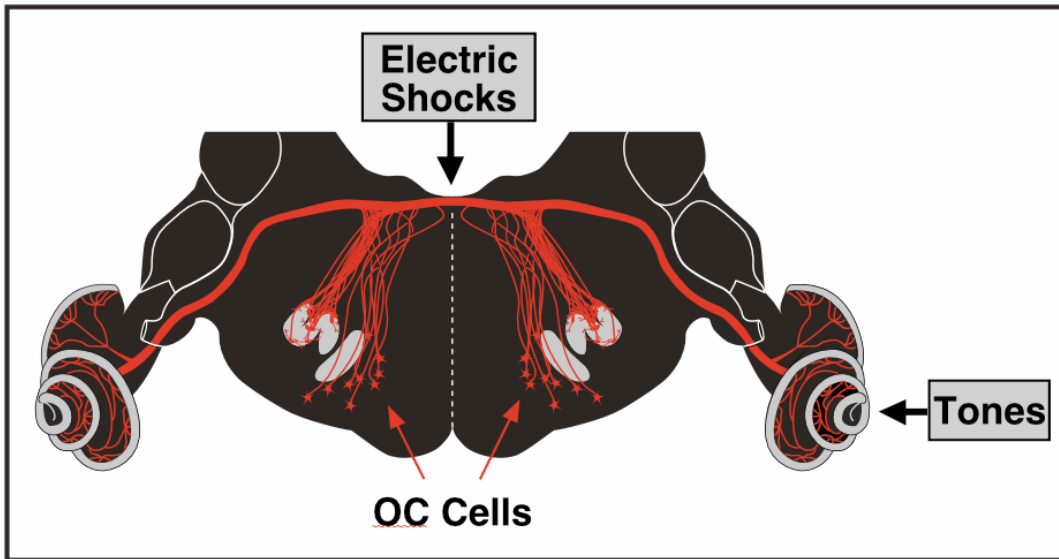
Contralaterally Evoked Reflex

MOC Effects: CAP Suppression



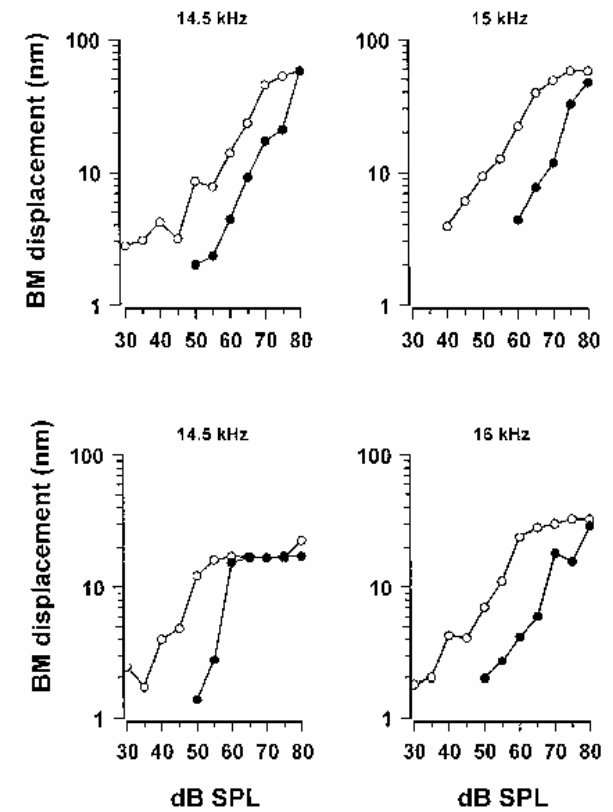
MOC Effects: Biggest at High Frequencies





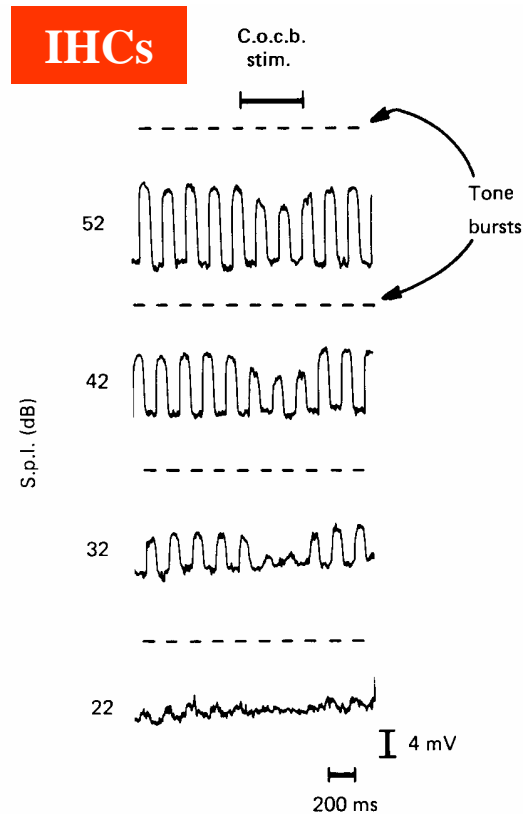
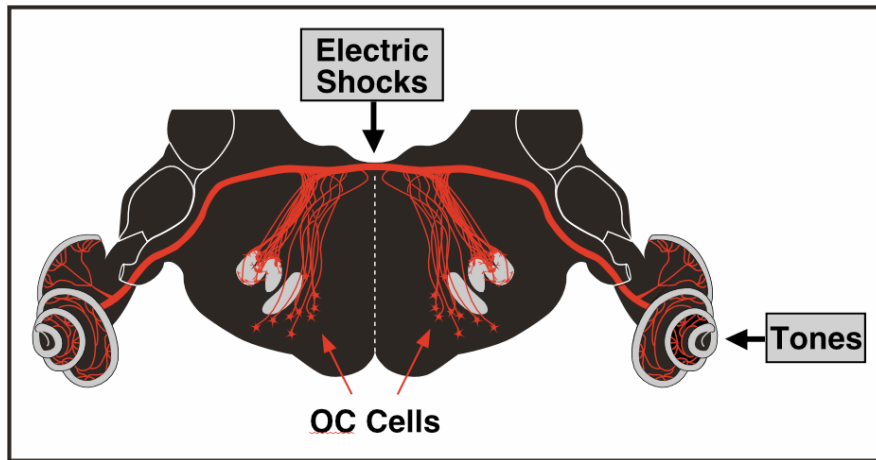
MOC Effects: Suppression in BM motion

Basilar Membrane

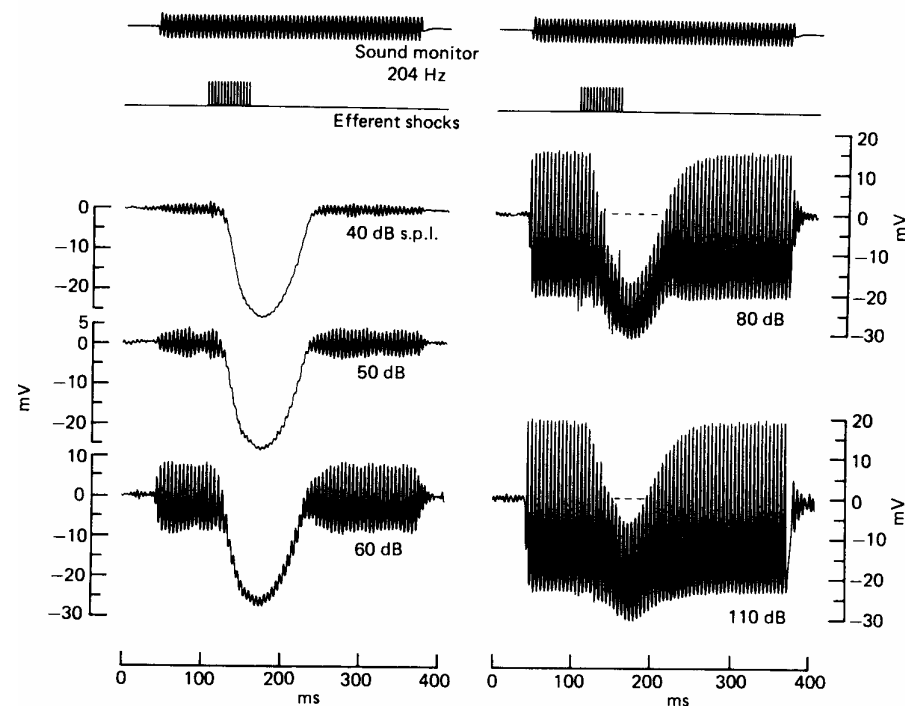


Murugasu E and Russell IJ (1996). The effect of efferent stimulation on basilar membrane displacement in the basal turn of the guinea pig cochlea. *J. Neuroscience* 16: 325-332.

MOC Effects: Differences between IHCs & OHCs

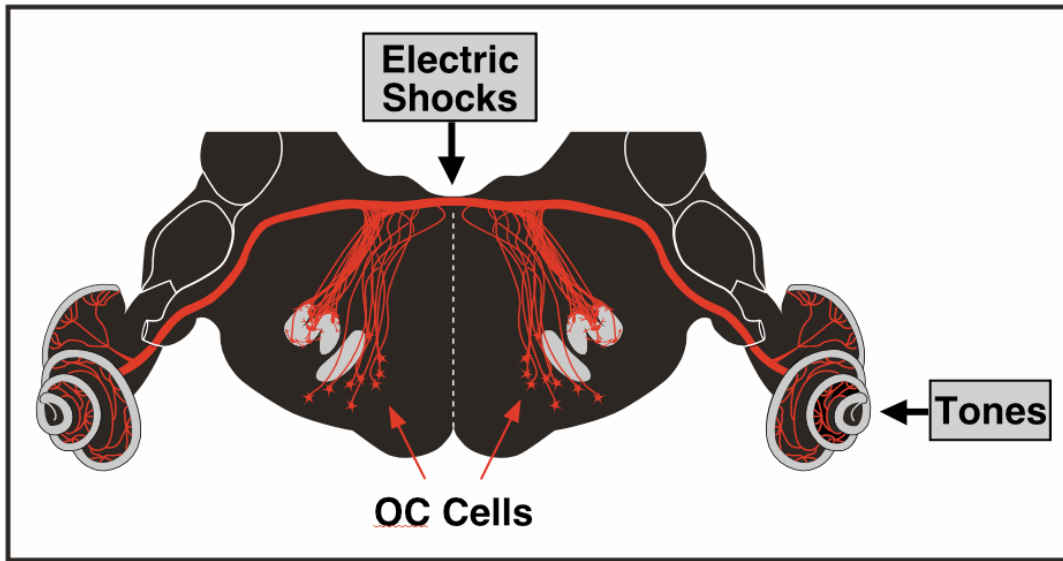


Turtle Hair Cells (OHC homologues)

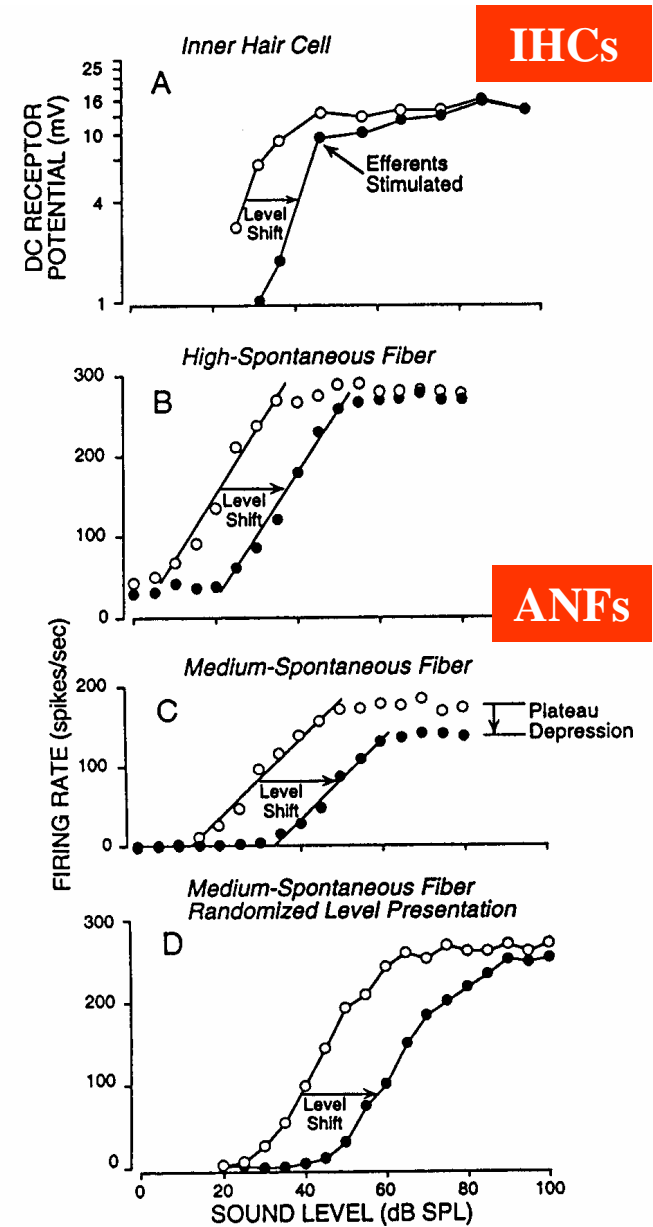
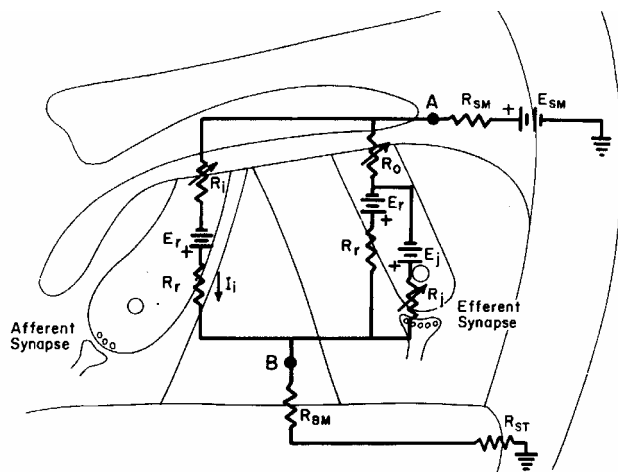


Brown MC and Nuttall AL (1984). Efferent control of cochlear inner hair cell responses in the guinea pig. *J. Physiol.* 354: 625-646.

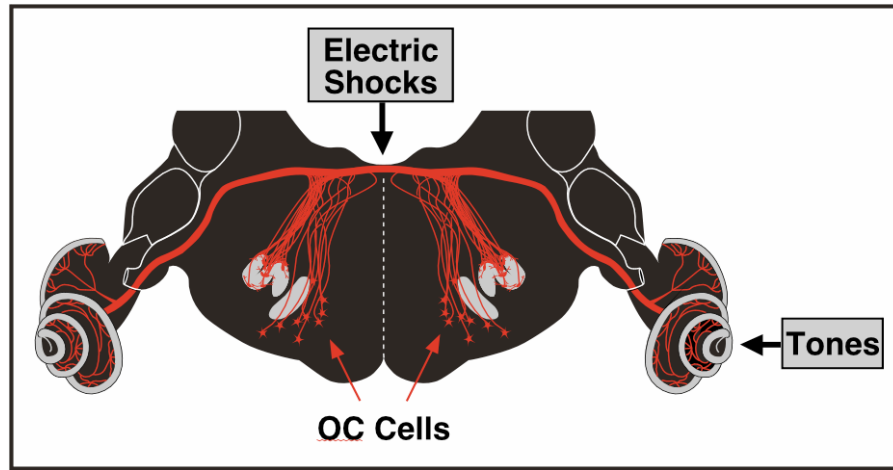
Art JJ, Fettiplace R and Fuchs PA (1984) Synaptic hyperpolarization of turtle cochlear hair cells. *J Physiol.* 356:525-550.



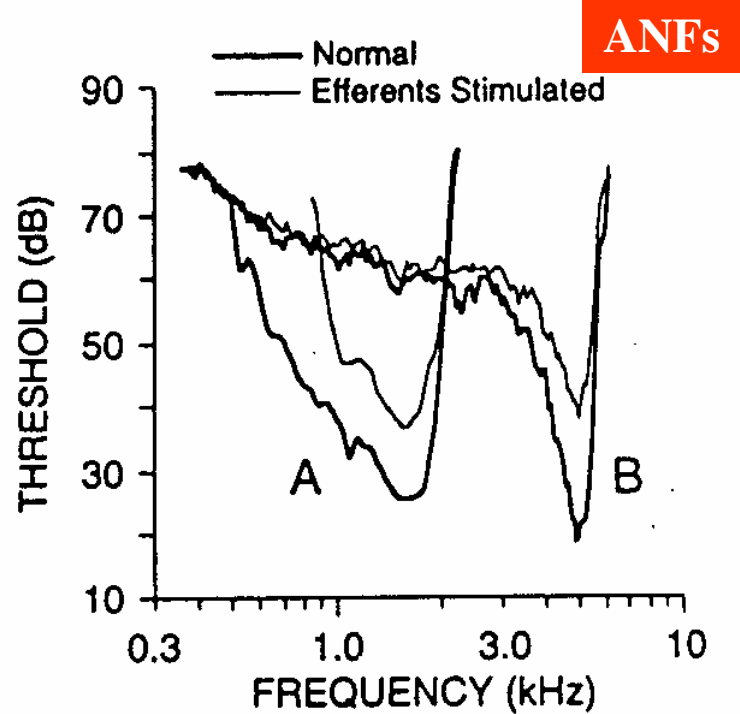
MOC Effects: Suppression in IHCs and ANFs



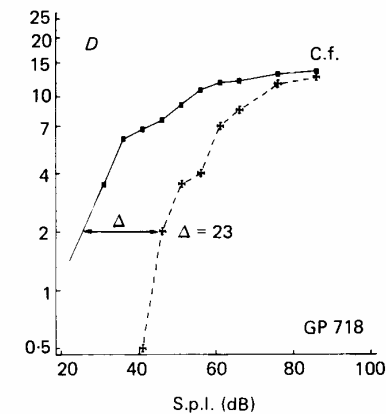
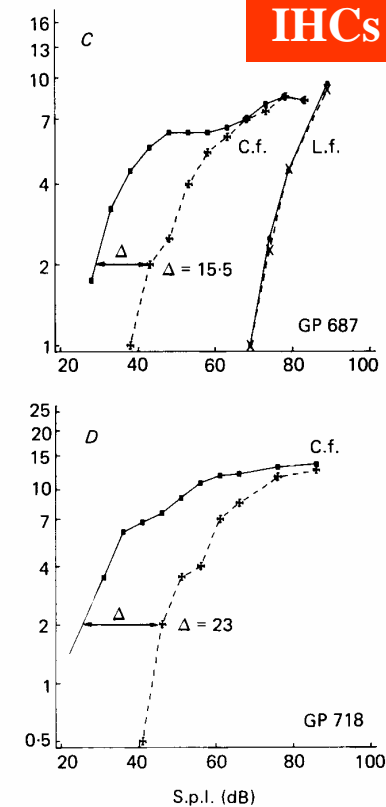
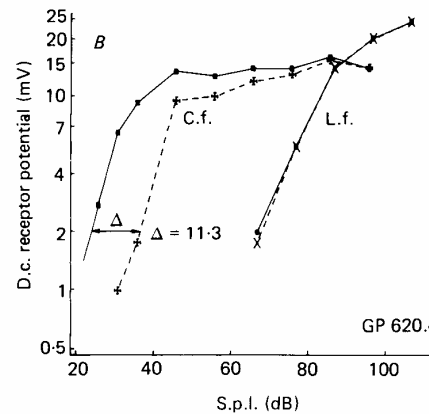
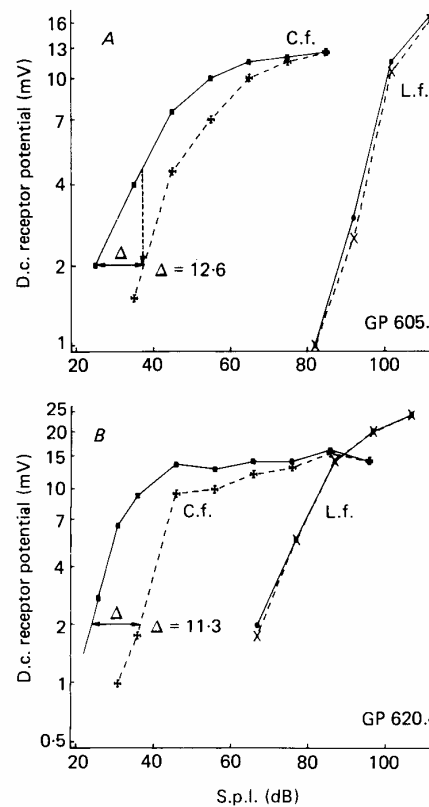
Guinan, JJ (1996) Physiology of Cochlear Efferents. In "The Cochlea", eds. P Dallos, AN Popper and RR Fay. New York, Springer, pp 435-500.



MOC Effects: Biggest at CF

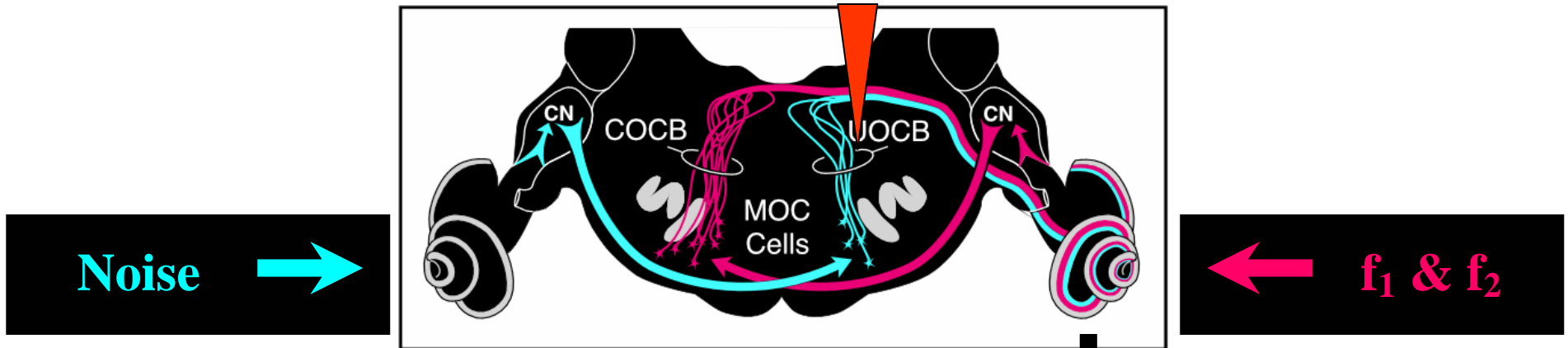


Guinan, JJ (1996) Physiology of Cochlear Efferents. In "The Cochlea", eds. P Dallos, AN Popper and RR Fay. New York, Springer, pp 435-500.

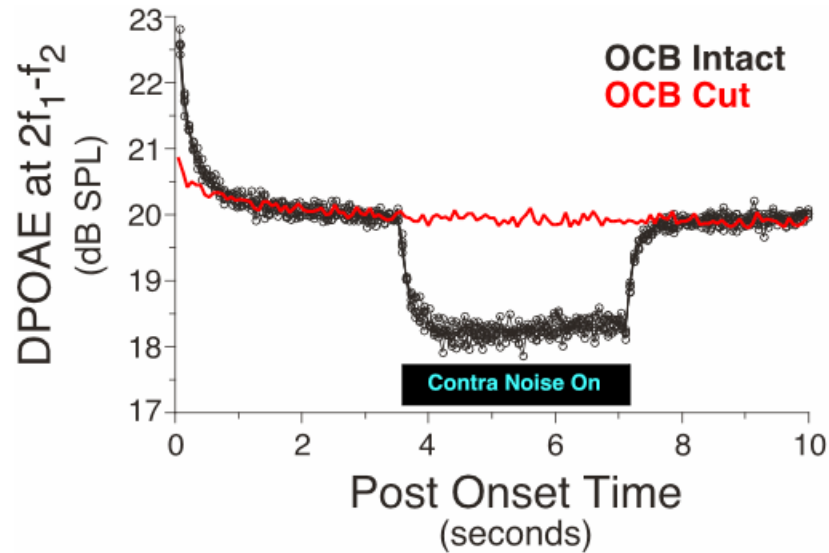


Brown MC and Nuttall AL (1984). Efferent control of cochlear inner hair cell responses in the guinea pig. J. Physiol. 354: 625-646.

MOC Reflex Affects OtoAcoustic Emissions

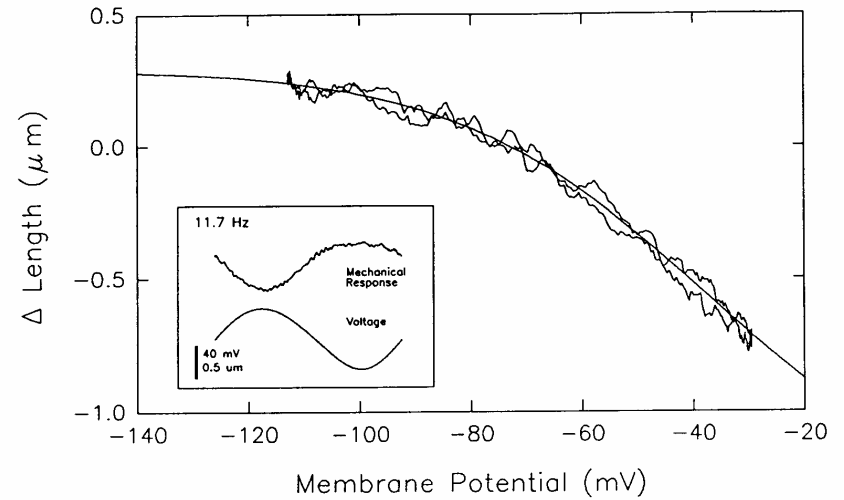
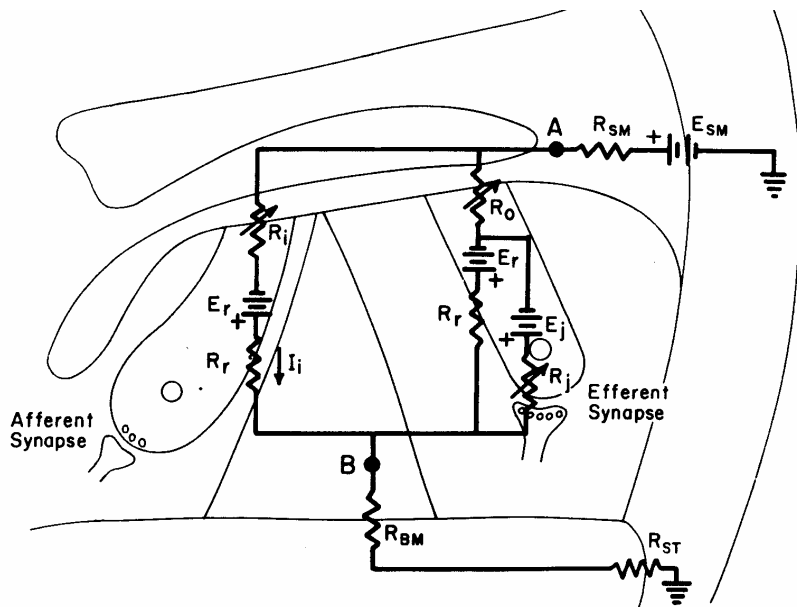


Distortion Product: $2f_1-f_2$



How MOC activity suppresses cochlea ?

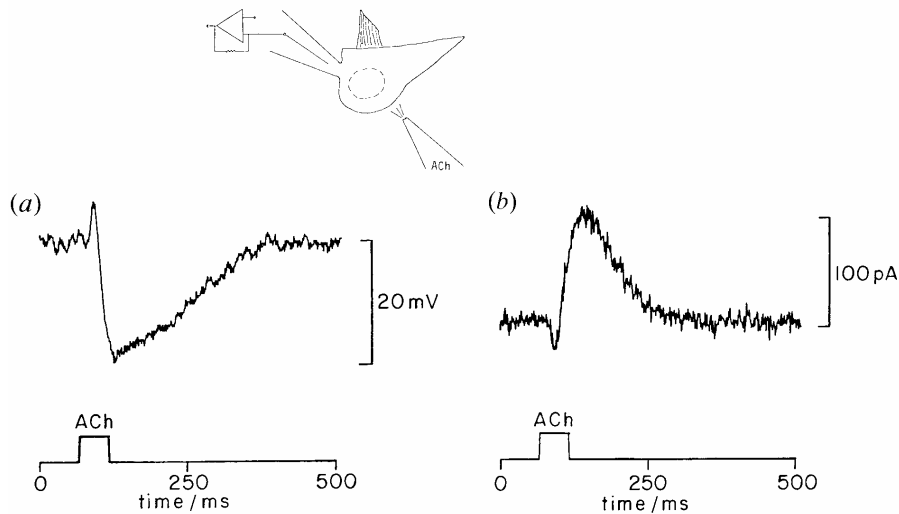
OHC motility decreases w/ Hyperpolarization



Santos-Sacchi, J. (1991) Reversible inhibition of voltage dependent outer hair cell motility and capacitance. *J. Neurosci.* 11: 3096-3110

What channels are involved in MOC effects?

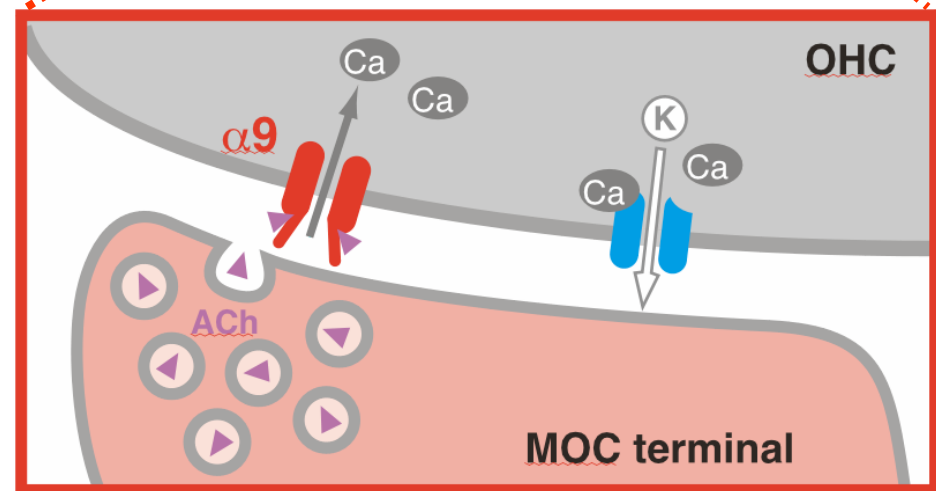
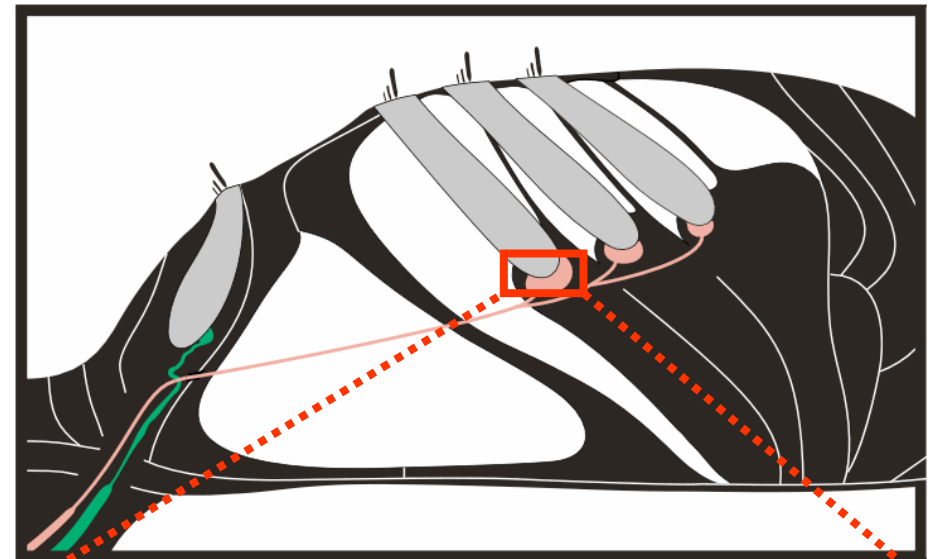
Chick Hair Cells in vitro



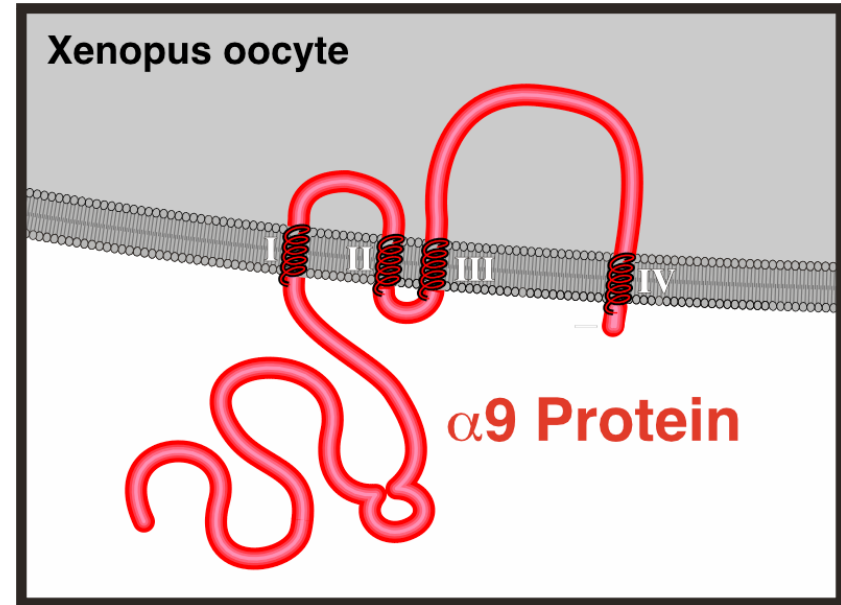
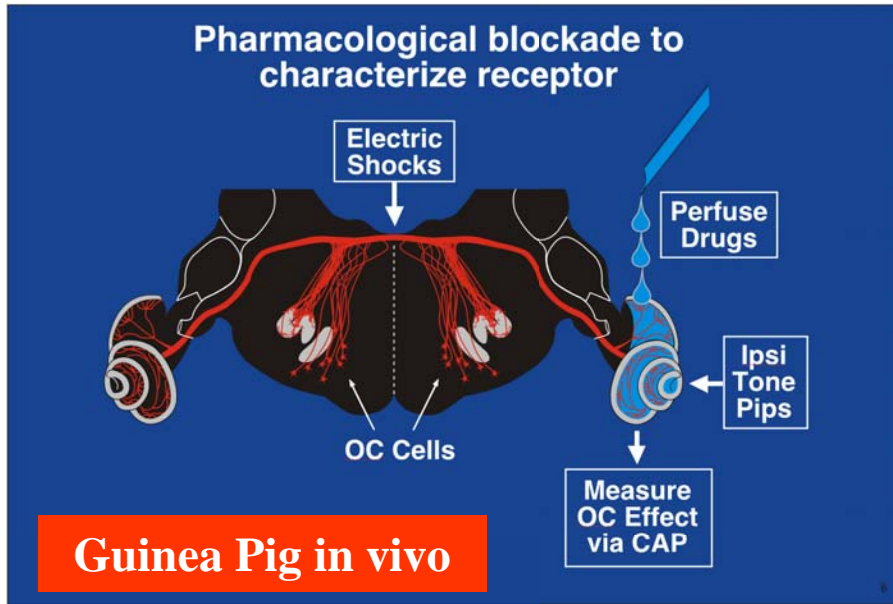
Fuchs PA, Murrow BW (1992) A novel cholinergic receptor mediates inhibition of chick cochlear hair cells. Proc R Soc Lond B Biol Sci 248:35-40.

Hypothesis: Two stage effect

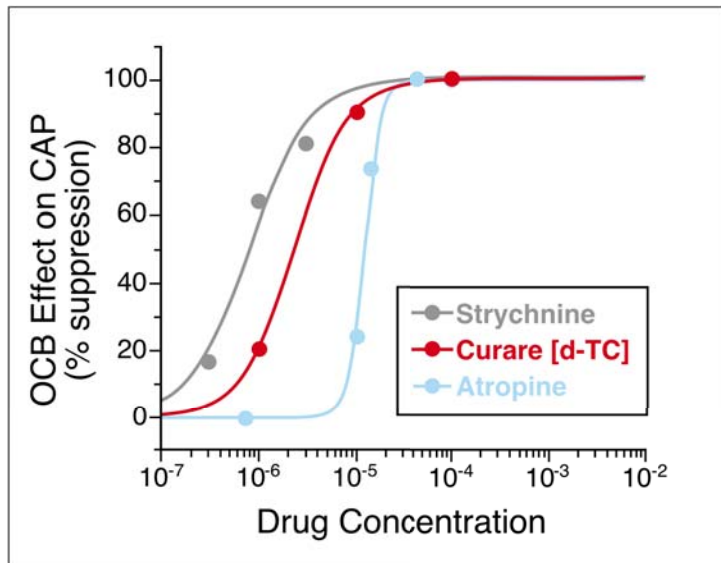
1. ACh activated Ca entry
2. Ca-activated K entry



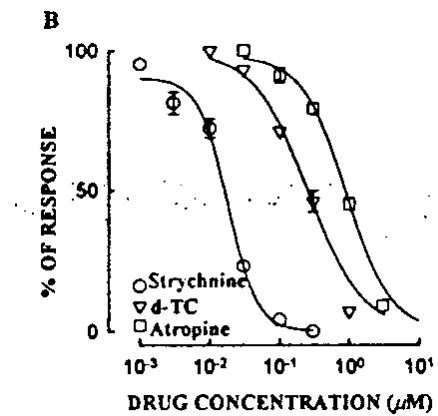
$\alpha 9$ nicotinic ACh receptor



In vitro: heterologous expression

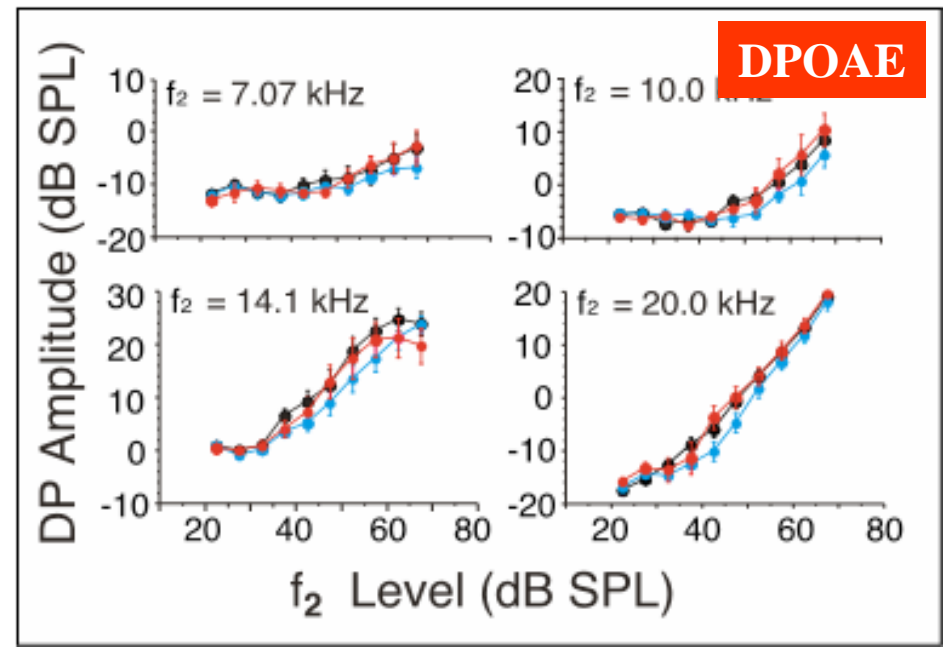
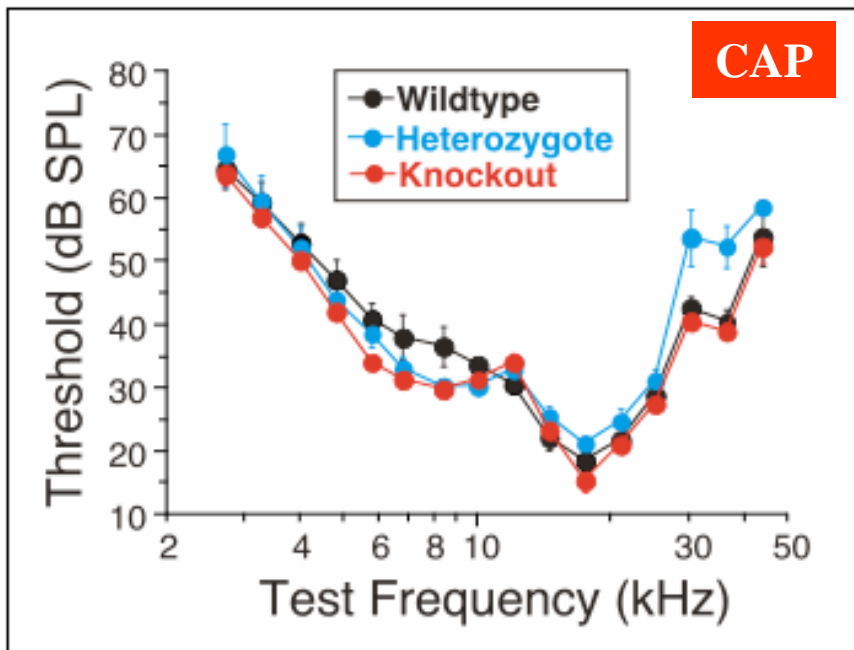
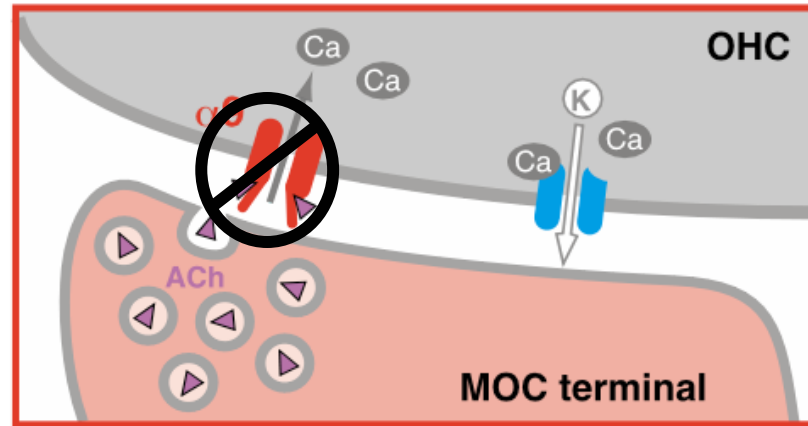


Sridhar T, Liberman MC, Brown MC and Sewell WF. A novel cholinergic "slow effect" of efferent stimulation on cochlear potentials in the guinea pig. *J. Neurosci.* 1995; 15:3667-3678.

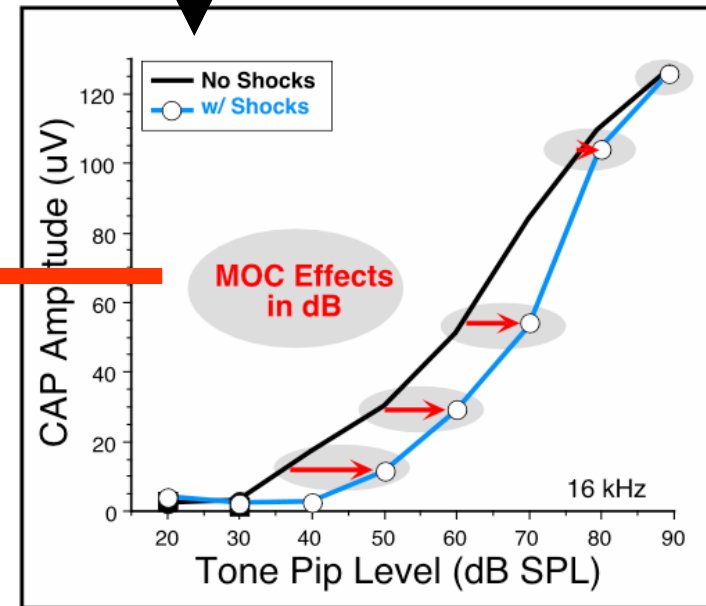
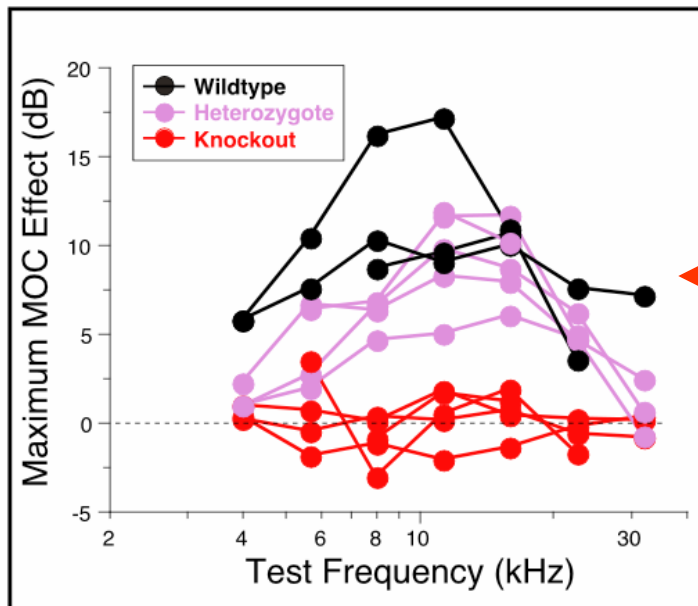
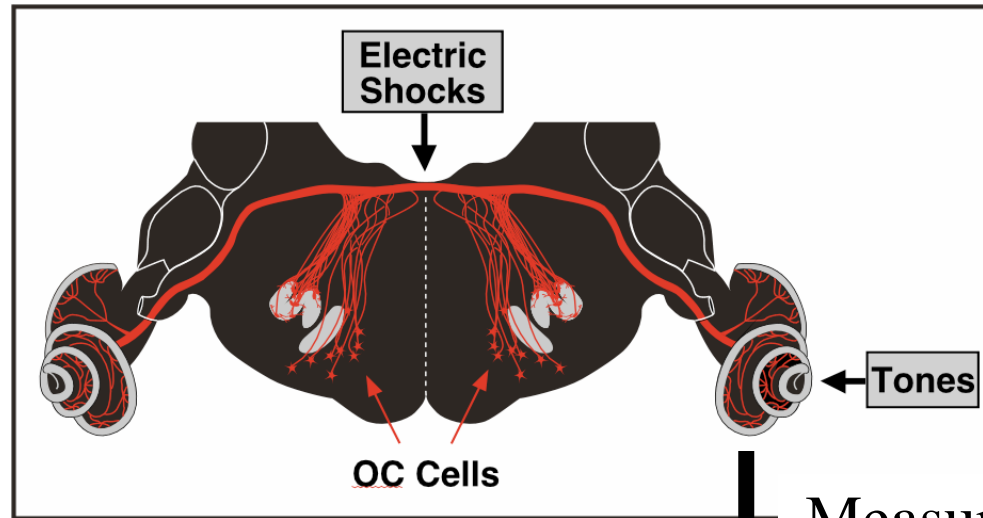


Elgoyhen AB et al. (1994) Alpha 9: an acetylcholine receptor with novel pharmacological properties expressed in rat cochlear hair cells. *Cell* 79:705-715.

$\alpha 9$ Knockout: Cochlear function normal



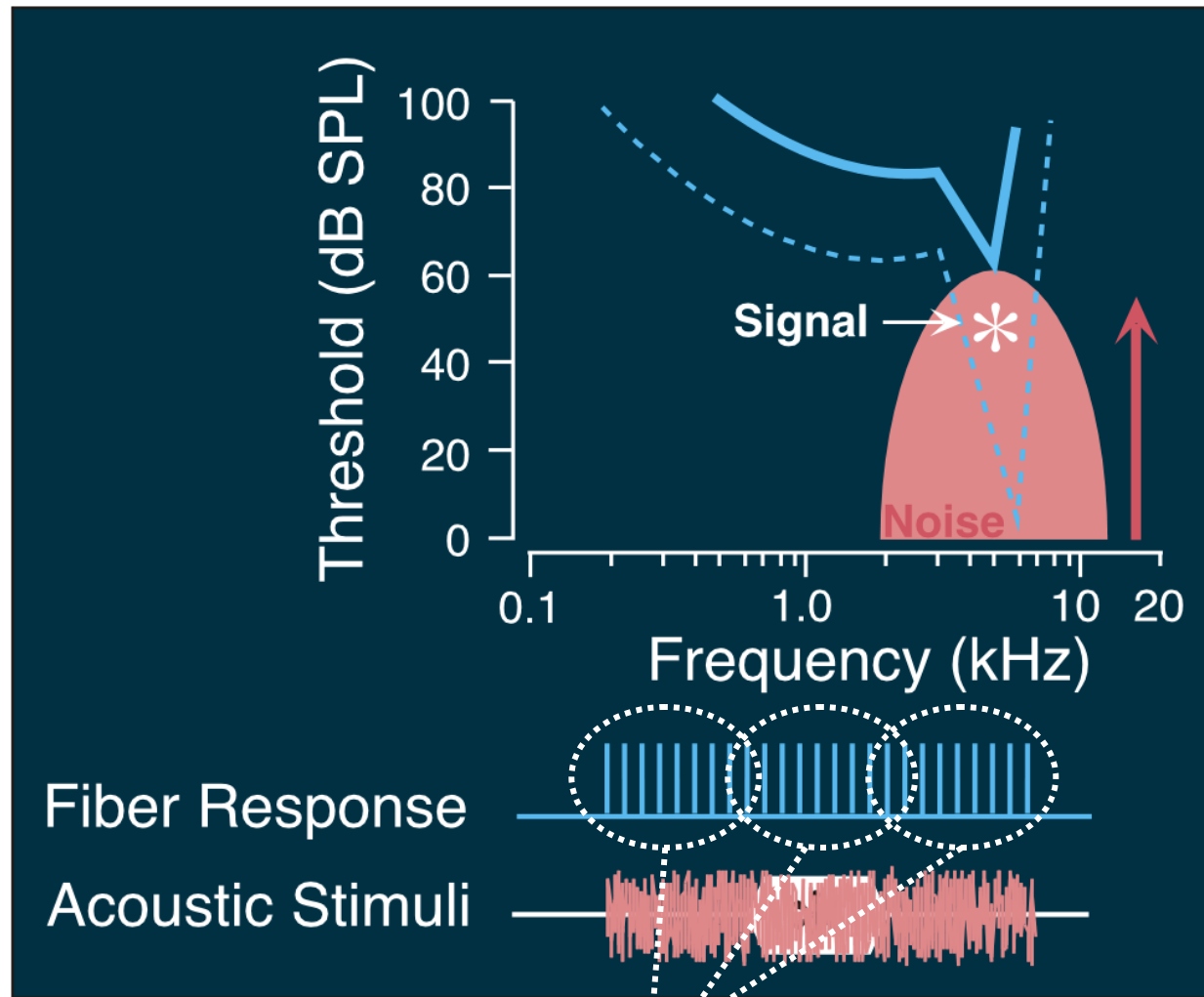
$\alpha 9$ Knockout is functionally de-efferented



Hypotheses for MOC Function

- 1. Extend dynamic range - a gain control system**
- 2. Mediate selective attention: auditory vs. visual
high vs. low frequency**
- 3. Control masking from background noise**
- 4. Shape normal cochlear development**
- 5. Protect the inner ear from acoustic injury**

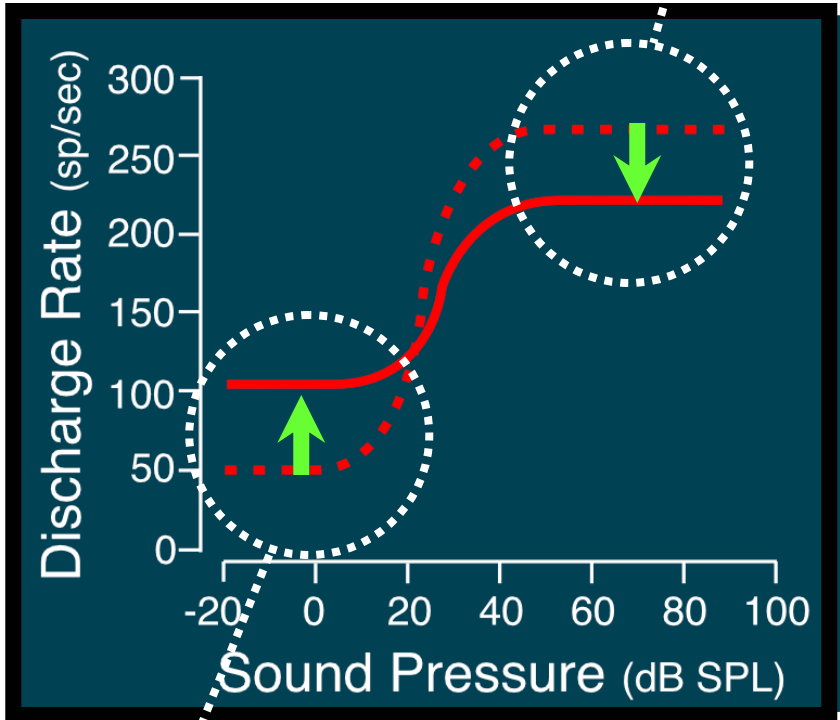
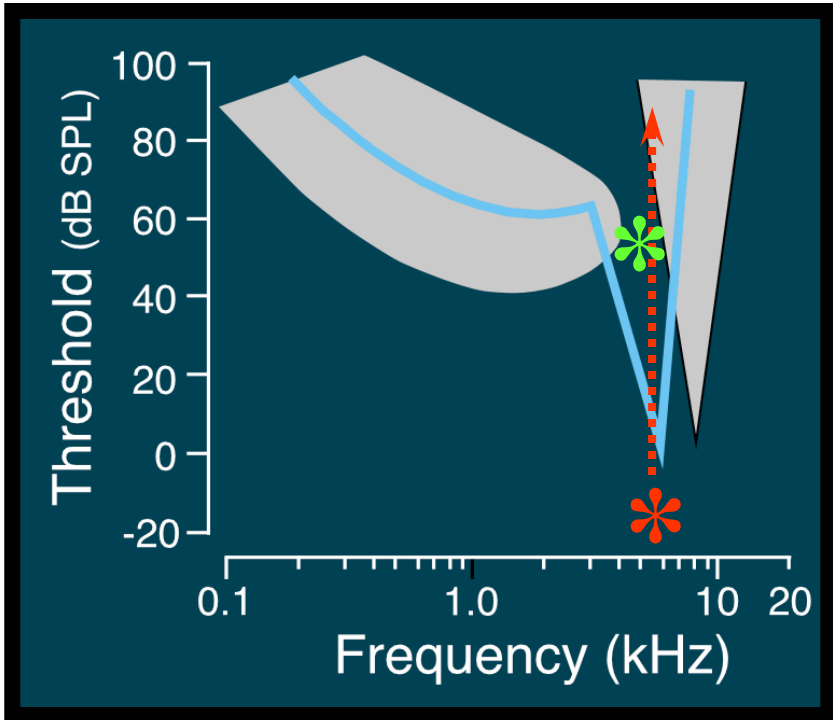
Excitatory Masking



Response to signal decreases (adaptation)

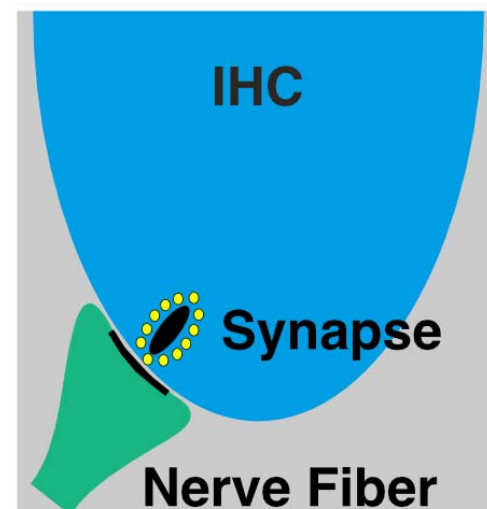
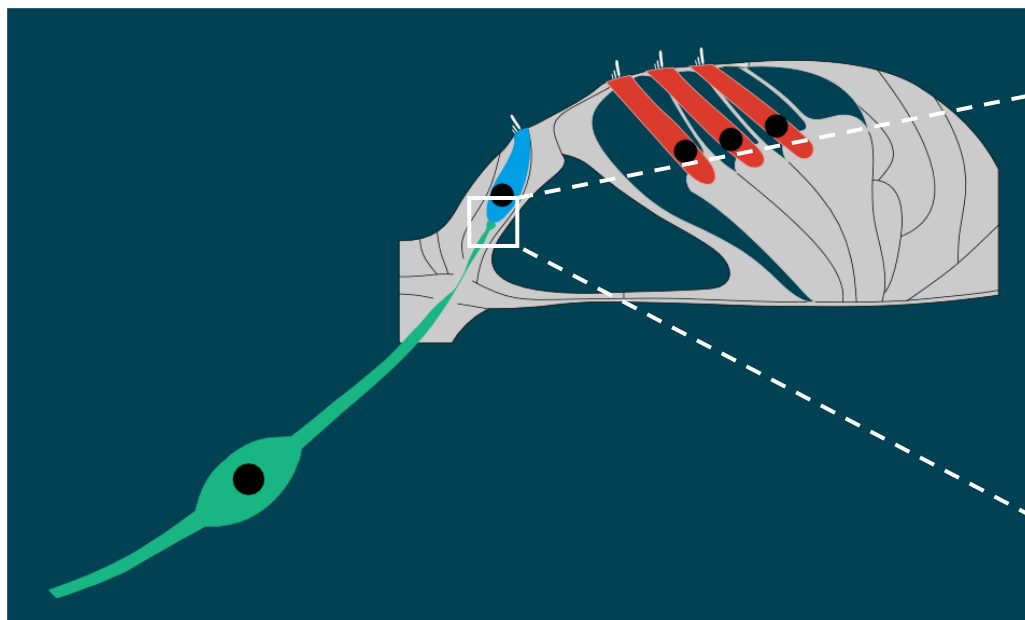
Excitatory Masking - Line Busy + Adaptation

Adaptation



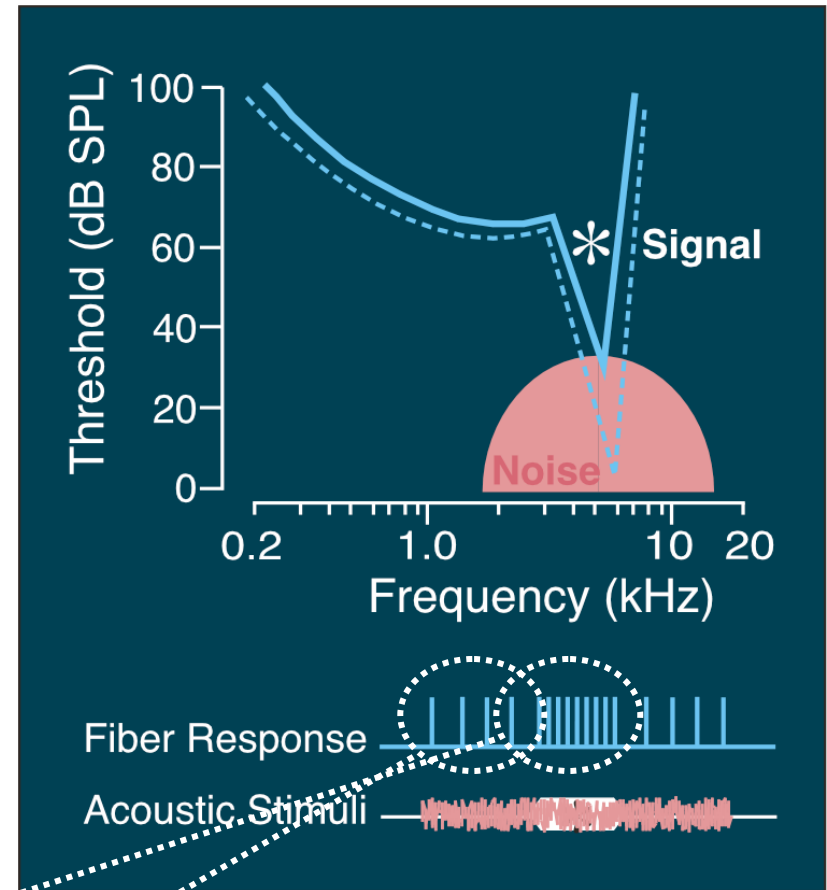
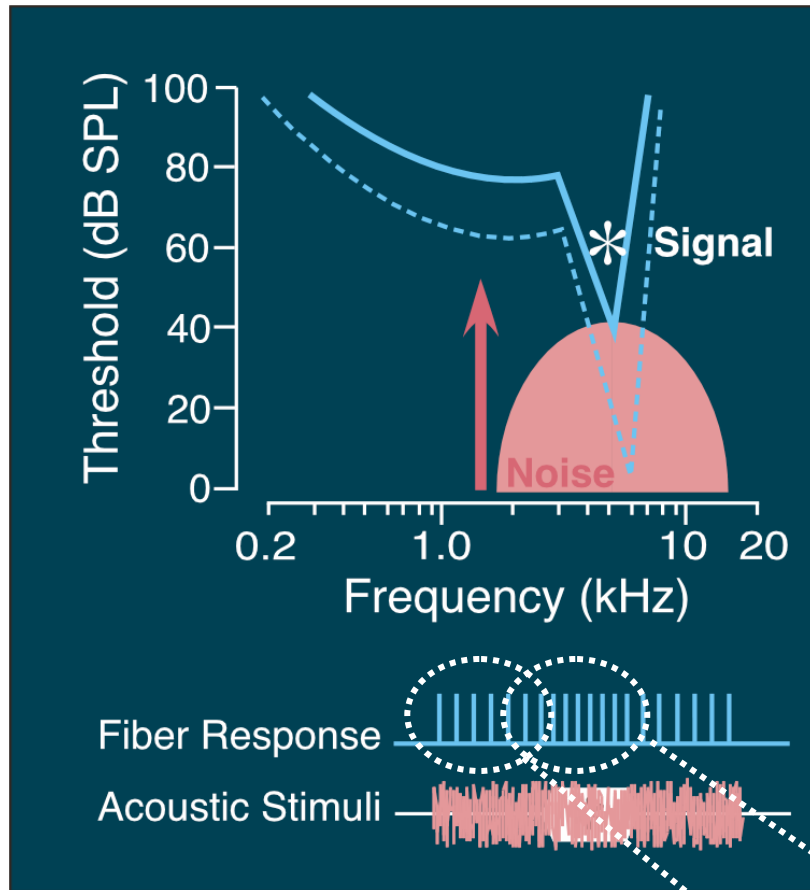
Line Busy

Adaptation component of Excitatory Masking



- **Arises at the synapse - vesicle depletion**
- **Does not require stimulus simultaneity**
- **Mechanism underlying forward masking**

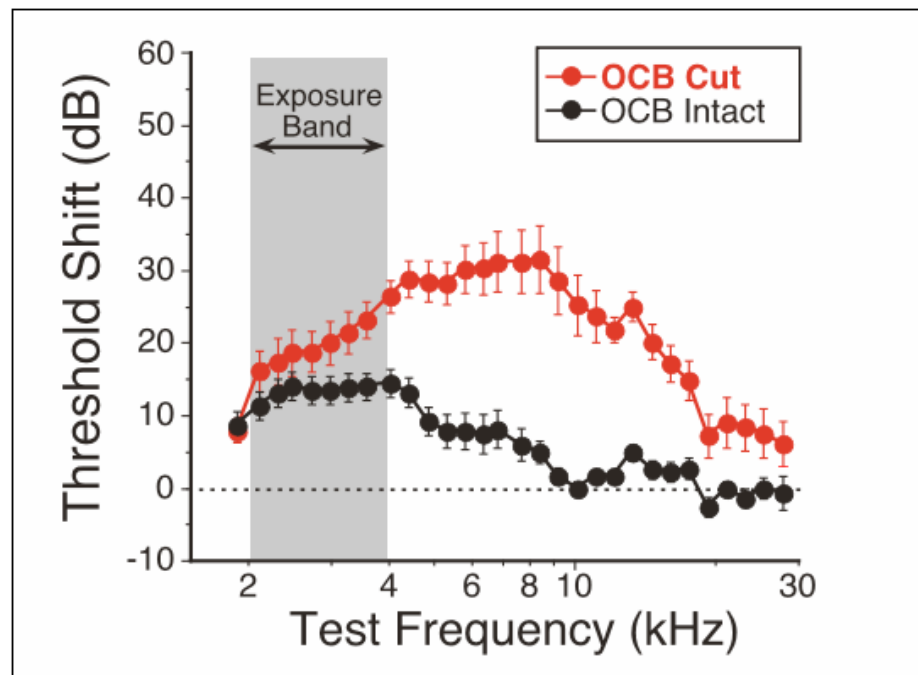
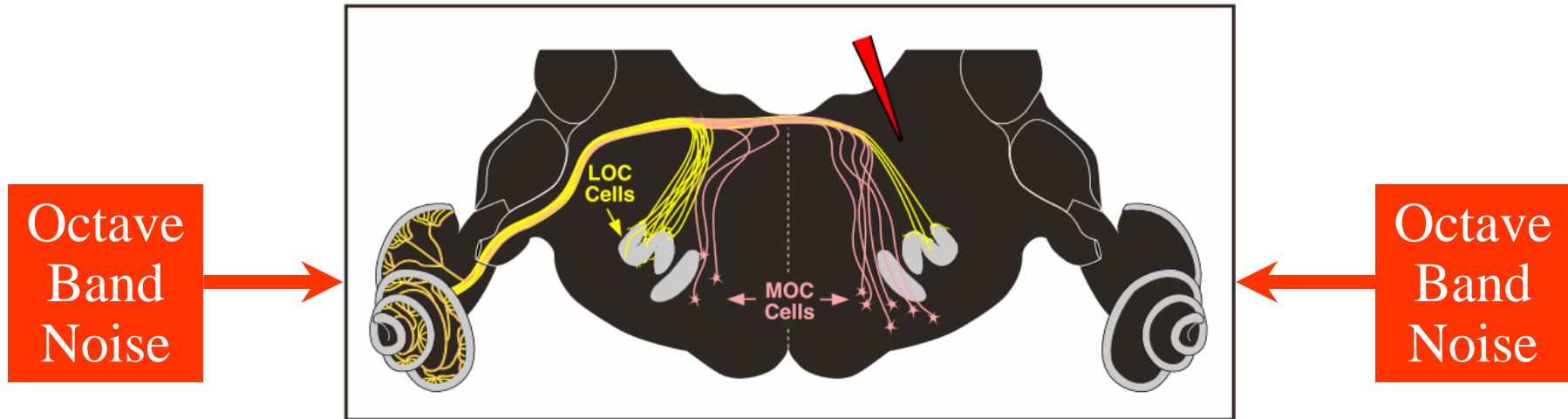
MOC counteracts excitatory masking



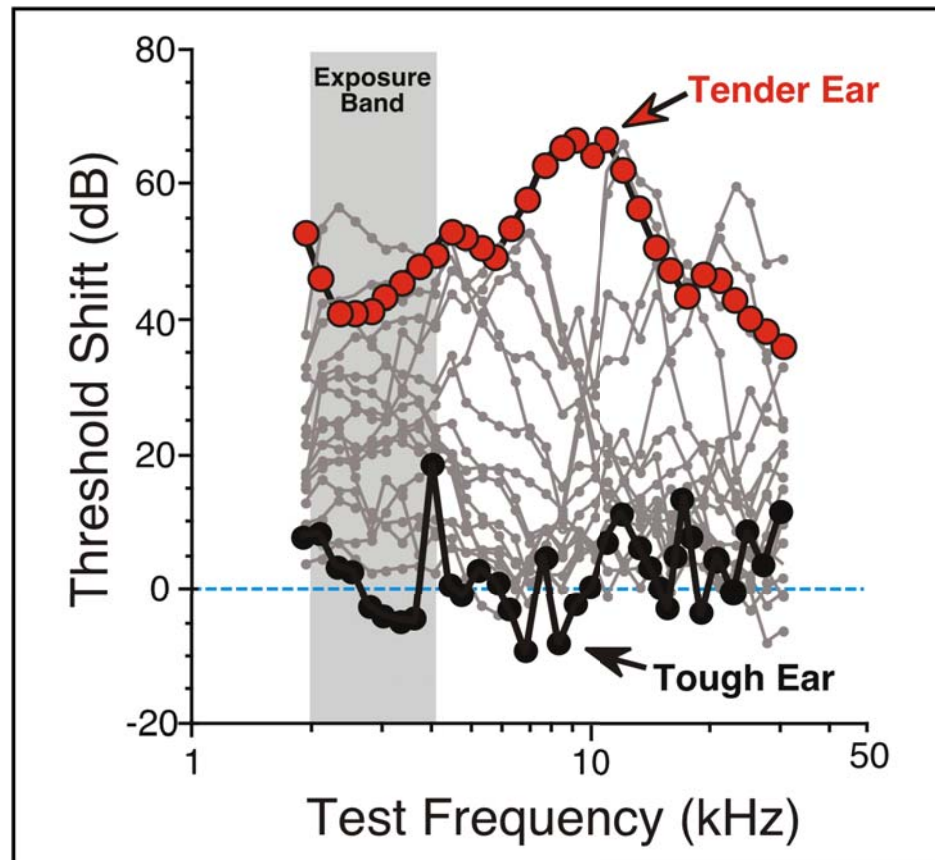
MOC reflex increases response to transients

By reducing response to steady noise

Unilateral De-efferentiation and Acoustic Injury



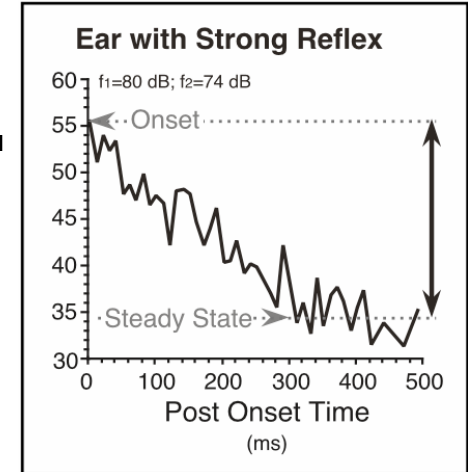
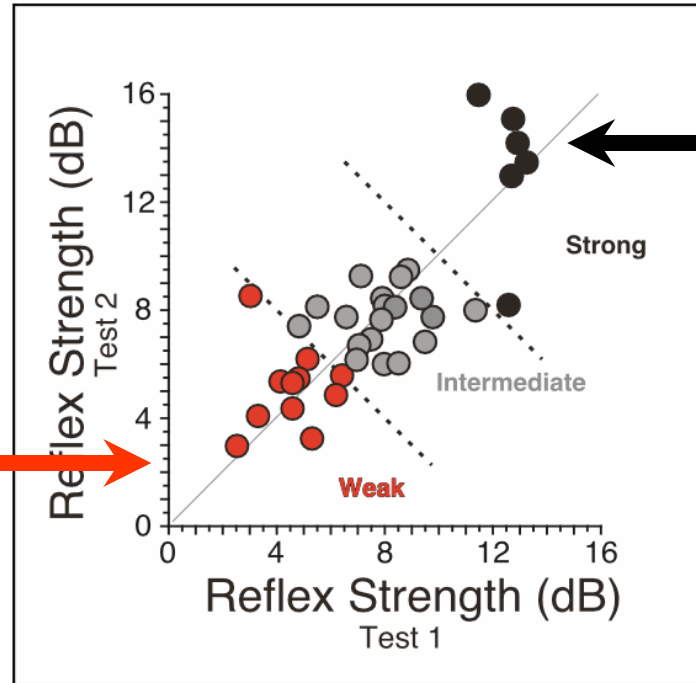
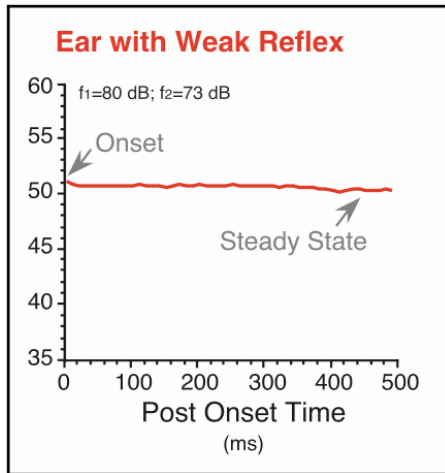
Intersubject Variability in Vulnerability to Acoustic Injury



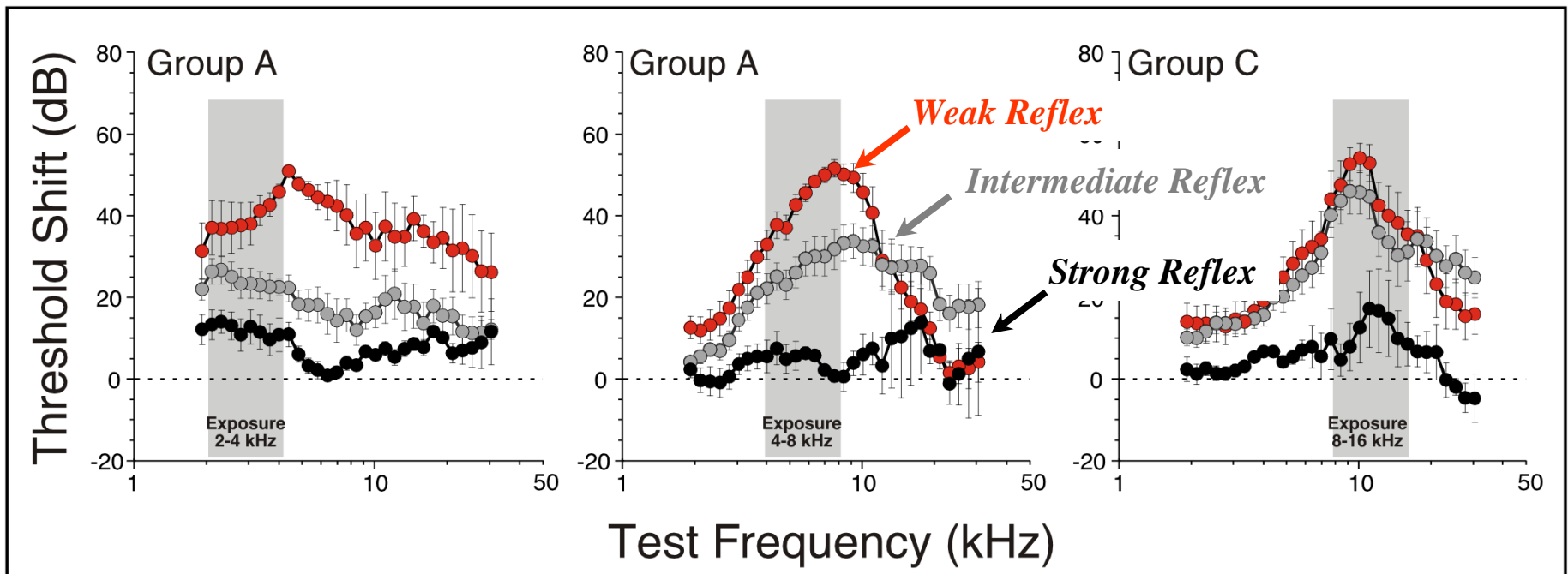
Is Variability due to Differences in MOC Reflex Strength?

Assaying MOC Reflex Strength

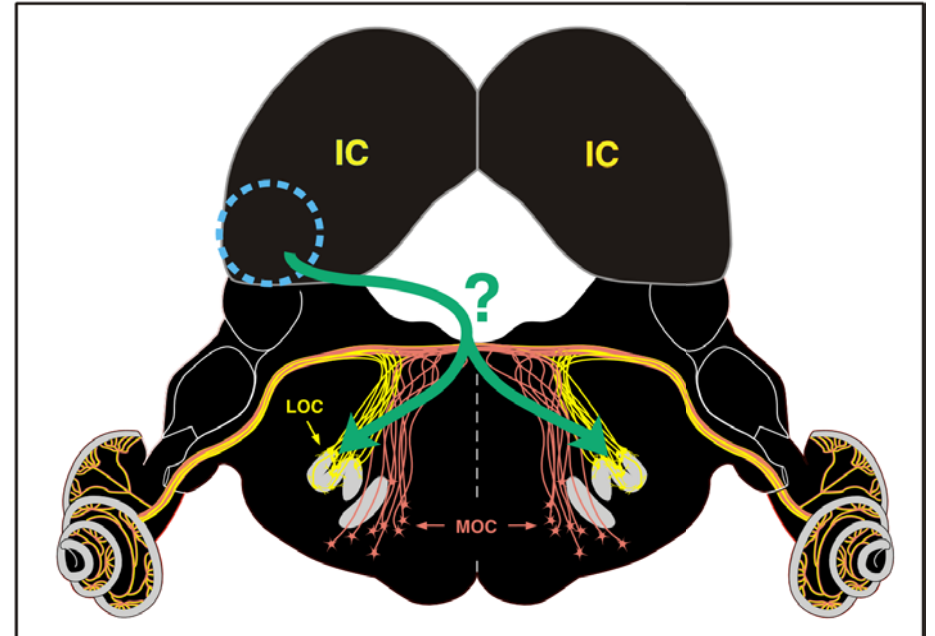
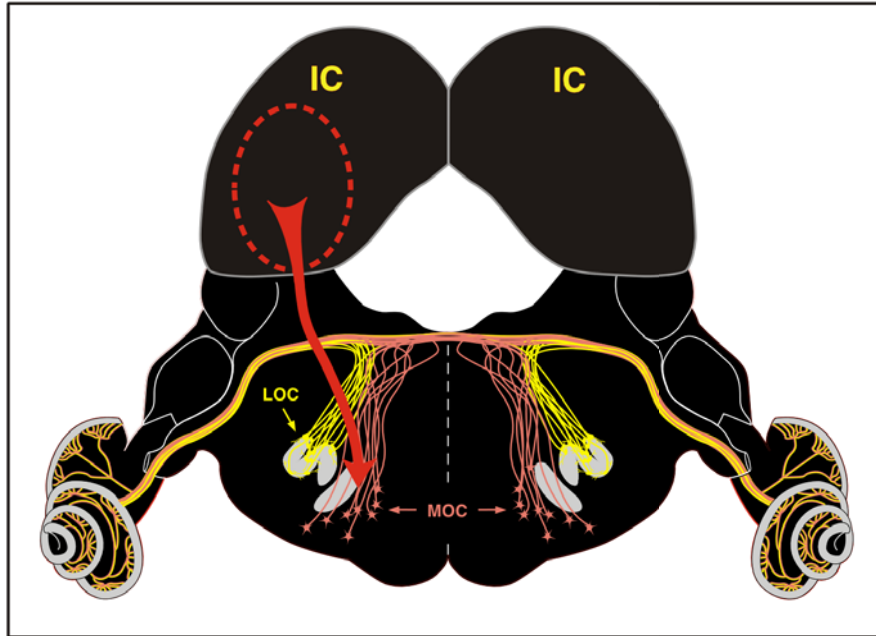
Measure Post-Onset Adaptation in 36 Awake Guinea Pigs



MOC Reflex Strength predicts vulnerability



Activating the LOC via IC stimulation



Activating the LOC via IC stimulation

