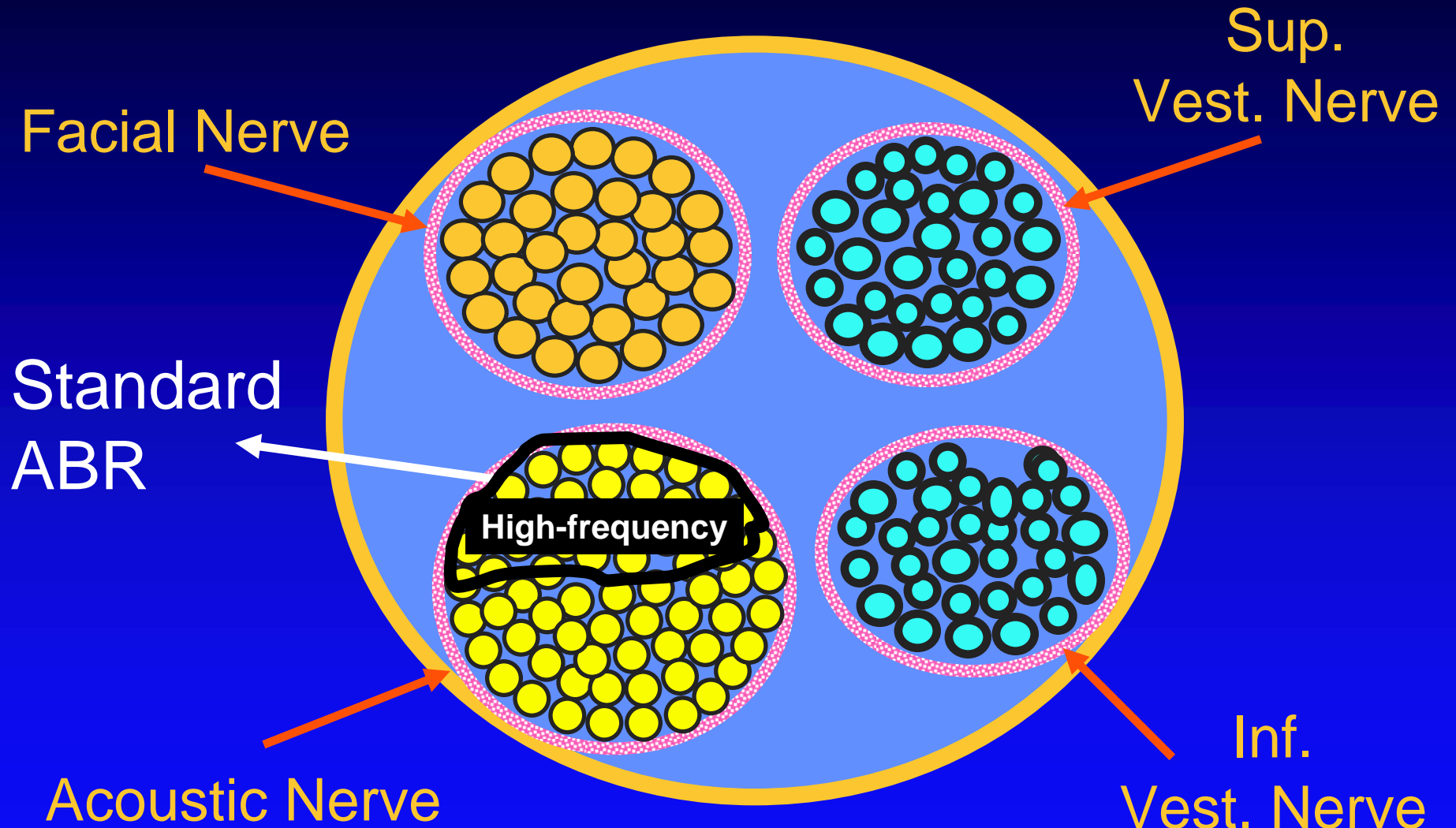


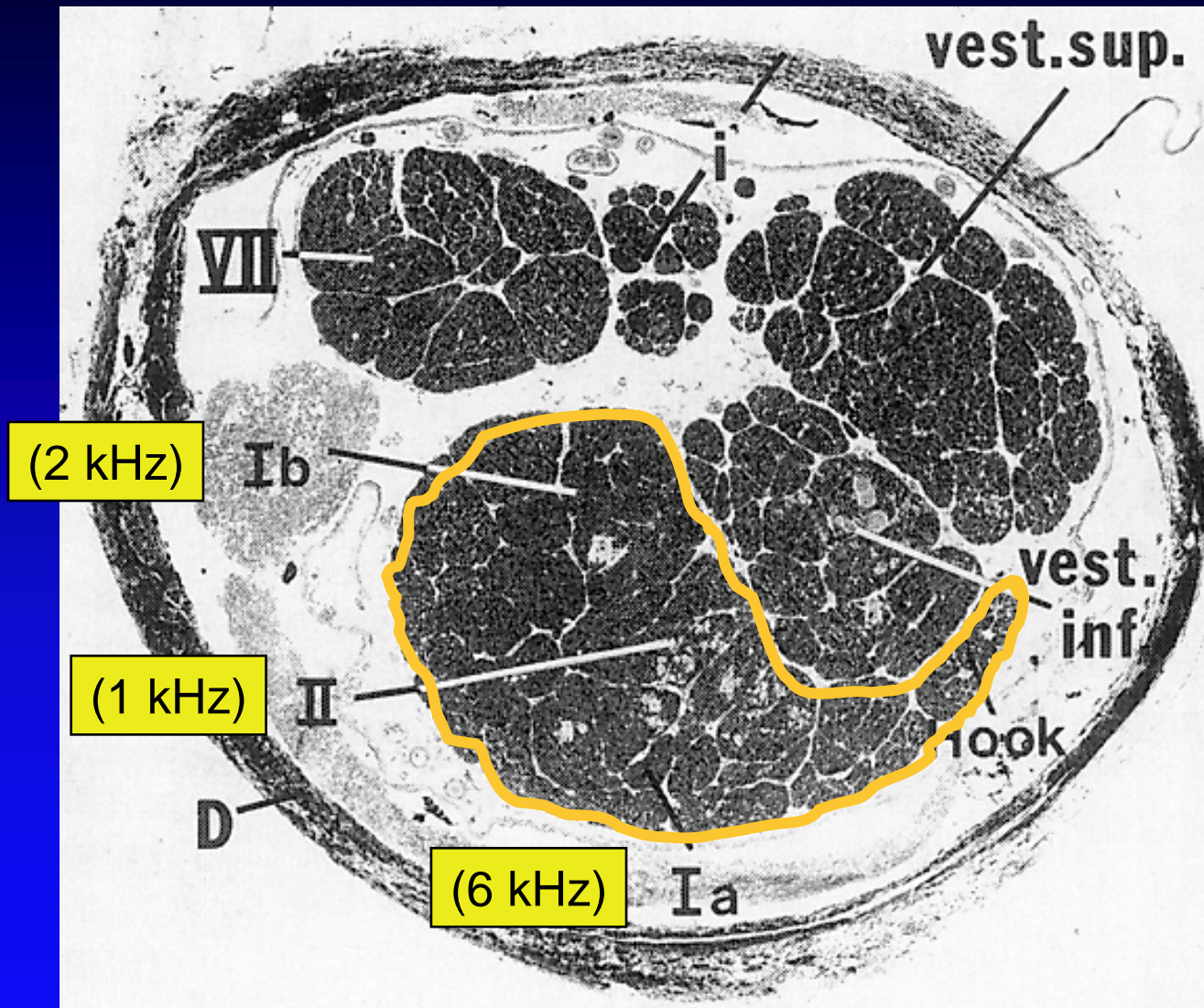
A Brief Introduction to Stacked ABR and Cochlear Hydrops Analysis Masking Procedure (CHAMP)

Prepared for Bio-logic Systems Corp. by
Manuel Don, Ph.D. / Betty Kwong, M.S.
Electrophysiology Department
House Ear Institute, Los Angeles, CA

Normal Internal Auditory Canal (IAC)

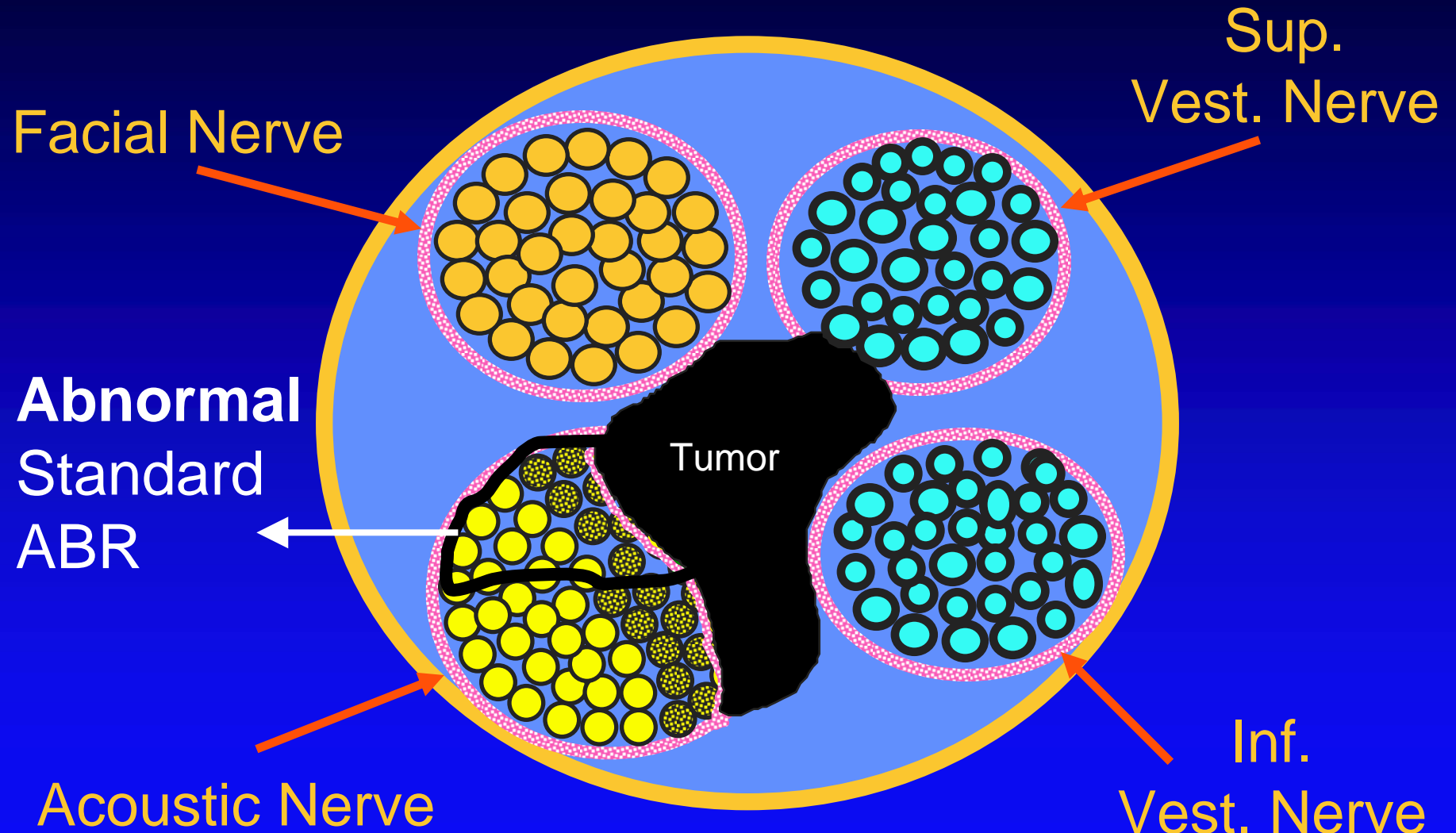


Cross Section: Human Auditory Meatus

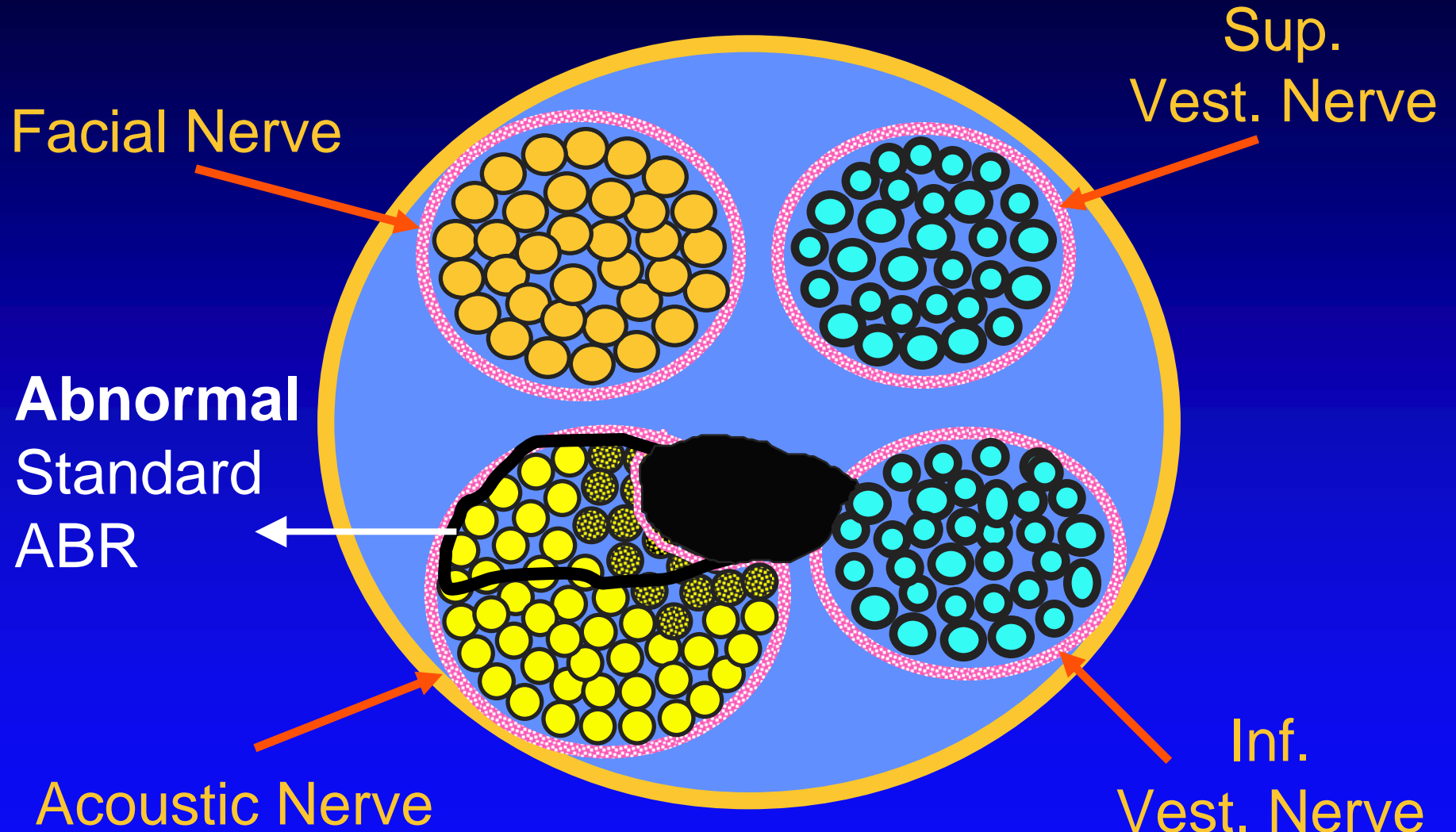


Spoendlin and Schrott (1989)

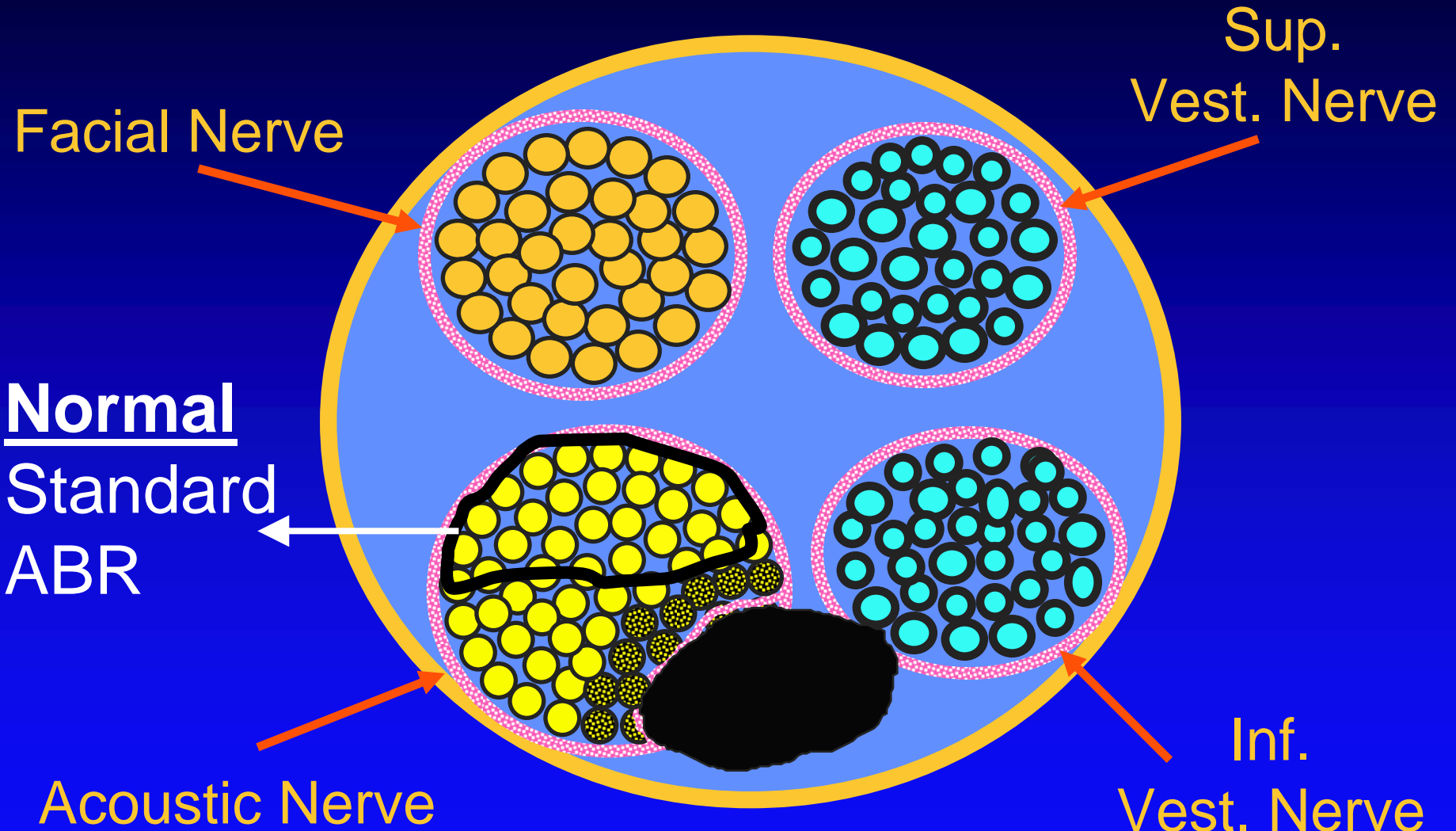
Medium or Large Tumor in IAC



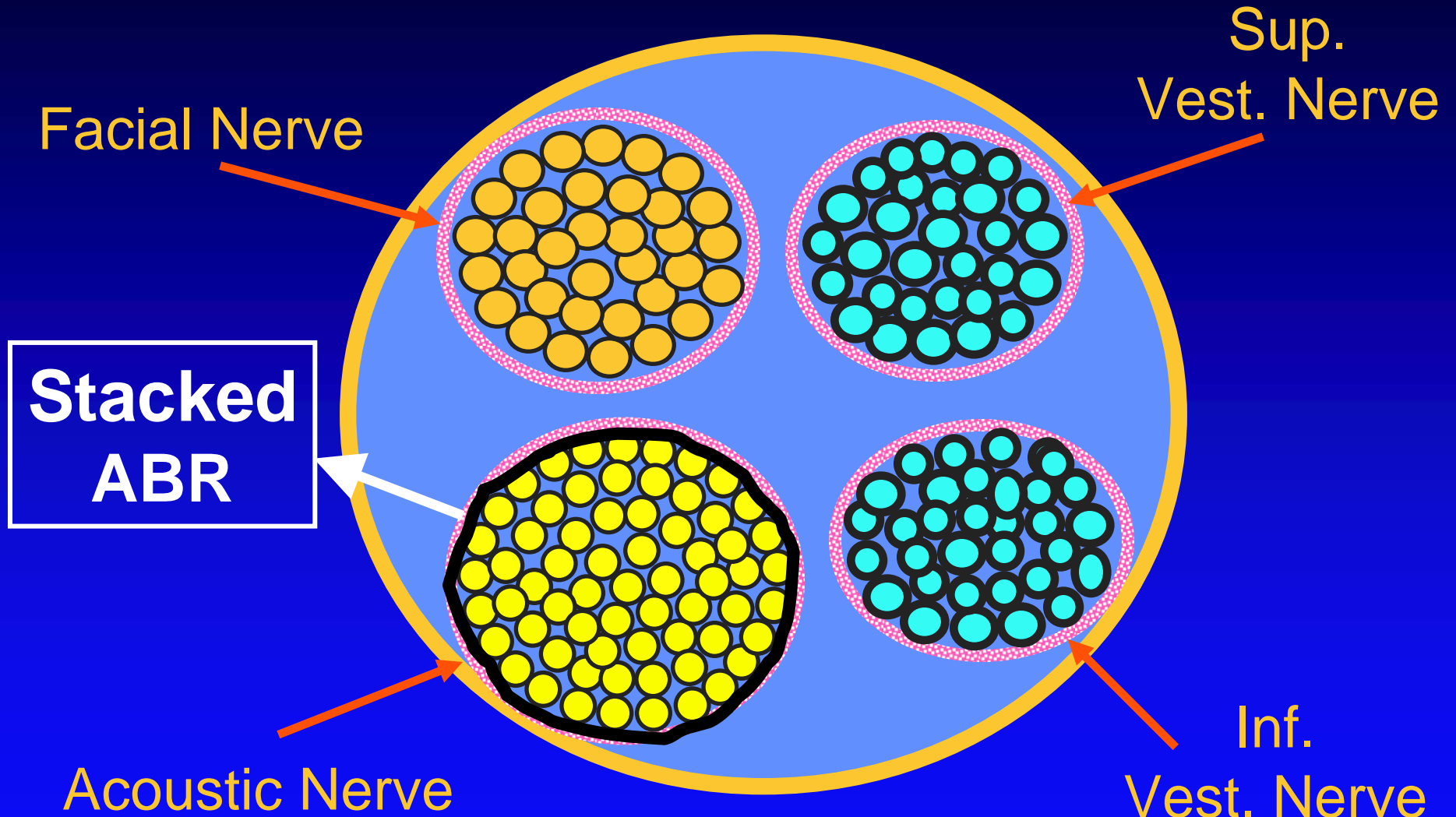
Small Tumor in IAC



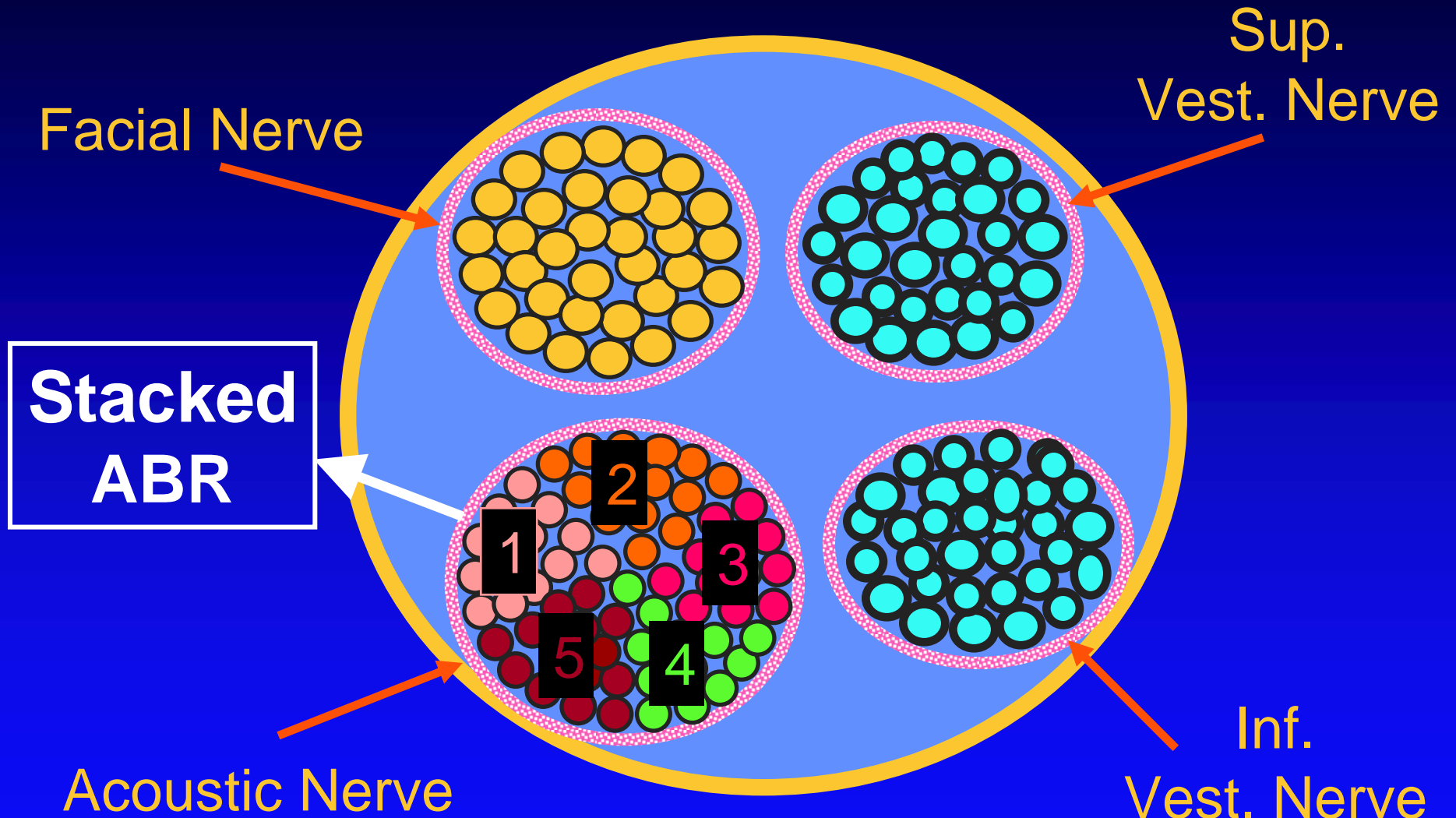
Small Tumor in IAC



Normal IAC



Normal IAC



Diagnostic Test: If you add the activity from each of the five areas, is the amplitude normal?

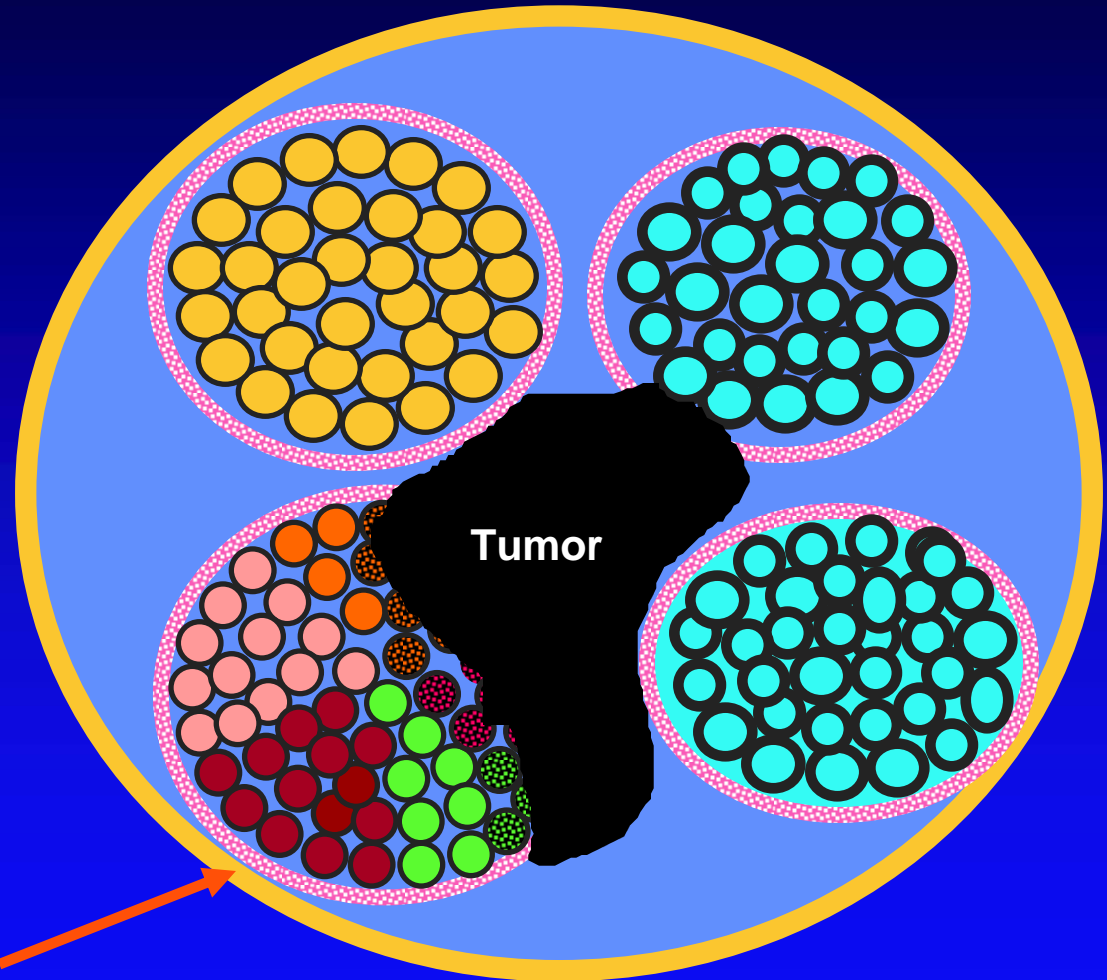
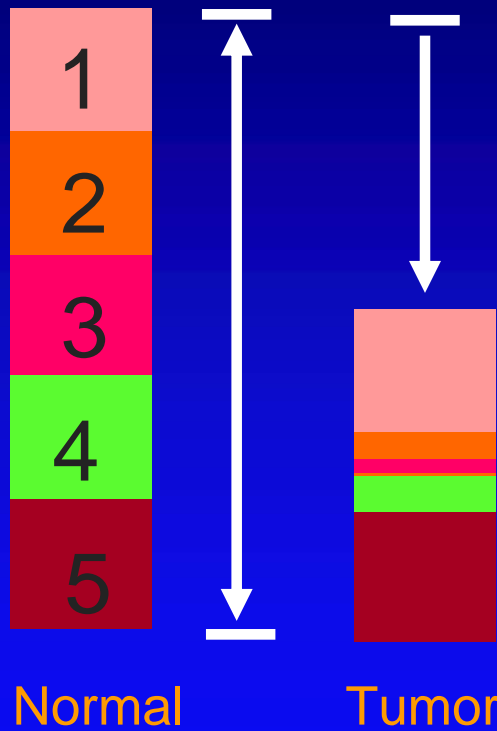
Activity from area 1
+
Activity from area 2
+
Activity from area 3
+
Activity from area 4
+
Activity from area 5



Normal Amplitude

Medium or Large Tumor in IAC

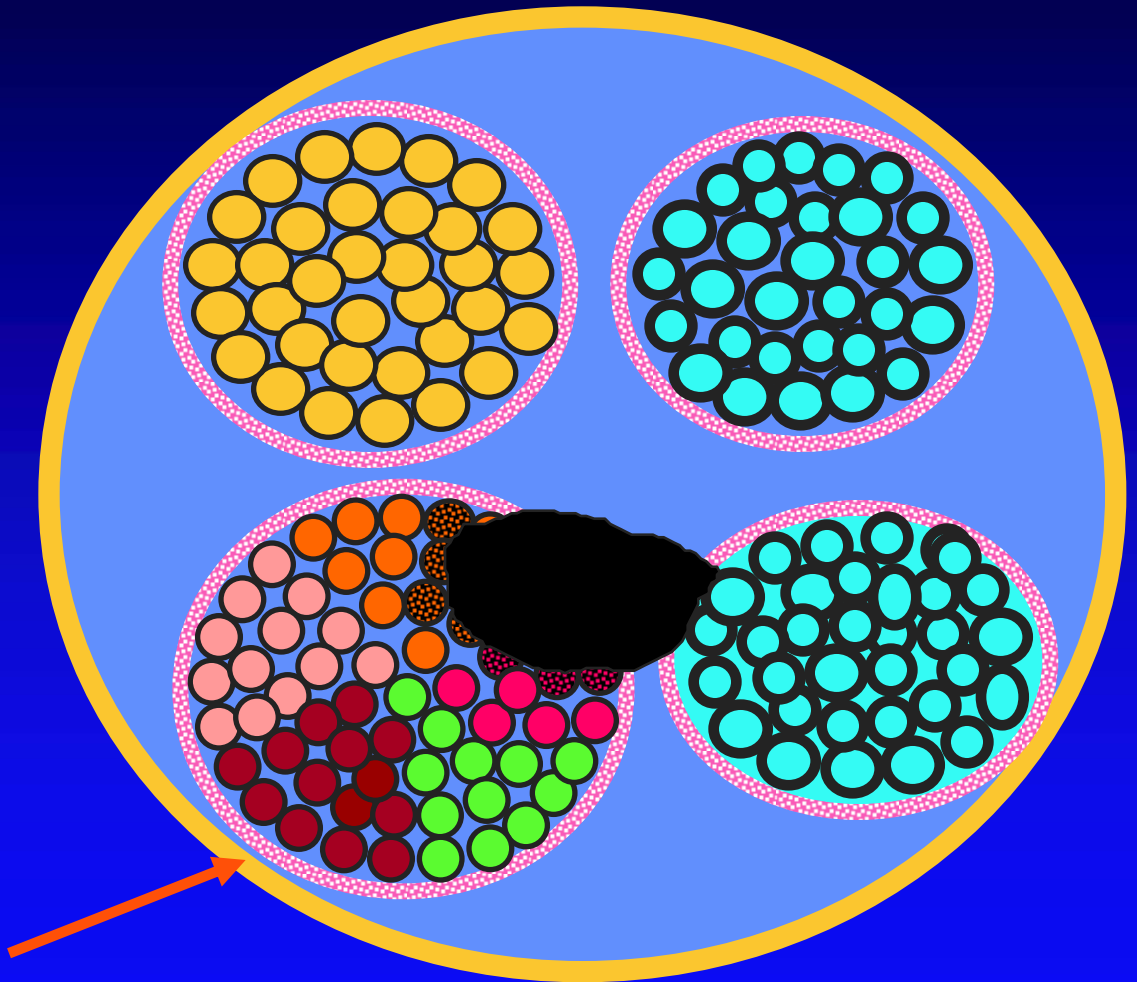
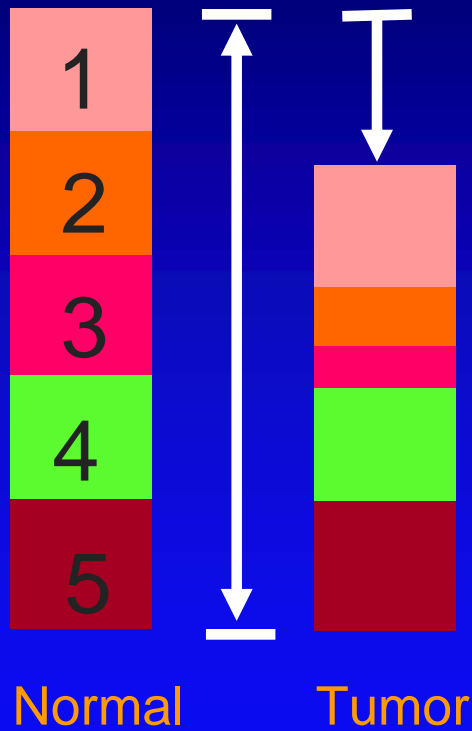
Abnormal
Stacked ABR



Acoustic Nerve

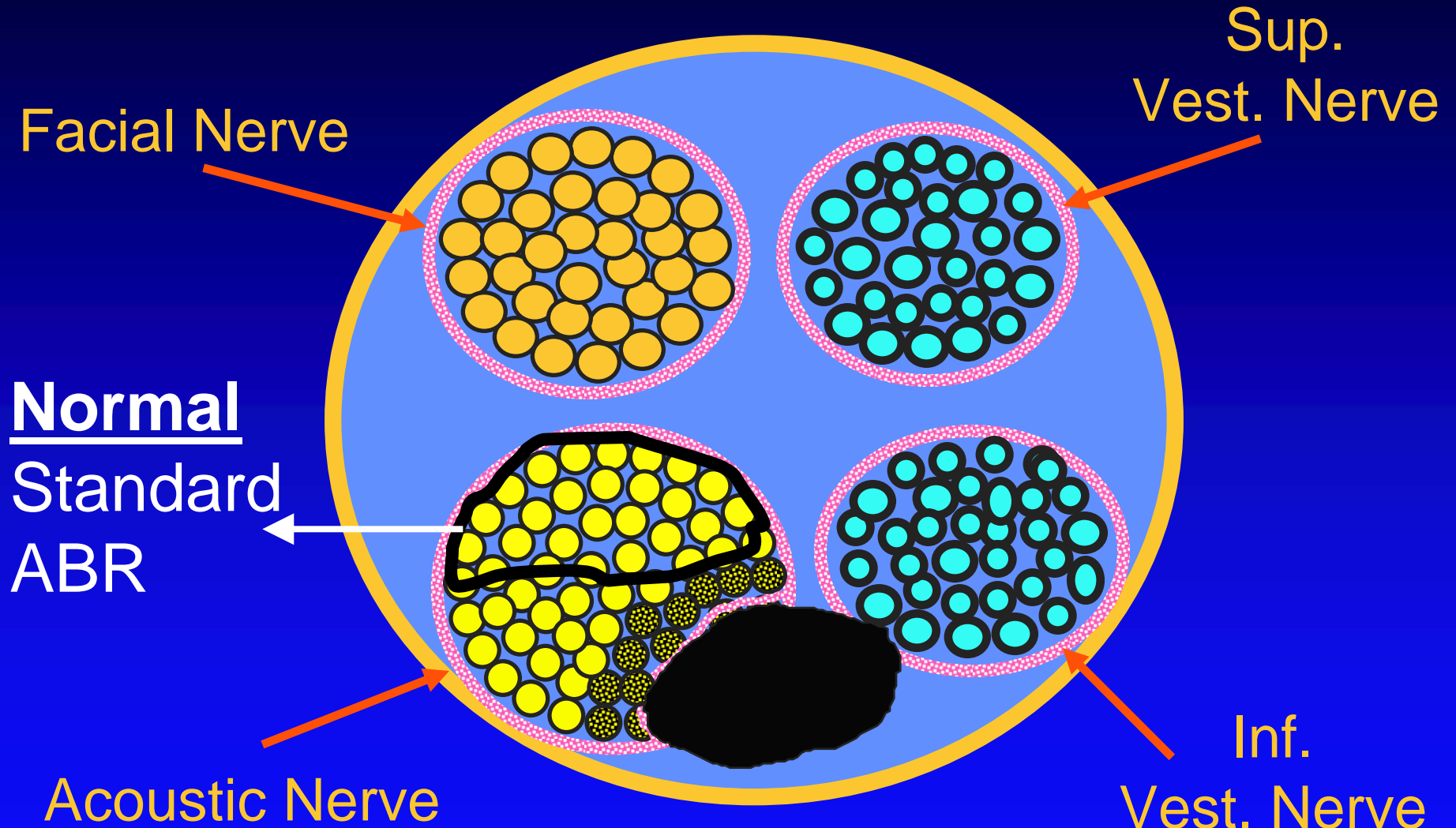
Small Tumor in IAC

Abnormal
Stacked ABR



Acoustic Nerve

Small Tumor in IAC Missed by Standard ABR



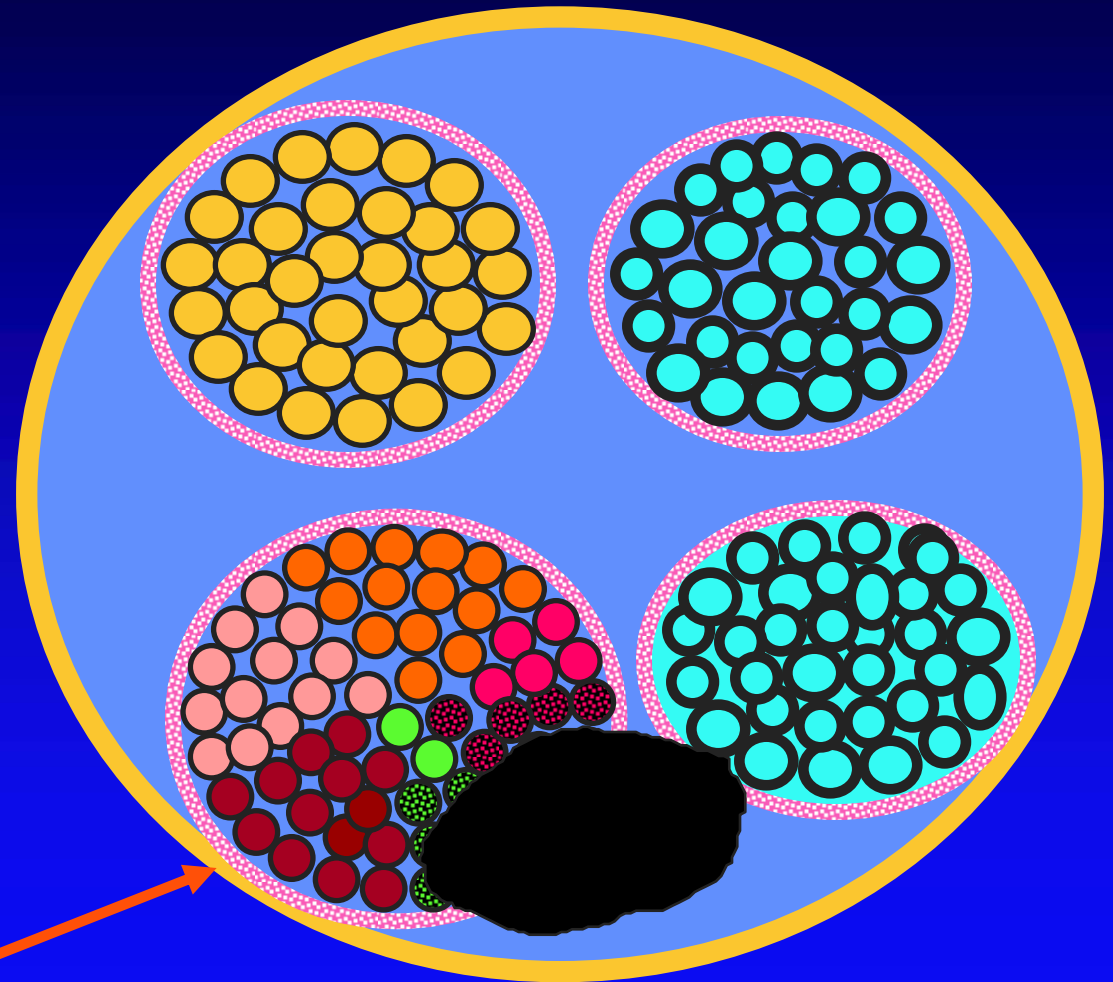
Small Tumor in IAC

Abnormal
Stacked ABR



Normal

Tumor



Acoustic Nerve

Stacked ABR Measure

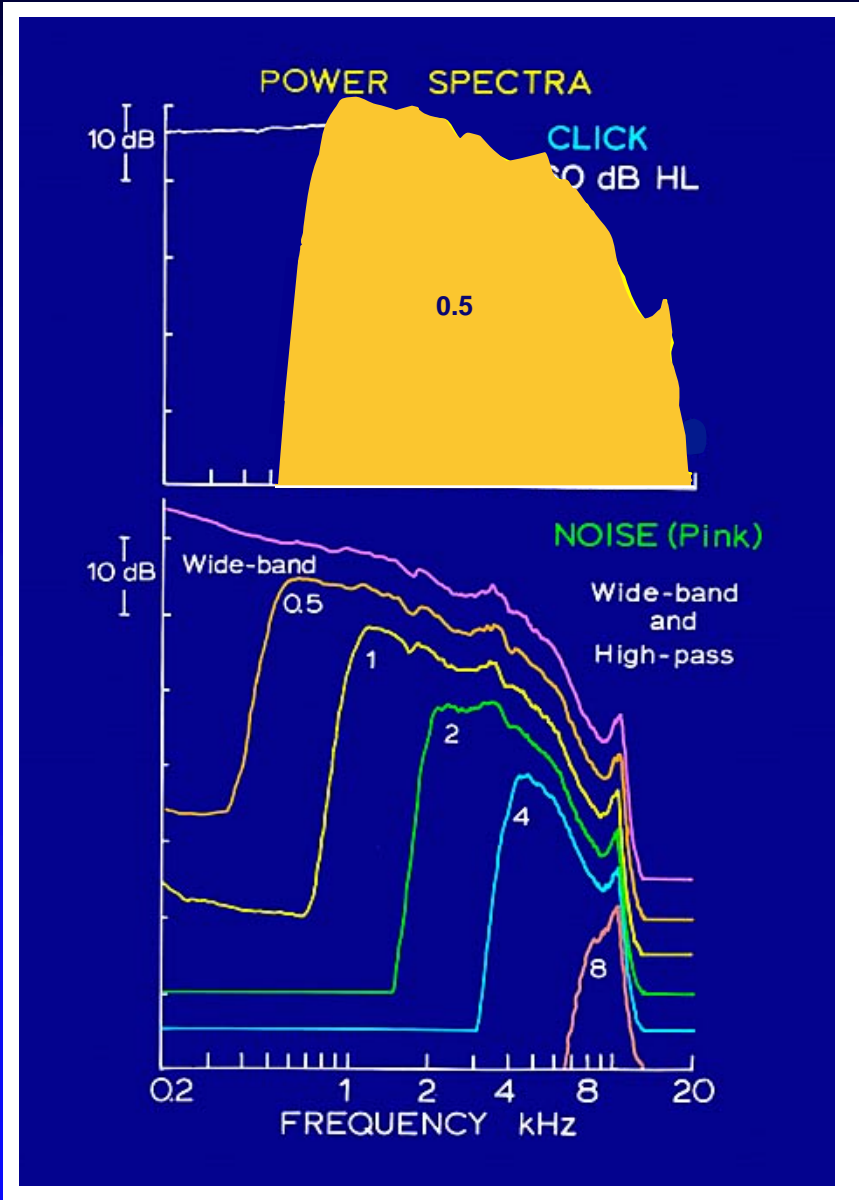
Requirements

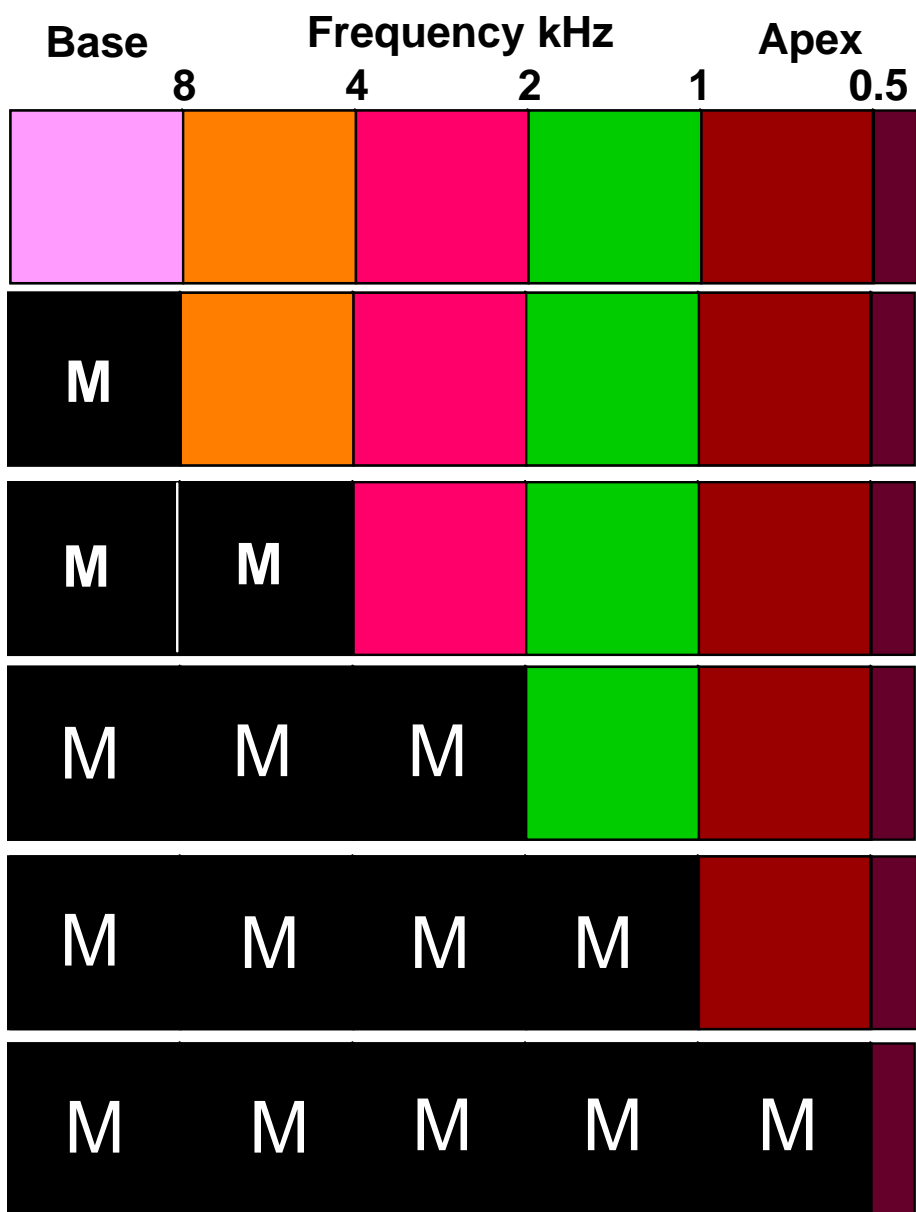
Proposed Methods

- | | | |
|---|----|--------------------------------|
| 1. An auditory signal that stimulates essentially all frequency regions of the cochlea | => | Wide-band Click |
| 2. A method for separating the responses from different frequency regions of the cochlea | => | The Derived-band ABR Technique |
| 3. A procedure for summing the responses to approximate total neural activity | => | The Stacking Technique |

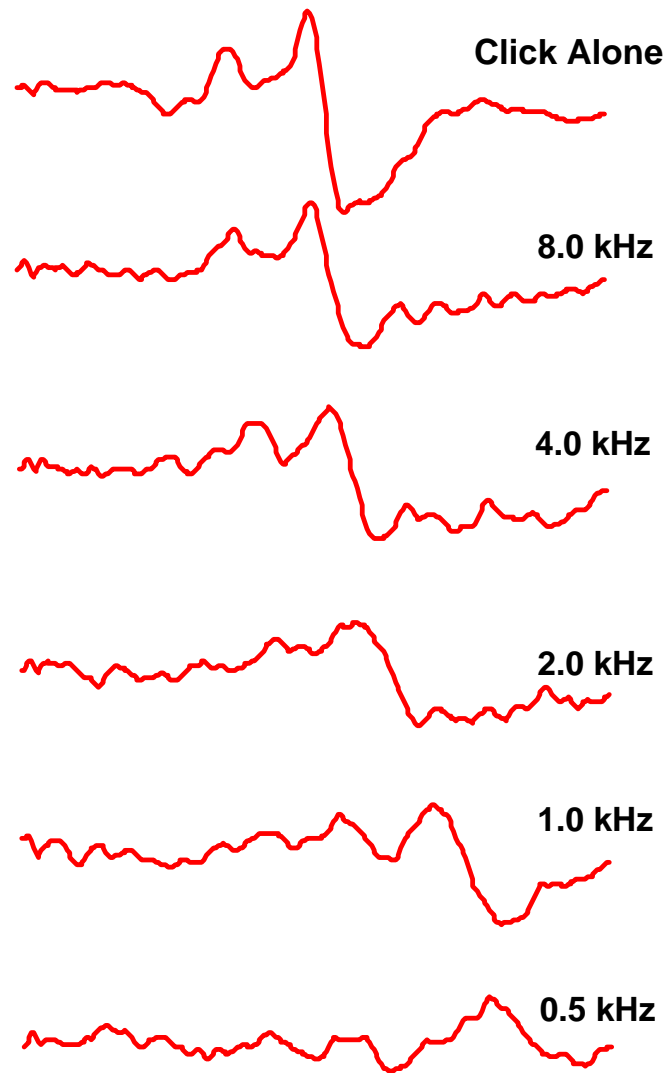
Click

High-pass
Masking Noise
(8.0, 4.0, 2.0, 1.0, and 0.5 kHz)



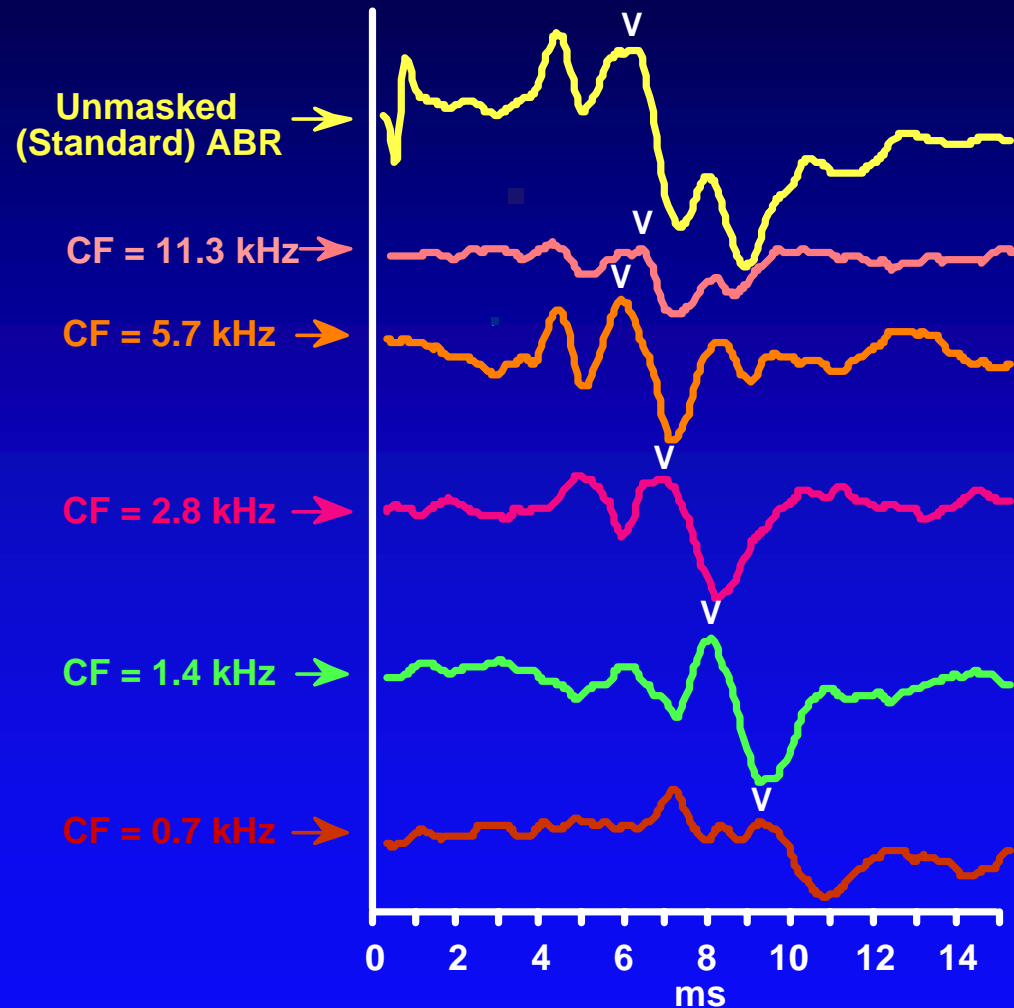


Click Alone and High Pass Noise Responses



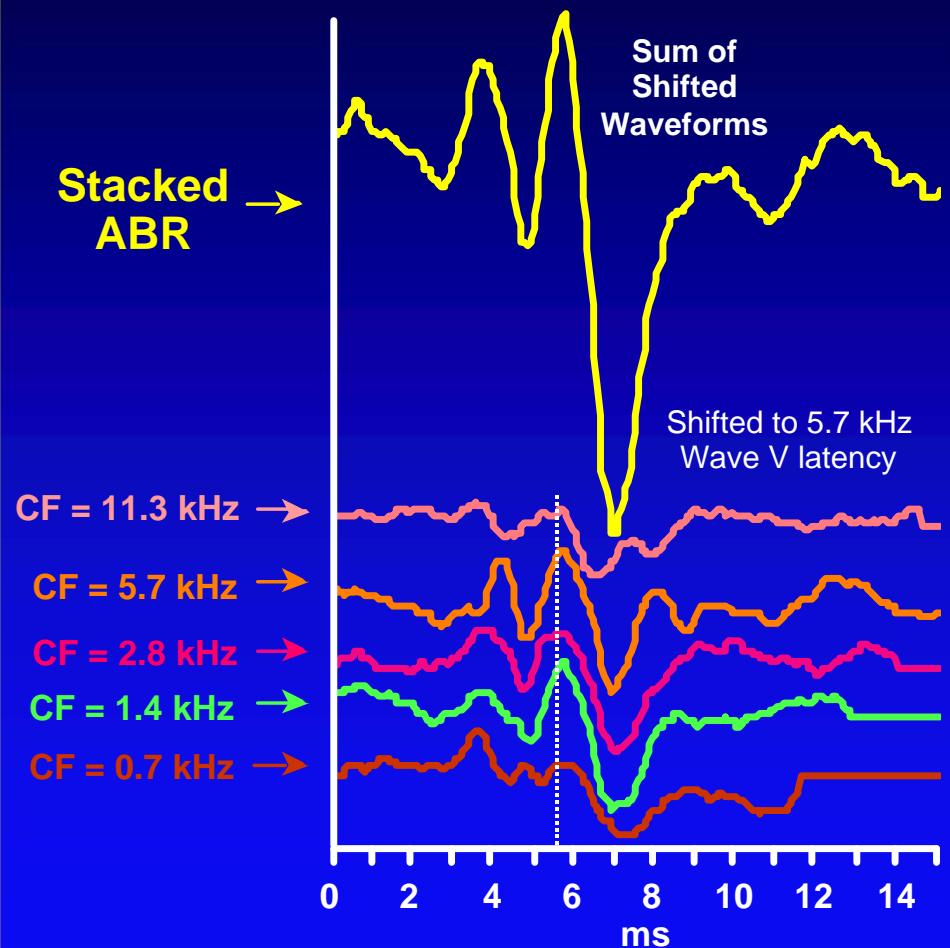
The Derived-band ABR Technique

The derived-band technique uses subtraction of the click alone response and the five different high pass responses to obtain five derived-band ABRs that reflect the neural contributions from five different octave-wide frequency regions of the cochlea.

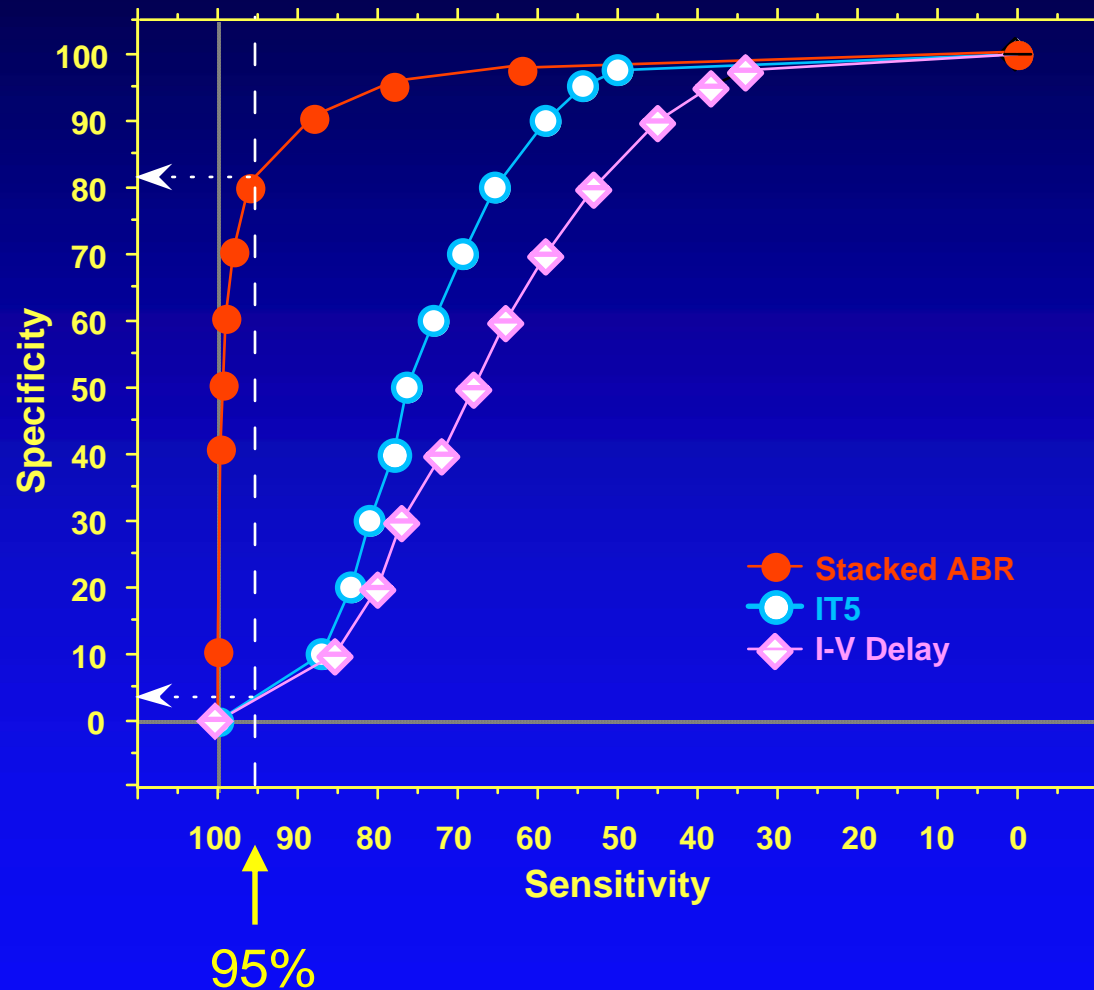


The Stacking Technique

- The **Stacked ABR** is formed by first temporally **aligning wave V** of the derived-band ABRs, then **summing the responses**.
- Aligning the derived-band ABRs eliminates phase cancellation of lower frequency activity. Thus, the Stacked ABR amplitude reflects activity from **all** frequency regions of the cochlea, not just the high frequencies.
- Reduction of **any** neural activity due to a tumor, even a small tumor, will result in a reduction of the Stacked ABR amplitude.



Standard vs Stacked ABR Measures



For 95% sensitivity

(that is, for correct identification of 95 out of every 100 small tumors):

The **IT5** and **I-V Delay** measures have less than 5% specificity

(that is, the IT5 and I-V Delay correctly identify less than 5 out of every 100 non-tumor patients),

But the Stacked ABR has 83% specificity

(that is, the Stacked ABR correctly identifies 83 out of every 100 non-tumor patients)!

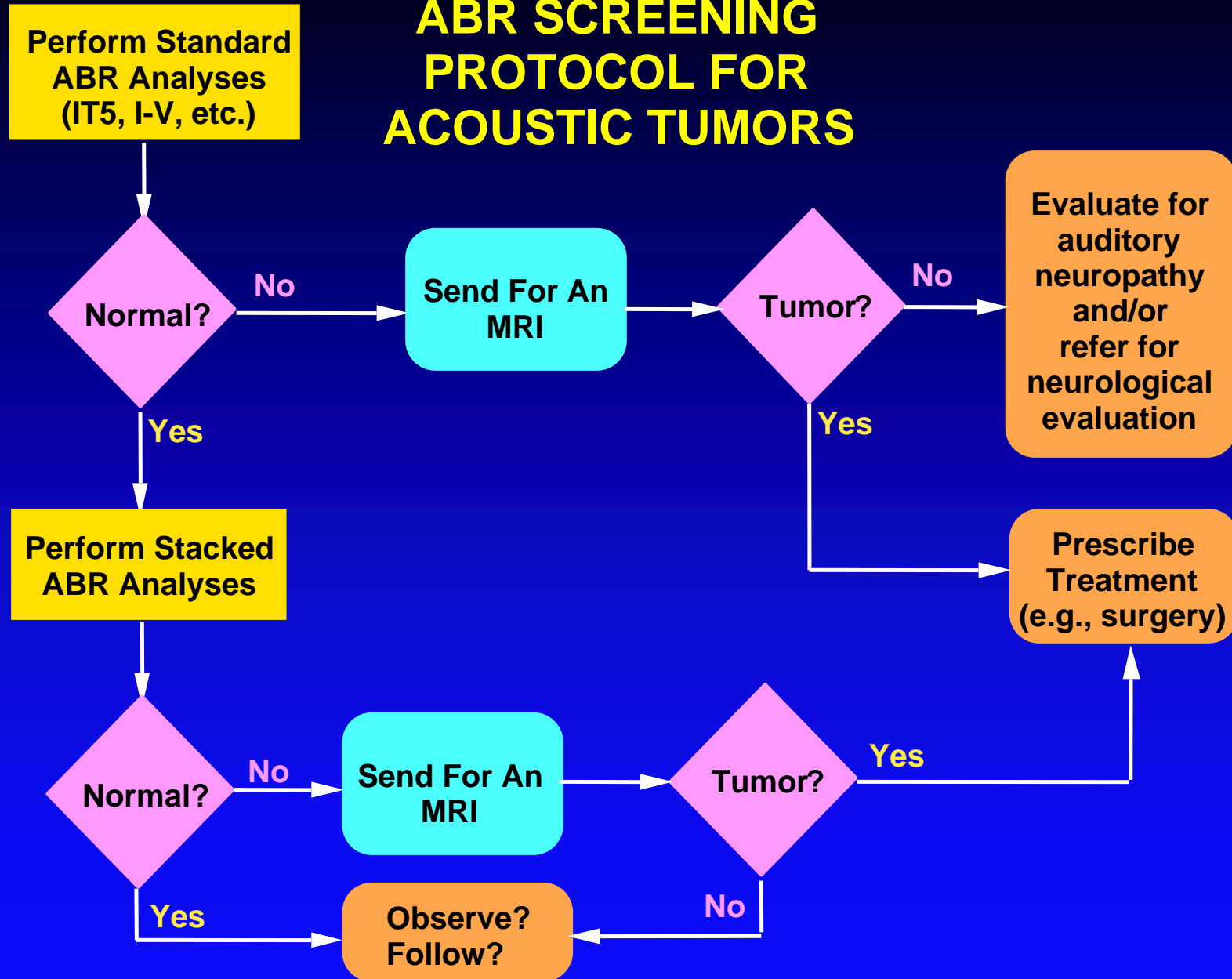
CONCLUSION

The Stacked ABR appears to have better sensitivity and specificity than the standard ABR for small (≤ 1 cm) tumors.

In other words, the Stacked ABR is better at :

1. detecting small tumors, and
2. decreasing the number of misdiagnosed non-tumor patients (i.e., decreasing the number of false-positives referred for MRI).

ABR SCREENING PROTOCOL FOR ACOUSTIC TUMORS



Breaking News: Meniere's Disease

- Symptoms of early Meniere's disease and small acoustic tumors are similar.
- There's a pattern in the high pass responses of patients with Meniere's disease that may be used to help with the differential diagnosis.

Endolymphatic Hydrops

```
graph TD; A[Endolymphatic Hydrops] --> B[Alters Basilar Membrane Parameters (e.g., stiffness, fluid column height, etc.)]; B --> C[Changes how cochlea processes auditory stimuli];
```

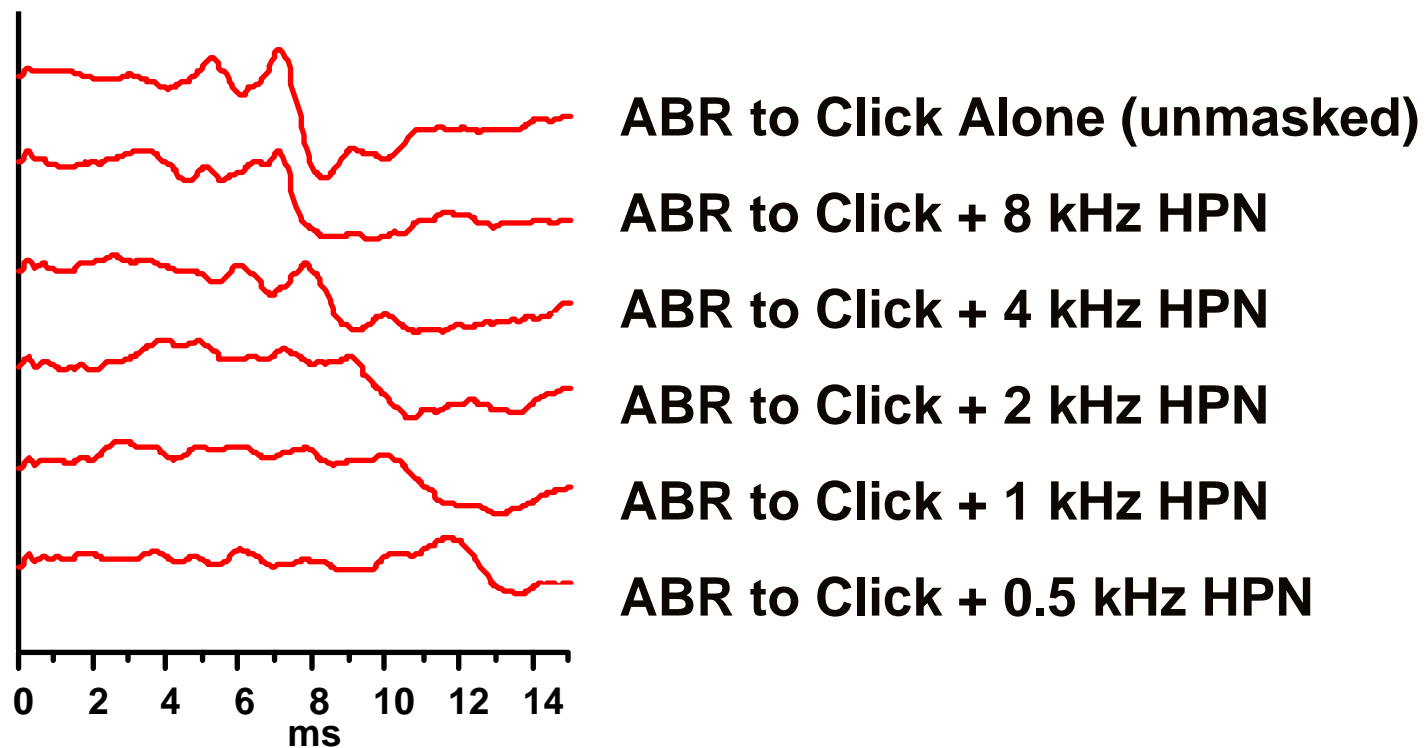
Alters Basilar Membrane Parameters
(e.g., stiffness, fluid column height, etc.)

Changes how cochlea processes
auditory stimuli

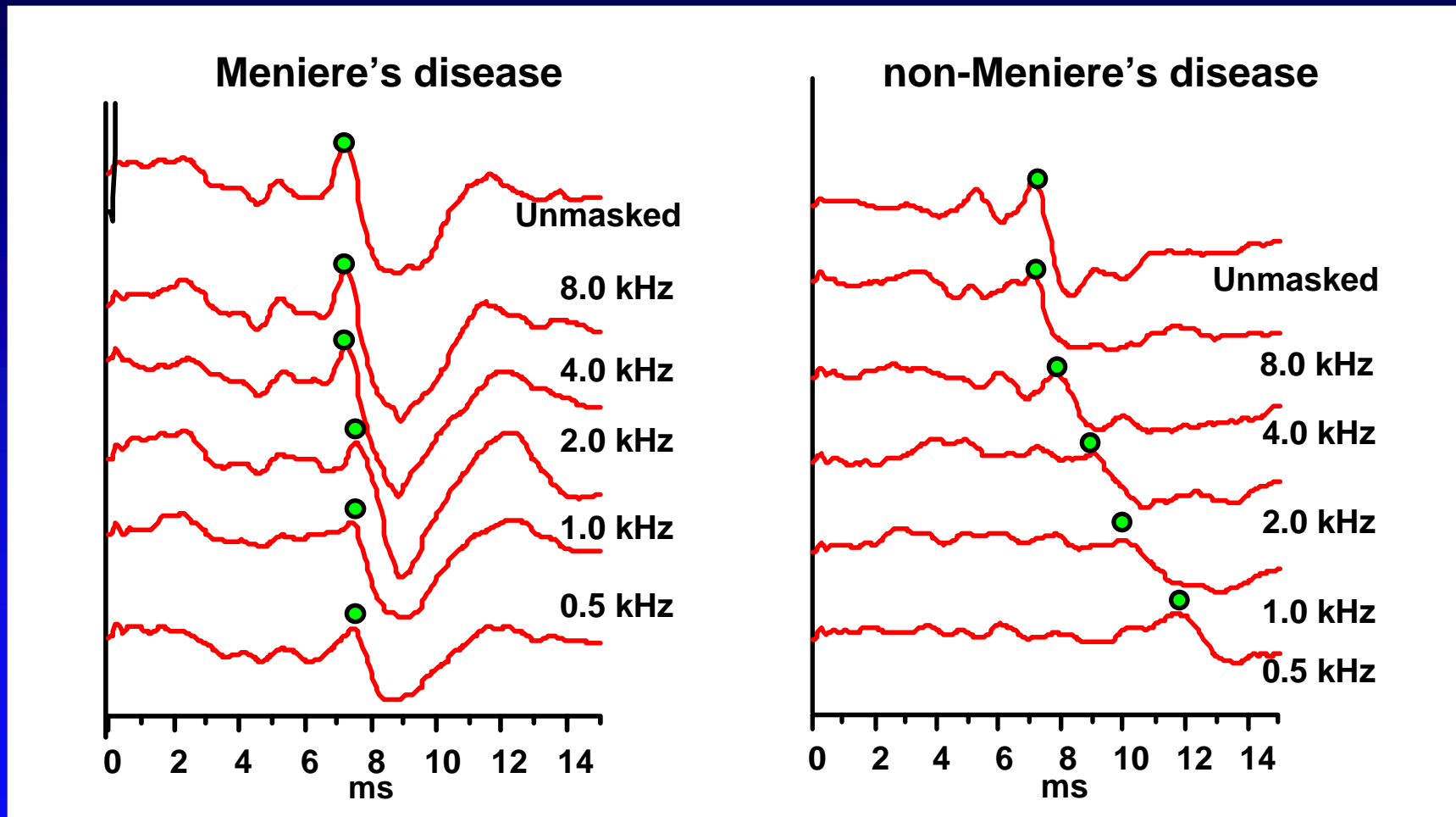
In Meniere's disease, we think that:

- Cochlear hydrops alters the response properties of the basilar membrane.
- Low frequency masking noise is less effective for masking activity in higher frequency regions.
- Thus, we observe undermasking in the high pass responses.

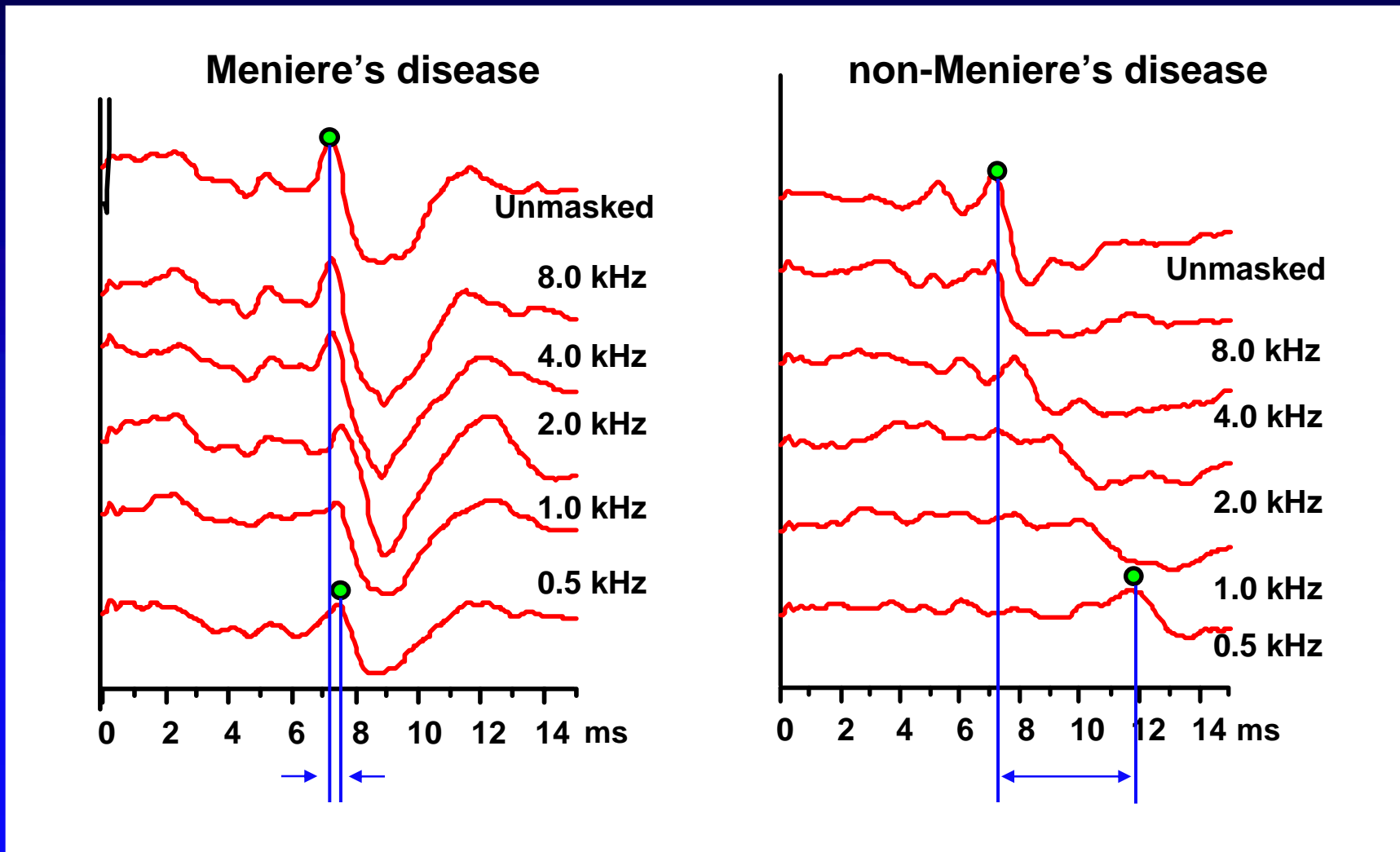
Click Alone (Unmasked) and High Pass Noise (HPN) Responses



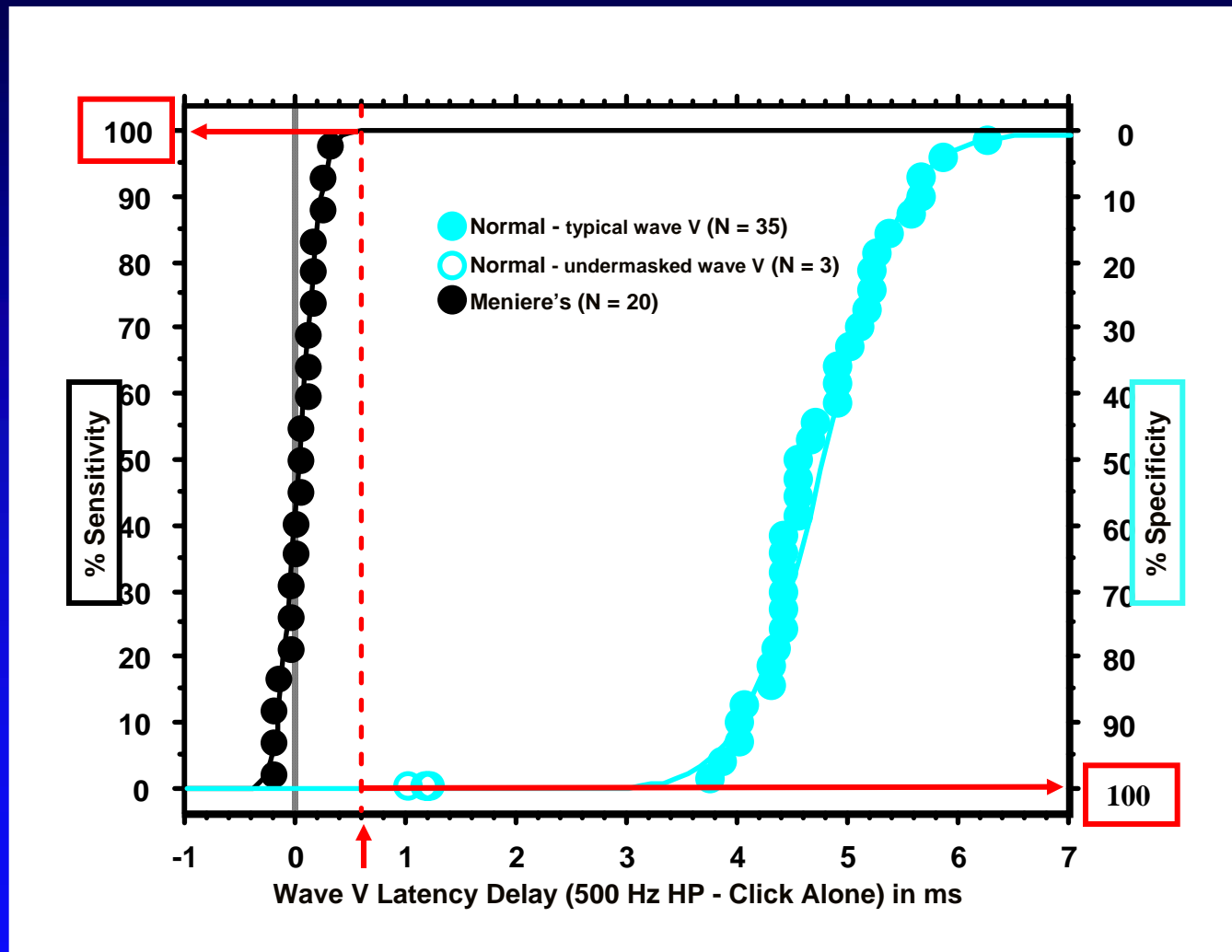
Undermasking in Meniere's Disease



Wave V Latency Delay (500 Hz HP – Click Alone)



Wave V Latency Delay (500 Hz HP – Click Alone)



Current Status

- In addition to the wave V latency delay, we are investigating other measures of this undermasking phenomenon.
- We are also analyzing data from non-Meniere's disease subjects with hearing loss and patients diagnosed with cochlear hydrops, not Meniere's disease.
- Preliminary results show very good separation of Meniere's disease/cochlear hydrops patients and non-Meniere's disease subjects.

IMPORTANT!

- Do not confuse the Stacked ABR method with this method for evaluating Meniere's disease.
- The Stacked ABR is for small tumor detection and is not used for Meniere's disease assessment.
- Stacked ABR uses the sum of the aligned derived-band (subtracted) ABRs while the Meniere's test uses only the high-passed noise masked responses to clicks.

Staff Acknowledgements

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Support

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NIH/NIDCD 2R44 DC04141 Raviv (PI)

NIH/NIDCD R01 DC03592 Don (PI)

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Don M, Masuda A, Nelson RA, and Brackmann DE (1997). Successful Detection of Small Acoustic Tumors Using the Stacked Derived Band ABR method. *Am J Otolaryngol.*; 18: 608-621.

Don M and Kwong B (2002). Auditory Brainstem Response: Differential Diagnosis. In: Katz J, Eds. *Handbook of Clinical Audiology*, Fifth Edition. Pennsylvania: Lippincott Williams & Wilkins Publishing; pp. 274-297.

Don M (2002). Auditory brainstem response testing in acoustic neuroma diagnosis. *Current Opinion in Otolaryngology & Head and Neck Surgery* 10:376-381.

Don M, Kwong B, Tanaka C, Brackmann DE, Nelson RA (2005) The Stacked ABR: A Sensitive and Specific Screening Tool for Detecting Small Acoustic Tumors (*Audiology & Neurotology* 10: 274-290)

Don M, Kwong B, Tanaka C (2005) A Diagnostic Test for Meniere's Disease and cochlear Hydrops: Impaired High-pass Noise Masking ABRs. (*Otology & Neurotology* 26: 711-722.)