
Longitudinal Changes in Hearing and Speech Perception in Older Adults

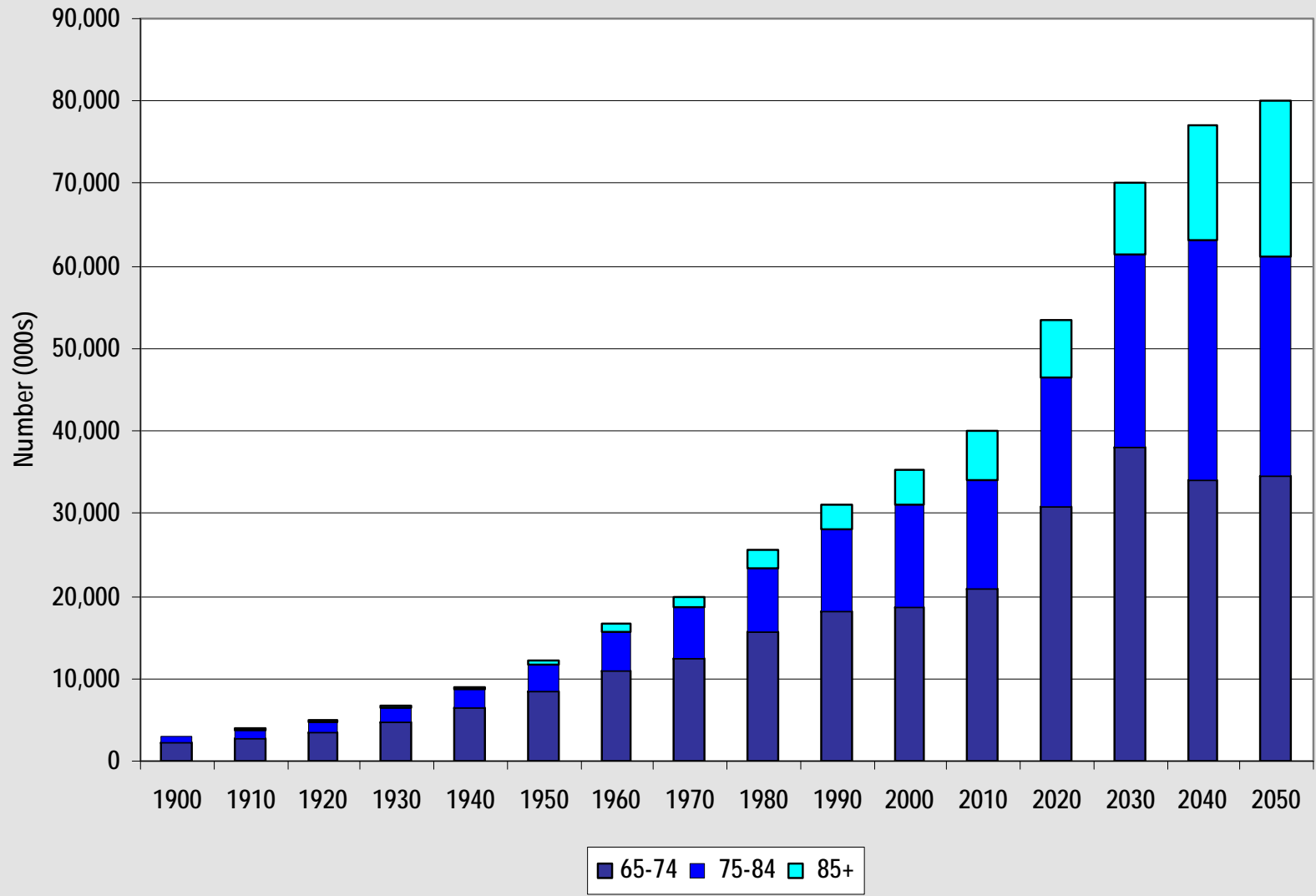
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Older Population by Age: 1900-2050

Source: U.S. Bureau of the Census



Program on Age-Related Hearing Loss

- Established as NIH Program Project Grant in 1987 to evaluate age-related changes in auditory function in **humans** and a **gerbil animal model**
 - NIH/NIDCD Clinical Research Center now focused exclusively on hearing in **older humans**
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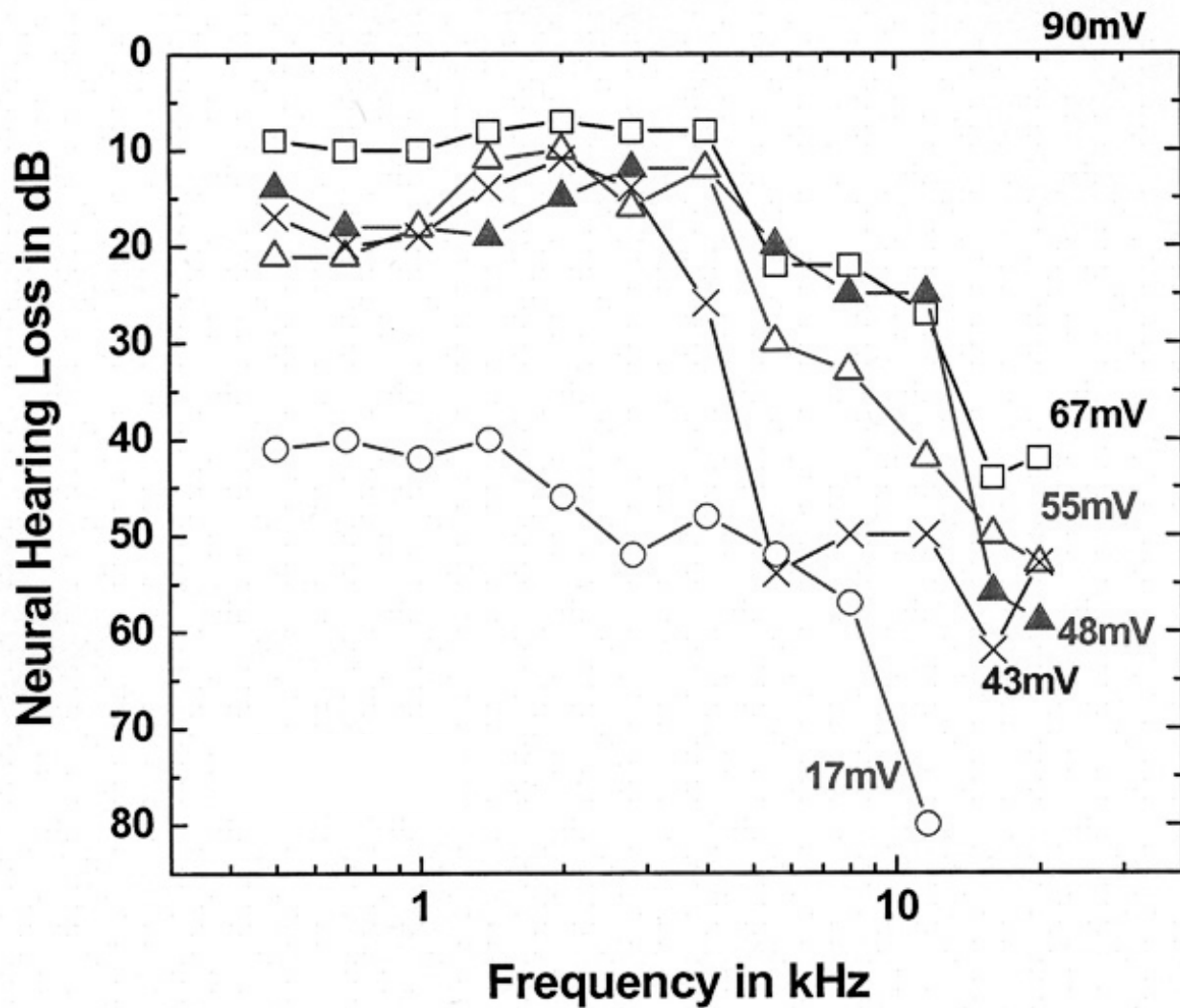
Outline

- Briefly review results from gerbil animal model of age-related hearing loss
 - Human subject database (21+ years)
 - **Cross-sectional** and **longitudinal** changes in hearing in older adults
 - **Longitudinal** changes in speech recognition in older adults, independent of changes in hearing
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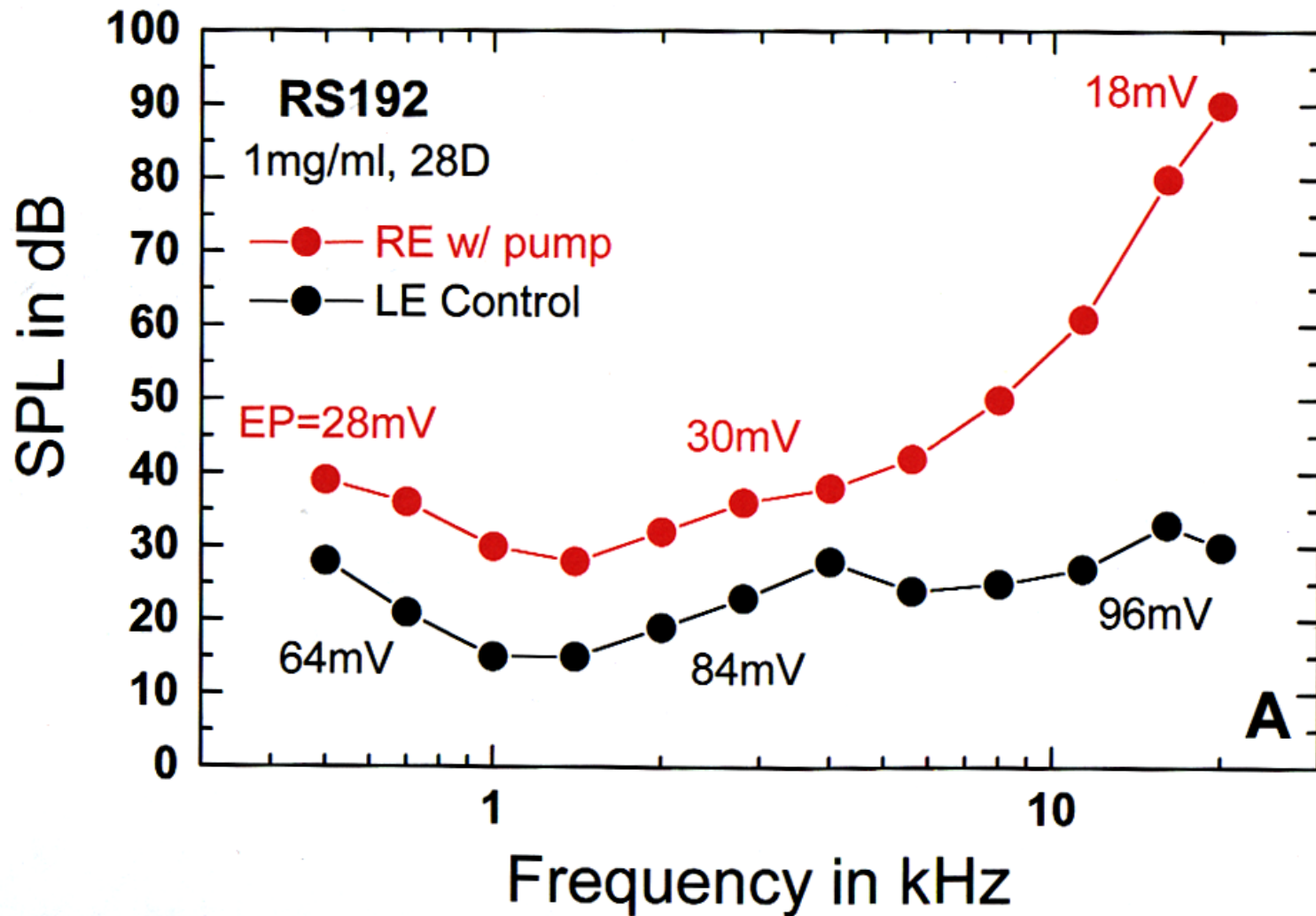
Gerbil animal model of presbycusis

- Only scattered hair cell loss (sensory presbycusis)
 - Primary degeneration of spiral ganglion neurons (neural presbycusis)
 - Systematic degeneration of cochlear lateral wall (stria vascularis, spiral ligament)
 - Metabolic presbycusis
 - Lateral wall responsible for production and maintenance of endocochlear potential (EP)
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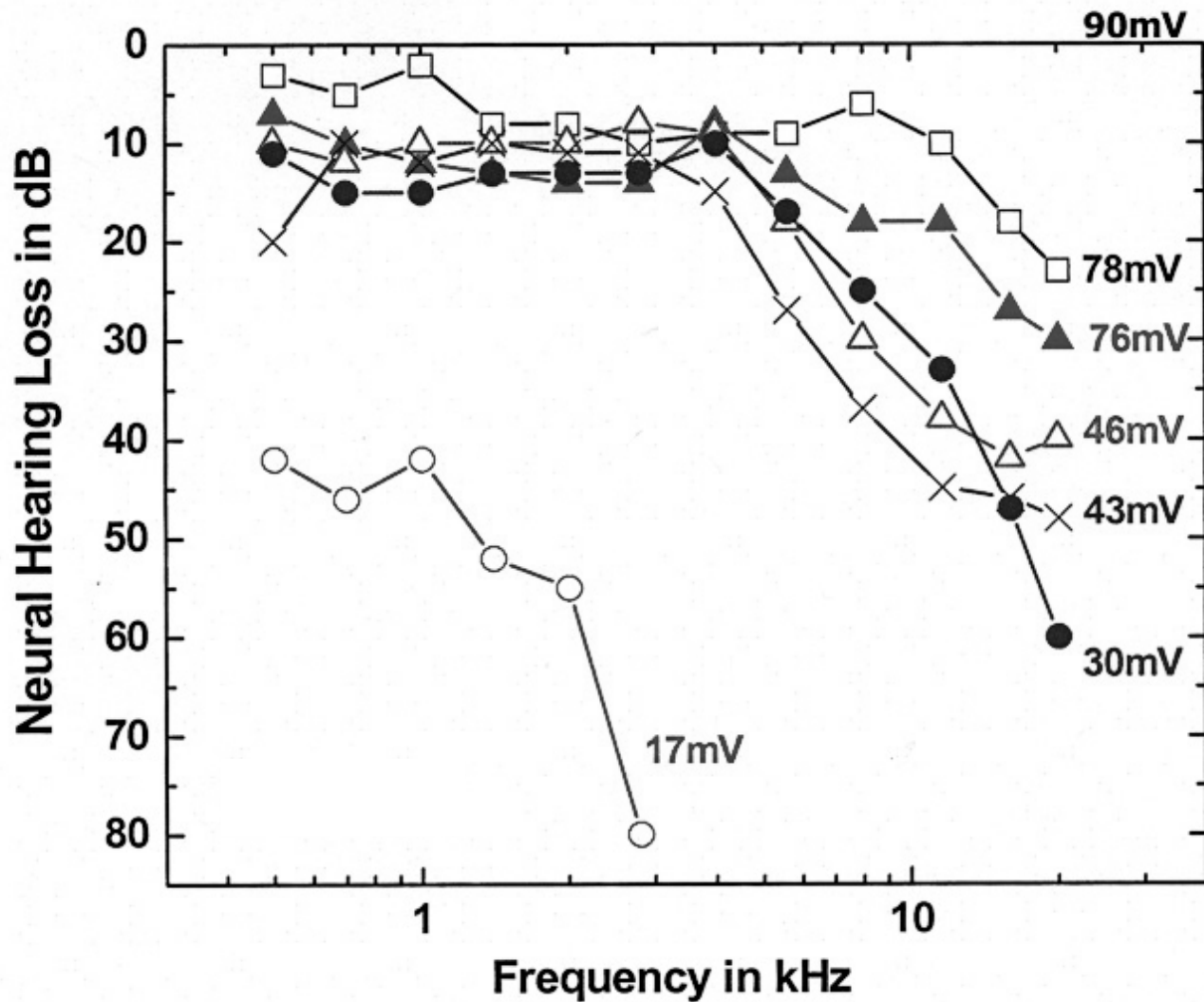
Hearing Loss of Quiet-Aged Gerbils



CAP Thresholds



Hearing Loss of Furosemide-Treated Gerbils



Gerbil animal model of presbycusis

- Decline in endocochlear potential (EP)
 - Reduces voltage available to cochlear amplifier
 - Reduces cochlear amplifier gain
 - Low frequencies: as much as 20 dB
 - High frequencies: as much as 60 dB
 - EP declines result in the characteristic audiogram of older gerbils (metabolic presbycusis)
 - Is this the case for older humans?

Human Subject Database

- Inclusion and Exclusion Criteria
 - 60 years or older (now 18 or older)
 - Hearing ability to provide measurable results
 - In good general health
 - Screened with Mini-Mental State Exam (MMSE)
 - No evidence of conductive hearing loss
 - No evidence of active otologic disease
-

Human Subject Protocol

- Audiometric Measures
 - Hearing for pure tones, including extended high frequencies
 - Ability to understand speech in quiet and in noise
 - Otoacoustic emissions
 - Upward and downward spread of masking
 - Middle ear function
 - Auditory brainstem responses
-

Human Subject Protocol

- Cognitive Measures
 - Attention
 - Working memory
 - Processing speed
 - Perceived workload
 - Brain imaging while listening to and understanding:
 - Low-pass filtered speech
 - Speech in background noise
-

Human Subject Protocol

- Questionnaires
 - Medical history
 - Prescription and over-the-counter drugs
 - Noise history
 - Hearing aid history
 - Hearing handicap (HHIE)
 - Tinnitus
 - Smoking
 - Handedness
 - Family pedigree for hearing loss
 - Otologic examination
-

Human Subject Protocol

- Blood measures
 - Clinical chemistries
 - Lipid profile
 - Hematology panel
 - Hormones (Estradiol, Progesterone – Female subjects)
 - C-reactive protein
 - Electrolyte panel - Discontinued
 - Immunoglobulin panel - Discontinued
 - Thyroid function – Discontinued
 - DNA extracted
 - To identify and characterize genes that are under- or over-expressed with age
-

Human Database Participants

	Total with any data		Total with longitudinal data		Currently Active	
Age Range	18-59	60-98	18-59	60-95	18-59	60-93
Female	85	483	10	235	14	201
Male	76	385	12	200	15	125
Total	161	868	22	435	29	326
Grand Total	1,029		457		355	

- Measures are made yearly or every 2-3 years

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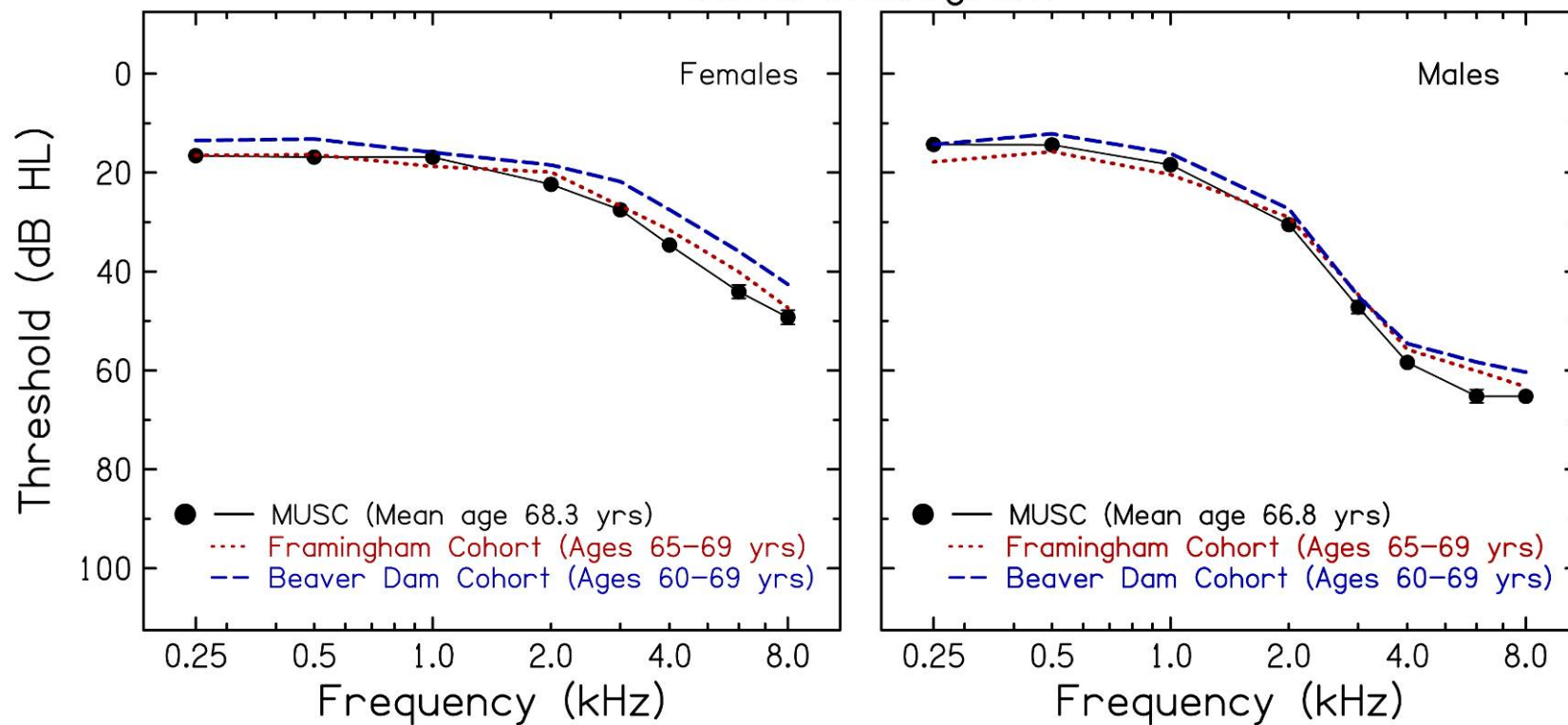
Longitudinal Study Design

- Advantages
 - Participants act as their own controls
 - Minimizes effects of uncontrollable factors
 - noise history
 - occupation
 - nutrition
 - Pre-existing health conditions
 - Measures age-related changes for groups and individuals (cross-sectional designs – groups only)
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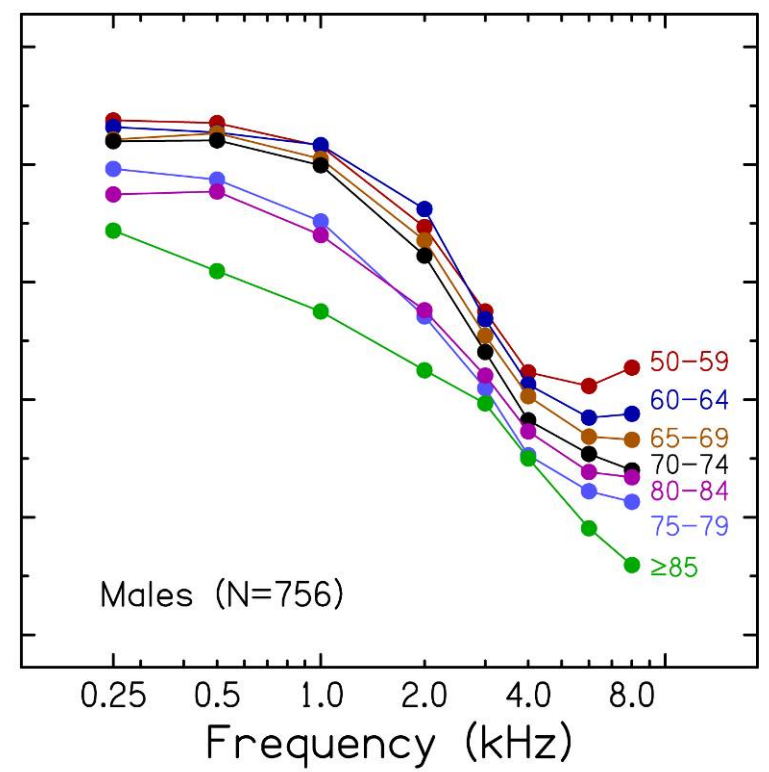
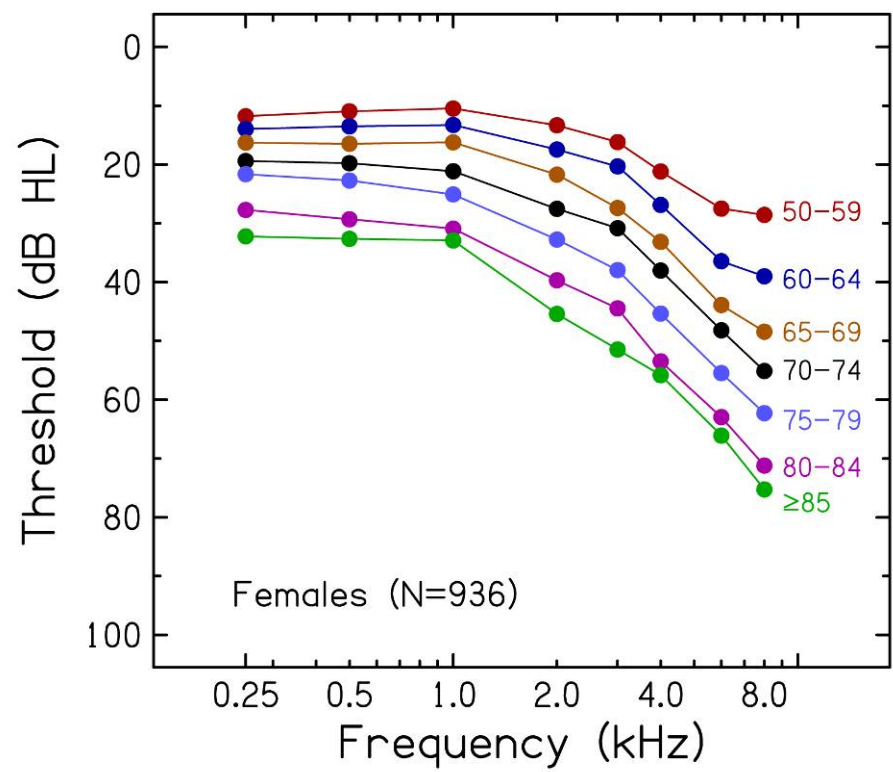
Longitudinal Study Design

- Disadvantages
 - Data collection takes many years
 - Must retain participants for long periods of time
 - Recruitment more difficult
 - Selective attrition
 - For longitudinal studies of aging
 - Healthier or higher performing participants may remain in the study longer
 - High cost
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Initial Audiogram

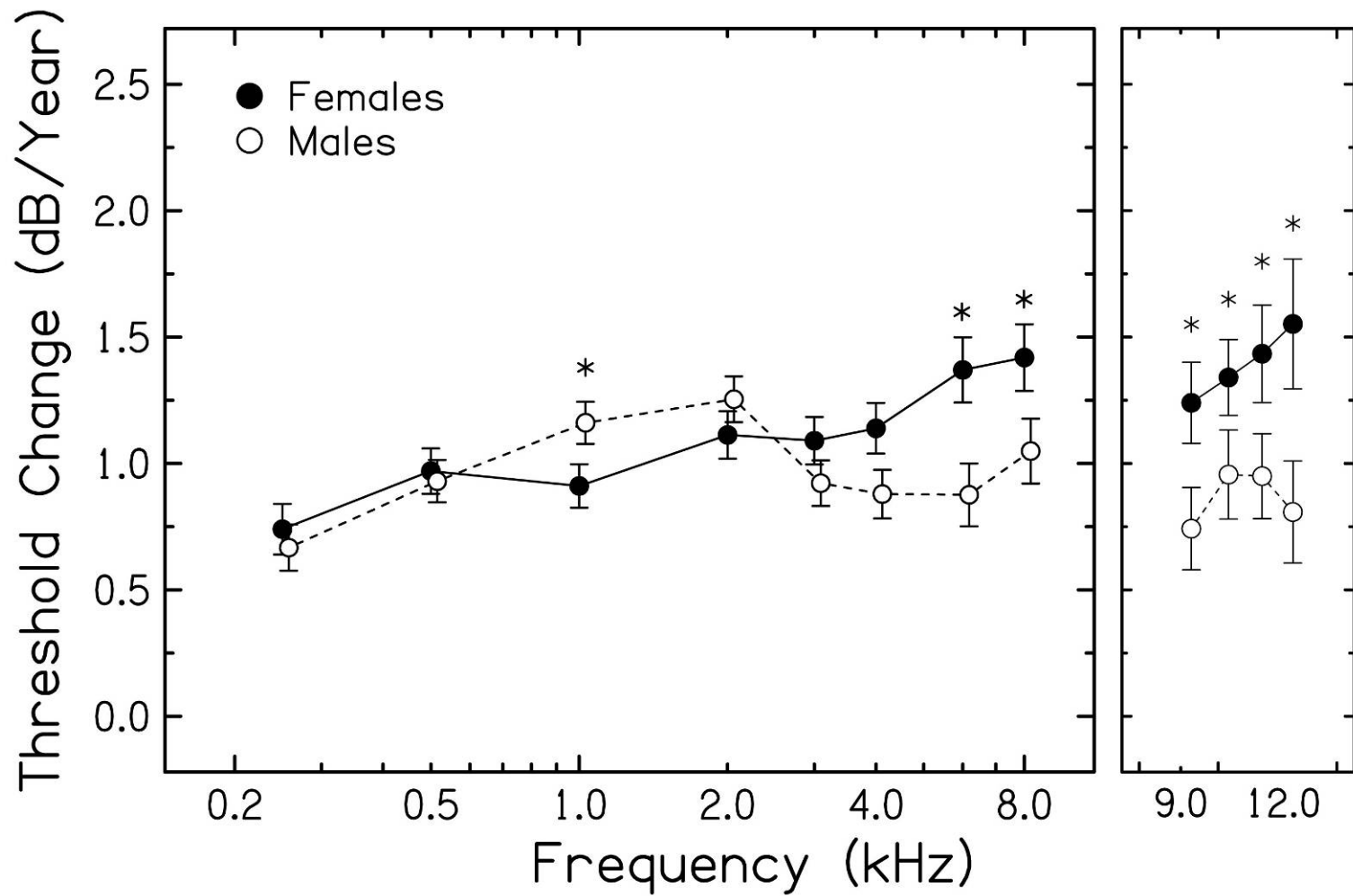


Cross Sectional



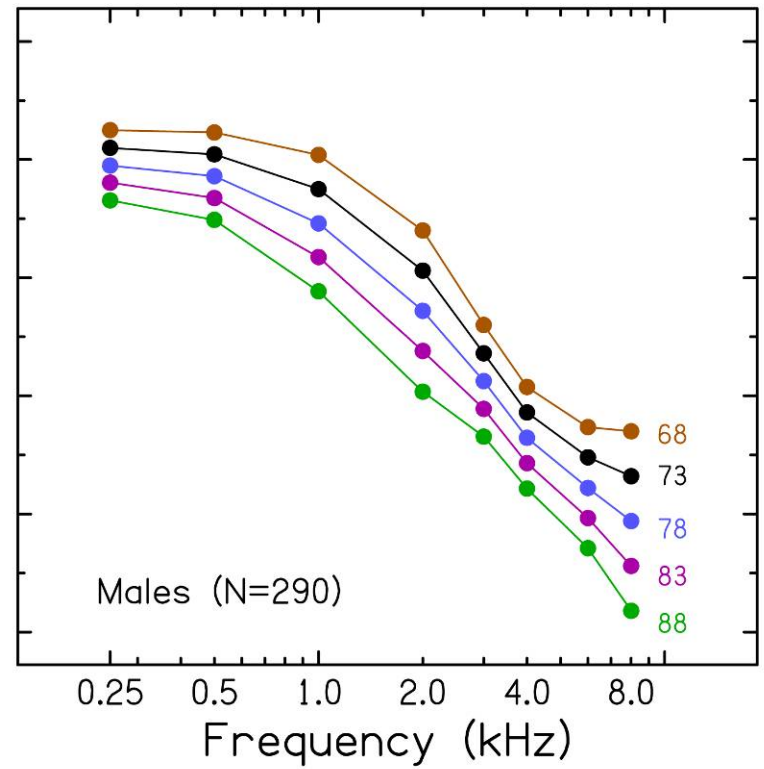
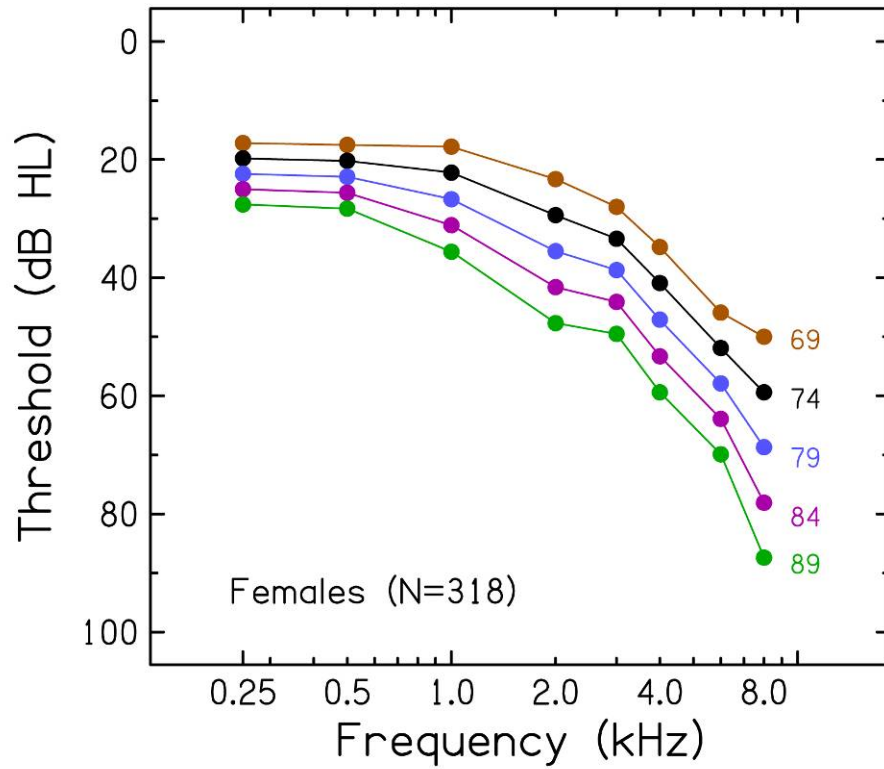
Longitudinal Changes in Thresholds

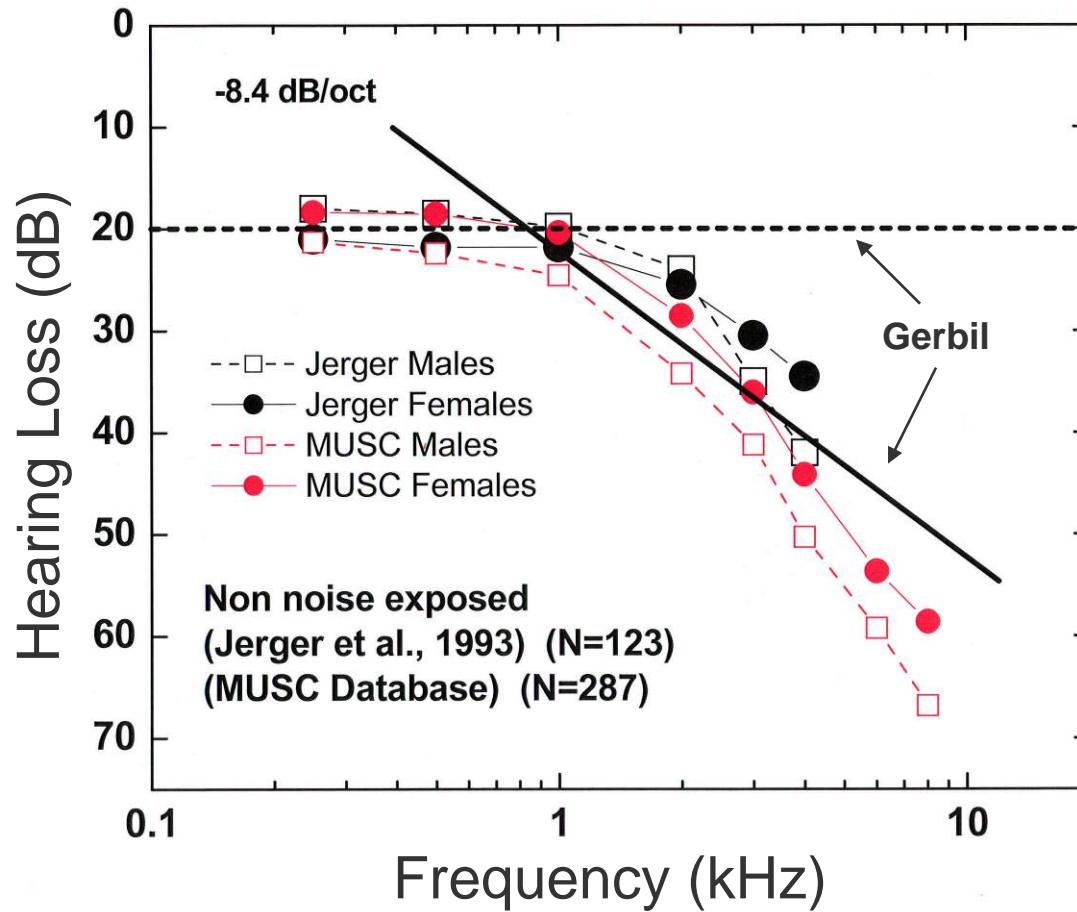
- Serial audiograms obtained over ≥ 3 years
 - 3,690 audiograms from 376 ears
 - Mean number of audiograms per subject: 9.8 (Range 2-21)
 - Mean time span: 6.4 years (Range 3-12)
 - Mean age at entry: 68.1 years (Range 60-81)
 - Mean current age: 74.5 years (Range 64-89)
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Lee F, Matthews L, Dubno J, Mills J (2005). Ear Hear 26, 1-11.

Longitudinal

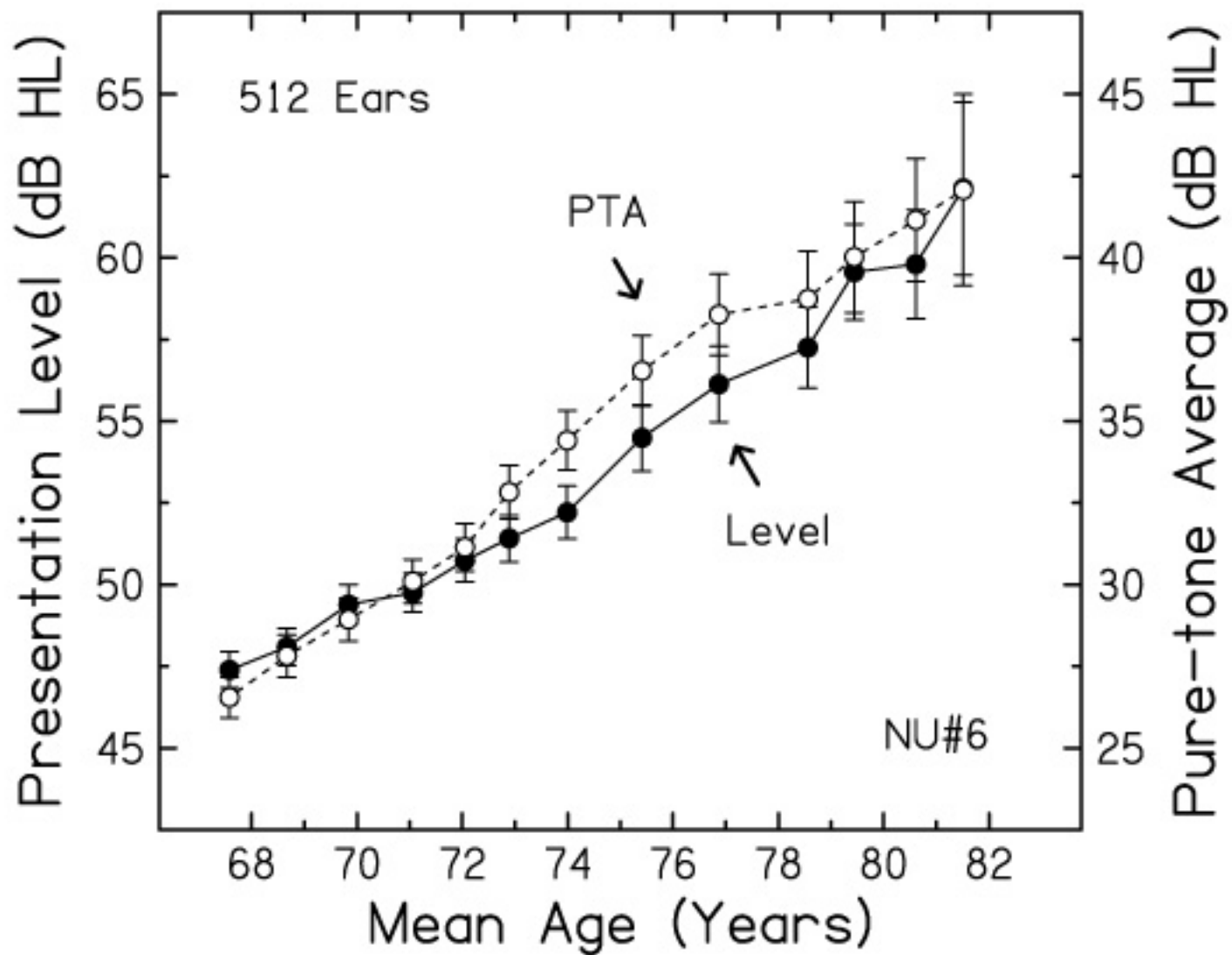




Schmiedt R, Lang H, Okamura H, Schulte B (2002). J Neurosci 22, 9643-9650.

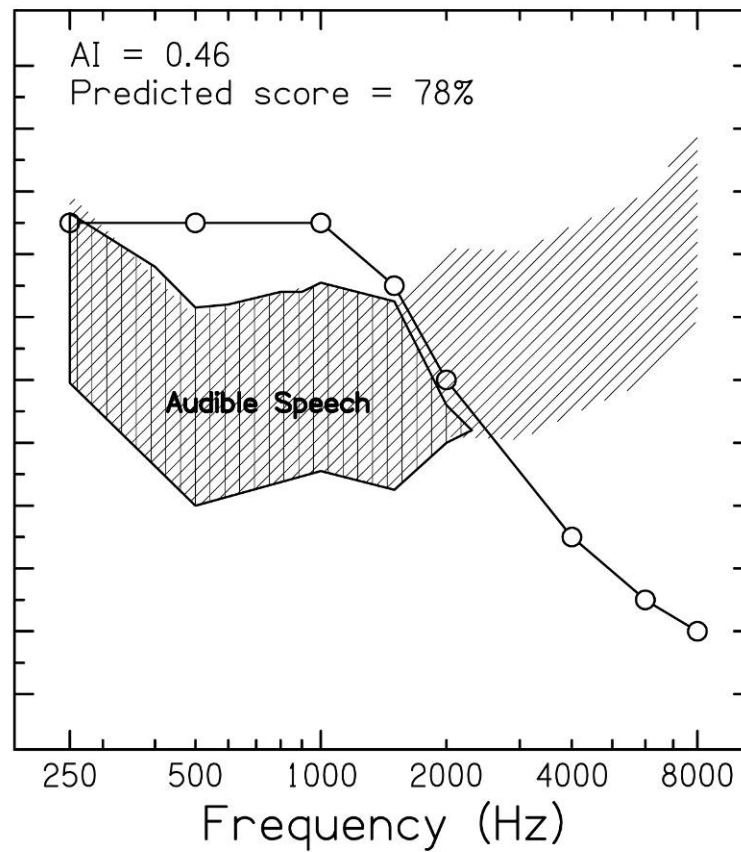
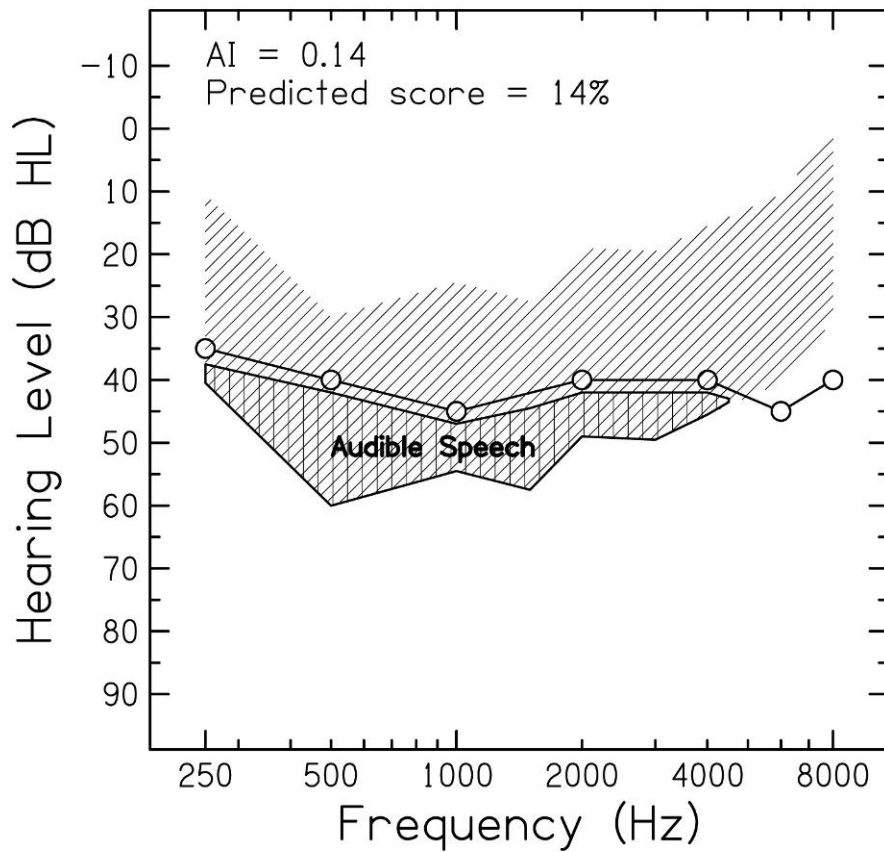
Longitudinal Changes in Speech Recognition

- SRT
 - Word recognition in quiet (NU-6)
 - Maximum word recognition (NU-6)
 - Recognition of sentences in noise (Speech Perception in Noise Test, SPIN)
 - Binaural word recognition (SSW)
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Rationale

- To assess age-related declines in word recognition
 - Using AI (weighted average speech audibility), predict word recognition scores for each subject for each time point
 - Compare measured and predicted scores for each time point
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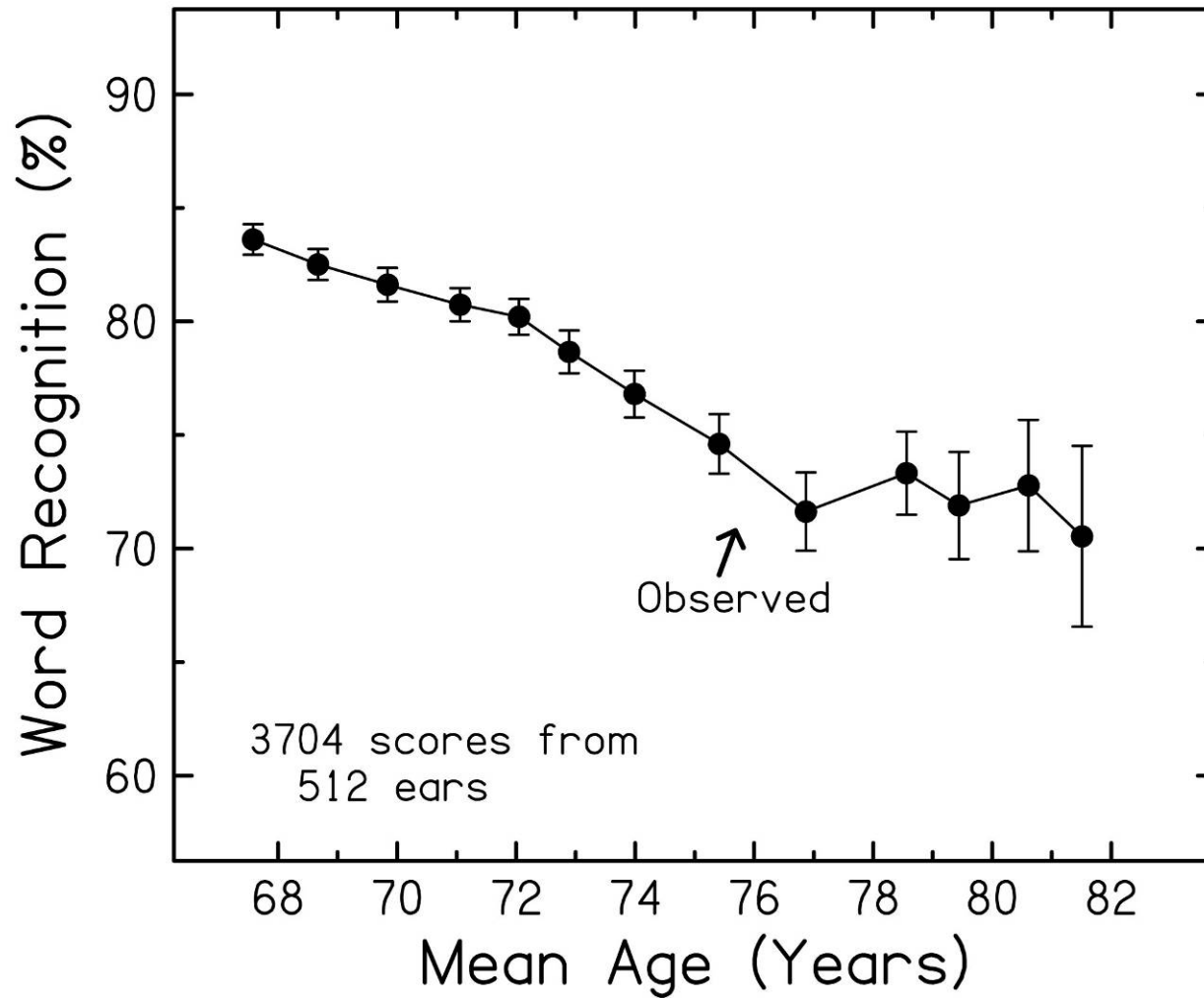


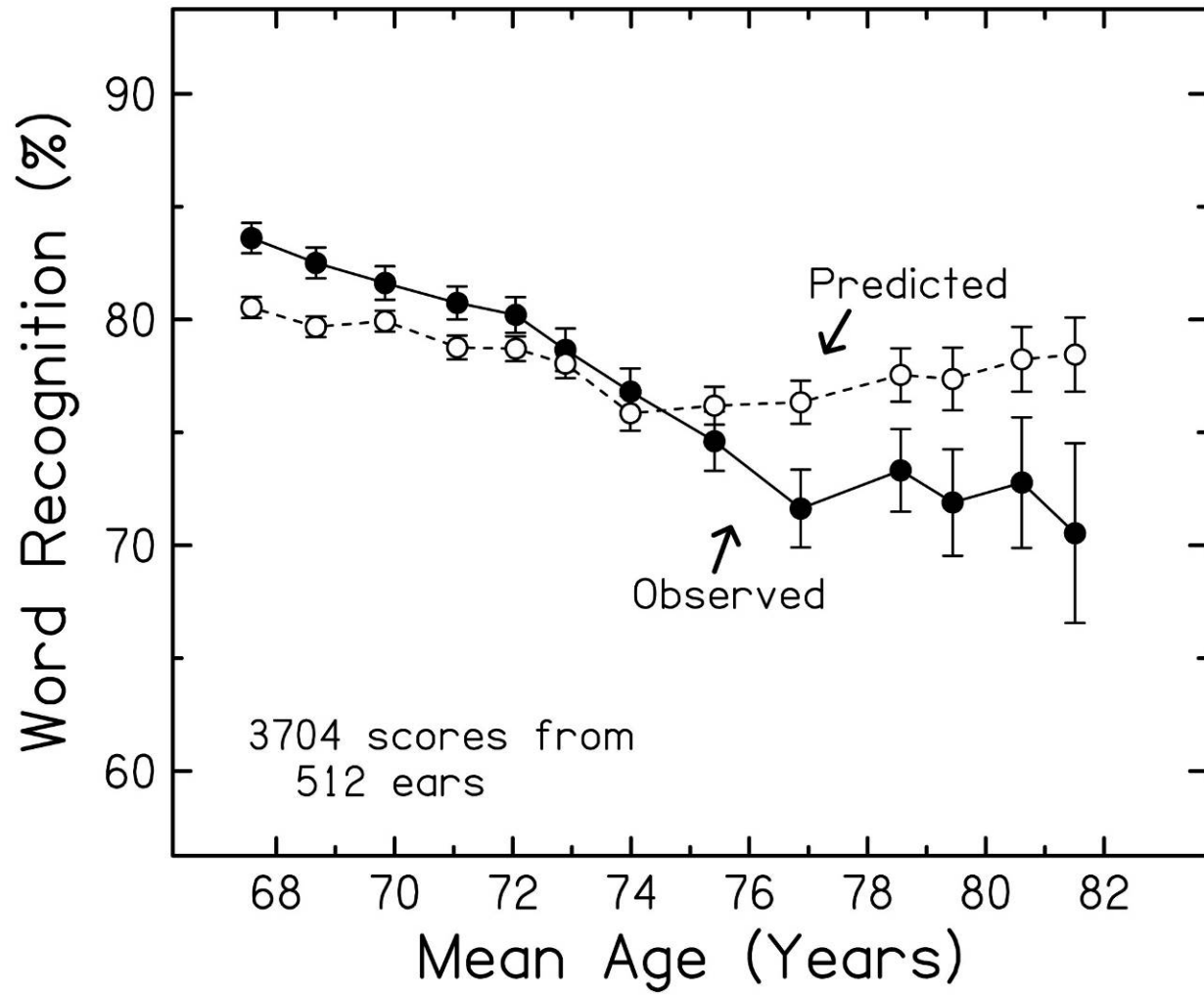
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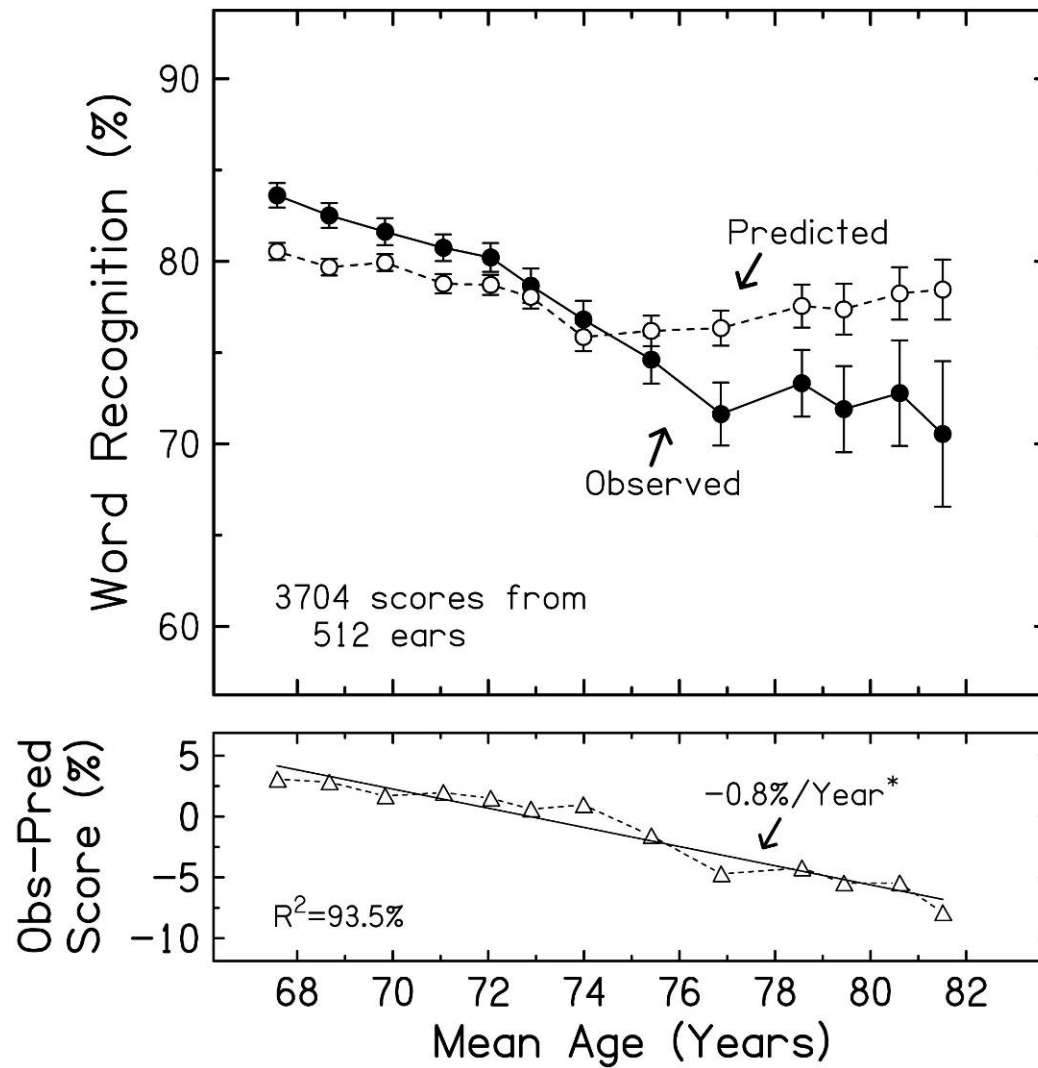
- To assess age-related declines in word recognition
 - If declines in word recognition over time are similar to predicted declines, poorer hearing (reduced audible speech) accounts for these changes, rather than increasing age
 - If declines are faster than predicted, poorer hearing does not entirely account for declines in word recognition – the remainder may be attributed to other factors, such as increasing age
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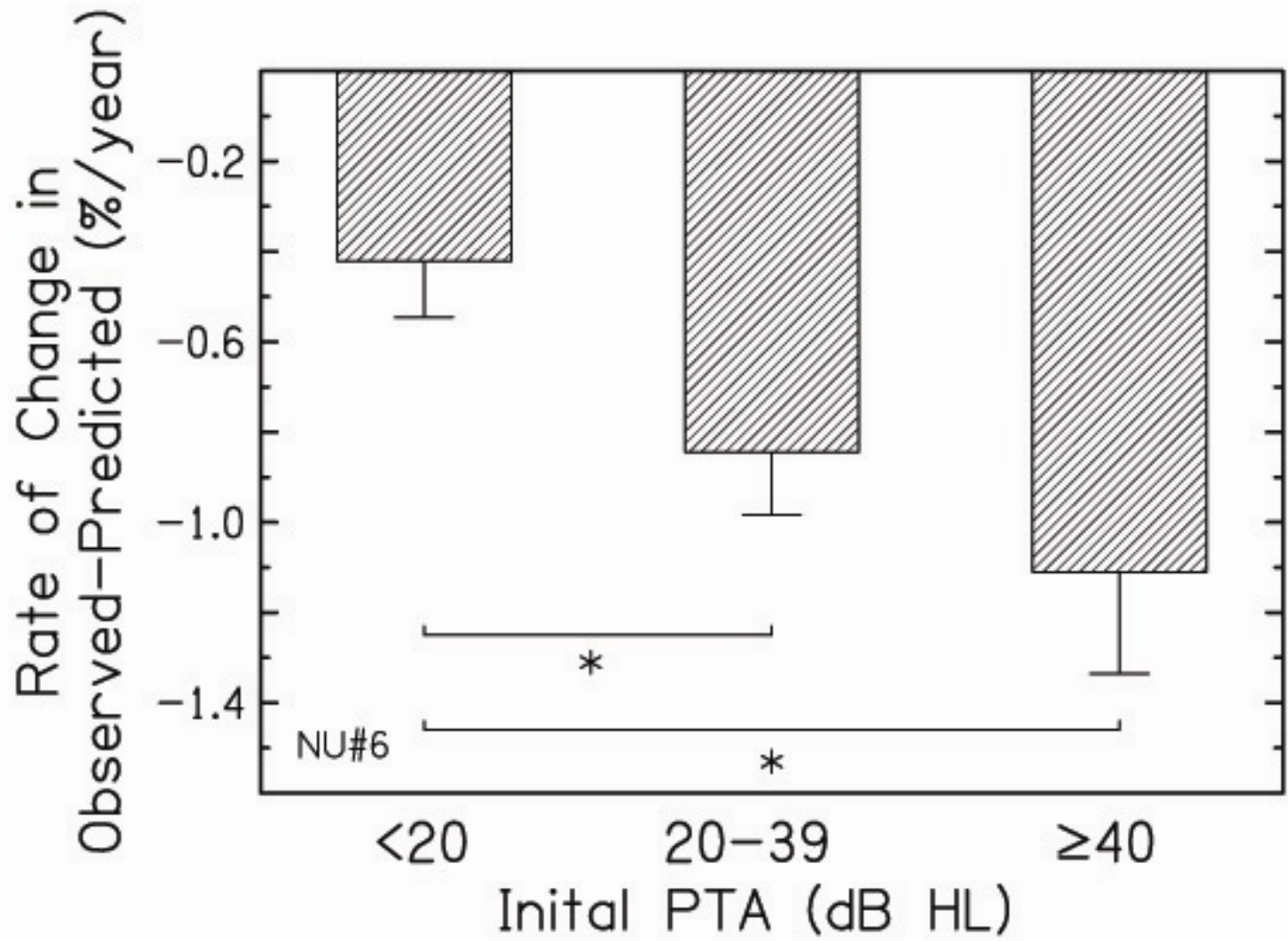
Word Recognition in Quiet

- Serial measures obtained over ≥ 3 years
 - Each subject had a minimum of 3 NU-6 scores
 - 3,704 scores from 512 ears
 - Mean number of scores per subject: 7.2
(Range 3-18)
 - Mean time span: 7.3 years (Range 3-15)
 - Mean age at entry: 67.6 years (Range 50-82)
 - Mean current age: 75.0 years (Range 60-91)
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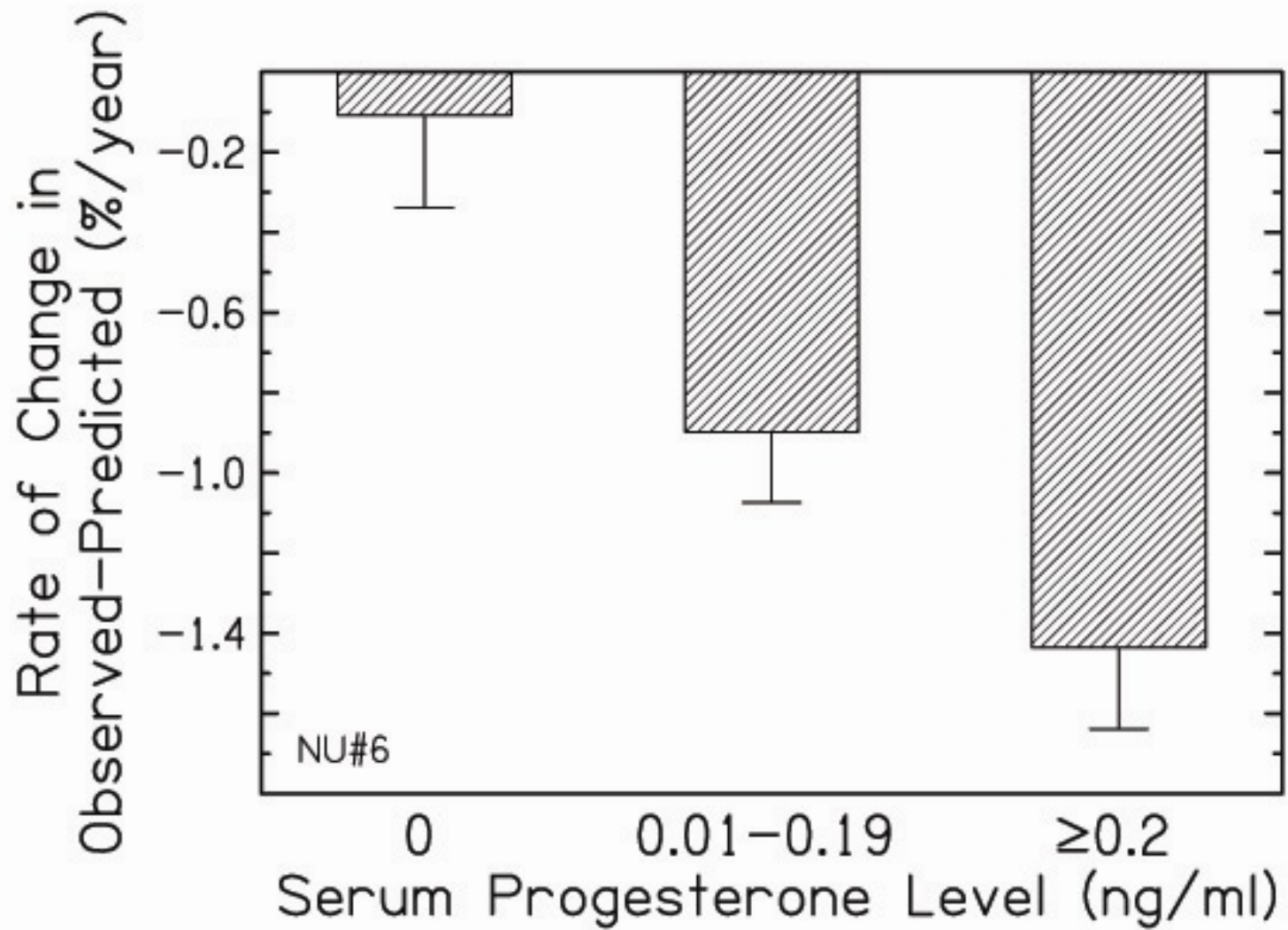






Effect of Initial Hearing Loss

- More severe injury to the auditory system resulted in faster declines in word recognition as subjects aged
 - Not related simply to more-elevated thresholds
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Effect of Serum Progesterone

- Consistent with the negative effect of hormone therapy that includes progestin reported by Guimaraes et al. (PNAS 2006)
 - Consistent with a biochemical mechanism that relates progesterone to activation of inhibitory neurotransmitters, such as GABA, in the aging auditory system
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Summary and Conclusions

- Pure-tone thresholds increase with age by an average of 1 dB/year (10 dB/decade)
 - Rate of decline in high-frequency hearing increases for females but decreases for males
 - Word recognition in quiet declines with age, even after accounting for reductions in audible speech due to poorer hearing
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Summary and Conclusions

- Rate of decline is faster for individuals with more severe hearing loss
 - Rate of decline is faster for females with higher levels of progesterone in their blood
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Summary and Conclusions

- Audiogram shapes and longitudinal changes in hearing are consistent with the view of age-related hearing loss as a **metabolic, vascular, neural** disorder rather than a sensory disorder
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Cognitive Test Battery

- Connections Test (Salthouse)
 - Visual Search and Attention Task (VSAT)
 - Stroop Neuropsychological Screening Test
 - Abbreviated Wechsler Memory Scale (WMS-III)
 - Wechsler Abbreviated Scale of Attention (WASI)
 - Mini-Mental Status Exam (MMSE)
 - Edinburgh Handedness Scale
 - NASA Task Load Index (Workload)
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