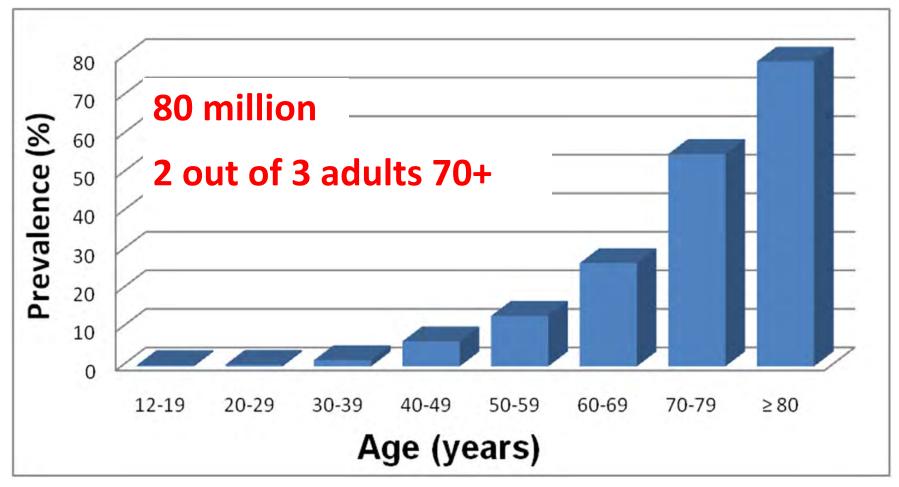
Hearing Loss and Dementia: Public Health Implications for Older Adults

Nicholas S. Reed, AuD

Cochlear Center for Hearing and Public Health Johns Hopkins Bloomberg School of Public Health nreed9@jhmi.edu

Prevalence of Hearing Loss by Age, United States, 2001-2008

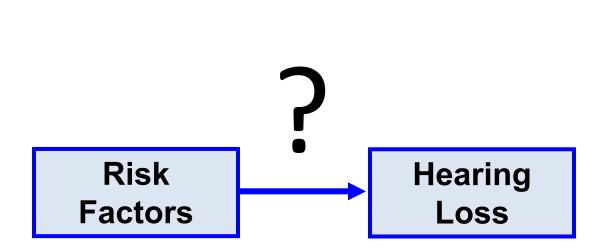


Hearing loss defined as a better-ear PTA of 0.5-4kHz tones > 25 dB

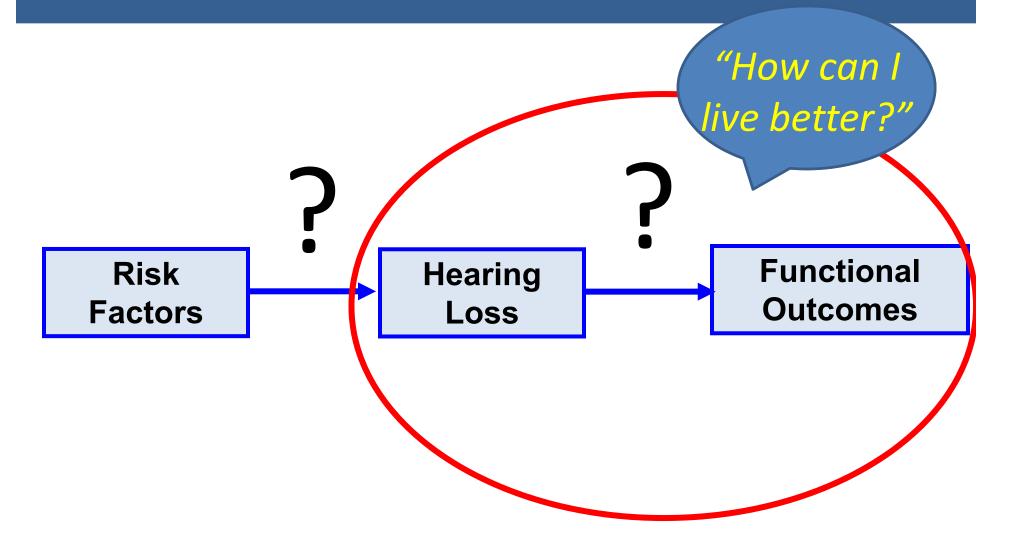
Lin et al., Arch Int Med. 2011

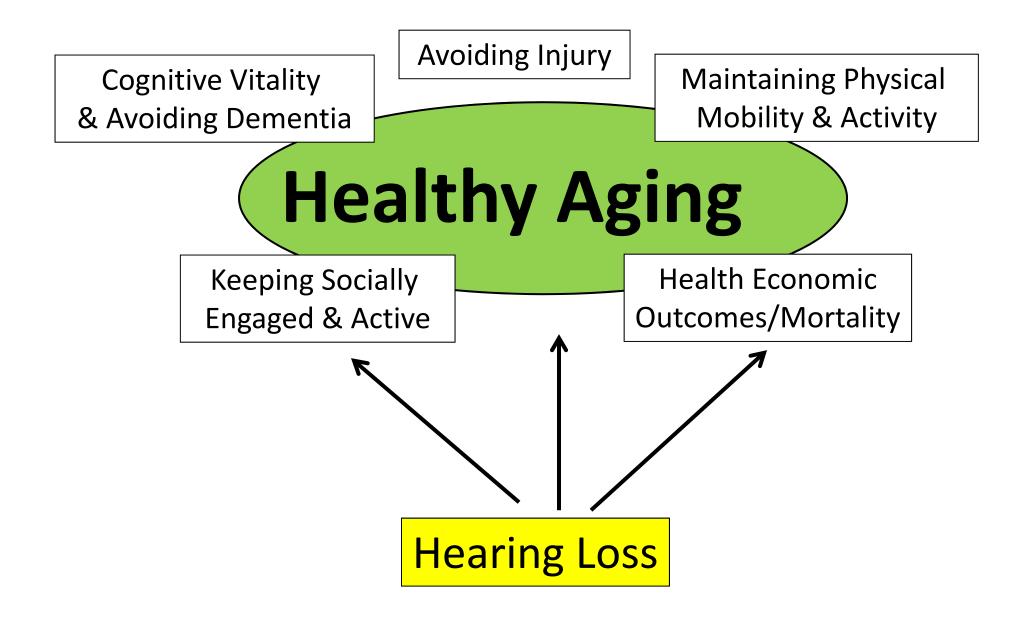
Hearing Loss & Function

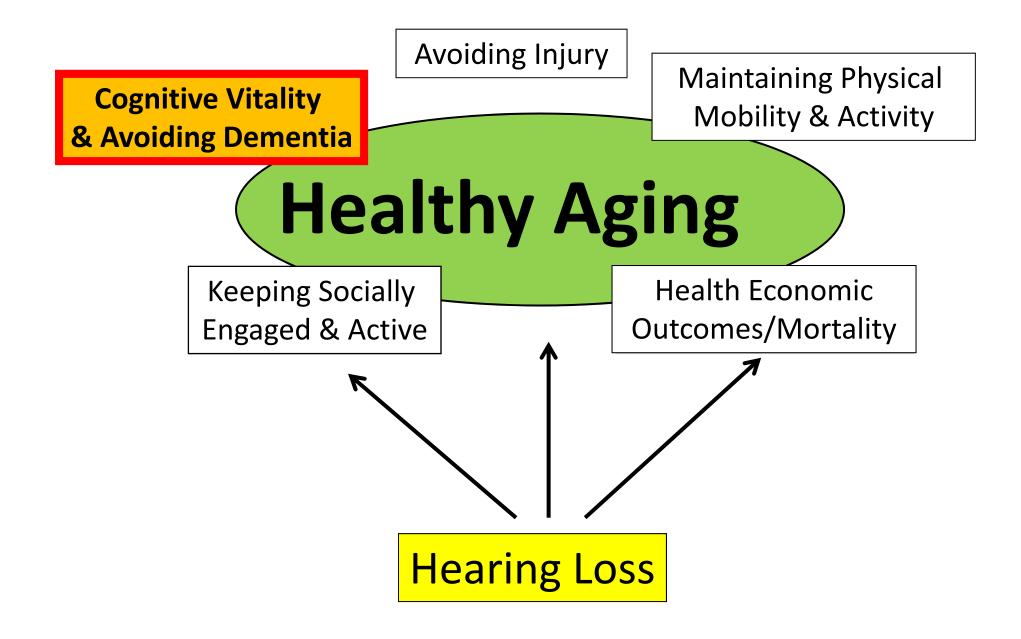
"How can I live better?"



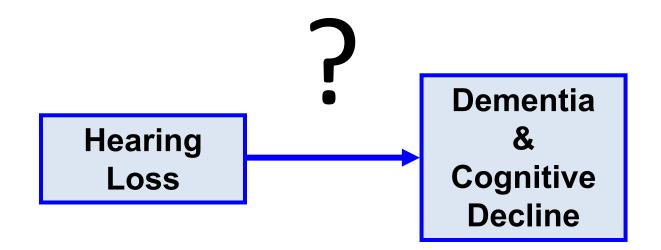
Hearing Loss & Function







Hearing Loss & Cognition



Objectives

- Introduce important cognitive outcomes in geriatrics/gerontology dementia
- Present epidemiologic evidence for a relationship between hearing impairment and cognitive decline and dementia

What is cognitive function?

- Collection of mental processes controlled by the brain
- Includes attention, memory, language production and understanding, learning, reasoning, problem-solving and decisionmaking

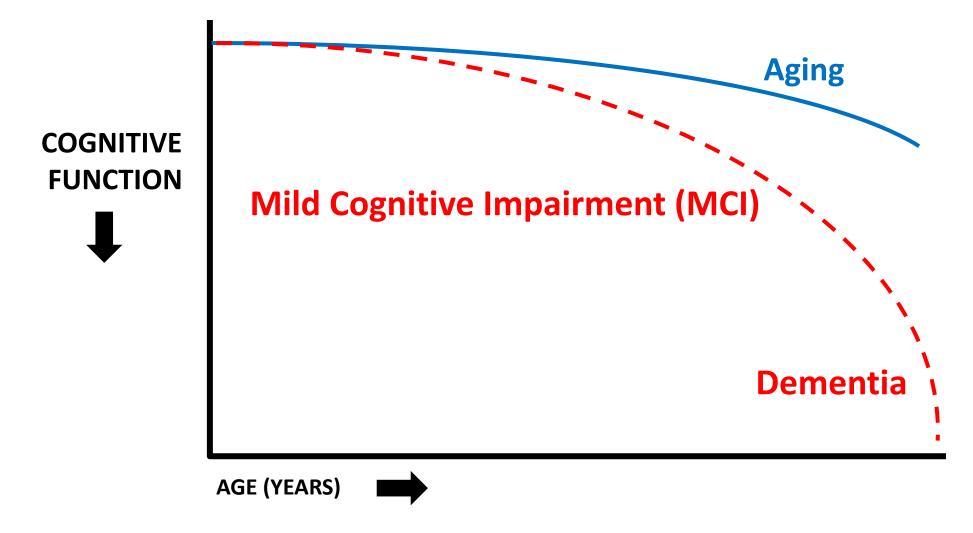
Dementia

- Cognitive or behavioral (neuropsychiatric) symptoms that:
- 1. Interfere with the ability to function at work or at usual activities; and
- 2. Represent a decline from previous levels of functioning and performing; and
- 3. Are not explained by delirium or major psychiatric disorder

Dementia

- ≥ 2 domains of cognitive function
 - e.g., memory, language, executive function
- Significant interference in the ability to function at work or in usual daily activities
 - Differentiates dementia from MCI

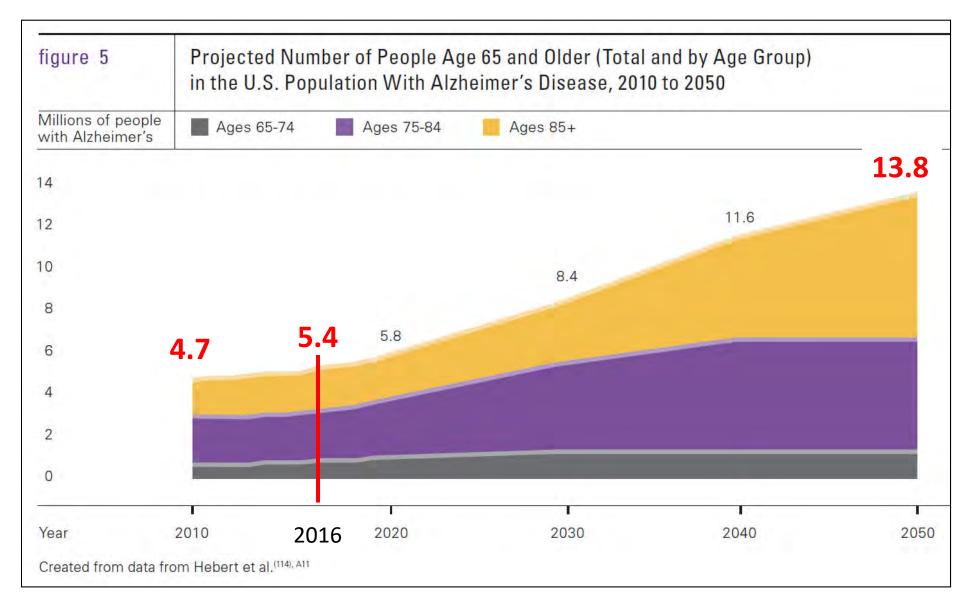
The Continuum of Alzheimer's disease



Adapted from Sperling et al., Alzheimer's and Dementia (2011) 7:280-292.

Alzheimer's disease

- Insidious onset
- Clear-cut worsening of cognition
- Initial/most prominent cognitive complaints:
 - Memory & learning (amnestic)
 - Non-amnestic (language, visuospatial, executive function)
- No evidence of other causes (e.g., cerebrovascular disease)



https://www.alz.org/downloads/Facts_Figures_2014.pdf

Hebert & al. Neurology May 7, 2013 vol. 80 no. 19 1778-1783

Dementia Incidence Declining? Temporal Trends in the Framingham Heart Study

Table 2. Temporal Trends in the Incidence of Dementia.*								
Subtype	No. of Cases	Total No. o Observation Periods		5-Yr Hazard Ra	tio (95% Cł);	P Value for Trend		
			Epoch 2	Epoch 3	Epoch 4			
Overall dementia	371	9015	0.78 (0.59–1.04)	0.62 (0.47–0.83)	0.56 (0.41-0.77)	<0.001		
Alzheimer's disease	264	9015	1.00 (0.70–1.43)	0.88 	0.70 (0.48–1.03)	0.052		
Vascular dementia	84	9014	0.89 (0.51–1.56)	0.46 (0.25–0.86)	0.45 (0.23–0.87)	0.004		
			late 1980's – early 90's	late 90's – early 2000's	late 2000's – early 2010's			
			Compared to late 1970's – early 80's					

Global Burden of Dementia

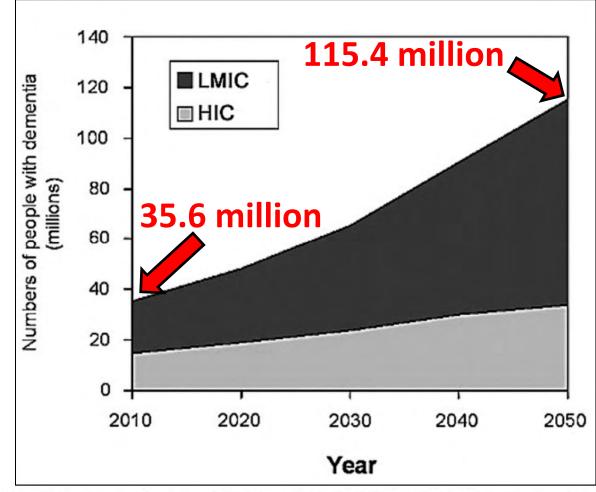


Fig. 2. The growth in numbers of people with dementia in high-income (HIC) and low- and middle-income countries (LMIC).

M. Prince et al. / Alzheimer's & Dementia 9 (2013) 63-75

The Cost of Dementia

Costs for Dementia Care Far Exceeding Other Diseases, Study Finds

By GINA KOLATA OCT. 26, 2015



Conclusion: Health care expenditures among persons with dementia were substantially larger than those for other diseases, and many of the expenses were uncovered (uninsured). This places a large financial burden on families, and these burdens are particularly pronounced among the demographic groups that are least prepared for financial risk.

Alicia Joseph, a home health aide with Partners in Care, helps Naomi Wallace, a dementia patient. Many of the costs of caring for dementia patients are not covered by Medicare. Sam Hodgson for The New York Times

Annals of Internal Medicine

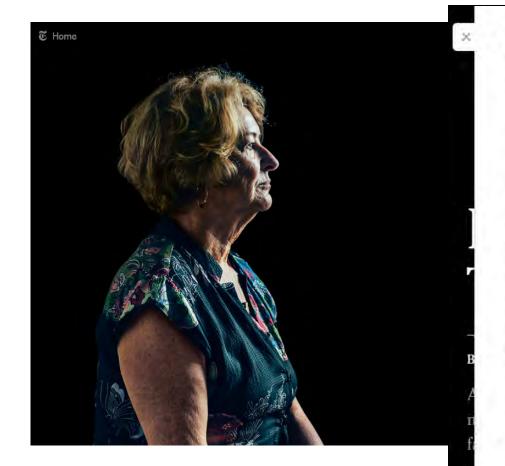
ORIGINAL RESEARCH

The Burden of Health Care Costs for Patients With Dementia in the Last 5 Years of Life

Amy S. Kelley, MD, MSHS; Kathleen McGarry, PhD; Rebecca Gorges, MA; and Jonathan S. Skinner, PhD

Ann Intern Med. Published online 27 October 2015 doi:10.7326/M15-0381 http://www.nytimes.com/2015/10/27/health/costs-for-dementia-care-far-exceeding-other-diseases-study-finds.html?_r=1

The Cost of Dementia



439 COMMENTS

Readers shared their thoughts on this article.

The comments section is closed. To submit a letter to the editor for publication, write to letters@nytimes.com.

All 139 Readers' Picks 318

NYT Picks 31 NYT Replies 11

John Princeton - April 86, 2016

I notice that I lose names, sometimes names that I have mentioned in conversation just moments ago. And it isn't so much the loss, I can usually come up with a suitable pronoun, but it is the discouraging feeling that the name lies just on the far side of a tiny membrane but could be deep in space in terms of its availability. It sometimes comes back to me within seconds, sometimes minutes, occasionally hours but, so far, always.

On the other hand I can always tell you that I lost a spelling bee to Juanita Meyers in the fifth grade; the word was necessary, I misspelled it with two "c's."

🖒 63 Recommend - 🚮 🔰

🏓 Flag

Patrick Foreman San Francisco · April 10 2016

My great-grandfather, grandfather, and now early 60s father are suffering from Alzheimer's. In our experience, my mom and I noticed the signs before my dad did. In 2012 my dad started asking the same questions over and over again, and really struggled with dates and times specifically. Next went the ability to operate basic kitchen appliances like the toaster or coffee maker. It

Treatments for Alzheimer's disease

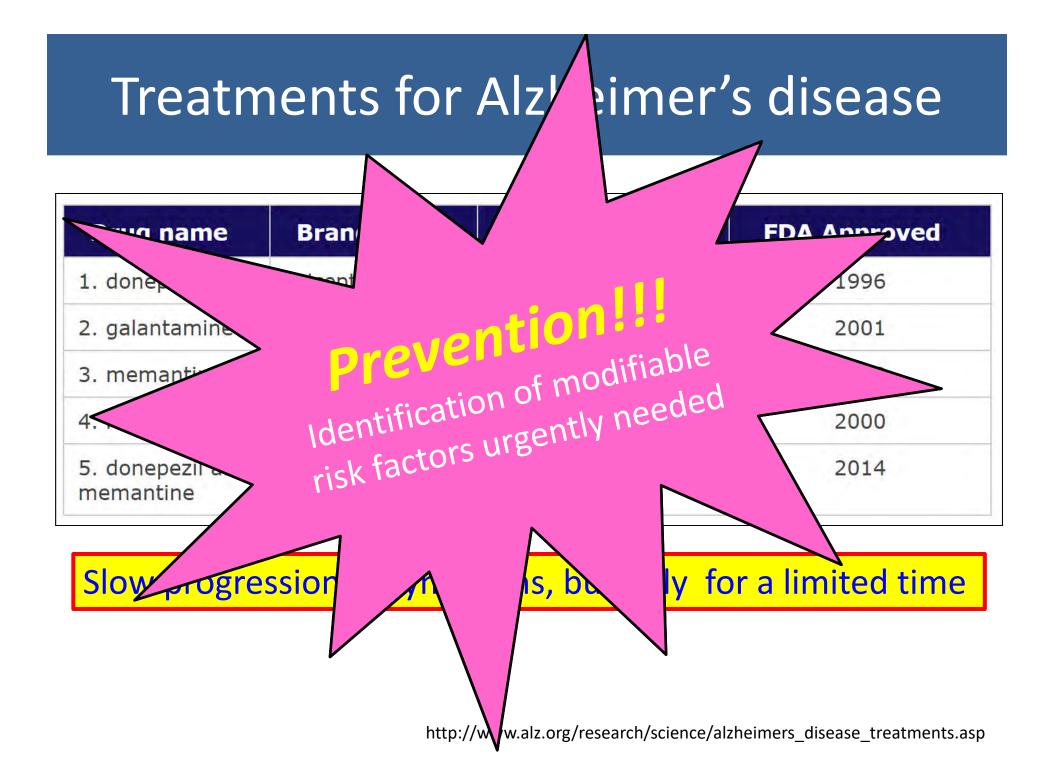
Drug name	Brand name	Approved For	FDA Approved
1. donepezil	Aricept	All stages	1996
2. galantamine	Razadyne	Mild to moderate	2001
3. memantine	Namenda	Moderate to severe	2003
4. rivastigmine	Exelon	All stages	2000
5. donepezil and memantine	Namzaric	Moderate to severe	2014

Treatments for Alzheimer's disease

Drug name	Brand name	Approved For	FDA Approved
1. donepezil	Aricept	All stages	1996
2. galantamine	Razadyne	Mild to moderate	2001
3. memantine	Namenda	Moderate to severe	2003
4. rivastigmine	Exelon	All stages	2000
5. donepezil and memantine	Namzaric	Moderate to severe	2014

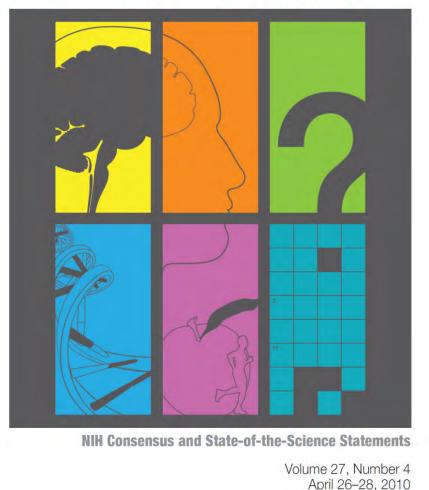
Slow progression of symptoms, but only for a limited time

http://www.alz.org/research/science/alzheimers_disease_treatments.asp

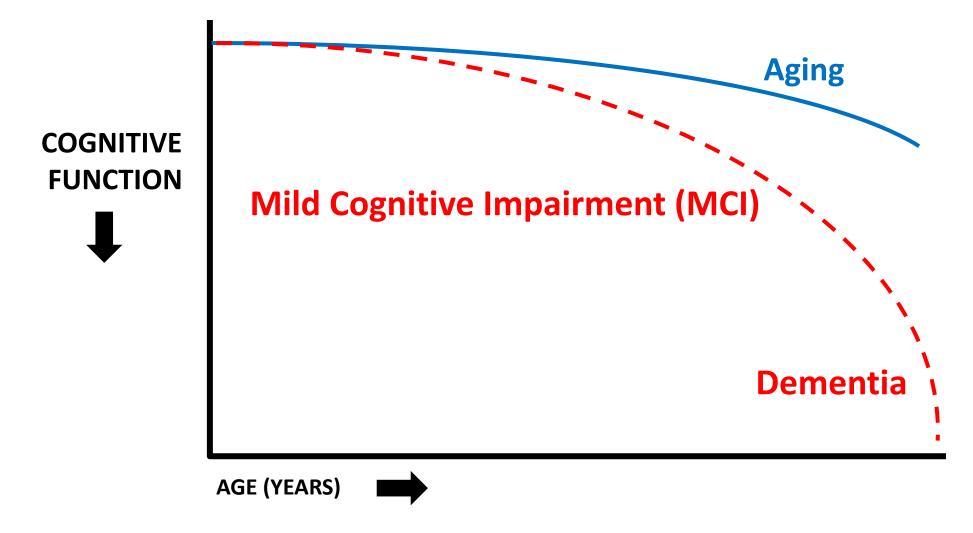


State of Prevention

"Currently, firm conclusions cannot be drawn about the association of any modifiable risk factor with cognitive decline" NIH State-of-the-Science Conference Statement on Preventing Alzheimer's Disease and Cognitive Decline



The Continuum of Alzheimer's disease

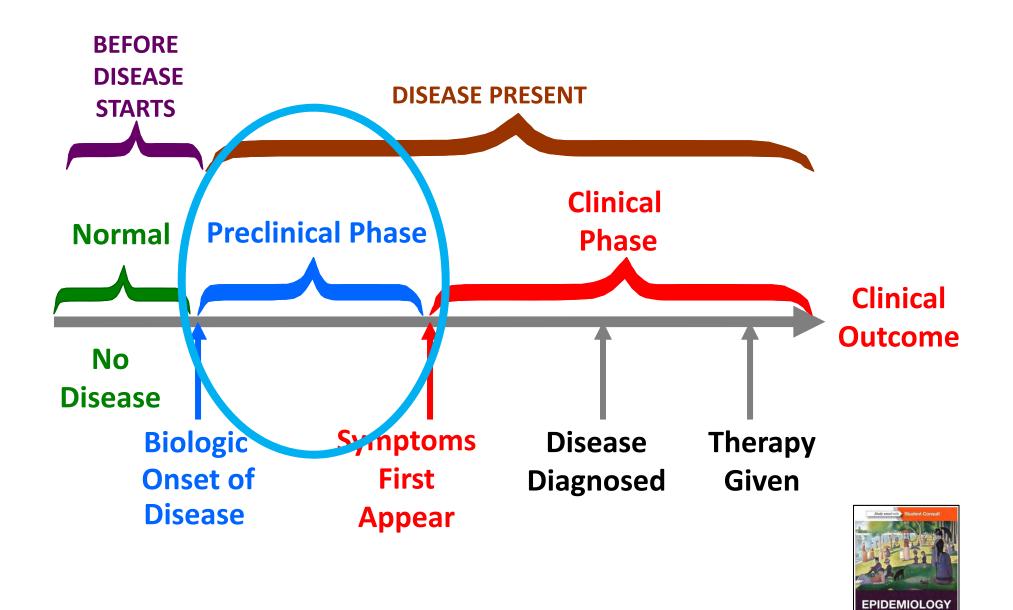


Adapted from Sperling et al., Alzheimer's and Dementia (2011) 7:280-292.

The Continuum of Alzheimer's disease Timing of intervention Aging Mild Cognitive Impairment (MCI Dementi AGE (YEARS) VS.

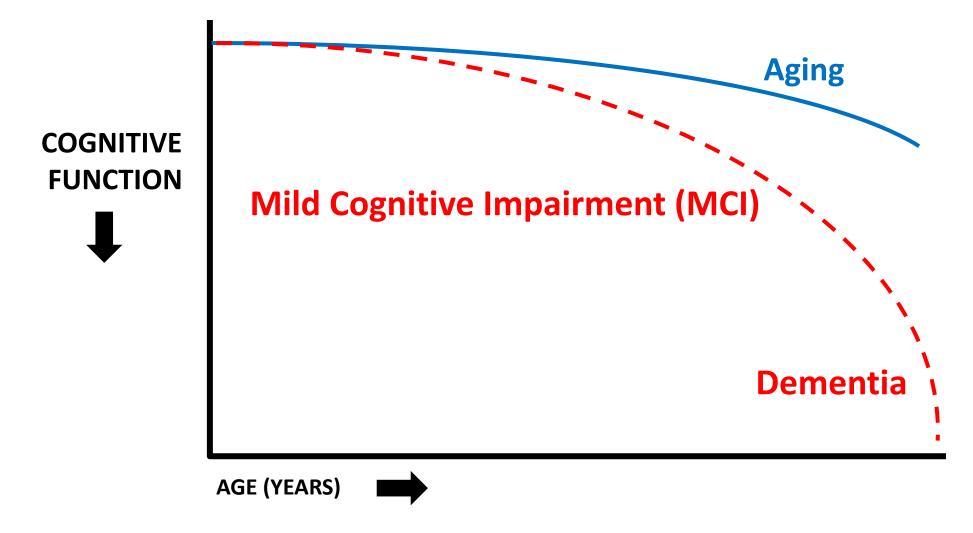
Does the effect of a factor differ by point on the continuum?

Adapted from Sperling et al., Alzheimer's and Dementia (2011) 7:280-292.

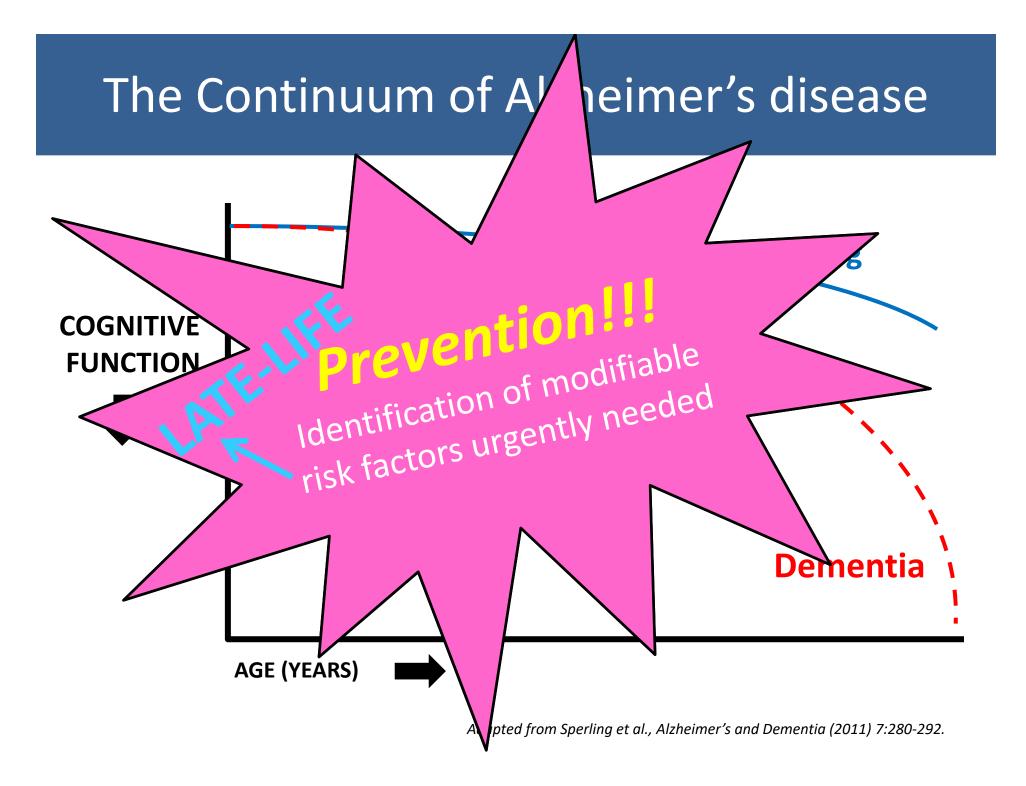


LEON GORDIS

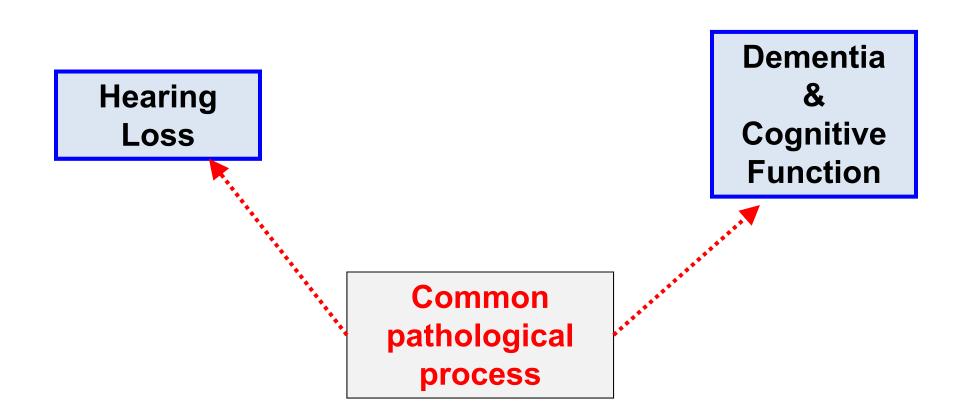
The Continuum of Alzheimer's disease



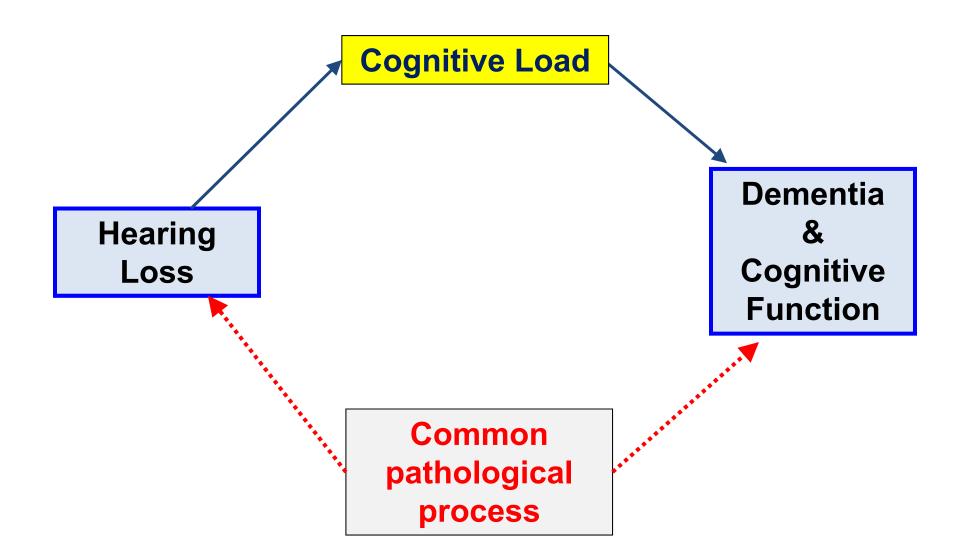
Adapted from Sperling et al., Alzheimer's and Dementia (2011) 7:280-292.

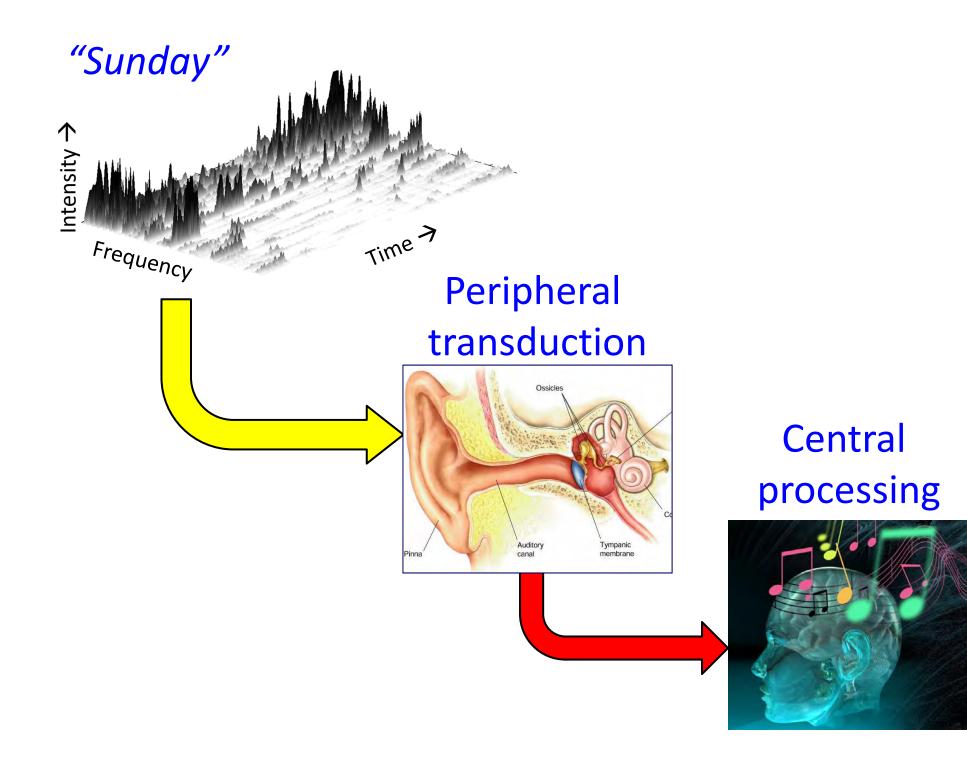


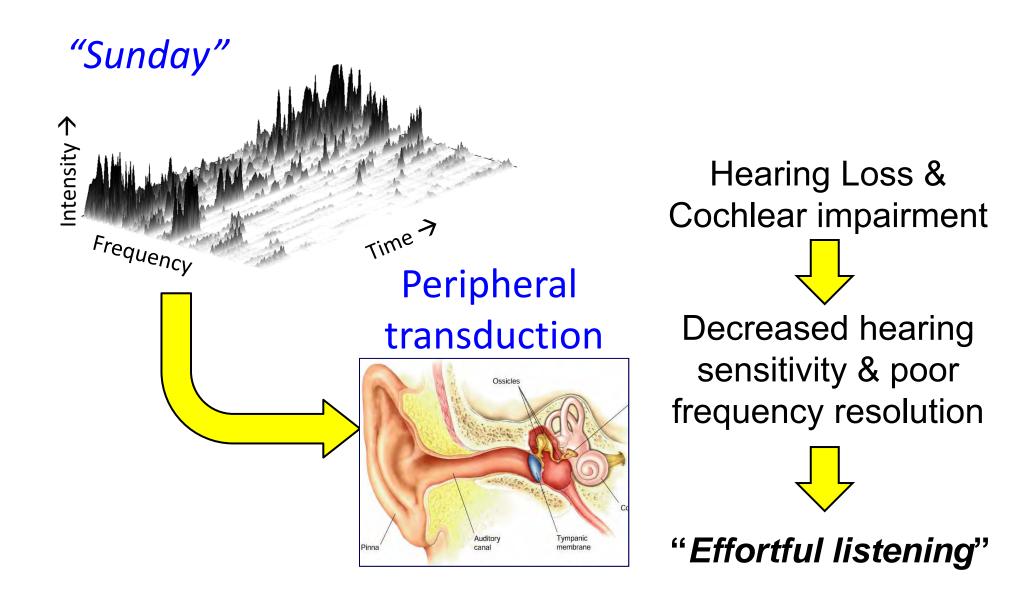
Hearing Loss & Cognition



Hearing Loss & Cognition







Hearing Loss & Cognitive Load

Kahneman model of shared attention and resource capacity (D. Kahneman, Attention & Effort, 1973)

Cognitive Resource Capacity

Auditory Perceptual Processing Requirements	Available Cognitive Resources For Performance of Tasks	Age-Related Decline
--	---	------------------------

Hearing Loss & the Brain

Poorer hearing is associated with:

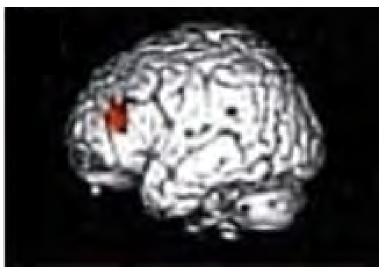
A. Reduced language-driven activity in primary auditory pathways

B. Increased
compensatory
language-driven
activity in pre-frontal
cortical areas

A Decreased language-driven speech activity in poorer hearers

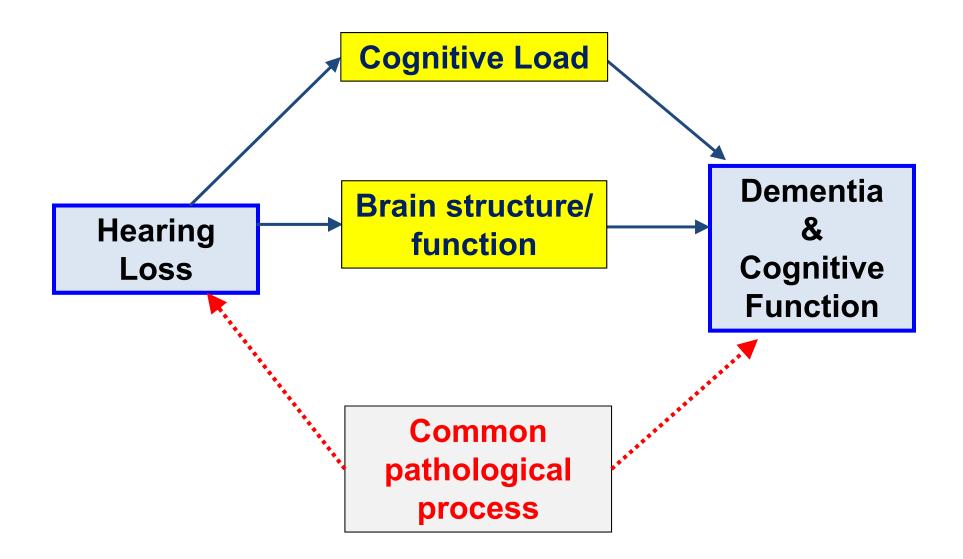


Peelle et al, J. Neurosci, 2011

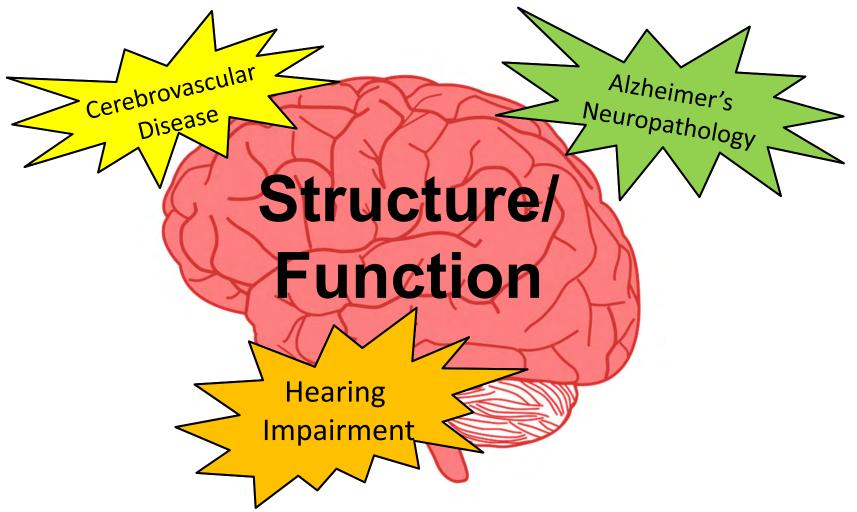


Grossman et al, Brain Lang, 2002

Hearing Loss & Cognition



Double Hit Theoretical Model Hearing Loss & Brain Structure/Function

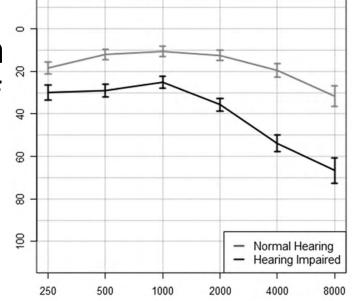


F. Lin & M. Albert, Aging & Mental Health, 2014

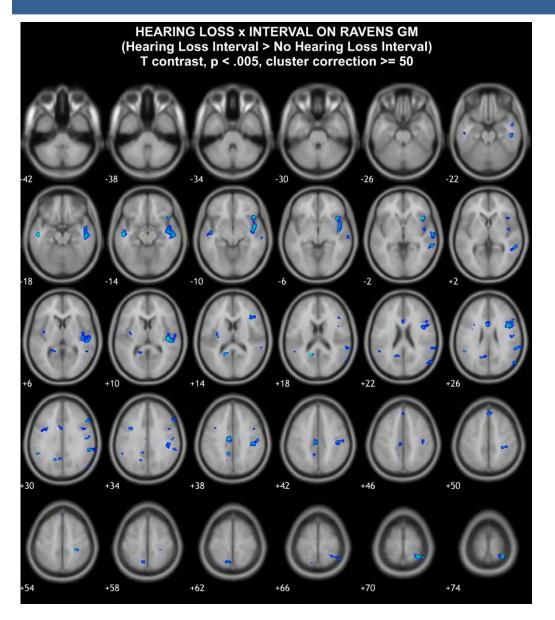
Hearing Loss & Incident Dementia

Dementia incidence in 1889 adults followed for 9 years in HealthABC

- **Hypothesis**: Hearing loss is associated with accelerated atrophy in the superior, middle, and inferior temporal gyri
- 126 participants (56-86 yrs) in the neuroimaging substudy of the BLSA
 - Mean follow-up duration of 6.4 years
 - 1.5T MRI performed annually

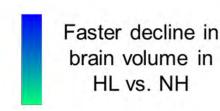


Hearing Loss & Accelerated Brain Volume Decline



Voxel-Based Analyses

Difference in mean gray matter volume change in those with HL vs. NH



Lin et al. Neuroimage 2014

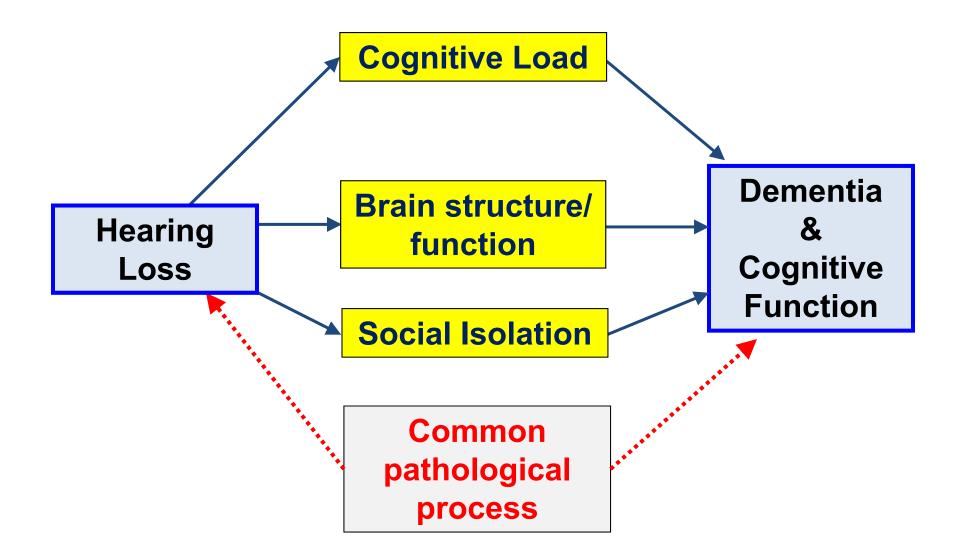
Estimated Annual Rates of Change in Brain Volume (cm³/year), Baltimore Longitudinal Study on Aging (BLSA)

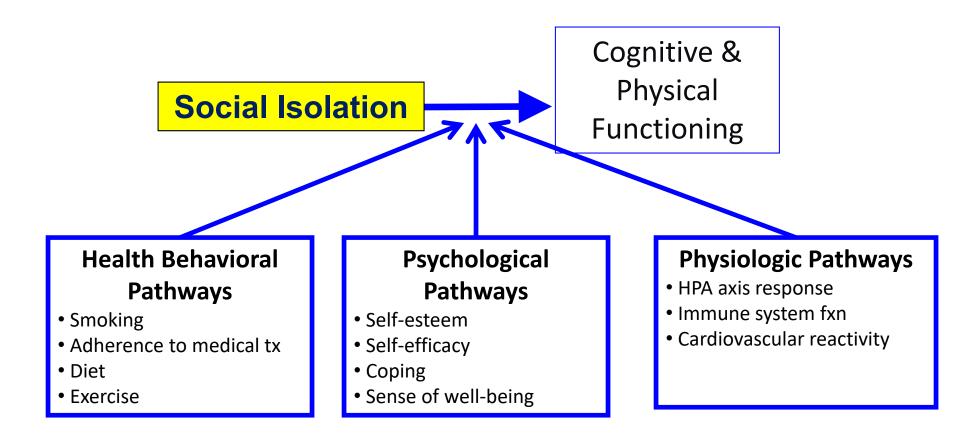
	Normal Hearing mean (SE)	Hearing Loss, mean (SE)	Difference mean (SE)	
Global measures	N=75	N=51		
Whole brain	-7.21 (0.27)**	-8.33 (0.36)**	-1.13 (0.45) ⁺ p = .015	
vCSF	1.30 (0.10) **	1.28 (0.14)**	-0.020 (0.18)	
White matter	-4.14 (0.31)**	-4.99 (0.39)**	-0.85 (0.39) ⁺ p = .031	N=126 participants
Gray matter	-2.63 (0.22)**	-3.38 (0.28)**	-0.76 (0.36) ⁺ p = .036	aged 56-86 years Annual MRI for up to 10 yrs
Lobar measures Gray matter			-	Mean follow-up = 6.4 years
Frontal	-0.96 (0.11)**	-1.11 (0.14)**	-0.16 (0.14)	
Temporal	-0.46 (0.096)**	-0.71 (0.12)**	-0.25 (0.12) ⁺ p = .036	
Parietal	-0.71 (0.051)**	-0.74 (0.066)**	-0.044 (0.081)	
Occipital	-0.54 (0.057)**	-0.50 (0.073)**	0.047(0.071)	
Regional Volumes	**	**		Cound & speech
Superior temporal gyrus	-0.20 (0.023)**	-0.31 (0.030)**	-0.11 (0.038) [*] p = .0046	Sound & speech
Middle temporal gyrus	-0.15 (0.033)**	-0.30 (0.042)**	-0.15 (0.054) * p = .0065	But also memory,
Inferior temporal gyrus	-0.048 (0.015)*	-0.12 (0.020)**	-0.067 (0.025) * p = .0093	sensory integration
Hippocampus	-0.019 (0.0051)**	-0.031 (0.0065)	-0.012 (0.0002)	, <u> </u>

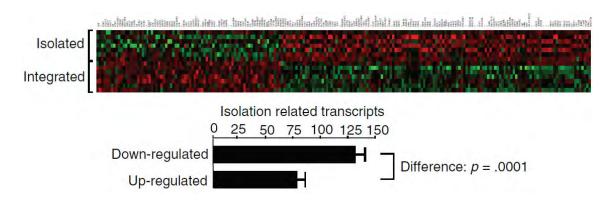
+ < .05; * < .01; ** <.001

Lin et al., Neuroimage 2013

Hearing Loss & Cognition







Cole & Cacioppo, Genome Biology, 2007 Cole & Cacioppo, PNAS, 2011 Social isolation is associated with upregulation of proinflammatory genes & increased inflammation

Hearing Loss & Healthy Aging Cross-sectional Datasets for Epidemiologic Analyses

NHANES: National Health and Nutritional Examination Survey

Cross-sectional, representative sample of U.S. population



Hearing Loss & Healthy Aging Longitudinal Datasets for Epidemiologic Analyses

ARIC: Atherosclerosis Risk in Communities



- Prospective, population-based study of 15,792 men and women from 4 US communities aged 45-64 years at baseline (1987-89)
- BLSA: Baltimore Longitudinal Study on Aging
 - Ongoing prospective study of older adults since 1958
- Health ABC: Health, Aging, & Body Composition
 - Prospective, population-based study of ~3000 adults 70 years and older



Cognitive Assessment

- Global cognitive function
 - Mini-Mental State Exam (Folstein 1975)
- Cognitive domains
 - Address different cognitive abilities
 - e.g.,
 - Memory
 - Executive Function
 - Cognitive abilities that control & regulate other abilities & behaviors
 - Goal-directed behavior, planning, initiating, inhibiting

Cognitive Assessment

- Why study cognitive domains?
 - Hierarchy of decline across domains
 - Speed 30's
 - Vocabulary intact into 70's
 - Domain-specific decline associated with:
 - Transition to dementia
 - Disability
 - Mortality

Global Function: The Mini-Mental State Exam

Patient		Examiner	Date
Maximum	Score		
		Orientation	
5	()	What is the (year) (season) (date) (day) (month)?	
5	()	Where are we (state) (country) (town) (hospital) (floor)?	
		Registration	
3	()	Name 3 objects: 1 second to say each. Then ask the pati	
		all 3 after you have said them. Give 1 point for each	
		Then repeat them until he/she learns all 3, Count tri	als and record.
		Trials	
		Attention and Calculation	
5	()	Serial 7's. 1 point for each correct answer. Stop after 5	answers.
		Alternatively spell "world" backward.	
		Recall	
3	()	Ask for the 3 objects repeated above. Give 1 point for ea	ch correct answer.
		Language	
2	()	Name a pencil and watch.	
1	()	Repeat the following "No ifs, ands, or buts"	
3	1 1	Follow a 3-stage command:	
		"Take a paper in your hand, fold it in half, and put it	on the floor."
1	()	Read and obey the following: CLOSE YOUR EYES	
1	()	Write a sentence.	
1	()	Copy the design shown.	
		\wedge	
		$\langle \rangle \rangle$	
		$\langle \langle \langle \rangle \rangle$	
		X /	
		Total Score	
		ASSESS level of consciousness along a continuum	
		Alert Drowsy Stup	or Coma

Executive Function: Digit Symbol Substitution Test (DSST)

				1 		2	3 	4	2	5	6	7 ⊏	8		9				
2	1	3	1	2	1	3	1	4	2	4	2	5	1	4	3	5	2	6	2
1	6	5	2	4	7	3	5	1	7	6	3	8	5	3	6	4	2	1	8
9	2	7	6	3	5	8	3	6	5	4	9	7	1	8	5	3	6	8	2
7	1	9	3	8	2	5	7	4	1	6	7	4	5	8	2	9	6	4	3

Executive Function: Stroop Mixed

Look at the list below and say the **<u>COLOR</u>** not the word.

YELLOW	BLUE	ORANGE
BLACK	RED	GREEN
PURPLE	YELLOW	RED
ORANGE	GREEN	BLACK
BLUE	RED	PURPLE
GREEN	BLUE	ORANGE

Left - Right Conflict

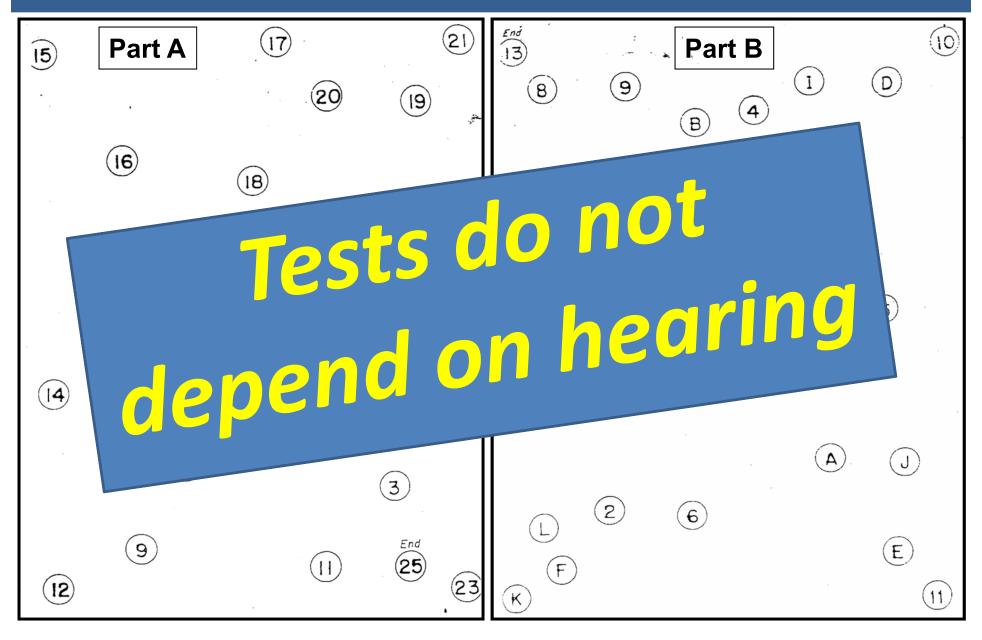
Your right brain tries to say the color but Your left brain insists on reading the word.

http://cdn.lifeinthefastlane.com/wp-content/uploads/2015/06/stroop-effect.jpg

Executive Function: Trail Making Tests

End 13 (21) (17) (10)Part A Part B 15 (\mathbf{l}) (D)(9) (8) (20) (19) (4) B) (16) (18) 3 (22) 4 5 Begin (7)(6) (13) 5) Ή) C Begin (24) 7 12) (14) G 2 J 10 (8) 3) (2) 6 End (25) 9 (E) (Π) F (12) 23 11 K

Executive Function: Trail Making Tests



Hearing Loss and Cognition Cross-Sectional Studies

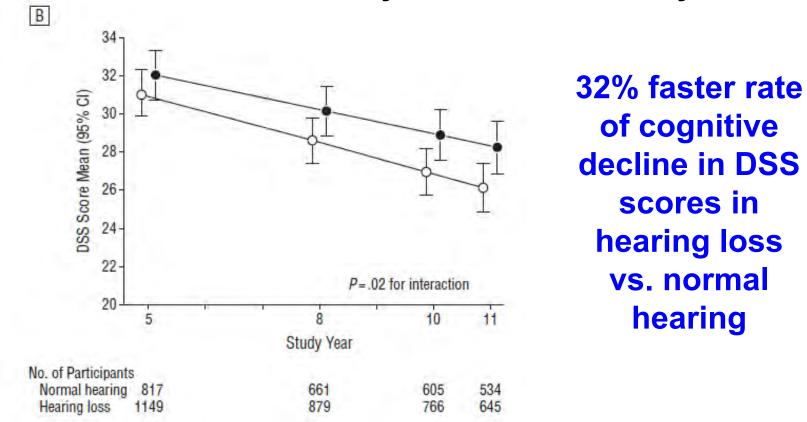
NHANES	N = 605 adı	ults 60-69	years Li	in, J. Gero	ont. Med. Sci., 201
		Age er year) <i>P</i>	Hearing loss (per 25 dB) β ^b (95% CI)		∆ Age (years) equivalent to 25 dB of hearing loss
Digit Symbol Substitution Test	-0.55	<.01	-3.86	.02	7.0
Substitution Test	(-0.920.18)		(-7.15 – -0.56)		
BISA	N = 347 ad	lults >60 v	aars		

BLSA N = 347 adults >60 years				Lin et al., Neuropsych., 201			
Stroop Mixed	-0.33 (-0.480.18)	<.001	-2.27 (-4.140.40)	.02	6.8		
Trail Making B	-0.00011 (-0.000180.000044)	.001	00074 (-0.0015 - 2.74x10 ⁻⁶)	.05	6.7		

Models adjusted for age, sex, race, education, diabetes, smoking, hypertension

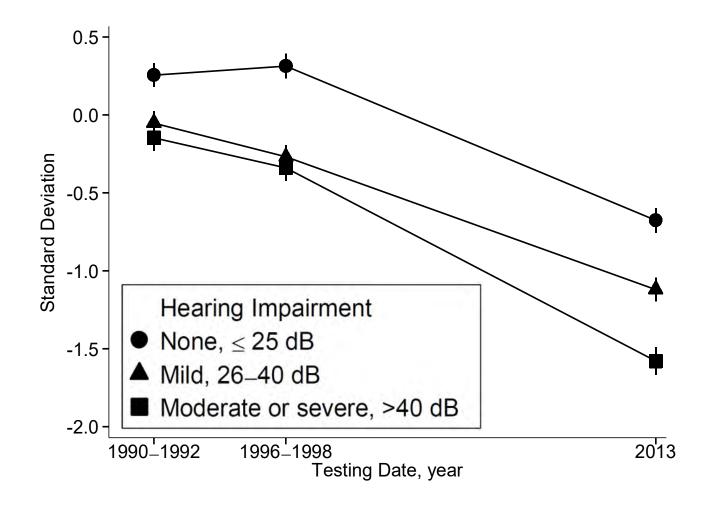
Hearing Loss & Cognitive Decline HealthABC

Adjusted DSS scores by years of follow-up and hearing loss status in 1,966 adults > 70 years followed for 6 years



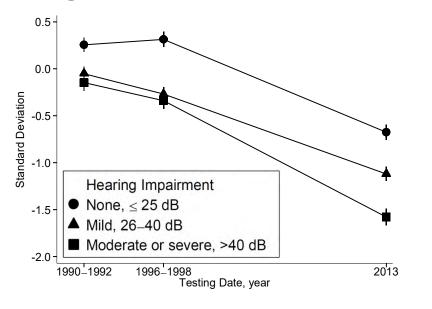
Adjusted for age, sex, race, education, study site, smoking status, hypertension, diabetes, and stroke history Lin et al. JAMA Int Med. 2013

Hearing Loss & Memory Decline ARIC, 1990-2013



* Adjusted for age, age², sex, education, smoking status, diabetes, hypertension, and Wide Range Achievement Test (WRAT)

Change in MEMORY,* 1990-2013

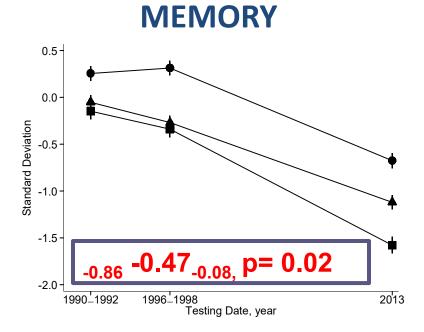


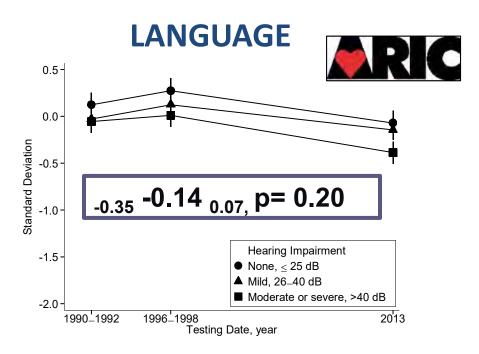


* Adjusted for age, age², sex, education, smoking status, diabetes, hypertension, and Wide Range Achievement Test (WRAT)

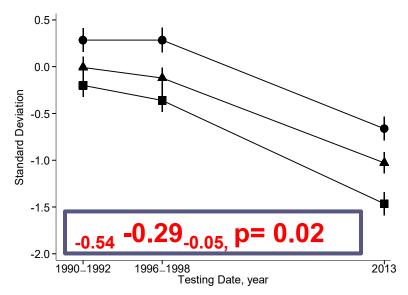
Rates of 20-Year Change* in MEMORY, ARIC

Hearing status:	Estimate _{95% CI}	P-value
Moderate/severe	_{-1.66} -1.35 _{-1.05}	<0.0001
Mild	-1.24 -1.02 -0.80	<0.0001
None	-1.13 -0.88 -0.64	<0.0001
Difference Comparing:		
Mild to None	_{-0.47} -0.14 _{0.20}	0.411
Moderate/severe to None	-0.86 -0.47 -0.08	0.018

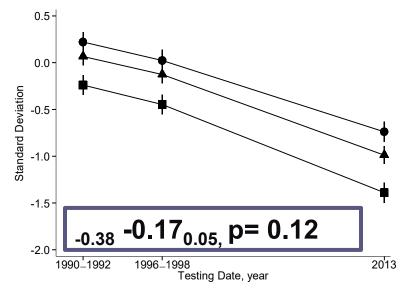


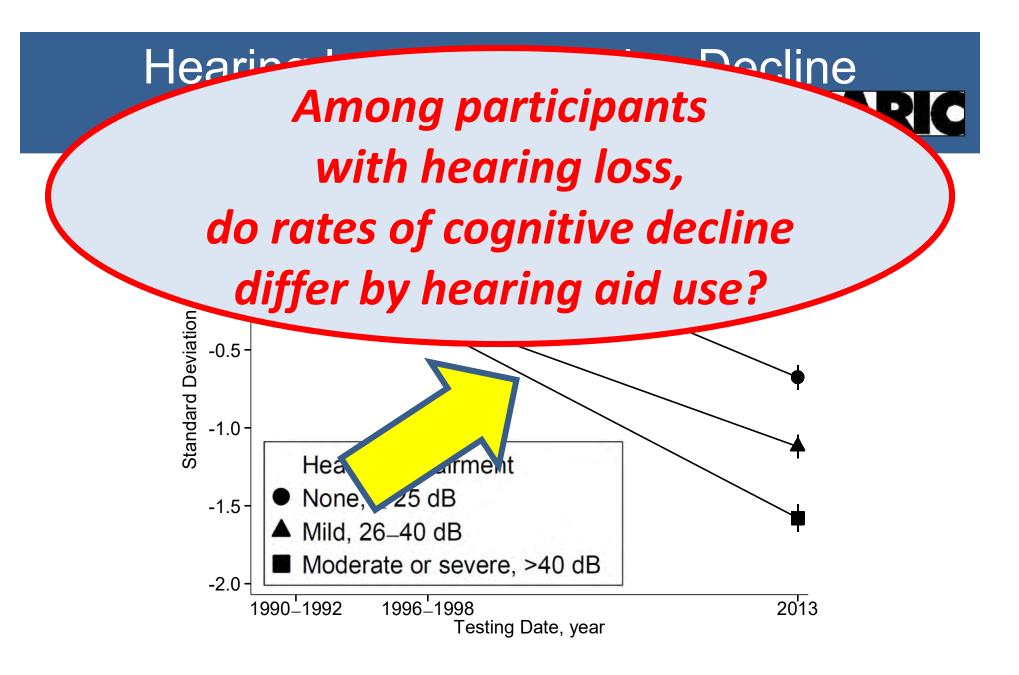


GLOBAL FUNCTION



SPEED & ATTENTION





* Adjusted for age, age², sex, education, smoking status, diabetes, hypertension, and Wide Range Achievement Test (WRAT)

Rates of Cognitive Change (1990-2013) by Hearing Aid Use* (2013), N=85

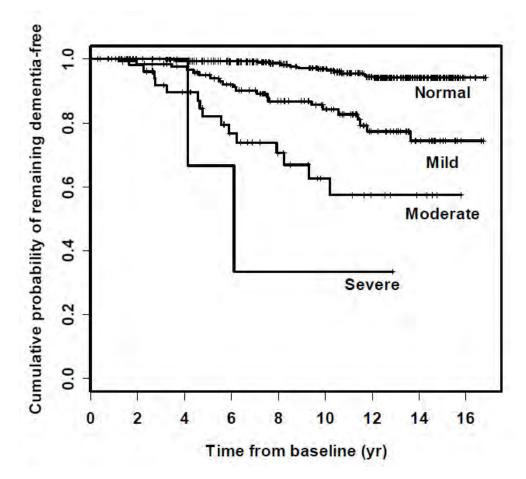
MEMORY	Estimate 95% CI	P-value
No hearing aid, N=42	_{-2.28} -1.84 _{-1.39}	< 0.0001
Hearing aid, N=43	-1.25 -0.89 -0.52	< 0.0001
Difference	-1.53 -0.95 _0.38	0.001

GLOBAL COMPOSITE		
No hearing aid, N=42	_{-1.70} -1.45 _{-1.20}	<0.0001
Hearing aid, N=43	-1.21 -0.97 -0.74	<0.0001
Difference	-0.83 -0.48 -0.14	0.006

*Among participants with moderate/severe hearing loss

Hearing Loss & Incident Dementia

Dementia incidence in 639 adults followed for >10 years in the BLSA



Risk of incident all-cause dementia (compared to normal hearing)^a

	<u>HR</u>	<u>95% CI</u>	p
Mild	1.89	1.00 – 3.58	.05
Moderate	3.00	1.43 - 6.30	.004
Severe	4.94	1.09 – 22.4	.04

^a Adjusted for age, sex, race, education, DM, smoking, & hypertension

Lin et al., Arch Neuro., 2011

Question to be Answered

 If we treat hearing impairment, do we delay or prevent functional decline in older adults?

ACHIEVE Randomized Trial

- Aging, Cognition, and Hearing Evaluation in Elders
- ACHIEVE-Feasibility
 - 20 participants
 - Hearing intervention
- ACHIEVE-Pilot
 - 40 participants



- Randomized: Hearing vs. Successful Aging
- 6-mo follow-up



Hearing Intervention





Theresa Chisolm, PhD



Michelle Arnold AuD, PhD, CCC-A



Victoria Sanchez AuD, PhD, CCC-A

Courtney Matthews

- Goal: eliminate or minimize activity limitations & participation restrictions from HL
- Individual needs assessment & goal-setting, development of selfmanagement abilities
- 4 sessions (~1 hr each, over 2-3 mos)
- Hearing aids & other technologies

Successful Aging Intervention





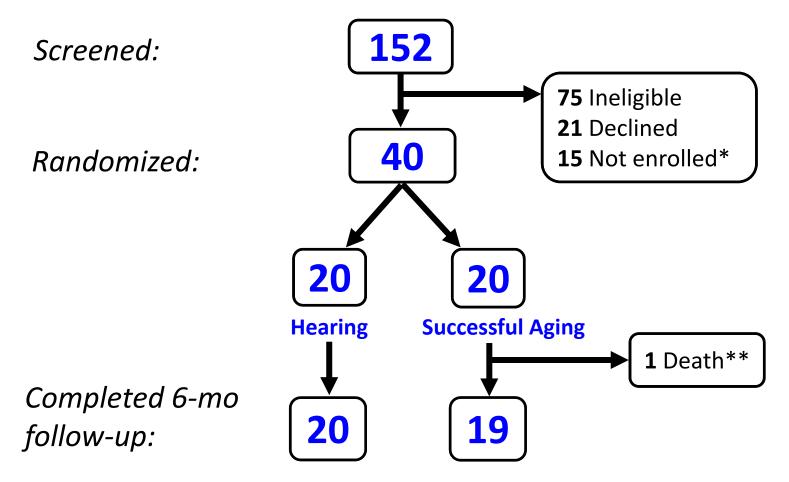
Nancy W Glynn, PhD

• Elizabeth Rogers

- Based on the 10 Keys[™] to Healthy Aging program (Center for Aging and Population Health Prevention Research Center)
- 4 sessions (~1 hr each, over 2-3 mos)
- Aging Successfully With Pain RCT



ACHIEVE-P Eligibility, Randomization and Follow-up



- * Eligible but not enrolled because recruitment targets had been reached
- ** Unrelated to study intervention

ACHIEVE-P

6-Month Change in Standardized Proximal Outcomes, N=40

Outcome	Hearing Intervention	Successful Aging Intervention
	Mean (SD), N=20	Mean (SD), N=20
Perceived handicap due to HI*+	-1.40 (0.96)	0.02 (0.68)
Loneliness*	-0.19 (0.87)	0.22 (0.94)
Social Network**		
Number of people‡	0.17 (0.65)	-0.42 (0.66)
Diversity	0.15 (1.25)	-0.12 (0.70)
Social Function ^{**}	0.00 (0.65)	-0.26 (0.91)
Mental Function ^{**}	0.26 (0.80)	-0.14 (0.60)
Physical Function**	0.11 (0.76)	-0.07 (0.40)

* Lower scores are better; **Higher scores are better +p<0.0001; +p<0.01

ACHIEVE-P

6-Month Change in Standardized Cognitive Domain Score, N=40

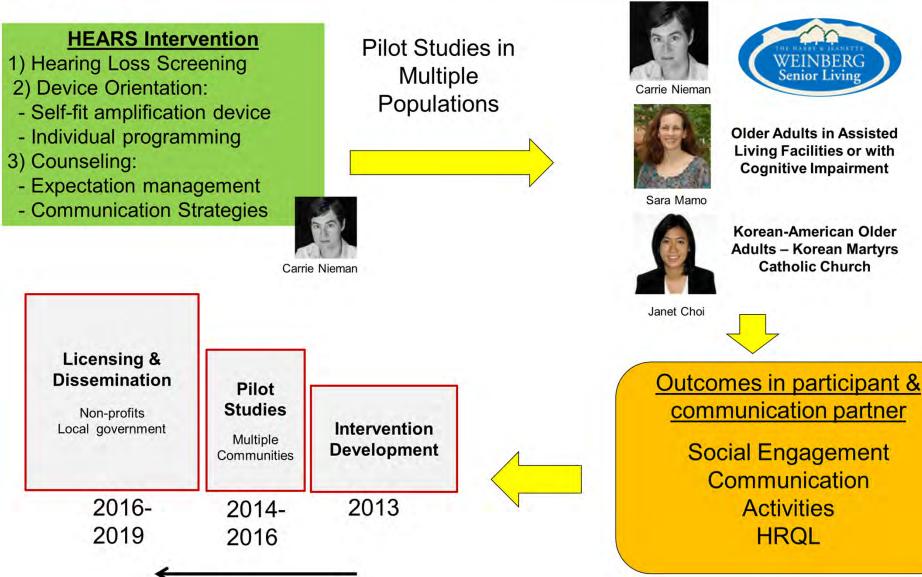
Cognitive Domain	Hearing Intervention	Successful Aging Intervention
	Mean (SD), n=20	Mean (SD), n=20
Memory	0.48 (0.69)	0.19 (0.66)
Language	0.05 (0.38)	0.00 (0.42)
Executive Function	0.03 (0.42)	0.17 (0.47)
Global Function	0.16 (0.42)	0.14 (0.39)

Full ACHIEVE Randomized Trial

- 750 participants: 70-84 year-old cognitively-normal older adults with hearing loss
- 1:1 randomization hearing vs. successful aging
- Follow-up at 6 mos & then annually for 3 years
- Primary outcome: change in global function
 - Powered to detect a minimum of a 0.30 SD difference
- Proximal outcomes: speech/audibility understanding, hearing aid use
- Secondary outcomes: domain-specific cognitive function, social fxn, physical fxn, physical activity

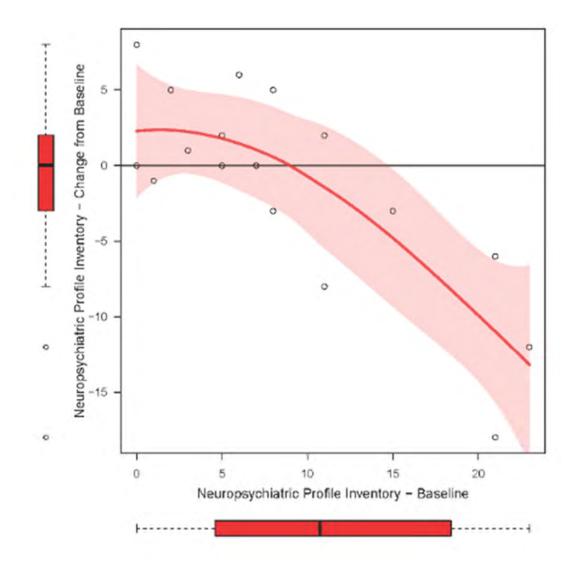


Access HEARS: Hearing care Equality through Accessible Research & Solutions



Memory Clinic HEARS

Results (N=20) of Neuropsychiatric Inventory Scores Post Intervention





Acknowledgements



BLSA BALTIMORE LONGITUDINAL STUDY OF AGING

- Frank R. Lin, MD, PhD
- Jennifer Deal, PhD
- Sara Mamo, AuD PhD
- Marilyn Albert, PhD
- Josef Coresh, MD PhD
- Luigi Ferrucci, MD PhD
- Tamara Harris, PhD
- Elizabeth Helzner, PhD
- David Knopman, MD
- Karen Bandeen-Roche, PhD
- Thomas H. Mosley, PhD
- Sheila Pratt, PhD
- Susan Resnick, PhD
- Suzanne Satterfield, MD DrPH
- A. Richey Sharrett, MD DrPH
- Eleanor Simonsick, PhD
- Lisa M. Wruck, PhD
- Kristine Yaffe, MD

- Eleanor Schwartz Charitable Foundation
- National Heart, Lung, and Blood Institute (NHLBI) contracts
 HHSN268201100005C,
 HHSN268201100006C,
 HHSN268201100007C,
 HHSN268201100008C,
 HHSN268201100009C,
 HHSN268201100010C,
 HHSN268201100011C,
 HHSN268201100012C.
- NIDCD K23DC011279
- National Institute on Aging (NIA) Contracts N01-AG-6-2101; N01-AG-6-2103; N01-AG-6-2106; NIA grant R01-AG028050, and NINR grant R01-NR012459. This research was supported in part by the Intramural Research Program of the NIH, National Institute on Aging

Thank you!

nreed9@jhmi.edu