Mechanisms of the Hearing–Cognition Relationship

Susan E. Fulton, Ph.D., CCC-A,¹ Jennifer J. Lister, Ph.D., CCC-A,² Aryn L. Harrison Bush, Ph.D.,² Jerri D. Edwards, Ph.D.,² and Ross Andel, Ph.D.²

ABSTRACT

This review provides a description of age-related changes in hearing and cognition, the relationship between hearing and cognition, and several potential mechanisms that underlie the relationship. Several studies have shown a significant relationship between peripheral hearing loss and cognitive impairment/decline but other studies have not. Furthermore, poor performance on measures of central auditory processing has been significantly associated with cognitive impairment. Important to understanding these relationships are the nature of the underlying mechanisms. Possible mechanisms are overdiagnosis, widespread neural degeneration, sensory degradation/deprivation, cognitive resource allocation/depletion, and social isolation/depression. Overdiagnosis occurs when hearing loss impacts tests of cognitive function or vice versa. Widespread neural degeneration can impact hearing, cognition, or both. Sensory degradation/deprivation due to hearing loss can result in neural degradation and reduced cognitive function. Increased demands due to hearing loss can result in changes in neural resource allocation, reducing available resources for cognitive function. Finally, hearing difficulties can cause social isolation and even depression, increasing the risk for cognitive decline. Data from our laboratory provide support for cognitive resource allocation/depletion. Understanding all five of these mechanisms will advance the development of effective interventions and treatments, thereby enhancing the quality of life of older adults.

KEYWORDS: Hearing loss, peripheral hearing, central auditory processing, cognition, older adults

Cognition and the Aging Auditory System; Guest Editor, Lindsey E. Jorgensen, Au.D., Ph.D.

Semin Hear 2015;36:140–149. Copyright © 2015 by Thieme Medical Publishers, Inc., 333 Seventh Avenue, New York, NY 10001, USA. Tel: +1(212) 584-4662. DOI: http://dx.doi.org/10.1055/s-0035-1555117. ISSN 0734-0451.

¹Department of Communication Sciences and Disorders, College of Arts and Sciences, University of South Florida Sarasota-Manatee, Sarasota, Florida; ²University of South Florida, Tampa, Florida.

Address for correspondence: Susan E. Fulton, Ph.D., CCC-A, University of South Florida Sarasota-Manatee, Speech-Language Sciences, College of Arts and Sciences, 8350 N. Tamiami Trail, B214, Sarasota, FL 34243 (e-mail: sefulton@sar.usf.edu).

Learning Outcomes: As a result of this activity, the participant will be able to (1) list the five key hearingcognition mechanisms; (2) discuss the impact of hearing loss on cognition.

Research suggests a significant, albeit weak, relationship between hearing and cognition. Less is known about the mechanisms of this relationship or how this relationship informs clinical audiology practice. This review will provide a description of age-related changes in hearing and cognition, the relationship between hearing and cognition, and several potential mechanisms that underlie the relationship.

Hearing may be described as having two domains: (1) peripheral hearing-the transmission of sound through the auditory periphery (i.e., outer ear, middle ear) and encoding of sound by the cochlea; and (2) central auditory processing (CAP)-the processing and decoding of sound through higher levels of the central auditory nervous system (i.e., brainstem, midbrain, auditory cortex). With age, declines in both peripheral hearing and CAP occur. In fact, peripheral hearing loss is the third most common chronic health condition in older adults,¹ and CAP disorder affects ~23 to 76% of older adults.^{2,3} Actual numbers may be larger, as many individuals with hearing difficulties do not pursue a hearing evaluation or diagnosis.^{4,5} Hearing impairment in either domain negatively impacts communication, health, quality of life, confidence, and self-image and has been associated with feelings of frustration, loneliness, depression, anxiety, paranoia, and emo-tional turmoil.⁶⁻⁹

Another condition associated with aging is cognitive impairment. Age-related cognitive decline can progress to mild cognitive impairment (MCI) and further deteriorate into dementia. Dementia (e.g., Alzheimer disease [AD]) and the often precursory stage of MCI are considered to be the most feared and devastating conditions that older adults face.^{10,11} MCI is estimated to affect 22.6% of older adults, and individuals with MCI are at an elevated risk of AD or related dementia.¹¹⁻¹³ The number of individuals with dementia is expected to double every 20 years, resulting in over 65 million affected people worldwide by 2030.14 Most important for this review is the long-established relationship between impairment in both domains of hearing and impaired cognition.

THE AUDITORY SYSTEM AND COGNITION

Peripheral Hearing and Cognition

Peripheral hearing loss has been associated with poorer cognitive performance and accelerated cognitive decline.^{15–17} Specifically, peripheral hearing loss has been associated with poorer performance on global cognitive assessments,^{15,16,18–20} as well as cognitive measures memory,^{16,19,21–23} executive of function,^{15,16,19,24} and speed of processing.¹⁹ In addition, the risk of cognitive decline increases linearly with hearing loss severity.15 As the degree of hearing loss increases in older adults, the risk for dementia is thought to significantly increase.²⁵ Thus, there is potential to intervene before dementia onset. To this end, understanding the relationship between hearing and cognitive decline is of critical importance.

Still, most of the evidence examining the relationship between peripheral hearing and cognitive impairment is correlational and cross-sectional in nature. Thus, the cause-effect direction of the relationship is not clear. For example, cognitive changes in older adults, such as processing speed, memory, and attention, can impact the processing of auditory information. Indeed, cognitive decline has been associated with faster rates of decline in peripheral hearing sensitivity.^{20,26,27} Kiely et al used established cohort data to follow audiological thresholds (0.5, 1, 2, 3, 4, 6, and 8 kHz), health data (e.g., blood pressure, hypertension, stroke, etc.), and global cognitive status (Mini Mental State Exam [MMSE]) for 4,221 participants (mean age = 73.6 years) for up to 4 sessions over an 11-year span.²⁰ Multivariate-adjusted linear mixed models indicated factors that were predictive of hearing sensitivity decline. After adjusting for socioeconomic status and health variables, cognitive status was associated with

poorer pure tone average (PTA) and faster rates of change in PTA. These relationships were evident while controlling for age. The authors suggest that these results reflect effects of topdown processing such that impaired cognitive functioning may impact hearing performance.

Other studies have found no relationship between peripheral hearing loss and cognitive decline.^{28–31} Lin et al examined the association between visual impairment (visual acuity worse than 20/40), hearing impairment (2 kHz threshold greater than 40-dB hearing level [HL]), and global cognitive status (greater than 1 standard deviation [SD] change in MMSE) across 4 years.³¹ Visual impairment and combined sensory impairment were associated with cognitive decline. Peripheral hearing loss alone was not associated with cognitive decline. However, both the use of an insensitive measure of global cognitive status (MMSE) and the limited hearing measurement (threshold at 2 kHz only) could have been factors in the null results of this study.

Although large, longitudinal studies may offer greater insight into the time course of the relationship between hearing loss and cognitive impairment, few have been completed. One exception is a study by Lin et al in which 1,162 older adults with hearing loss (mean age 77.9 years) and 822 older adults without hearing loss (mean age = 76.8 years) were followed for 6 years.¹⁵ Normal hearing was defined as a 4 frequency PTA (0.5, 1, 2, and 4 kHz) of 25-dB HL or better, measured at baseline. Cognition was assessed using the Modified Mini-Mental State Test (3MS; a measure of global cognitive status) and the Digit Symbol Substitution Test (DSST; a measure of cognitive speed of processing).^{32,33} The 3MS is a global cognitive screening tool that briefly assesses orientation, concentration, language, praxis, and memory.³² The DSST is a nonverbal test of speed of processing and also may tap executive function.³³ Using these two measures, adults with hearing loss were found to have annual rates of decline that were 30 to 40% higher than older adults with normal hearing. Rates of cognitive decline were linearly related to baseline hearing loss severity. Note that hearing sensitivity was measured only at baseline; subsequent audiometric assessments were not administered beyond the baseline assessment. In addition, only two cognitive measures (3MS and DSST) were used and one was a global screening measure (3MS); thus, only one cognitive domain (speed of processing) was assessed.

To more fully examine the relationship between peripheral hearing and the many facets of cognition, a variety of cognitive measures tapping different cognitive domains should be used. Bush et al examined the cross-sectional association between peripheral hearing (PTA for 0.5, 1, and 2 kHz), 10 measures of cognition, and one global measure of cognitive status. Measures of cognition were used to assess speed of processing (five tests), executive function (two tests), and memory (three tests).¹⁹ Participants (n = 894; mean age = 73.47 years) from the Staying Keen in Later Life (SKILL) study cohort were used. Multiple regression analysis showed PTA in the better-hearing ear to be related to a broad range of cognitive measures even after controlling for known risk factors for peripheral hearing loss (i.e., age, race, sex, education, diabetes, hypertension, stroke, heart disease, and depression). When examining a subsample of individuals with hearing loss (PTA > 25-dB HL), PTA accounted for a small but significant amount of the variance in measures of global cognitive status, executive function (one measure), and memory (three measures). However, no relationship between PTA and processing speed was found when examining the subsample of those with hearing loss.

Discrepancies exist regarding the relationship between peripheral hearing and cognition. Although some studies suggest a relationship between peripheral hearing and cognition,^{15–17,24} other studies indicate no relationship.^{28–31} Small, but significant relationships were found when large population samples were used,^{15,17,24} but no significant relationships were found in smaller studies.^{29,30,34} The cognitive outcome evaluated (cognitive performance, cognitive status, dementia diagnosis) also may have contributed to the fact that differing conclusions have been reported.^{17,31,35}

Central Auditory Processing and Cognition

Unlike peripheral hearing, decreased CAP abilities have been associated with poor cognitive

performance and cognitive decline in relatively small samples. MCI and AD have been associated with decreased performance on tests of CAP, such as speech in noise^{28,29,34,36} and dichotic digits.^{30,35,37,38} As performance on many CAP measures is related to the ability to understand speech in background noise,³⁹⁻⁴² the relationship between performance on CAP tasks and cognition may help identify sources of speech-in-noise difficulties often reported by older listeners. Anderson et al examined peripheral hearing, CAP, multiple cognitive skills (auditory attention, auditory short-term memory, and auditory working memory), and life experience (e.g., socioeconomic status, education level, physical activity level, and musical training) in 120 older adults (mean age = 69.89).⁴³ Cognitive and CAP abilities were significantly related to speech understanding in noise but peripheral hearing was not, supporting several other small sample studies that found no relationship between peripheral hearing loss and cognitive function when both peripheral and CAP factors were examined in concert.^{29,30,34,36} Gates et al examined PTA (0.5, 1, and 2 kHz), performance on competing sentence tasks, and MMSE scores.²⁹ In another study, Gates et al examined pure tone thresholds (250 to 8,000 Hz), otoacoustic emissions, performance on competing sentence tasks, and scores on the Clinical Dementia Rating Scale.44,45 Both studies showed significant relationships between CAP and cognitive function but not between peripheral hearing and cognitive function.

Recent studies in our laboratory have examined the relationship between peripheral hearing sensitivity, CAP (using both speech and nonspeech stimuli), and multiple measures of cognitive function. 46,47 Individuals with MCI performed significantly worse than individuals without MCI on competing sentence tasks (Synthetic Sentence Identification⁴¹ and Dichotic Sentence Identification³⁹) and temporal gap detection (Adaptive Tests of Temporal Resolution⁴⁰), but peripheral hearing sensitivity did not differ between groups for this small sample (n = 30). The results of these studies suggest that CAP and cognition are related more strongly than are peripheral hearing and cognition. Additionally, CAP measures may be useful tools to help identify or assess cognitive impairment.

Auditory Event-Related Potentials and Cognition

The bulk of research examining hearing and cognition has relied heavily on behavioral measures, which have limitations such as listener attention and fatigue, and examiner subjectivity. A less subjective means of examining the hearing-cognition relationship may be auditory event-related potentials (ERPs). ERPs provide an objective, noninvasive method of examining both sensory and cognitive processes with high temporal resolution. Neural transmission time (latency) and strength of response (amplitude) can be recorded in response to various stimuli. ERPs can be effectively used to examine waveform latency and amplitude changes related to cognitive impairment and dementia. 48-50 Recent data have shown that among older adults with sloping high frequency hearing loss, the P1-N1-P2 complex is longer in latency and lower in amplitude for those individuals with MCI compared with individuals without MCI.47 Increased P1 latency and amplitude and longer P3 latency also has been found for those with MCI.⁴⁹ In addition, the hemispheric activation pattern appears more diffuse for those with MCI.⁴⁷ These results suggest that the auditory signal traveling superiorly in the system appears to be degraded in individuals with MCI as compared with those without. Interestingly, ERPs may be useful indicators of emergent MCI or AD. Golob et al found ERP latencies (N1) to be longer in individuals with gene mutations consistent with familial AD than in individuals without the gene mutation, in the absence of any behavioral signs of MCI or AD.⁵¹

Regardless of the form they take, auditory identifiers of early cognitive impairment would be beneficial in guiding treatment in asymptomatic or prodromal stages of the disease. However, separation of peripheral, central, and cognitive processes is very difficult.⁵² These processes are clearly interrelated and decreased functioning in one area may impact another as the degraded signal travels along the auditory pathway to the brain.

HEARING-COGNITION MECHANISMS

Despite the long history of behavioral studies and the recent history of ERP studies, little is known about the underlying mechanisms of the hearing-cognition relationship. Given the large number of Americans who have hearing loss and cognitive impairment, a better understanding of the hearing-cognition mechanisms is crucial. Several potential mechanisms have been suggested, including overdiagnosis, 23,53-55 widespread neural degeneration,^{18,56,57} sensory degradation/deprivation,^{26,58,59} cognitive resource allocation/depletion,⁵⁹⁻⁶² and social isolation/depression.⁶³⁻⁶⁶ It is possible that the mechanism is the combination of any or all of these factors (e.g., multilevel model⁶¹). Furthermore, the relationship between hearing and cognition may be bidirectional in nature.²⁰ The potential mechanisms are discussed below.

Overdiagnosis

Many older adults do not seek diagnosis or treatment for hearing-related difficulties.⁴ In addition, many older listeners with hearing aids or assistive devices may not use them routinely.⁴ Older adults have been known to pretend they understand instructions or test prompts, rather than request clarification. Such behavior may reduce performance on cognitive measures and masquerade as clinically significant cognitive impairment.^{53,67} In this case, treatment of the hearing impairment should resolve the appearance of cognitive impairment.

One of the most common tests used to assess global cognitive status and screen for cognitive impairment is the MMSE. This test is administered verbally, often in a physician's office with background noise present. Not only is the listener receiving degraded instructions and test prompts due to hearing loss, but he or she is also contending with distractions and distortions in the signal. Background noise can affect a speech signal by masking high-frequency cues, filling silent gaps, and temporally distorting the signal.^{68–70}

To quantify the impact of aging and hearing loss on MMSE performance, Jorgensen examined the effects of simulated hearing loss on MMSE scores in 125 young adults.⁵⁵ Five audibility levels were used: no simulation (i.e., normal hearing) and four levels of simulated sloping hearing losses (mild to moderately severe, mild to severe, moderate to severe, and severe to profound). The MMSE was administered at all five levels. Results showed that MMSE performance was significantly affected by audibility, with incorrect identification of cognitive decline when hearing loss was simulated. At the highest audibility level used, a mild to moderately severe hearing loss, 16% of the young adults were misdiagnosed as having dementia. The rate of misdiagnosis increased with lower levels of audibility. The results suggest that hearing loss should be examined prior to cognitive assessment so that audibility may be ensured when administering cognitive assessments; this will reduce the chances of misdiagnosis.55

Widespread Neural Degeneration

Lindenberger and Baltes suggested a common factor between age-related changes in the brain and hearing, vision, and cognitive function.⁷¹ Additionally, Anstey et al found age, speed, vision, and hearing comprised a common factor in age-related changes in cognition.⁵⁶ This common cause mechanism suggests that age-related changes have a central or neural component in addition to any peripheral or end organ source.¹⁸

Indeed, a genetic association between hearing loss and dementia, such as hereditary sensory autonomic neuropathy with dementia and hearing loss, has been found. Klein et al found gene mutations of DNA methyltransferase 1 (DNMT1) were related to both hearing loss and dementia.⁷² DNMT1 is an enzyme involved in development, maintenance, and connectivity of neural cells throughout the central nervous system. Individuals found with DNMT1 mutations developed progressive hearing loss and sensory neuropathy when young (midteens to 30 years) and cognitive decline later in life (30 to 50 years). This association suggests that a widespread failing of neural function may first be evidenced by hearing loss and only later by clinical onset of dementia.

Sensory Degradation/Deprivation

Hearing loss results in a degradation of and loss of input to the brain. Peripheral hearing loss

impacts the audibility and clarity of sound; thus, the information reaching the cortex is degraded and the cortex is deprived of quality sensory information.²⁶ Over time, this degradation/ deprivation results in changes to the structure and function of the central auditory and cognitive systems, as shown in studies of animals and humans.^{59,73–77} Lin et al found neuroimaging evidence of brain atrophy and volumetric decline (structural changes) in the right temporal lobe of individuals with hearing loss, as compared with individuals with normal hearing, across ~ 6 years.¹⁶ Behaviorally, degraded auditory signals have been associated with functional changes such as decreased speech-in-noise performance,²⁶ comprehension and memory for spoken language,^{78,79} and cognitive impairment.⁵⁹

Another way to assess the impact of auditory sensory deprivation is to examine the cognitive changes that occur when hearing loss is treated with amplification. Studies have shown that hearing aid use improves signal detection and speech understanding; however, no strong evidence for long-term cognitive changes due to hearing aid use has been found.^{4,80} Although they did not directly study cognition, Palmer et al assessed behavioral changes in individuals with MCI or AD after short-term use of amplification.⁸¹ Caregivers reported greater compliance with hearing aid use, reduced problem behaviors, and reduced perceived hearing handicap after 2 months of amplification use. Many listeners wait years before seeking treatment for hearing loss; thus, the subsequent effects of long-term sensory deprivation may not be easily reversed. As with most diseases and disorders, early hearing loss identification leads to the best treatment effects. It is important to note that hearing aid use will not compensate for all the aforementioned effects of sensory degradation/deprivation; other accommodations may still be necessary when assessing cognitive function in individuals with hearing impairment.

Cognitive Resource Allocation/ Depletion

When an incoming auditory signal is degraded due to hearing loss or external noise, a greater number of neural resources must be allocated

for auditory processing of the degraded signal and, as a result, fewer resources remain for other, higher-level cognitive processes (e.g., working memory, recall).^{16,26,82} Imaging studies have shown that brain activation patterns change when processing tasks are challenging.^{83,84} In addition, decreased gray matter density in the primary auditory areas of individuals with hearing loss suggests a reorganization of processing systems when auditory stimulation is decreased.⁷⁶ MCI can impact resource allocation, as well. A recent study in our laboratory showed hemispheric activation during an auditory ERP task to be different and more widespread for individuals with MCI, suggesting a greater number of resources allocated to auditory detection.47

One way to examine allocation/depletion is to measure cognitive capacity following resource allocation in difficult listening situations (cognitive spare capacity). Mishra and colleagues examined the cognitive spare capacity of 24 older listeners (mean age = 69, SD = 4.7) with hearing loss (mean 4 frequency PTA = 34.5, SD = 3.6).⁸⁵ The Cognitive Spare Capacity Test was administered in quiet and in noise.⁸⁶ In addition, tests of cognitive ability (reading span, text reception threshold, letter memory, delayed free recall of the reading span test, and the Simon task) were administered. Amplification was used with the older listeners to compensate for hearing loss. Results were compared with similar, previous testing with young adults with normal hearing.⁸⁶ No difference in cognitive spare capacity was found between groups when the listening conditions were optimal. However, when presented in background noise, older listeners performed worse on the Cognitive Spare Capacity Test and cognitive tests. The background noise significantly affected older adults' performance, perhaps by decreasing the cognitive resources available to complete the tasks successfully. Therefore, cognitive function may be diminished in older listeners due to reallocation of valuable resources in an effort to process degraded auditory signals.

Social Isolation/Depression

As discussed, hearing difficulties place increased demands on cognitive processes and

resources. Many social gatherings are held in meeting rooms, restaurants, and social halls where a high level of background noise is present, increasing the effort required to communicate and creating challenging situations for any listener. Many individuals with hearing loss and/or cognitive impairment withdraw from these situations. The result is social isolation or a reduction in association with family, friends, or colleagues. As the degree of hearing loss increases, so do the odds of being socially isolated.⁶⁶ Hearing loss has been associated with anxiety, depression, self-esteem, and well-being.⁸⁷ Specifically, older adults with hearing loss are more likely to experience depressive symptoms, lower self-efficacy and mastery, more feelings of loneliness, and a smaller social network than normally hearing peers.63-65 Furthermore, studies have found that older adults with depressive symptoms (i.e., feelings of social isolation or loneliness) have an increased risk of cognitive decline.⁸⁸⁻⁹¹ This research implies a cascade from hearing loss to social isolation/depression to cognitive impairment. However, there is little, if any, research to help us understand how hearing, psychosocial, and cognitive factors interact. Such research is essential to design strategies to intervene at the point of hearing problems, social isolation, and/or cognitive difficulties.

CONCLUSIONS

Hearing loss may be a promising indicator of the prodromal stages of cognitive decline. A better understanding of the relationship between hearing and cognition is critical, as intervention prior to dementia onset may be possible. Currently, conflicting evidence exists regarding the respective contributions of peripheral hearing and CAP to cognitive function. More research is needed to tease apart the interaction between peripheral and/or central auditory function and cognitive decline.

Key to understanding this relationship is expanding our knowledge of the underlying mechanisms. Several potential mechanisms have been identified. Overdiagnosis can occur when undiagnosed hearing loss affects cognitive test results or vice versa, when undiagnosed cognitive impairment affects hearing evaluation results. Widespread neural degeneration due to aging can impact both sensory and cognitive function. In fact, certain genetic mutations are manifested first by hearing loss and then by cognitive decline. Sensory degradation/deprivation due to hearing loss can result in neural degradation and reduced function. Cognitive resource allocation occurs when a degraded sensory signal requires rearrangement of cognitive resources for processing, to the detriment of higher-order thinking and memory. Hearing loss can lead to social isolation and depression, resulting in an increased risk of cognitive decline. All five of these mechanisms demonstrate the delicate relationship between auditory and cognitive function. A better understanding of these mechanisms will advance the development of effective interventions and treatments, thereby enhancing the quality of life of older adults.

REFERENCES

- Cruickshanks KJ, Wiley TL, Tweed TS, et al; The Epidemiology of Hearing Loss Study. Prevalence of hearing loss in older adults in Beaver Dam, Wisconsin. Am J Epidemiol 1998;148(9):879–886
- Cooper JC Jr, Gates GA. Hearing in the elderly the Framingham cohort, 1983–1985: Part II. Prevalence of central auditory processing disorders. Ear Hear 1991;12(5):304–311
- Golding M, Carter N, Mitchell P, Hood LJ. Prevalence of central auditory processing (CAP) abnormality in an older Australian population: the Blue Mountains Hearing Study. J Am Acad Audiol 2004;15(9):633–642
- Wong LLN, Yu JKY, Chan SS, Tong MCF. Screening of cognitive function and hearing impairment in older adults: a preliminary study. Biomed Res Int 2014;2014:867852
- van Hooren SAH, Anteunis LJ, Valentijn SAM, et al. Does cognitive function in older adults with hearing impairment improve by hearing aid use? Int J Audiol 2005;44(5):265–271
- Arlinger S. Negative consequences of uncorrected hearing loss—a review. Int J Audiol 2003;42(2, Suppl 2):S17–S20
- Crews JE, Campbell VA. Vision impairment and hearing loss among community-dwelling older Americans: implications for health and functioning. Am J Public Health 2004;94(5): 823–829
- Gates GA, Mills JH. Presbycusis. Lancet 2005; 366(9491):1111-1120

- The National Council on the Aging. The Consequences of Untreated Hearing Loss in Older Persons. Washington, DC: NCOA; 1999:1–9
- Corner L, Bond J. Being at risk of dementia: fears and anxieties of older adults. J Aging Stud 2004; 18:143–155
- Morris JC, Storandt M, Miller JP, et al. Mild cognitive impairment represents early-stage Alzheimer disease. Arch Neurol 2001;58(3):397–405
- Lopez OL, Jagust WJ, DeKosky ST, et al. Prevalence and classification of mild cognitive impairment in the Cardiovascular Health Study Cognition Study: part 1. Arch Neurol 2003; 60(10):1385–1389
- Petersen RC, Smith GE, Waring SC, Ivnik RJ, Tangalos EG, Kokmen E. Mild cognitive impairment: clinical characterization and outcome. Arch Neurol 1999;56(3):303–308
- Prince M, Bryce R, Albanese E, Wimo A, Ribeiro W, Ferri CP. The global prevalence of dementia: a systematic review and metaanalysis. Alzheimers Dement 2013;9(1):63–75.e2
- Lin FR, Yaffe K, Xia J, et al; Health ABC Study Group. Hearing loss and cognitive decline in older adults. JAMA Intern Med 2013;173(4): 293–299
- Lin FR, Ferrucci L, Metter EJ, An Y, Zonderman AB, Resnick SM. Hearing loss and cognition in the Baltimore Longitudinal Study of Aging. Neuropsychology 2011;25(6):763–770
- Lin FR, Metter EJ, O'Brien RJ, Resnick SM, Zonderman AB, Ferrucci L. Hearing loss and incident dementia. Arch Neurol 2011;68(2): 214–220
- Baltes PB, Lindenberger U. Emergence of a powerful connection between sensory and cognitive functions across the adult life span: a new window to the study of cognitive aging? Psychol Aging 1997;12(1):12–21
- Bush AL, Lister JJ, Lin FR, Betz J, Edwards JD. Peripheral hearing and cognition: Evidence from the Staying Keen in Later Life (SKILL) study. Ear Hear 2015 DOI: 10.1097/AUD.000000000000142
- Kiely KM, Gopinath B, Mitchell P, Luszcz M, Anstey KJ. Cognitive, health, and sociodemographic predictors of longitudinal decline in hearing acuity among older adults. J Gerontol A Biol Sci Med Sci 2012;67(9):997–1003
- Pearman A, Friedman L, Brooks JO, Yesavage JA. Hearing impairment and serial word recall in older adults. Exp Aging Res 2000;26(4):383–391
- 22. van Boxtel MPJ, van Beijsterveldt CEM, Houx PJ, Anteunis LJ, Metsemakers JFM, Jolles J. Mild hearing impairment can reduce verbal memory performance in a healthy adult population. J Clin Exp Neuropsychol 2000;22(1):147–154
- Valentijn SAM, van Boxtel MPJ, van Hooren SAH, et al. Change in sensory functioning predicts

change in cognitive functioning: results from a 6year follow-up in the Maastricht aging study. J Am Geriatr Soc 2005;53(3):374–380

- Lin FR. Hearing loss and cognition among older adults in the United States. J Gerontol A Biol Sci Med Sci 2011;66(10):1131–1136
- Uhlmann RF, Larson EB, Rees TS, Koepsell TD, Duckert LG. Relationship of hearing impairment to dementia and cognitive dysfunction in older adults. JAMA 1989;261(13):1916–1919
- Pichora-Fuller MK, Schneider BA, Daneman M. How young and old adults listen to and remember speech in noise. J Acoust Soc Am 1995;97(1): 593–608
- Gordon-Salant S. Hearing loss and aging: new research findings and clinical implications. J Rehabil Res Dev 2005;42(4, Suppl 2):9–24
- Gates GA, Beiser A, Rees TS, D'Agostino RB, Wolf PA. Central auditory dysfunction may precede the onset of clinical dementia in people with probable Alzheimer's disease. J Am Geriatr Soc 2002;50(3):482–488
- Gates GA, Cobb JL, Linn RT, Rees T, Wolf PA, D'Agostino RB. Central auditory dysfunction, cognitive dysfunction, and dementia in older people. Arch Otolaryngol Head Neck Surg 1996; 122(2):161–167
- Idrizbegovic E, Hederstierna C, Dahlquist M, Kämpfe Nordström C, Jelic V, Rosenhall U. Central auditory function in early Alzheimer's disease and in mild cognitive impairment. Age Ageing 2011;40(2):249–254
- Lin MY, Gutierrez PR, Stone KL, et al; Study of Osteoporotic Fractures Research Group. Vision impairment and combined vision and hearing impairment predict cognitive and functional decline in older women. J Am Geriatr Soc 2004;52(12): 1996–2002
- Teng EL, Chui HC. The Modified Mini-Mental State (3MS) examination. J Clin Psychiatry 1987; 48(8):314–318
- Wechsler D. Wechsler Adult Intelligence Scale– Revised. San Antonio, TX: Psychological Corporation; 1981
- Strouse AL, Hall JW III, Burger MC. Central auditory processing in Alzheimer's disease. Ear Hear 1995;16(2):230–238
- Gates GA, Anderson ML, Feeney MP, McCurry SM, Larson EB. Central auditory dysfunction in older persons with memory impairment or Alzheimer dementia. Arch Otolaryngol Head Neck Surg 2008;134(7):771–777
- Zekveld AA, George ELJ, Houtgast T, Kramer SE. Cognitive abilities relate to self-reported hearing disability. J Speech Lang Hear Res 2013;56(5): 1364–1372
- Gates GA, Anderson ML, McCurry SM, Feeney MP, Larson EB. Central auditory dysfunction as a

harbinger of Alzheimer dementia. Arch Otolaryngol Head Neck Surg 2011;137(4):390–395

- Idrizbegovic E, Hederstierna C, Dahlquist M, Rosenhall U. Short-term longitudinal study of central auditory function in Alzheimer's disease and mild cognitive impairment. Dement Geriatr Cogn Dis Extra 2013;3(1):468–471
- Fifer RC, Jerger JF, Berlin CI, Tobey EA, Campbell JC. Development of a dichotic sentence identification test for hearing-impaired adults. Ear Hear 1983;4(6):300–305
- Lister JJ, Roberts RA, Shackelford J, Rogers CL. An adaptive clinical test of temporal resolution. Am J Audiol 2006;15(2):133–140
- Jerger J, Speaks C, Trammell JL. A new approach to speech audiometry. J Speech Hear Disord 1968; 33(4):318–328
- 42. Moore DR, Edmondson-Jones M, Dawes P, et al. Relation between speech-in-noise threshold, hearing loss and cognition from 40-69 years of age. PLoS ONE 2014;9(9):e107720
- Anderson S, White-Schwoch T, Parbery-Clark A, Kraus N. A dynamic auditory-cognitive system supports speech-in-noise perception in older adults. Hear Res 2013;300:18–32
- 44. Gates GA, Karzon RK, Garcia P, et al. Auditory dysfunction in aging and senile dementia of the Alzheimer's type. Arch Neurol 1995;52(6):626–634
- Morris JC. The Clinical Dementia Rating (CDR): current version and scoring rules. Neurology 1993; 43(11):2412–2414
- 46. Brandino A, Matthews C, Valdés EG, et al. Central and peripheral auditory processing in individuals with and without mild cognitive impairment. Paper presented at: American Academy of Audiology Conference; March 27– 29 2014; Orlando, FL
- 47. Fluegel BA, Matthews C, Harrison Bush AL, Lister JJ, Edwards JD, Andel R. Neurophysiological indicators of early-stage cognitive decline. Paper presented at: American Academy of Audiology Conference; March 27–29 2014; Orlando, FL
- Golob EJ, Irimajiri R, Starr A. Auditory cortical activity in amnestic mild cognitive impairment: relationship to subtype and conversion to dementia. Brain 2007;130(Pt 3):740–752
- Golob EJ, Johnson JK, Starr A. Auditory eventrelated potentials during target detection are abnormal in mild cognitive impairment. Clin Neurophysiol 2002;113(1):151–161
- Irimajiri R, Golob EJ, Starr A. Auditory brainstem, middle- and long-latency evoked potentials in mild cognitive impairment. Clin Neurophysiol 2005;116(8):1918–1929
- Golob EJ, Ringman JM, Irimajiri R, et al. Cortical event-related potentials in preclinical familial Alzheimer disease. Neurology 2009;73(20): 1649–1655

- 52. Humes LE, Dubno JR, Gordon-Salant S, et al. Central presbycusis: a review and evaluation of the evidence. J Am Acad Audiol 2012;23(8):635–666
- Arlinger S, Lunner T, Lyxell B, Pichora-Fuller MK. The emergence of cognitive hearing science. Scand J Psychol 2009;50(5):371–384
- Herbst KG, Humphrey C. Hearing impairment and mental state in the elderly living at home. BMJ 1980;281(6245):903–905
- 55. Jorgensen LE. The Potential Impact of Undiagnosed Hearing Loss on the Diagnosis of Dementia [Doctoral dissertation]. Pittsburgh, PA: University of Pittsburgh; 2012
- Anstey KJ, Luszcz MA, Sanchez L. Two-year decline in vision but not hearing is associated with memory decline in very old adults in a population-based sample. Gerontology 2001; 47(5):289–293
- Lindenberger U, Baltes PB. Sensory functioning and intelligence in old age: a strong connection. Psychol Aging 1994;9(3):339–355
- Oster C. Sensory deprivation in geriatric patients. J Am Geriatr Soc 1976;24(10):461–464
- Schneider BA, Pichora-Fuller MK. Implications of perceptual deterioration for cognitive aging research. In: Craik FIM, Salthouse TA, eds. The Handbook of Aging and Cognition. Vol. 2. Mahwah, NJ: Lawrence Erlbaum Associates; 2000: 155–220
- Tun PA, Wingfield A, Lindfield KC. Motor-speed baseline for the Digit-Symbol Substitution Test. Clin Gerontol 1997;18:47–51
- Li KZ, Lindenberger U. Relations between aging sensory/sensorimotor and cognitive functions. Neurosci Biobehav Rev 2002;26(7):777–783
- Li KZ, Lindenberger U, Freund AM, Baltes PB. Walking while memorizing: age-related differences in compensatory behavior. Psychol Sci 2001;12(3): 230–237
- 63. Gopinath B, Hickson L, Schneider J, et al. Hearing-impaired adults are at increased risk of experiencing emotional distress and social engagement restrictions five years later. Age Ageing 2012; 41(5):618–623
- 64. Gopinath B, Schneider J, McMahon CM, Teber E, Leeder SR, Mitchell P. Severity of age-related hearing loss is associated with impaired activities of daily living. Age Ageing 2012;41(2):195–200
- Kramer SE, Kapteyn TS, Kuik DJ, Deeg DJ. The association of hearing impairment and chronic diseases with psychosocial health status in older age. J Aging Health 2002;14(1):122–137
- Mick P, Kawachi I, Lin FR. The association between hearing loss and social isolation in older adults. Otolaryngol Head Neck Surg 2014;150(3): 378–384
- 67. Gordon-Salant S, Fitzgibbons PJ. Profile of auditory temporal processing in older

listeners. J Speech Lang Hear Res 1999;42(2): 300-311

- Lister JJ, Koehnke JD, Besing JM. Binaural gap duration discrimination in listeners with impaired hearing and normal hearing. Ear Hear 2000;21(2): 141–150
- Gordon-Salant S, Fitzgibbons PJ. Temporal factors and speech recognition performance in young and elderly listeners. J Speech Hear Res 1993;36(6): 1276–1285
- Houtgast T, Steeneken HJH. The modulation transfer function in room acoustics as a predictor of speech intelligibility. Acoustica 1973;28:66–73
- Lindenberger U, Baltes PB. Intellectual functioning in old and very old age: cross-sectional results from the Berlin Aging Study. Psychol Aging 1997; 12(3):410–432
- Klein CJ, Bird T, Ertekin-Taner N, et al. DNMT1 mutation hot spot causes varied phenotypes of HSAN1 with dementia and hearing loss. Neurology 2013;80(9):824–828
- Billings CJ, Tremblay KL, Stecker GC, Tolin WM. Human evoked cortical activity to signalto-noise ratio and absolute signal level. Hear Res 2009;254(1–2):15–24
- Kakigi A, Hirakawa H, Harel N, Mount RJ, Harrison RV. Tonotopic mapping in auditory cortex of the adult chinchilla with amikacin-induced cochlear lesions. Audiology 2000;39(3):153–160
- Schwaber MK, Garraghty PE, Kaas JH. Neuroplasticity of the adult primate auditory cortex following cochlear hearing loss. Am J Otol 1993; 14(3):252–258
- Peelle JE, Troiani V, Grossman M, Wingfield A. Hearing loss in older adults affects neural systems supporting speech comprehension. J Neurosci 2011;31(35):12638–12643
- Gussekloo J, de Craen AJM, Oduber C, van Boxtel MPJ, Westendorp RGJ. Sensory impairment and cognitive functioning in oldest-old subjects: the Leiden 85+ Study. Am J Geriatr Psychiatry 2005;13(9): 781–786
- Tun PA, Williams VA, Small BJ, Hafter ER. The effects of aging on auditory processing and cognition. Am J Audiol 2012;21(2):344–350
- Wingfield A, Tun PA. Cognitive supports and cognitive constraints on comprehension of spoken language. J Am Acad Audiol 2007;18(7):548–558

- Kalluri S, Humes LE. Hearing technology and cognition. Am J Audiol 2012;21(2):338–343
- Palmer CV, Adams SW, Bourgeois M, Durrant J, Rossi M. Reduction in caregiver-identified problem behaviors in patients with Alzheimer disease post-hearing-aid fitting. J Speech Lang Hear Res 1999;42(2):312–328
- Tun PA, McCoy S, Wingfield A. Aging, hearing acuity, and the attentional costs of effortful listening. Psychol Aging 2009;24(3):761–766
- Zarahn E, Rakitin B, Abela D, Flynn J, Stern Y. Age-related changes in brain activation during a delayed item recognition task. Neurobiol Aging 2007;28(5):784–798
- Holtzer R, Rakitin BC, Steffener J, Flynn J, Kumar A, Stern Y. Age effects on load-dependent brain activations in working memory for novel material. Brain Res 2009;1249:148–161
- Mishra S, Stenfelt S, Lunner T, Rönnberg J, Rudner M. Cognitive spare capacity in older adults with hearing loss. Frontiers in Aging Neurosci 2014;6:1–13
- Mishra S, Lunner T, Stenfelt S, Rönnberg J, Rudner M. Seeing the talker's face supports executive processing of speech in steady state noise. Front Syst Neurosci 2013;7:96
- Tambs K. Moderate effects of hearing loss on mental health and subjective well-being: results from the Nord-Trøndelag Hearing Loss Study. Psychosom Med 2004;66(5):776–782
- Wilson RS, Barnes LL, Aggarwal NT, et al. Cognitive activity and the cognitive morbidity of Alzheimer disease. Neurology 2010;75(11): 990–996
- Verdelho A, Madureira S, Moleiro C, et al; LADIS Study. Depressive symptoms predict cognitive decline and dementia in older people independently of cerebral white matter changes: the LADIS study. J Neurol Neurosurg Psychiatry 2013;84(11): 1250–1254
- Green RC, Cupples LA, Kurz A, et al. Depression as a risk factor for Alzheimer disease: the MIRAGE Study. Arch Neurol 2003;60(5): 753–759
- Shimada H, Park H, Makizako H, Doi T, Lee S, Suzuki T. Depressive symptoms and cognitive performance in older adults. J Psychiatr Res 2014;57:149–156