

Auditory Pathway Encoding and Neural Plasticity in Children with Learning Problems

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Key Words

Auditory pathway · Perception · Learning · Evoked responses · Neural plasticity · Learning disorders

Abstract

An inability to process auditory information, especially speech, characterizes many children with learning and attention problems. Our working hypothesis is that these speech-sound perception problems arise, at least in some cases, from faulty representation of the speech signal in central auditory centers. Preconscious neurophysiologic representation of sound structure by central auditory pathway neurons can be reflected by subcortical and cortical aggregate neural responses. These neurophysiologic responses can be modified by perceptual learning. Our research has shown that some children with learning problems demonstrate abnormal perception and neural representation of certain speech sounds. Differences between normal and learning-impaired groups can be attributable to aspects of neural synchrony that are reflected in aggregate neural responses. Deficiencies in neural synchrony in these children are apparent in subcortical (as well as cortical) representations of speech-sound structure, and these timing deficits are related to performance on speech-sound perception and learning measures. Moreover, impaired perception and

neurophysiologic encoding of speech sounds can be improved with cue enhancement and can be modified by perceptual learning associated with auditory training.

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Introduction

Understanding speech during everyday listening places many demands on the auditory system. Among these demands are the accurate representation of rapidly changing spectral information comprising the speech signal, and the separation of speech from background noise. Our research is aimed at understanding the basic biologic processes underlying speech-sound perception in quiet and noise. Specifically, we are interested in how neural activity gives rise to these processes in normal school-age children and in children with auditory learning problems. In addition, we are investigating the neurobiological processes involved in the perceptual learning of speech sounds in order to impact the design of training regimens that may assist those individuals who have difficulty perceiving speech sounds. Our group has used behavioral and neurophysiologic measures to investigate biologic processes involved in speech-sound perception and to delineate the nature and origin of auditory deficits affecting communication [Bradlow et al., 1999; Carrell et al., 1999; Cunningham et al., 2000b, 2001; Koch et al., 1999; King et al., 1999; Kraus et al., 1994a, b, 1996, 1998, 1999, 2000; McGee et al., 1996].

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Evoked potentials can reflect level-specific neural synchrony to speech-sound elements. The auditory brainstem response (ABR) depends on a high degree of synchronized firing between neurons. If there is excessive neural jitter, which might occur in an impaired auditory system, the separation of individual neural responses by even a fraction of a millisecond could cause responses to cancel each other out. The frequency following response (FFR) also depends on a high degree of neural synchrony. It reflects brainstem-generated, phase-locked responses to the speech stimulus' fundamental frequency and, to a lesser extent, its harmonics. Cortical responses reflect stimulus-locked synchronous firing across neural ensembles. P1/N1/N2 are elicited by stimuli presented in a simple repetitive sequence, whereas the mismatch negativity (MMN) is elicited by an acoustic change in a repetitive sequence. P1/N1/N2 and MMN each arise from different anatomic sources [Kraus et al., 1994a; Näätänen and Picton, 1987; Sams et al., 1991; Scherg et al., 1989] and represent different aspects of auditory function. That is, P1/N1/N2 and MMN (largely) reflect primary and nonprimary auditory pathway activity, respectively, and differ in their time course of maturation, patterns of hemispheric symmetry and responses to sound in background noise [Bellis et al., 2000; Cunningham et al., 2000b; Kraus et al., 1999; Ponton et al., 2000; Sharma et al., 1997; Martin et al., 1997]. Fine-grained speech-sound discrimination is associated with MMN [Kraus et al., 1993; Sams et al., 1985], whereas P1/N1/N2 are associated with other, more global aspects of auditory function such as the perception of syllables, words and sentences, and auditory short-term memory [Conley et al., 1999; Cunningham et al., 2000b]. Taken together, these aggregate neural responses can be used to acquire knowledge about speech-sound perception.

Speech Perception and Learning Problems

Almost 10% of children exhibit learning and reading disabilities [Torgeson, 1991]. Recent research has suggested that a subset of these children have difficulty with perception of certain fundamental acoustic differences within speech sounds [Brandt and Rosen, 1980; De Weirtdt, 1988; Elliot et al., 1989; Kraus et al., 1996; Leonard et al., 1992; Mody et al., 1997; Stark and Heinz, 1996a; Sussman, 1993; Tallal and Piercy, 1974; Tallal and Stark, 1981; Werker and Tess, 1987]. Those perceptual deficits are associated with poor phonologic processing and poor reading skills [Fletcher et al., 1994; Godfrey et al., 1981; McBride-Chang, 1996; Reed, 1989]. Our hypothesis has

been that, for some children, the difficulty in perceiving fundamental acoustic parameters stems from abnormalities in the neural representation of speech that occurs after peripheral sensory encoding but prior to conscious perception. Consistent with this view, we have demonstrated a fundamental, biologic basis for perceptual deficits in some children with learning problems at cortical and brainstem levels of the auditory pathway [Bradlow et al., 1999; Cunningham et al., 2000b, 2001; Kraus et al., 1996]. We have also shown that preattentive neural responses to sound can be improved with acoustically enhanced signals [Bradlow et al., 2000, 2001; Cunningham et al., 2001] and modified by short-term perceptual training [Hayes et al., 2001; Kraus et al., 1995; Tremblay et al., 1997, 1998].

Many individuals with learning problems (LP) demonstrate particular communication difficulties in noise [Bellis, 1996; Breedin et al., 1989; Chermak and Musiek, 1997; Jerger et al., 1987; Katz, 1992; Welsh et al., 1996], when stimuli are rapidly presented [Cestnick and Jerger, 2000; Farmer and Klein, 1995; Hari and Kiesila, 1996; Livingstone et al., 1991; Nagarajan et al., 1999; Tallal and Piercy, 1974], or when fine-grained discrimination is required [Bradlow et al., 1999; Elliot et al., 1989; Kraus et al., 1996; Wright et al., 1997]. Despite general acknowledgment that these factors excessively tax perception in this population, recent studies are just beginning to reveal neurobiological differences between normal and LP children when signals are presented in these challenging listening situations [Bradlow et al., 1999; Cestnick and Jerger, 2000; Cunningham et al., 2001; Kraus et al., 1996; Nagarajan et al., 1999; Wible et al., 2001].

Project on Listening, Learning and the Brain

In an ongoing project, we are investigating the correspondence between electrophysiologic responses and behavioral abilities in a large population of both normal and impaired children. Specifically examined is the relationship among psychophysical speech discrimination (listening), standardized measures of learning ability and academic achievement (learning), and neurophysiology (the brain).

Psychophysical Perception and Neurophysiologic Representation of Speech-Sound Differences

We have hypothesized that there is a biological basis for perceptual deficits in some of the LP children, and that disruption occurs in the representation of sound at a

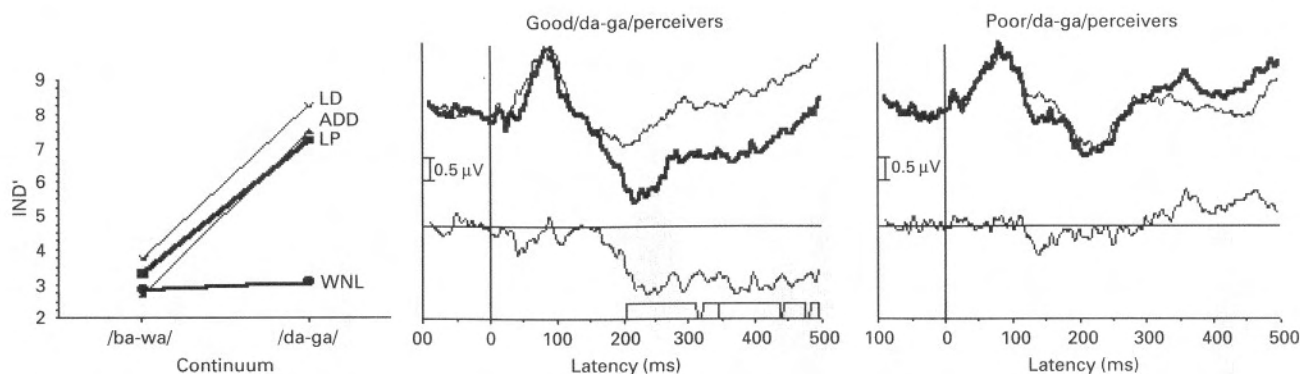


Fig. 1. Left: In LP children, perception was selectively worse for /da-ga/ than for /ba-wa/. Middle/right: MMN was evident for 'good' and absent for 'poor' /da-ga/ perceivers. Boxes below indicate the latency range over which a significant MMN occurs [from Kraus et al., *Science*, 1996].

preconscious, preattentive level. Our results support these hypotheses and provide evidence of neural asynchrony in sound representation at both brainstem and cortical levels [Bradlow et al., 1999; Cunningham et al., 2000b, 2001; King et al., 2001; Kraus et al., 1996].

On a behavioral task assessing fine-grained perception along /da-ga/ and /ba-wa/ continua, LP subjects performed significantly worse than normal subjects [Kraus et al., 1996]. Perception was selectively worse for /da-ga/ (change in third-formant onset frequency) than for /ba-wa/ (change in formant transition duration). This pattern indicates that LP children are better able to discriminate synthetic speech stimuli that differ in the temporal domain than stimuli that differ spectrally at stimulus onset. The discrepancy between /da-ga/ and /ba-wa/ in LP subjects is important in that the better performance on /ba-wa/ indicates that attention-motivation factors did not preclude good performance on the task. Overall, perceptual deficits cut across diagnostic categories, occurring in LP children with diagnoses of learning disability, attention deficit hyperactivity disorder, combined learning/attention disorder, and dyslexia. This supports the notion that there is a common perceptual deficit in a subset of children with various clinical diagnoses.

An association between perception and neurophysiologic mechanisms has been established as illustrated in figure 1. Good perception of /da-ga/ is associated with robust cortical responses to stimulus change (MMN), whereas poor discrimination is associated with diminished responses [Bradlow et al., 1999; Kraus et al., 1996]. Children in both groups had MMNs in response to the /ba-wa/ stimulus contrast, consistent with their good dis-

crimination of those stimuli. The psychophysical and electrophysiologic data together provide evidence for a preattentive, biological basis for learning problems in some children.

Brainstem and Cortical Asynchrony to Speech Sounds

Our data on normal (NL) and LP children indicate that brainstem responses to speech syllables differ for NL compared to LP children [King et al., 2001]. In addition, we have investigated brainstem and cortical encoding of speech sounds presented in background noise [Cunningham et al., 2001]. Subjects were children with LP and age-matched normal controls. LP subjects performed significantly worse than normal children on measures of auditory processing, reading, spelling, and fine-grained discrimination along a /da-ga/ continuum. Speech perception in noise (discrimination along an /ada-aga/ continuum) was significantly worse in the LP children.

Results are summarized in figure 2. ABR, FFR and P1/N1/N2 were elicited by /da/, presented in quiet and in background noise [Cunningham et al., 2001]. In noise, LP children exhibited significantly prolonged wave V latencies. A fast Fourier transform of the FFR revealed reduced energy in certain frequency bands (250–750 Hz) in the LP children compared to normals. Correlations between the stimulus and response waveforms in noise were significantly lower for the LP group. Overall, the data indicate that synchrony of auditory brainstem neurons differs between NL and LP children. Consequently certain learning deficits may originate from a disorder in auditory neural timing already seen at the brainstem level. In addition, cortical responses revealed that LP chil-

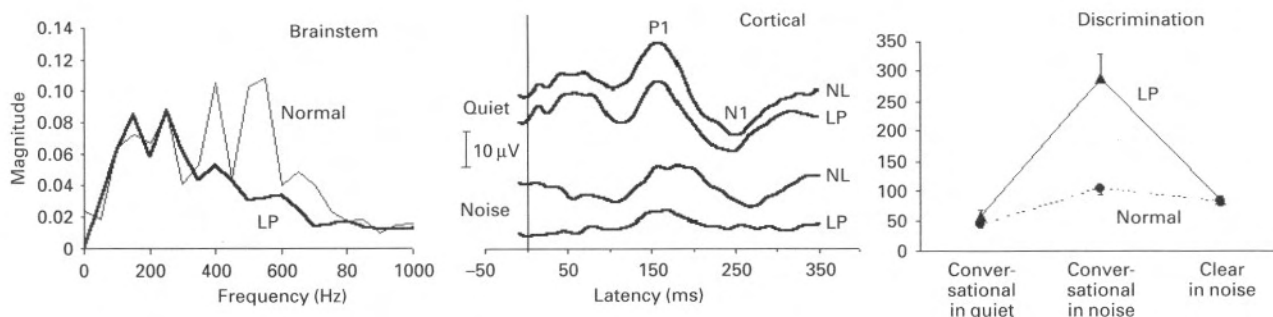


Fig. 2. Left: Magnitude of the spectral content of FFR shown by FFT in NL and LP children. Groups differed significantly in noise (between 450 and 750 Hz), not in quiet. Middle: Cortical potentials did not differ between NL and LP children in quiet, but were significantly smaller in LP subjects in noise. Right: Mean JNDs for NL and LP children along /ada-aga/ continua. LP children had poorer speech discrimination in noise and regained normal performance with cue-enhanced 'clear' stimuli [from Cunningham et al., Clin Neurophysiol, 2001].

dren showed significantly reduced P2-N2 amplitude to stimuli in noise. This demonstration of deficient neural representation of speech-in-noise at brainstem and cortical levels in LP children indicates that deficiencies in neural representation exist at multiple levels of the auditory pathway. In another study [Bradlow et al., 2000], LP children performed significantly worse than NL children in the perception of sentences in noise. Finally, preliminary data describe the effects, both singular and combined, of repeated stimulus repetition and background noise on cortical potentials [Wible et al., 2001]. The neural representation of repeated speech stimuli has been found to be diminished in LP children and was related to behavioral measures of auditory processing.

Cue Enhancement

The perceptual benefits of 'clear' or acoustically enhanced speech have been established [Jerger, 1999; Payton et al., 1994; Picheny et al., 1985; Smith and Levitt, 1999; Uchanski et al., 1996], and some features have been incorporated into commercially available auditory training programs designed for LP children. The effects of 'clear' speech in noise were investigated in LP and NL children [Cunningham et al., 2001]. The acoustic cues that are enhanced during 'clear' speech include an increased amplitude of consonant burst and a lengthened stop-gap duration [Picheny et al., 1986].

ABR, FFR and P1/N1/N2 potentials were obtained to enhanced /da/ stimuli (with amplification of the release burst intensity) presented in background noise [Cunningham et al., 2001]. Cue enhancement elicited normal

cortical responses in those LP children with deficient responses to unenhanced signals. This implies that the neural representation of acoustic events can be improved by specific cue enhancement.

Behaviorally, fine-grained discrimination in noise (along an /ada-aga/ continuum) was significantly worse in LP than normal children [Cunningham et al., 2001]. Yet, when 'clear' speech characteristics are added to the stimuli, perception improved. When each of the 'clear' speech enhancements was studied in isolation, increased release burst intensity was found to be a more important acoustic cue manipulation than lengthened stop gap duration. In addition, perception of sentences in noise in LP children improved dramatically when the same sentences are produced with 'clear' speech [Bradlow et al., 2000, 2001], expanding our findings to more real-world listening situations.

Plasticity and Speech-Sound Perceptual Learning

Speech perception abilities in humans are modified both by long-term experience with one's native language [Aslin et al., 1981; Cheour et al., 1998; Dehaene-Lambertz and Baillet, 1998; Jusczyk et al., 1984; Kuhl et al., 1992; Mehler et al., 1978; Näätänen et al., 1997; Werker et al., 1981], and by short-term directed auditory training in a laboratory or clinical environment [Bradlow and Pisoni, 1999; Pisoni et al., 1982], and speech perception can be modified by auditory training in language-im-

paired children [Ball and Blanchman, 1991; Bradley and Bryant, 1983; Merzenich et al., 1996; Schankweiler et al., 1995; Tallal et al., 1996]. Little is known about neural plasticity associated with perceptual learning in humans, although animal experiments have demonstrated that sensory cortex becomes restructured with training [Jenkins et al., 1990; Kraus et al., 1982; Merzenich and Jenkins, 1993; Recanzone et al., 1992, 1993]. Our research has demonstrated physiologic changes associated with auditory perceptual learning in humans. These changes are preattentive and can precede behavioral learning [Kraus et al., 1995; Tremblay et al., 1997, 1998, 2001].

Speech-Sound Training in Children with LP

Interest in perceptual training programs for people with LP has persisted throughout the years in the fields of education, psychology, speech and hearing, and learning disabilities [Orton, 1937]. More recently, there has been much interest in interactive computer-based auditory training programs [Diehl, 1999; Morrison, 1998; Tallal et al., 1998]. However, the efficacy of these programs is not uniform across children, and it is unclear which kind of training, for which profile of deficits, results in perceptual improvements. Studying children who undergo this training provides important insights into the neurophysiologic and perceptual changes associated with perceptual learning. It is important to determine which children might benefit from training, and how training may alter the neural representation of sound at various levels of the auditory pathway.

We are investigating the effects of commercial computer-based auditory training programs on behavioral and neurophysiologic measures in children with LP [Hayes et al., 2001; Zhang et al., 2000]. Subjects are tested before and after participation in these programs on learning, perceptual, and neurophysiologic measures. Preliminary results indicate improvement on measures of perception and learning, and changes in cortical potentials to stimuli presented in quiet and in noise. These changes did not occur in an untrained control group. Results have direct bearing on the interpretation of brain/behavior changes associated with auditory training programs and require considerable additional systematic study.

Animal Studies

The distinctive roles of the auditory midbrain, thalamus and cortex have been directly explored in an animal model using identical stimuli used in human studies. First, elemental acoustic parameters of synthetic speech stimuli were reflected in auditory pathway responses

[McGee et al., 1996]. Second, the encoding of stimulus change (mismatch negativity) has been shown to occur predominantly in the nonprimary thalamocortical pathway. Third, different acoustic contrasts appear to be encoded at distinct places along the auditory pathway [Kraus et al., 1994a, b; Sharma et al., 1994]. Processing certain rapid spectrotemporal difference appears to require the auditory cortex. Fourth, when stimuli were presented in background noise, neural representation of the consonant portion was affected to a greater extent than the vowel portion of the stimulus [Cunningham et al., 1999, 2000a, 2001]. The percent decrease in the onset response that can be attributed to the effects of noise was greatest at cortical compared to subcortical locations. At the midbrain, noise reduced the magnitude of low frequency spectral components (FFT analysis), whereas higher frequency components remained unchanged, mirroring FFT findings in NL children using the same stimuli [Cunningham et al., 2001]. Fifth, the effects of 'clear' speech parameter manipulations were assessed in neurophysiologic responses in the midbrain, thalamus, and epidural cortex [Cunningham et al., 2000a]. Onset response amplitudes increased as the stop gap duration or burst intensity alone were varied. Combined manipulations resulted in maximal effects and were not a simple linear sum of the response of each manipulation alone. In background noise, 'clear' speech stimuli elicited an onset response (absent in unenhanced signals). The increase in the onset response that can be attributed to the effect of cue enhancement was greater at cortical than subcortical levels. Cue enhancements did not affect the representation of steady-state portions of the response. Overall, animal data on the neural representation of speech in noise complement the human studies and provide information about physiologic mechanisms underlying the perception of speech in NL and LP children.

Summary

LP children demonstrate abnormal perception and neural representation of fine-grained stimulus differences. Differences between normal and LP groups can be attributable to aspects of neural synchrony that are reflected in aggregate neural responses. Deficiencies in neural synchrony in these children are already apparent in subcortical representation of speech-sound structure, and these timing deficits are related to speech-sound perception and learning measures. Thus, neural representation of certain speech features – independent of cognitive and attentional influences – underlies certain learning and reading disorders. Moreover, impaired perception and neurophysiologic encoding of speech sounds in LP children can be improved with cue enhancement and perceptual learning.

References

- Aslin RN, Pisoni DB, Hennessy BL, Perey AJ: Discrimination of voice onset time by human infants: New findings and implications for the effects of early experience. *Child Dev* 1981;52:1135-1145.
- Ball EW, Blanchman BA: Does phoneme awareness training in kindergarten make a difference in early recognition and developmental spelling. *Reading Res Q* 1991;26:49-66.
- Bellis TJ: Assessment and Management of Central Auditory Processing Disorders in the Educational Setting: From Science to Practice. San Diego, Singular Publishing, 1996.
- Bellis TJ, Nicol T, Kraus N: Aging affects hemispheric asymmetry in the neural representation of speech sounds. *J Neurosci* 2000;20:791-797.
- Bradley L, Bryant PE: Categorizing sounds and learning to read - a causal connection. *Nature* 1983;301:419-421.
- Bradlow AR, Kraus N, Hayes E: Speaking clearly for learning-disabled children: Sentence perception in noise (abstract 108). *J Acoust Soc Am* 2000.
- Bradlow AR, Kraus N, Nicol TG, McGee TJ, Cunningham J, Zecker SG, Carrell TD: Effects of lengthened formant transition duration on discrimination and neural representation of synthetic CV syllables by normal and learning-disabled children. *J Acoust Soc Am* 1999;106:2086-2096.
- Bradlow AR, Pisoni DB: Recognition of spoken words by native and non-native listeners: Talker-, listener-, and item-related factors. *J Acoust Soc Am* 1999;106:2074-2085.
- Bradlow AR, Kraus N, Hayes E: Speaking clearly for learning-impaired children: Sentence perception in noise. Submitted.
- Brandt J, Rosen JJ: Auditory phonemic perception in dyslexia: Categorical identification and discrimination of stop consonants. *Brain Lang* 1980;9:324-337.
- Breedin SD, Martin RC, Jerger S: Distinguishing auditory and speech-specific perceptual deficits. *Ear Hear* 1989;10:311-317.
- Carrell TD, Bradlow AR, Nicol TG, Koch DB, Kraus N: Interactive software for evaluating auditory discrimination (letter). *Ear Hear* 1999;20:175-176.
- Cestnick L, Jerger J: Auditory temporal processing and lexical/nonlexical reading in developmental dyslexics. *J Am Acad Audiol* 2000;11:501-513.
- Cheour M, Ceponiene R, Lehtokoski A, Luuk A, Allik J, Alho K, Naatanen R: Development of language-specific phoneme representations in the infant brain. *Nat Neurosci* 1998;1:351-353.
- Chermak GD, Musiek FE: Central Auditory Processing Disorders: New Perspectives. San Diego, Singular Publishing, 1997.
- Conley EM, Michalewski HJ, Starr A: The N100 auditory cortical evoked potential indexes scanning of auditory short-term memory. *Clin Neurophysiol* 1999;110:2086-2093.
- Cunningham J, King C, Nicol T, Bradlow AR, McGee T, Kraus N: Neurophysiologic representation of 'clear' speech in the midbrain, thalamus, and cortex. *Assoc Res Otolarygol* 1999.
- Cunningham J, Nicol T, Kraus N: Neurophysiologic representation of clear speech in noise. *Assoc Res Otolarygol* 2000a.
- Cunningham J, Nicol T, Zecker SG, Kraus N: Neurobiologic responses to speech in noise in children with learning problems: Deficits and strategies for improvement (article and accompanying editorial). *Clin Neurophysiol* 2001;112:758-767.
- Cunningham J, Nicol T, Zecker S, Kraus N: Speech-evoked neurophysiologic responses in children with learning problems: Development and behavioral correlates of perception. *Ear Hear* 2000b;21:554-568.
- Dehaene-Lambertz G, Baillet S: A phonological representation in the infant brain. *Neuroreport* 1998;9:1885-1888.
- De Weirdt W: Speech perception and frequency discrimination in good and poor readers. *Appl Psycholinguist* 1988;9:163-183.
- Diehl S: Listen and learn? A software review of earbics, language, speech and hearing services in schools. *Asha* 1999;30:108-116.
- Elliott LL, Hammer MA, Scholl ME: Fine-grained auditory discrimination in normal children and children with language-learning problems. *J Speech Hear Res* 1989;32:112-119.
- Farmer ME, Klein R: The evidence for a temporal processing deficit linked to dyslexia: A review. *Psychonorm Bull Rev* 1995;2:460-493.
- Fletcher JM, Shaywitz SE, Shankweiler DP, Katz L, Liberman IY, Stuebing KK, Francis DJ, Fowler AE, Shaywitz BA: Cognitive profiles of reading disability: Comparisons of discrepancy and low achievement definitions. *J Educ Psychol* 1994;86:6-23.
- Geisler CD, Gamble T: Responses of 'high-spontaneous' auditory-nerve fibers to consonant-vowel syllables in noise. *J Acoust Soc Am* 1989;85:1639-1652.
- Godfrey JJ, Syrdal-Lasky AK, Millay KK, Knox CM: Performance of dyslexic children on speech perception tests. *J Exp Child Psychol* 1981;32:401-424.
- Hari R, Kiesila P: Deficit of temporal auditory processing in dyslexic adults. *Neurosci Lett* 1996;205:138-140.
- Hayes E, Nicol T, Zecker S, Kraus N: Auditory processing and neural plasticity with auditory perceptual training in children with learning problems. *Assoc Res Otolarygol* 2001.
- Jenkins WM, Merzenich MM, Ochs MT, Allard T, Guic-Robles E: Functional reorganization of primary somatosensory cortex in adult owl monkeys after behaviorally controlled tactile stimulation. *J Neurophysiol* 1990;63:82-104.
- Jerger J: Consonant enhancement: Promising findings. *J Am Acad Audiol* 1999;10:410.
- Jerger S, Martin RC, Jerger J: Specific auditory perceptual dysfunction in a learning disabled child. *Ear Hear* 1987;8:78-86.
- Jusczyk PW, Shea SL, Aslin RN: Linguistic experience and infant speech perception: A re-examination of Eilers, Gavin & Oller (1982). *J Child Lang* 1984;11:453-466.
- Katz J: Classification of auditory disorders; in Katz J, Stecker N, Henderson D (eds): Central Auditory Processing: A Transdisciplinary View. St Louis, Mosby, 1992.
- King C, Cunningham J, Nicol T, Warrier C, Kraus N: Speech-evoked cochlear (CM) and brainstem (ABR) responses in normal and learning impaired children. *Assoc Res Otolarygol* 2001.
- King C, Nicol T, McGee T, Kraus N: Thalamic asymmetry is related to acoustic signal complexity. *Neurosci Lett* 1999;267:89-92.
- Koch DB, McGee TJ, Bradlow AR, Kraus N: Acoustic-phonetic approach toward understanding neural processes and speech perception. *J Am Acad Audiol* 1999;10:304-318.
- Kraus N, Bradlow AR, Cheatham MA, Cunningham J, King C, Koch DB, Nicol T, McGee T, Stein S, Wright B: Consequences of neural asynchrony: A case of auditory neuropathy. *J Assoc Res Otolaryngol* 2000;1:33-45.
- Kraus N, Koch DB, McGee TJ, Nicol TG, Cunningham J: Speech-sound discrimination in school-age children: Psychophysical and neurophysiologic measures. *J Speech Lang Hear Res* 1999;42:1042-1060.
- Kraus N, McGee T, Carrell T, King C, Littman T, Nicol T: Discrimination of speech-like contrasts in the auditory thalamus and cortex. *J Acoust Soc Am* 1994a;96:2758-2768.
- Kraus N, McGee T, Carrell T, King C, Tremblay K: Central auditory system plasticity associated with speech discrimination training. *J Cogn Neurosci* 1995;7:27-34.
- Kraus N, McGee TJ, Carrell TD, Zecker SG, Nicol TG, Koch DB: Auditory neurophysiologic responses and discrimination deficits in children with learning problems. *Science* 1996;273:971-973.
- Kraus N, McGee TJ, Koch DB: Speech sound representation, perception, and plasticity: A neurophysiologic perspective. *Audiol Neurotol* 1998;3:168-182.
- Kraus N, McGee T, Littman T, Nicol T, King C: Nonprimary auditory thalamic representation of acoustic change. *J Neurophysiol* 1994b;72:1270-1277.
- Kraus N, McGee T, Micco A, Sharma A, Carrell T, Nicol T: Mismatch negativity in school-age children to speech stimuli that are just perceptibly different. *Electroencephalogr Clin Neurophysiol* 1993;88:123-130.
- Kraus N, Disterhoft JF: Response plasticity of single neurons in rabbit auditory association cortex during tone-signalled learning. *Brain Res* 1982;246:205-215.
- Kuhl PK, Williams KA, Lacerda F, Stevens KN, Lindblom B: Linguistic experience alters phonetic perception in infants by 6 months of age. *Science* 1992;255:606-608.
- Leonard LB, McGregor KK, Allen GD: Grammatical morphology and speech perception in children with specific language impairment. *J Speech Hear Res* 1992;35:1076-1085.

- Livingstone MS, Rosen GD, Drislane FW, Galaburda AM: Physiological and anatomical evidence for a magnocellular defect in developmental dyslexia. *Proc Natl Acad Sci USA* 1991; 88:7943-7947.
- McBride-Chang C: Models of speech perception and phonological processing in reading. *Child Dev* 1996;67:1836-1856.
- McGee T, Kraus N, King C, Nicol T, Carrell TD: Acoustic elements of speechlike stimuli are reflected in surface recorded responses over the guinea pig temporal lobe. *J Acoust Soc Am* 1996;99:3606-3614.
- Martin BA, Sigal A, Kurtzberg D, Stapells DR: The effects of decreased audibility produced by high-pass noise masking on cortical event-related potentials to speech sounds /ba/ and /da/. *J Acoust Soc Am* 1997;101:1585-1599.
- Mehler J, Bertoncini J, Barriere M: Infant recognition of mother's voice. *Perception* 1978;7:491-497.
- Merzenich MM, Jenkins WM: Reorganization of cortical representations of the hand following alterations of skin inputs induced by nerve injury, skin island transfers, and experience. *J Hand Ther* 1993;6:89-104.
- Merzenich MM, Jenkins WM, Johnston P, Schreiner C, Miller SL, Tallal P: Temporal processing deficits of language-learning impaired children ameliorated by training. *Science* 1996; 271:77-81.
- Mody M, Studdert-Kennedy M, Brady S: Speech perception deficits in poor readers: Auditory processing or phonological coding? *J Exp Child Psychol* 1997;64:199-231.
- Morrison S: Computer Applications: Earobics, Child Language Teaching and Therapy. London, Arnold, 1998.
- Näätänen R, Lehtokoski A, Lennes M, Cheour M, Huotilainen M, Iivonen A, Vainio M, Alku P, Ilmoniemi RJ, Luuk A, Allik J, Sinkkonen J, Alho K: Language-specific phoneme representations revealed by electric and magnetic brain responses. *Nature* 1997;385:432-434.
- Näätänen R, Picton T: The N1 wave of the human electric and magnetic response to sound: A review and an analysis of the component structure. *Psychophysiol* 1987;24:375-425.
- Näätänen R, Schroger E, Karakas S, Tervaniemi M, Paavilainen P: Development of a memory trace for a complex sound in the human brain. *Neuroreport* 1993;4:503-506.
- Nagarajan S, Mahneke H, Salz T, Tallal P, Roberts T, Merzenich MM: Cortical auditory signal processing in poor readers. *Proc Natl Acad Sci USA* 1999;96:6483-6488.
- Orton ST: Reading, Writing and Speech Problems in Children. New York, Norton, 1937.
- Payton KL, Uchanski RM, Braid LD: Intelligibility of conversational and clear speech in noise and reverberation for listeners with normal and impaired hearing. *J Acoust Soc Am* 1994;95: 1581-1592.
- Picheny MA, Durlach NI, Braid LD: Speaking clearly for the hard of hearing. I: Intelligibility differences between clear and conversational speech. *J Speech Hear Res* 1985;28:96-103.
- Picheny MA, Durlach NI, Braid LD: Speaking clearly for the hard of hearing. II: Acoustic characteristics of clear and conversational speech. *J Speech Hear Res* 1986;29:434-446.
- Pisoni DB, Aslin RN, Perey AJ, Hennessy BL: Some effects of laboratory training on identification and discrimination of voicing contrasts in stop consonants. *J Exp Psychol Hum Percept Perform* 1982;8:297-314.
- Ponton CW, Eggermont JJ, Kwong B, Don M: Maturation of human central auditory system activity: Evidence from multi-channel evoked potentials. *Clin Neurophysiol* 2000;111:220-236.
- Recanzone GH, Schreiner CE, Merzenich MM: Plasticity in the frequency representation of primary auditory cortex following discrimination training in adult owl monkeys. *J Neurosci* 1993;13:87-103.
- Recanzone GH, Merzenich MM, Jenkins WM: Frequency discrimination training engaging a restricted skin surface results in an emergence of a cutaneous response zone in cortical area 3a. *J Neurophysiol* 1992;67:1057-1070.
- Reed MA: Speech perception and the discrimination of brief auditory cues in reading disabled children. *J Exp Child Psychol* 1989;48:270-292.
- Sams M, Kaukoranta E, Hämäläinen M, Näätänen R: Cortical activity elicited by changes in auditory stimuli: Different sources for the magnetic N100m and mismatch responses. *Psychophysiology* 1991;28:21-29.
- Sams M, Paavilainen P, Alho K, Näätänen R: Auditory frequency discrimination and event-related potentials. *Electroencephalogr Clin Neurophysiol* 1985;62:437-448.
- Scherg M, Vajsar J, Picton T: A source analysis of the human auditory evoked potentials. *J Cogn Neurosci* 1989;3:336-355.
- Shankweiler DP, Crain S, Katz L, Fowler AE, Liberman AM, Brady S, Thorton R, Lundquist E, Dreyer L, Fletcher JM, Stuebing KK, Shaywitz SE, Shaywitz BA: Cognitive profiles of reading-disabled children: Comparison of language skills in phonology, morphology, and syntax. *Psychol Sci* 1995;6:149-156.
- Sharma A, Kraus N, McGee TJ, Nicol TG: Developmental changes in P1 and N1 central auditory responses elicited by consonant-vowel syllables. *Electroencephalogr Clin Neurophysiol* 1997;104:540-545.
- Sharma A, Kraus N, Nicol T, Carrell T, McGee T: Physiological bases of pitch and place of articulation perception: A case study. *J Acoust Soc Am* 1994;95:3011.
- Smith LZ, Levitt H: Consonant enhancement effects on speech recognition of hearing-impaired children. *J Am Acad Audiol* 1999;10:411-421.
- Stark RE, Heinz JM: Perception of stop consonants in children with expressive and receptive-expressive language impairments. *J Speech Hear Res* 1996a;39:676-686.
- Sussman JE: Perception of formant transition cues to place of articulation in children with language impairments. *J Speech Hear Res* 1993; 36:1286-1299.
- Tallal P, Miller SL, Bedi G, Byma G, Wang X, Nagarajan SS, Schreiner C, Jenkins WM, Merzenich MM: Language comprehension in language-learning impaired children improved with acoustically modified speech. *Science* 1996;271:81-84.
- Tallal P, Piercy M: Developmental aphasia: Rate of auditory processing and selective impairment of consonant perception. *Neuropsychologia* 1974;12:83-93.
- Tallal P, Stark RE: Speech acoustic-cue discrimination abilities of normally developing and language-impaired children. *J Acoust Soc Am* 1981;69:568-574.
- Tallal P, Merzenich MM, Miller S, Jenkins W: Language learning impairments: Integrating basic science, technology, and remediation. *Exp Brain Res* 1998;123:210-219.
- Torgeson JK: Learning disabilities: Historical and conceptual issues; in Wong BYL (ed): *Learning about Learning Disabilities*. San Diego, Academic Press, 1991.
- Tremblay K, Kraus N, Carrell TD, McGee T: Central auditory system plasticity: Generalization to novel stimuli following listening training. *J Acoust Soc Am* 1997;102:3762-3773.
- Tremblay K, Kraus N, McGee T: The time course of auditory perceptual learning: Neurophysiological changes during speech-sound training. *Neuroreport* 1998;9:3557-3560.
- Tremblay K, Kraus N, McGee TJ, Ponton CW, Otis B: Central auditory plasticity: Changes in the N1-P2 complex following speech-sound training. *Ear Hear* 2001;22:79-90.
- Uchanski RM, Choi SS, Braid LD, Reed CM, Durlach NI: Speaking clearly for the hard of hearing. IV. Further studies of the role of speaking rate. *J Speech Hear Res* 1996;39:494-509.
- Welsh LW, Welsh JJ, Healy MP: Learning disabilities and central auditory dysfunction. *Ann Otol Rhinol Laryngol* 1996;105:117-122.
- Werker JF, Frost PE, McGurk H: La langue et les lèvres: Cross-language influences on bimodal speech perception. *Can J Psychol* 1992;46: 551-568.
- Werker JF, Gilbert JH, Humphrey K, Tees RC: Developmental aspects of cross-language speech perception. *Child Dev* 1981;52:349-355.
- Werker JF, Tees RC: Speech perception in severely disabled and average reading children. *Can J Psychol* 1987;41:48-61.
- Wible B, Nicol T, Kraus N: Effects of stimulus repetition and background noise on neural representation of speech sounds in normal and learning impaired children. *Assoc Res Otolaryngol* 2001;Wright BA, Lombardino LJ, King WM, Puranik CS, Leonard CM, Merzenich MM: Deficits in auditory temporal and spectral resolution in language-impaired children. *Nature* 1997;387:176-178.
- Zhang J, Hayes E, Cunningham J, Wible B, Bradlow AR, Zecker SG, Kraus N: Brain/behavior changes and auditory training programs. *Assoc Res Otolaryngol* 2000.