Time Course of the Suppression Effect on Transient Evoked Otoacoustic Emissions by Prolonged Contralateral Acoustic Stimulation

Hyun Woo Kang, Hyun Joon Shim, Seong Jun Song, Seong Hee Lee and Sang Won Yoon
Department of Otolaryngology, Eulji Medical Center, Eulji University School of Medicine, Seoul, Korea

Background and Objectives: Although the suppressive effect of the medial efferent acoustic reflex is well known, the time course of this effect over prolonged periods has yet to be fully evaluated. We assessed time-dependent change in the suppression of transient evoked otoacoustic emissions (TEOAEs) by the medial efferent acoustic reflex over a relatively long period.

Subjects and Methods: We measured TEOAEs in the right ear before contralateral acoustic stimulation (CAS), and then measured serial TEOAEs in the right ear at four intervals during a total of 16 minutes of continuous CAS, followed by three more recordings after termination of CAS. Results: TEOAE amplitudes were reduced with CAS during a certain period (from the immediate period to 10 minutes depending on frequency) and subsequently recovered. TEOAE suppression values in the mean amplitudes for overall frequency were 0.76 dB at the initial recording, 0.35 dB at 5 minutes, 0.44 dB at 10 minutes, and 0.33 dB at 15 minutes during CAS. The initial suppression value was significantly larger than other suppression values of 5, 10, and 15 minutes (p<0.05). In recordings obtained after CAS, TEOAE amplitude exceeded pre-acoustic amplitudes at 1 kHz, 1.5 kHz, and 2 kHz. Conclusions: The present results show the existence of the medial efferent acoustic reflex and demonstrate the time course that TEOAE suppressions present initially after CAS, showing fatigue over time. Overshooting of TEOAE was observed in recordings at several frequencies after termination of CAS.

KEY WORDS: Otoacoustic emissions · Acoustic reflex · Efferent fiber · Acoustic stimulation.

Introduction

Otoacoustic emissions (OAEs) have become a useful, non-invasive, objective measurement of outer hair cell response since their discovery by Kemp in 1978. OAEs are generated by the outer hair cells of the cochlea and can be recorded in the external auditory canal. OAEs are created by an outer hair cell active amplification mechanism, and these sounds can be reduced by contralateral noise. The medial olivocochlear bundle efferent pathway is composed of ipsilateral and contralateral neurons. Acoustic stimulation of a single side stimulates type I afferent fibers of the auditory nerve. These fibers project centrally to the cochlear nucleus, where they synapse. Reflex interneurons innervate the ipsilateral and contralateral superior olivary complex. The contralateral outer hair cells are then innervated by the bilateral medial olivocochlear fiber bundle to suppress the mechanical response of the outer hair cells. The medial olivocochlear bundle acts to protect the cochlea from acoustic overstimulation. Another possible function of the medial olivocochlear system is in facilitating selective attention. The central nervous system uses the medial efferent pathway to cause the outer hair cells to change the micromechanical properties of the cochlea during selective attention tasks.

Previous attempts to study the action duration and characteristics of the efferent effect of the medial olivocochlear system in animals and humans using different recording methods have delivered inconsistent results. Some studies showed that suppression reached a constant steady state during contralateral acoustic stimulation (CAS), whereas another reported a gradual release of suppression during CAS. In addition, when contralateral acoustic stimulation was terminated, some research-
Subjects and Methods

We used an experimental setting similar to that described by van Zyl, et al., with a larger study cohort.

Participants

Twenty adults (5 men, 15 women; age range, 23–35 years; mean age, 26.9±5.3 years) with normal hearing abilities were evaluated in this study. All subjects met the following criteria based on a subjective clinical interview: 1) no history of past otologic disease, 2) no familial history of hearing loss, and 3) no history of ototoxic drug use. All subjects presented with normal appearance of the tympanic membrane and external auditory canal. All subjects underwent audiologic testing, which revealed that all had hearing thresholds less than 20 dB between 250 and 8000 Hz, and had normal type A tympanogram and normal stapedial reflex at 90 dB HL.

Procedure

TEOAEs were recorded and analyzed using the Scout Sport system in Navigator pro (Bio-logic Systems Corp., Chicago, IL, USA) in a sound-treated audiological test booth. Linear broadband 80 μS clicks were presented at 80 dB peak equivalent SPL at a repetition rate of 40/s and recorded in the right ear of each subject. TEOAEs were evaluated at frequency bands of 1000, 1500, 2000, 3000, and 4000 Hz, and 260 times averages were completed to reduce the noise effect. The average duration of a TEOAE recording was 1 minute. CAS was presented through earphones at an intensity of 45 dB SL to the left ear of each subject. CAS was white noise (duration, 300 msec; rise-fall time, 0.5 ms; sampling, 44.1 kHz). TEOAE recordings were measured under experimental conditions without CAS, and then at 5-minute intervals over a period of 16 minutes with CAS. After 16 minutes of CAS, TEOAEs were recorded at 1, 3, and 5 minutes without any acoustic stimulation. Over the course of the experiments, TEOAEs were measured eight times in each subject (Fig. 1).

The mean TEOAE amplitudes of all subjects were compared between pre-acoustic stimulation and four times during acoustic stimulation at 1, 1.5, 2, 3, and 4 kHz and the mean TEOAE amplitudes of all subjects at 1, 1.5, 2, 3, and 4 kHz were compared between pre-acoustic stimulation and after termination of acoustic stimulation. Suppression values (difference between pre-acoustic stimulation amplitude and amplitude during acoustic stimulation) and enhancement values (difference between pre-acoustic stimulation amplitude and amplitude after acoustic stimulation) were calculated for the mean amplitudes of overall frequency, and suppression values were compared as a function of CAS duration.

Statistical analysis

Paired t-test was used to compare TEOAE amplitudes between pre-acoustic stimulation and four times during acoustic stimulation, and between pre-acoustic stimulation and post-acoustic stimulation, using SPSS software. Repeated ANOVA was applied to compare suppression values for overall frequency as a function of CAS duration and to compare suppression values among the different frequencies.

Results

A significant reduction in the mean amplitude of the 1 kHz TEOAEs by acoustic stimulation was detected initially, at 5
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minutes, and at 10 minutes after acoustic stimulation; in contrast, recorded values after 15 minutes showed no significant reduction. Recorded values at 1, 3, and 5 minutes after termination of acoustic stimulation showed a significant increase compared with the amplitudes recorded for pre-acoustic stimulation at 1 kHz (Fig. 2).

At 1.5 kHz, a significant reduction was present initially and at 5 minutes after acoustic stimulation; the recorded values at 1 and 3 minutes after termination of acoustic stimulation showed a significant increase compared with the amplitudes recorded for pre-acoustic stimulation (Fig. 2). At 2 kHz, a significant reduction was detected only for initial recordings with acoustic stimulation; the recorded values at 1 and 3 minutes after termination of acoustic stimulation showed a significant increase (Fig. 2). At 3 kHz, a significant reduction in the mean amplitude was present only for initial recordings with acoustic stimulation; there was no other significant difference in amplitude at any point when comparing pre-acoustic stimulation and after termination of acoustic stimulation (Fig. 2). As was found at frequencies of 3 kHz, acoustic stimulation at 4 kHz showed a significant reduction in the mean amplitude only in the initial recordings, and a significant increase was not shown at any recording after termination of acoustic stimulation (Fig. 2).

Suppression values in the mean amplitudes for overall frequency were 0.76 dB (12.82% suppression from pre-acoustic stimulation) at the initial recording, 0.35 dB (5.90% suppression) at 5 minutes, 0.44 dB (7.42% suppression) at 10 minutes,
and 0.33 dB (5.56% suppression) at 15 minutes during CAS. The initial suppression value was significantly larger than suppression values at 5, 10, and 15 minutes \((p<0.05)\) (Fig. 3). After CAS, the enhancement values in the mean amplitudes for overall frequency were 0.47 (7.93% enhancement from pre-acoustic stimulation) at 1 minute, 0.56 (9.44% enhancement) at 3 minutes, and 0.20 (3.37% enhancement) at 5 minutes. There were no significant differences among the three values after CAS (Fig. 3).

As frequency increased, a tendency for suppression to decrease was observed during the whole duration with acoustic stimulation; however, among the frequencies, no statistical difference was found with regard to suppression (Fig. 4).

**Discussion**

In the present study, we investigated the inhibitory effect of the medial olivocochlear system as measured by TEOAE in the presence of prolonged CAS in adults with normal hearing. Though the maintenance periods of inhibitory effects with CAS differed according to frequency, the amplitude of TEOAEs decreased during a certain period (from the immediate period to 10 minutes, depending on the frequency) before recovering. The TEOAE suppression value for overall frequency showed the highest score at the initial recording and fell abruptly after that point. These results are somewhat different to those of van Zyl, et al.\(^9\) in which amplitude suppressions of TEOAE were maintained at all frequencies during 16 minutes of acoustic stimulation and the overall TEOAE suppression values demonstrated no change between recording intervals during CAS. We consider that the present result showed efferent fatigue or sensory adaptation in the medial olivocochlear system. A human study published in 1997 reported that suppression of TEOAE remained constant throughout the entire period of CAS (from 10 to 180 seconds) and that there was no manifestation of significant efferent fatigue.\(^9\) A study that used distortion product OAE reported that maximal suppression occurred within 2.6 seconds after acoustic stimulation and that the medial olivocochlear effect was maintained for a duration of acoustic stimulation of 20 minutes.\(^9\)

In contrast, a human study using spontaneous OAE reported efferent fatigue or sensory adaptation, in agreement with the findings of the present study, and reported a rapid reduction in amplitude at the onset of acoustic stimulation followed by a lessening of this reduction with 20 seconds of CAS.\(^9\) To evaluate the efferent effect of the medial olivocochlear system, compound action potential (CAP) and ensemble background activity (EBA) have been used in animal studies. One study monitored cochlear CAP during 12 seconds of contralateral broadband sound in anesthetized cats and found that suppression reached a constant steady state 2 seconds after the onset of contralateral sound, followed by a constant state during contralateral sound. When the sound was turned off, suppression disappeared in less than 0.62 seconds.\(^10\) In another study using electrical stimulation instead of acoustic stimulation, CAP amplitude was reduced to less than 50% of control values within 100 ms after the onset of electrical stimulation, after which CAP amplitude continued to decline slowly by a further 20% within 1 minute after the onset of stimulation, and the decreased CAP amplitude remained until electrical stimulation ceased.\(^11\)

Sridhar, et al.\(^11\) analyzed the duration properties of the medial olivocochlear system reflex and identified two different effects: the fast effect (rapid

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**Fig. 3.** Mean TEOAE suppression values and enhancement values, in dBA HL, for overall frequency as a function of CAS duration. The vertical bars represent standard deviations. Suppression value is defined as the difference between pre-acoustic stimulation amplitude and the amplitude during acoustic stimulation. Enhancement value is defined as the difference between pre-acoustic stimulation amplitude and the amplitude after acoustic stimulation. The initial suppression value was significantly larger than suppression values at 5, 10, and 15 minutes \((p<0.05)\). TEOAE: transient evoked otoacoustic emission, CAS: contralateral acoustic stimulation.

**Fig. 4.** Difference in TEOAE amplitude suppression among frequency bands during contralateral acoustic stimulation. No significant difference in TEOAE suppression was found among the frequencies. TEOAE: transient evoked otoacoustic emission, CAS: contralateral acoustic stimulation.
decrease in amplitude after the onset of stimulation) and the slow effect (further decrease in amplitude after the fast effect). These effects have been measured previously in an animal study using EBA and CAS; the authors reported a rapid 75% decrease in the total decreased EBA amplitude within 10 ms after the onset of CAS (fast effect), and an additional slower decrease of 25% (slow effect) after the fast decrease remained constant during presentation of more than 2 hours of acoustic stimulation. Previous studies, however, were unable to fully explain the mechanism behind these two effects, and no studies have been performed that confirm these effects in humans.

In the present study, we detected a pattern of a longer time period of suppression at lower frequencies: at 1 and 1.5 kHz, OAE amplitude reduction remained until recordings at 10 and 5 minutes, respectively, with acoustic stimulation; whereas at 2, 3, and 4 kHz, amplitude reduction was present only at the immediate recording with acoustic stimulation. These results imply that the protective mechanism of the medial olivocochlear system is more active at low frequencies than at high frequencies, and may be correlated with the greater damage caused by noise in the high-frequency area in the cochlea. The tendency for greater suppression during acoustic stimulation at lower frequencies is in agreement with this assumption, and was also observed in the study of van Zyl, et al., who investigated 16-minute CAS in 10 adults with normal hearing. Following the period of suppression, an immediate recording with acoustic stimulation was performed, whereas at 2, 3, and 4 kHz, amplitude reduction was present only at the immediate recording with acoustic stimulation. In the study of van Zyl, et al.,9 there was a 75% decrease in the total decreased EBA amplitude within 20 minutes of CAS.

The authors reported that DPOAE amplitude remained suppressed at the recording obtained 1 minute after termination of acoustic stimulation, and that the depression recovered in subsequent recordings at 1-minute intervals, indicating residual suppression after the termination of acoustic sound. They attributed this phenomenon to the medial olivocochlear slow effect or to the persistence of the efferent effect after the termination of acoustic sound.

Even though we used a similar experimental setup to that of van Zyl, et al., we found several different phenomena: efferent fatigue during CAS, and overshooting and recovery at several frequencies after CAS. Consequently, more research is necessary to fully understand the underlying mechanisms of the suppressive effect.

The limitation of the current study is the absence of the control group whose TEOAEs were checked at the same time interval without CAS.

**Conclusion**

We revealed significant TEOAE amplitude reduction with CAS in adults with normal hearing, and verified the suppressive effect of the medial olivocochlear system. The present results demonstrate that TEOAE suppression occurs immediately after CAS, and indicate the possible manifestation of efferent fatigue or sensory adaptation. After the termination of acoustic stimulation, transient TEOAE amplitude enhancement (overshooting) and spontaneous recovery were observed.

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