## **JADARA**

Volume 49 Number 2 Winter 2015

Article 2

2-28-2015

# A Review of Ototoxic Medications: Implications for Professionals Working with Consumers with Hearing Loss

Shawn P. Saladin Ph.D, CRC, CPM The University of Texas-Pan American

Robert B. Perez The University of Texas-Pan American

Bianca Cruz Pharm.D The University of Texas-Pan American

Yasar Tasnif Pharm.D, BCPS The University of Texas-Pan American

Follow this and additional works at: https://repository.wcsu.edu/jadara



Part of the Social and Behavioral Sciences Commons

#### Recommended Citation

Saladin, S. P., Perez, R. B., Cruz, B., & Tasnif, Y. (2015). A Review of Ototoxic Medications: Implications for Professionals Working with Consumers with Hearing Loss. JADARA, 49(2). Retrieved from https://repository.wcsu.edu/jadara/vol49/iss2/2

This Article is brought to you for free and open access by WestCollections: digitalcommons@wcsu. It has been accepted for inclusion in JADARA by an authorized editor of WestCollections: digitalcommons@wcsu. For more information, please contact ir@wcsu.edu.

## A Review of Ototoxic Medications! Implications for Professionals Working with Consumers with Hearing Loss

Shawn P. Saladin, Ph.D., CRC, CPM Robert B. Perez Bianca Cruz, Pharm.D. Yasar Tasnif, Pharm.D., BCPS The University of Texas-Pan American

#### Abstract

Rehabilitation professionals work with a variety of people with various disabilities and disease states. Oftentimes, those disease states are treated with medications. An understanding of the most common types of medications that can cause ototoxicity, such as aminoglycides, cisplatin and loop diuretics can assist in the vocational rehabilitation process. Rehabilitation professionals are encouraged to establish positive communication with the consumers in order to foster a relationship of trust. Furthermore they are encouraged to establish relationships with pharmacists for an understanding of the implications of commonly used medications on hearing.

Keywords: deaf, ototoxic, drugs, hearing loss, treatment

#### Introduction

Both general and specialized rehabilitation professionals need to have an understanding of the effects of mediations on their consumers.. State, federal and private rehabilitation professionals can work with homogeneous or heterogeneous populations with regards to the type of disability reported in each population. For example, some homogeneous caseloads may focus on people with hearing loss, spinal cord injuries, traumatic brain injuries, or learning disabilities, to name a few. These are specialized caseloads in which, ideally, the rehabilitation professional is an expert. This is often the case with people who are blind, as is evident from several states having separate rehabilitation agencies to address the specialized needs of this population. Another common caseload focus is serving people who have hearing loss. Some states have a designated agency to serve the individualized needs of this population while others specify certain caseloads to best serve this population. However, the ability to set aside an entire agency, or even a caseload, is often not feasible, especially in sparsely populated states (Watson, Jennings, Tomlinson, Boone & Anderson, 2008). Sometimes, in rural areas, the distance the consumer and/or the rehabilitation professional

1

need to travel for basic services is excessive; therefore, it is not feasible to have a specialized counselor. In light of not all consumers being served by specialty counselors who focus on their type of disability, being served by a general rehabilitation professional is the likely scenario for the majority of consumers.

#### Ototoxic Medications

This paper focuses on commonly used medications that may cause ototoxicity and the implication for rehabilitation professionals serving the people who experience this side effect. Ototoxicity is a serious adverse drug reaction, and may result in cochleotoxocity, or permanent irreversible bilateral sensorineural hearing loss. It can also cause ataxia, nausea, and vertigo (vestibulotoxicity; Xie, Talaska, & Schact, 2011). These possible reactions warrant attention. The consumer may or may not start the caseload with severe irreversible hearing loss, but it is possible for this to occur because of the types of medications consumers are prescribed for the various other health conditions they may have.

### Aminoglycosides

Aminoglycosides (AGs) are a class of antibiotics (e.g., gentamicin, tobramycin, amikacin, neomycin, streptomycin) commonly used to treat infections. Ototoxic effects vary within this class. The ranking of AGs from most toxic to least toxic to the auditory system are as follows: neomycin, tobramycin, kanamycin, gentamicin and amikacin (Rybak, 2007).

When properly administered, AGs rapidly enter the inner ear (Fisman & Kaye, 2000). However, AG concentration levels do not appear to correlate well with incidence of ototoxicity. Prolonged use of AGs over five to six months has shown evidence of AGs in sensory cells (Barza, Ioannidis, Cappelleri, & Lau, 1996; Dulon, Aran, Zajic, & Schacht, 1986). This prolonged use may be the cause of damage to type 1 sensory hair cells in the vestibular organ and the outer hair cells in the origin of Corti (Rybak, 2007). Damage begins gradually from the base of the cells that detect high frequency sounds and progresses to the apex of the cochlea, which detects low frequency sound (Huizing & deGroot, 1987). Conversely, Hinojosa and Lerner (1987) discovered cochlear ganglion cell destruction without damage to the hair cells.

In addition to taking the medication, there are other factors that can increase the ototoxic effects of the AGs. Noise induction (Zimmerman & Lahav, 2012), concurrent use of other ototoxic medications (Ding & Salvi, 2010; Garcia, Martinez, Agustí, Mencía & Asenjo, 2001), mutations in an individual's sequences of mitochondrial ribosomal RNA (Fishel-Ghodsian, et al., 1997; Xing, Chen, & Cao, 2007) and other biomedical reasons (Guan, Fischel-Ghodsian, & Giuseppe, 2000) may exacerbate the ototoxic effects of the AGs.

## Cisplatin

Cisplatin is an antineoplastic agent used to treat cancers such as ovarian, testicular, and bladder cancer. This drug has been found to be ototoxic to the point of causing irreversible hearing loss in the cochlea (Schacht, Talaska, & Rybak, 2012). Similar to AGs, cisplatin affects the hair cells in the cochlea from the base to the apex. In other words, the ability to hear high frequencies will be affected first, and then the continuum of loss will progress to the lower frequencies (Church, Blakley, Burgio, & Gupta, 2004; Estivill, et al., 1998; Harris, Gilbert, Lormore, Musunuru, & Fritsch, 2011; Hinojosa, Riggs, Strauss, & Matz, 1995). Other factors, such as the dose administered, combined with noise exposure may also contribute to a higher risk of cisplatin-induced ototoxicity (Bokemeyer et al., 1998; Li, Womer, & Silber, 2004).

## **Loop Diuretics**

Edema, defined as swelling, and occurs as a result of excess fluid trapped in the interstitium. It is often caused by disease states such as heart failure, renal insufficiency, or liver cirrhosis. Edema is most commonly treated with loop diuretics that include furosemide, bumetanide, and ethacrynic acid. Furosemide, administered orally or intravenously, is most commonly associated with ototoxicity (Chiodo & Alberti, 1994). This is caused by the blockage of fluid movement out of the stria vasularis affecting the marginal cells. However, unlike AGs and cisplatin, loop diuretics have very little effect on the hair cells (Ikeda, Oshima, Hidaka, & Takasaka, 1997). The hearing loss is usually temporary and correlates with the amount received; the higher the amount of furosemide given, the greater the hearing loss (Chiodo & Alberti, 1994). Additionally, when the loop diuretic is discontinued, the hearing may be restored.

## Implications for Rehabilitation Professionals

Rehabilitation professionals whose primary responsibility is assisting consumers in obtaining and maintain employment need to have an understanding of the types of medications the consumers may take. The consumers may have issues in regards to adjusting to new hearing loss. For example, they may not know the types of technology available to them or how to use these technologies. Furthermore, in the event they are to interview, not knowing how to use technology or simple communication strategies may have negative effects in terms of employment outcomes. As rehabilitation professionals work on such cases, they are encouraged to be cognizant of changes in the consumers' health, and be knowledgeable of the medications the consumers are prescribed. Developing a positive supportive working relationship with the consumer is vital to the exchange of this type of information. Furthermore, it behooves the rehabilitation professional to establish a relationship with a pharmacist. This healthcare professional is better able to explain the effects of the consumer's medications. The rehabilitation professional can use this information, in conjunction with information from a comprehensive assessment, to develop a plan of services. Establishing this relationship may take a bit of effort, and one must be cautious not to share confidential information. There are several ways to do this, and each situation will be different; the salient factor is communication. The rehabilitation professional may simply make an appointment with an area pharmacist and discuss ototoxicity. Pharmacists may also be invited to speak at the rehabilitation professional's monthly quarterly or annual meetings or at local professional meetings.

If a more self-directed search is needed, several websites list the various medications that can cause hearing loss. A search of "organizations with ototoxic medication list 2014" resulted in about 275,000 results, with the first result being from the Northern Virginia Resource Center for Deaf and Hard of Hearing People (http://www.nvrc.org/hearing-loss-deafness/ototoxic-drugs/). Another reliable source is from the World Health Organization (WHO: http://www.who.int/mediacentre/factsheets/fs300/en/).

#### Conclusion

In addition to understanding the medications, rehabilitation professionals need to understand the potential ototoxic effects some medications can cause. As previously mentioned, a noisy environment (Zimmerman & Lahay,

2013) and other genetic issues (Fishel-Ghodsian, et al., 1997; Xing, Chen, & Cao, 2007) may compound hearing loss in consumers. An understanding of hearing loss types, whether permanent or temporary, and the severity of the loss, is important for working with consumers.

In addition to counseling, other psychological services may be necessary for those who expect a slow recovery or permanent loss. Consumers may need to learn coping strategies for communication in their usual environments, which could have service implications. The consumers may need training and counseling on the roles and functions of additional assistive technology, learn visual forms of communication, or even be trained for another career.

#### **Contact Information**

Shawn P. Saladin
College of Health Sciences and Human Services
The University of Texas-Pan American
1201 West University Drive
Edinburg, TX 78539
(956) 665-2291.
ssaladin@utpa.edu

#### References

- Barza, M., Ioannidis, J. P., Cappelleri, J. C., & Lau, J. (1996). Single or multiple daily doses of aminoglycosides: a meta-analysis. *British Medical Journal*, 312(7027), 338-344.
- Bokemeyer, B., Berger, C., Hartmann, J., Kollmannsberger, C., Schmoll, H., Kuczyk, M. et al. (1998). Analysis of risk factors for cisplatin-induced ototoxicity in patients with testicular cancer. *British Journal of Cancer*, 160(6), 2292-2293.
- Chiodo, A. A., & Alberti, P. W. (1994). Experimental, clinical and preventive aspects of ototoxicity. *European Archives of Oto-Rhino-Laryngology*, 251(7), 375-392.
- Church, M. W., Blakley, B. W., Burgio, D. L., & Gupta, A. K. (2004). WR-2721 (Amifostine) ameliorates Cisplatin-induced hearing loss but causes neurotoxicity in hamsters: Dose-dependent effects. *Journal of the Association for Research in Otolaryngology*, 5(3), 227-237.
- Ding, D., & Salvi, R. J. (2010). Mechanisms of rapid sensory hair-cell death following coadministration of gentamicin and ethacrynic acid. *Hearing Research*, 259, 16-23. doi: 10.1016/j. heares.2009.08.008.
- Dulon, D., Aran, J. M., Zajic, G., & Schacht, J. (1986). Comparative uptake of gentamicin, netilmicin, and amikacin in the guinea pig cochlea and vestibule. *Antimicrobial Agents and Chemotherapy*, 30(1), 96-100.
- Estivill, X., Durbano, L., Scozzari, R., Moral, L., Romero, E., Badenas, C. et al. (1998). Familial progressive sensorineural deafness is mainly due to the MtDNA A1555G mutation and is enhanced by treatment with Aminoglycosides. *The American Journal of Human Genetics*, 62(1), 27-35.
- Fischel-Ghodsian, N., Prezant, T. R., Chaltraw, W. E., Wendt, K. A., Nelson, R. A., Arnos, K. S., et al. (1997). Mitochondrial gene mutation is a significant predisposing factor in aminoglycoside ototoxicity. *American Journal of Otolaryngology*, 18(3), 173-178.

- Fisman, D. N., & Kaye, K. M. (2000). Once-daily dosing of aminoglycoside antibiotics. *Infectious Disease Clinics of North America*, 14(2), 475-487.
- Garcia, V. P., Martínez, F. A., Agustí, E. B., Mencía, L. A., & Asenjo, V. P. (2001). Drug-induced otoxicity: current status. *Acta Oto-Laryngologica*, 121(5), 569-572.
- Guan, M., Fischel-Ghodsian, N., & Giuseppe, A. (2000). A biochemical basis for the inherited susceptibility to aminoglycoside ototoxicity. *Human Molecular Genetics*, *9*(12), 1787-1793.
- Harris, M. S., Gilbert, J. L., Lormore, K. A., Musunuru, S. A., & Fritsch, M. H. (2011). Cisplatin ototoxicity affecting cochlear implant benefit. *Otology & Neurotology*, 32(6), 969-972. doi: 10.1097/MAO.0b013e3182255893
- Hinojosa, R., & Lerner, S. A. (1987). Cochlear neural degeneration without hair cell loss in two patients with aminoglycoside ototoxicity. *Journal of Infectious Diseases*, 156(3), 449-455.
- Hinojosa, R., Riggs, L. C., Strauss, M., & Matz, G. J. (1995). Temporal bone histopathology of cisplatin ototoxicity. *American Journal Otology*, 16(6), 731-740.
- Huizing, E. H., & deGroot, J. C. (1987). Human cochlear pathology in aminoglycoside ototoxicity—a review. *Acta Oto-Laryngologica*, *104*(s436), 117-125.
- Ikeda, K., Oshima, T., Hidaka, H., & Takasaka, T. (1997). Molecular and clinical implications of loop diuretic ototoxicity. *Hearing Research*, 107(1-2), 1-8.
- Li, Y., Womer, R. B., & Silber, J. H. (2004). Predicting cisplatin ototoxicity in children: the influence of age and the cumulative dose. *European Journal of Cancer*, 40(16), 2445-2451.
- Northern Virginia Resource Center for Deaf and Hard of Hearing People. (n.d.) Ototoxic drugs. Retrieved from http://www.nvrc.org/hearing-loss-deafness/ototoxic-drugs/

- Rybak, L. P. (2007). Mechanisms of cisplatin ototoxicity and progress in otoprotection. *Current Opinion in Otolaryngology & Head and Neck Surgery*, 15(5), 364-369.
- Schacht, J., Talaska, A. E., & Rybak, L. P. (2012). Cisplatin and aminoglycoside antibiotics: hearing loss and its prevention. *The Anatomical Record: Advances in Integrative Anatomy and Evolutionary Biology*, 295(11), 1837-1850. doi: 10.1002/ar.22578
- Watson, D., Jennings, T., Tomlinson, P., Boone, S., & Anderson, G. (2008). Model state plan for rehabilitation of persons who are deaf, deaf-blind, hard of hearing or late deafened. University of Arkansas RRTC for Persons who are Deaf or Hard of Hearing. Retrieved from http://humanservices.hawaii.gov/vr/files/2013/04/mspdeaf. pdf
- World Health Organization. (February 2014). Deafness and hearing loss. Retrieved from http://www.who.int/mediacentre/factsheets/fs300/en/Xie, J., Talaska, A. E., & Schacht, J. (2011). New developments in aminoglycoside therapy and ototoxicity. *Hearing Research*, 281(1-2), 28-37. doi: 10.1016/j.heares.2011.05.008
- Xing, G., Chen, Z., & Cao, X. (2007). Mitochondrial rRNA and tRNA and hearing function. *Cell Research*, 13, 227-239.
- Zimmerman, E., & Lahav, A. (2012). Ototoxicity in preterm infants: effects of genetics, aminoglycosides, and loud environmental noise. *Journal of Perinatology*, 33(1), 3-8. doi: 10.1038/jp.2012.105