# Common Aminoglycosides and Platinum-Based Ototoxic Drugs: Cochlear/Vestibular Side Effects and Incidence

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#### **ABSTRACT**

This is a reference chart that identifies 16 aminoglycoside antibiotics and platinum based drugs that could be cochleotoxic, vestibulotoxic or both. Using the most currently available data from published research from the National Library of Medicine's PubMed data base, incidence figures and risk factors are included in the chart along with the potential of permanence of reversibility of the impairment.

**KEYWORDS:** Cochleotoxic, vestibulotoxic, aminoglycoside antibiotics

Aminoglycosides and chemotherapy agents are two classes of drugs that are known to have hearing and vestibular side effects. The table below is offered as a quick resource to these classes of drugs, common uses, toxicity, and any information related to possible reversibility of the toxic effect.

NOTE: SEVERAL RESOURCES FOR DRUG INFORMATION INCLUDE

- 1. www.drugs.com
- 2. www.rxlist.com
- 3. Drug manufacturer's Web site

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## Reference Chart Incidence<sup>a</sup> of Aminoglycoside and Platin Hearing Loss and Vestibular Side Effects

Name	Uses	C. to XIC	Risk (%)	V. toxic	Risk (%)	Reversible?
Amikacin	Gram-negative infections, especially meningitis, respiratory infections, cystic fibrosis, and burns	Yes	71% <sup>1</sup>	No		Possibly
Carboplatin	Chemotherapeutic alkylating agent used for ovarian, testicular, and lung cancers, medulloblastoma	Yes <sup>b</sup>	8–10% <sup>2,4,5</sup>	No		No <sup>3</sup>
Cisplatin	Tx of gyn., genitourinary tract, lung, testicular, central nervous system, and head/neck cancers	Yes <sup>b</sup>	Up to 100% <sup>6</sup>	Yes (temp) <sup>7</sup>	D/N/A	No <sup>3</sup>
Erythromycin	Gram-positive and negative infections, diphtheria, tetanus, pertussis, also used for acute otitis media, sinusitis pharyngitis, tracheobronchitis, bronchitis, and bacterial laryngitis	Yes	~38% <sup>8</sup>	Yes <sup>9</sup>	D/N/A	Yes (hearing only)
Ethacrynic acid	Congestive heart failure, renal failure, cirrhosis, hypertension	Yes	<1% <sup>10</sup>	No <sup>11</sup>		Yes (adults) No (neonates)
- urosemide	Congestive heart failure, renal failure, cirrhosis, hypertension	Yes	Varies <sup>12</sup>	Possibly <sup>13</sup> Rare <sup>14</sup>		Yes (adults) No (neonates)
Gentamicin	Gram-negative infections, particularly meningitis, acute/chronic mastoiditis, necrotizing abscess, malignant otitis externa, chronic suppurative otitis media	Yes <sup>b</sup> No <sup>16</sup>	0–58% <sup>15</sup>	Yes <sup>15</sup>		D/N/A
Kanamycin	Gram-negative infections, including tuberculosis, hepatic coma, and for prophylaxis	Yes	Up to 41% <sup>16</sup>	No	D/N/A	Possibly
Neomycin	Otic and ototopical agent for chronic suppurative otitis media, acute dif- fuse otitis externa and	No (short term) <sup>17</sup> Yes (long term) <sup>17</sup>	D/N/A D/N/A	Yes	<1% <sup>18</sup>	No (long term) <sup>17</sup>
						(Continu

### (Continued)

Name	Uses	C. to XIC	Risk (%)	V. toxic	Risk (%)	Reversible?_
	use with hepatic coma					
	and as prophylaxis					
Netilmicin	Gram-negative infec-	Yes <sup>c</sup>	D/N/A	Yes <sup>19</sup>	D/N/A	No
	tions, particularly those					
	that are gentamicin					
	resistant					
Quinine	Historically to treat mala-	Yes <sup>20,21</sup>	~90% <sup>21</sup>	Yes <sup>20</sup>	D/N/A <sup>20</sup>	Possibly
	ria, with popular use for					
	leg cramps, also an					
	antipyretic					
Salicylates	Anti-inflammatory, anti-	Yes <sup>b</sup>	Dose	No	D/N/A	Yes w/in
	pyretic, analgesic used		dependent			48–72 h
	to treat patients with					after d/c
	arthritis, rheumatic fever,					
	and disorders of connec-					
	tive tissues, and widely					
	used for cardiac patients					
	with transient ischemic					
	attacks, stroke, unstable					
	angina, or myocardial					
Ctrantanavain	infarction	Yes <sup>22,23</sup>	33-59% <sup>24</sup>	Va 22,23	D/N/A <sup>b</sup>	Cochlear—
Streptomycin	· ·	res		res	D/N/A	Yes
	tions, tuberculosis, tula- remia plague, brucellosis		up to 100% <sup>b</sup>			vestibular—
	reifila piague, pruceilosis		100 %			Yes
Tobramycin	Gram-negative infec-	Yes <sup>25</sup>	11.5% <sup>25</sup>	Yes	D/N/A	Yes <sup>25</sup>
TODIATTIYCIII	tions, particularly menin-	162	11.570	162	DINA	162
	gitis, lower respiratory,					
	and urinary infections					
Vancomycin	Methicillin-resistant	Yes <sup>26</sup>	12% <sup>26</sup>	No	D/N/A	Possibly
vancomycin	(staph resistant) gram-	. 50	1270		2/14/16	. 2001017
	negative infections, com-					
	monly used for acute					
	mastoiditis and sinusitis,					
	orbital cellulitis, and					
	parotitis					
Viomycin	Gram-negative infec-	Yes <sup>27</sup>	D/N/A <sup>27,28</sup>	Yes <sup>27</sup>	D/N/A	D/N/A
	tions; tuberculosis		•	No <sup>28</sup>		

Abbreviation: D/N/A, data not available.

alnoidence figures are the highest percentages as reported in the most current articles in the PubMed database and other sources (see references).

bMultiple sources.

cIncludes late onset.

### **REFERENCES**

- Javadi MR, Abtahi B, Gholami K, Safari Moghadam B, Tabarsi P, Salamzadeh J. The incidence of amikacin ototoxicity in multidrug-resistant tuberculosis patients. Iran J Pharm Res 2011;10(04):905–911
- Fetoni AR, Ruggiero A, Lucidi D, et al. Audiological monitoring in children treated with platinum chemotherapy. Audiol Neurotol 2016;21(04): 203–211
- Clemens E, de Vries AC, Am Zehnhoff-Dinnesen A, et al. Hearing loss after platinum treatment is irreversible in noncranial irradiated childhood cancer survivors. Pediatr Hematol Oncol 2017;34(02):120–129
- Brooks B, Knight K. Ototoxicity monitoring in children treated with platinum chemotherapy. Int J Audiol 2018;57(Suppl 4):S34–S40
- Musial-Bright L, Fengler R, Henze G, Hernáiz Driever P. Carboplatin and ototoxicity: hearing loss rates among survivors of childhood medulloblastoma. Childs Nerv Syst 2011;27(03):407–413
- Paken J, Govender CD, Pillay M, Sewram V. Cisplatin-associated ototoxicity: a review for the health professional. J Toxicol 2016;2016:1809394
- Kobayashi H, Ohashi N, Watanabe Y, Mizukoshi K. Clinical features of cisplatin vestibulotoxicity and hearing loss. ORL J Otorhinolaryngol Relat Spec 1987;49(02):67–72
- Erythromycin and hearing loss. Available at: www. ehealthme.com/ds/erythromycin/hearing-loss. May 27, 2018
- Agustí C, Ferrán F, Gea J, Picado C. Ototoxic reaction to erythromycin. Arch Intern Med 1991; 151(02):380
- Matz GJ. The ototoxic effects of ethacrynic acid in man and animals. Laryngoscope 1976;86(08): 1065–1086
- Elidan J, Lin J, Honrubia V. The effect of loop diuretics on the vestibular system. Assessment by recording the vestibular evoked response. Arch Otolaryngol Head Neck Surg 1986;112(08):836–839
- 12. Rybak LP. Pathophysiology of furosemide ototoxicity. J Otolaryngol 1982;11(02):127–133
- Vestibular Disorders Association. Available at: https://vestibular.org/ototoxicity. Accessed 2018
- Frymark T, Leech H, Mullen R, Schooling T, Venediktov R, Wang B. Evidence-based systematic review: drug-induced hearing loss – gentamicin. Available at: www.drugs.com/sfx/lasix-side-effects. html. Accessed 2010

- Sequi Canet JM, Angelats Romero CM, Sequi Sabater JM, Miralles Torres A, Boronat Garcia M, Gomez Delgado M. Otoacoustic emissions in children treated with gentamicin in a secondary hospital. Acta Otorrinolaringol Esp 2018;69(05): 291–296
- Sharma V, Bhagat S, Verma B, Singh R, Singh S. Audiological evaluation of patients taking kanamycin for multidrug resistant tuberculosis. Iran J Otorhinolaryngol 2016;28(86):203–208
- Winterstein AG, Liu W, Xu D, Antonelli PJ. Sensorineural hearing loss associated with neomycin eardrops and nonintact tympanic membranes. Otolaryngol Head Neck Surg 2013;148(02):277–283
- Professional guide for Neomycin and Polymyxin B. Available at: www.drugs.com/ppa/neomycin.html. Accessed 2018
- American Medical Association. AMA Drug Evaluations. 3rd ed. Littleton, MA: PSG Publishing Co., Inc.; 1977:759
- Jarboe JK, Hallworth R. The effect of quinine on outer hair cell shape, compliance and force. Hear Res 1999;132(1-2):43–50
- Obasikene G, Adobamen P, Okundia P, Ogusi FO. Prevalence of ototoxicity in University of Benin Teaching Hospital, Benin city: a 5-year review. Niger J Clin Pract 2012;15(04):453–457
- Duggal P, Sarkar M. Audiologic monitoring of multi-drug resistant tuberculosis patients on aminoglycoside treatment with long term follow-up. BMC Ear Nose Throat Disord 2007;7:5
- Adeyemo AA, Oluwatosin O, Omotade OO. Study of streptomycin-induced ototoxicity: protocol for a longitudinal study. Springerplus 2016;5 (01):758
- Bitner-Glindzicz M, Rahman S. Ototoxicity caused by aminoglycosides. BMJ 2007;335(7624(:784–785
- Matz GJ. Aminoglycoside ototoxicity. Am J Otolaryngol 1986;7(02):117–119
- Forouzesh A, Moise PA, Sakoulas G. Vancomycin ototoxicity: a reevaluation in an era of increasing doses. Antimicrob Agents Chemother 2009;53 (02):483–486
- Lustig LR. Drug-induced ototoxicity. Merck Manual; 2018. Available at: www.merckmanuals. com/professional/ear,-nose,-and-throat-disorders/ inner-ear-disorders/drug-induced-ototoxicity. Accessed February 28, 2019
- 28. Brummett RE. Drug-induced ototoxicity. Drugs 1980;19(06):412–428