

Meningitis post-cochlear implant and role of vaccination

Ghadah A. Alanazi, MBChB, Aumnayat S. Alrashidi, MBChB, Khalid S. Alqarni, MD, Saeed A. Al khozaym, MD, Saad Alenzi, MD.

ABSTRACT

الأهداف: التحقيق في حدوث وعوامل الخطر وعلاج التهاب السحايا لدى مستخدمي غرسة القوقعة الصناعية (CI).

المنهجية: أجريت مراجعة منهجية باستخدام Scopus, PubMed و Web of Science و كوكرين. تم اعتبار المقالات ذات صلة إذا تم الإبلاغ عن أي بيانات عن حدوث، والحالات السريرية، ودور التطعيم، والعلاج، ونتائج التهاب السحايا بعد (CI).

النتائج: اشتملت الدراسة على 32 دراسة شملت 27358 مريضاً، وتم الإبلاغ عن التهاب السحايا في 202 حالة فقط. حدث التهاب السحايا في فترة تتراوح من يوم واحد إلى 72 شهراً بعد CI. تلقى مجموعه 55 مريضاً لقاح المكورات الرئوية، بينما تلقى 20 مريضاً لقاح المستدمية النزلية من النوع ب. ارتبط عدد كبير من المشاركين (العدد = 47) بتشوّهات تشريحية مرتبطة، بينما كان لدى 62 مشاركاً التهاب الأذن الوسطى قبل التهاب السحايا. مجموعه 24 حالة تتطلب جراحة مراجعة إلى جانب العلاج الطبي. كان الشفاء التام هو النتيجة التي أبلغت عنها الدراسات المشمولة في 19 مريضاً.

الخلاصة: يبدو أن مستخدمي غرسة القوقعة الصناعية معرضون لخطر الإصابة بالتهاب السحايا الجرثومي في أي وقت بعد الزرع، خاصة في ظل وجود عوامل الخطر، مثل التهاب الأذن الوسطى.

Objectives: To investigate the incidence, risk factors, and management of meningitis in cochlear implant (CI) users.

Methods: A systematic review was carried out using PubMed, Scopus, Web of Science, and Cochrane Central Register. Articles were considered relevant if reported any data on incidence, clinical presentations, the role of vaccination, management, and outcomes of meningitis after CI.

Results: A total of 32 studies including 27358 patients were included, and meningitis was reported in only 202 cases. Meningitis occurred in the period ranging from 1 day to 72 months after CI. A total of 55 patients received the pneumococcal vaccine, while 20 patients received the *Haemophilus influenzae* type B vaccine. A large number of participants (n=47) had associated anatomical malformations, while 62

had otitis media before meningitis. A total of 24 cases required revision surgery along with medical treatment. Full recovery was the outcome reported by the included studies in 19 patients.

Conclusion: Cochlear implant users seem to be at possible risk of bacterial meningitis at any time after implantation, especially in the presence of risk factors, such as otitis media and anatomical malformations of the cochlea.

Keywords: cochlear implant; bacterial meningitis, vaccine, otitis media

Saudi Med J 2022; Vol. 43 (12): 1300-1308
doi: 10.15537/smj.2022.43.12.20220426

From the Faculty of Medicine (Alanazi, Alrashidi), Tabuk University; from the Department of Otolaryngology (Alqarni, khozaym, Alenzi), King Fahad Specialist Hospital, Tabuk, Kingdom of Saudi Arabia.

Received 17th July 2022. Accepted 5th October 2022.

Address correspondence and reprint request to: Dr. Ghadah A. Alanazi, Faculty of Medicine, Tabuk University, Tabuk, Kingdom of Saudi Arabia. E-mail: Ghada.alanazi35@gmail.com
ORCID ID: <https://orcid.org/0000-0002-6474-1088>

Cochlear implants (CI) are commonly used interventions for children and adults with sensorineural hearing loss (SNHL). Cochlear implants utilisation among children began in 1980 and has been significantly increasing.¹ The CI devices carry electrical stimulation using specific electrodes to the fibres of the cochlear nerve which provides an efficacious and reliable method of rehabilitation for patients with SNHL.^{2,3}

Several adverse events may follow CI such as device malfunction, bleeding, and electrode migration, fascial nerve stimulation, and infection.³ It has been reported that CI may carry a higher risk of bacterial meningitis in comparison with the general population.⁴⁻⁸ In 2002, the United States Food and Drug Administration released a notification of receiving several reports, indicating a potential correlation between CI and the incidence of bacterial meningitis.⁹ Bacterial meningitis is a critical illness with remarkable morbidity and mortality rates. The most commonly identified meningitis organisms in a previous report were *Streptococcus pneumoniae* (*S. pneumoniae*) and *Haemophilus influenzae*. The death rate from pneumococci-induced bacterial meningitis has been estimated to range from 15% to 60%.¹⁰

There is a lack of knowledge regarding the direct effects of meningitis on CI; however, some reports have mentioned the failure of implanted devices that require reimplantation. Several risk factors may contribute to meningitis in CI users, including otitis media, head trauma, cochlear malformations, and cerebrospinal fluid (CSF) leaks.⁴⁻⁸ The clinical features of post-CI meningitis do not differ from those of classical meningitis which include high temperature, headache, neck stiffness, photophobia, sickness, and vomiting. However, CI users may experience further complaints such as vertigo.¹¹ It has been recommended that CI candidates and users should receive full immunization against organisms causing bacterial meningitis, especially *S. pneumoniae*, as it is more prevalent.^{12,13} The CDC recommendations for CI patients who never received pneumococcal vaccines include receiving of one shot of pneumococcal conjugate vaccine 15 followed by one shot of pneumococcal polysaccharide vaccine 23.¹⁴

There is a lack of systematic reviews that assemble published reports on meningitis among CI users and draw comprehensive recommendations or conclusions. This systematic review aimed to report meningitis occurrence among CI users in terms of patients' demographic characteristics, incidence, associated risk factors, possible immunization role, and management.

Methods. A systematic literature search was carried out following the Preferred Reporting Items for Systematic Reviews and Meta-Analysis

recommendations. The project protocol was written and registered in PROSPERO (CRD42021288471).

In November 2021, a systematic search was conducted in PubMed, Scopus, Web of Science, and Cochrane Central Register using these search terms (Meningitis AND [CI OR cochlear implantation]). We aimed to retrieve all search results without applying any search filters. All search results were combined into one Endnote library, and all duplicated references were removed. Articles were then transferred to an Excel spreadsheet for title/abstract screening for potential relevance. Articles were considered relevant if they were original studies reporting any data on the incidence, clinical presentations, role of vaccination, management, and outcomes of meningitis after cochlear implantation. There were no restrictions on the year or place of research, patient age, or study design. However, we did not include i) conference papers, comments, letters, review papers, or book chapters; ii) articles with overlapping data sets; iii) non-English articles; and iv) animal studies. Two authors independently screened articles according to the aforementioned criteria. Further screening rounds using full texts were also performed for the final decision on the inclusion or exclusion of specific reports.

Three reviewers independently extracted the data from the final full texts. Extracted data included information on individuals' demographic characteristics, sample size, the number of CI users with meningitis, and implant device type. It also comprises data on the number of episodes, causative organism, risk factors, immunization status, diagnostic method, management plan, and outcome. Any discrepancies were resolved through discussion with a fourth author.

Statistical analysis. Descriptive analysis of the extracted data was carried out using Microsoft Excel to estimate the numbers, percentages and means.

Results. A total of 1600 reports were obtained from the databases, of which 909 were removed as duplicate references. After title/abstract screening, only 48 remained for full-texts review. Manual reference checks retrieved more reports; eventually, 32 articles were eligible for analysis (Figure 1).

A total of 32 studies published between 1991 and 2020 and 27358 patients were included.^{4-6,8,10,14-40} Seventeen were case studies while 15 were retrospective observational studies. Most of the studies (37.5%) were carried out in the United States. The mean age of the patients with meningitis after CI ranged from 1 to 11 years. Among the 23 studies reporting patient gender, there were 45 (64.3%) male patients and 25 (35.7%)

Disclosure. Authors have no conflict of interests, and the work was not supported or funded by any drug company.

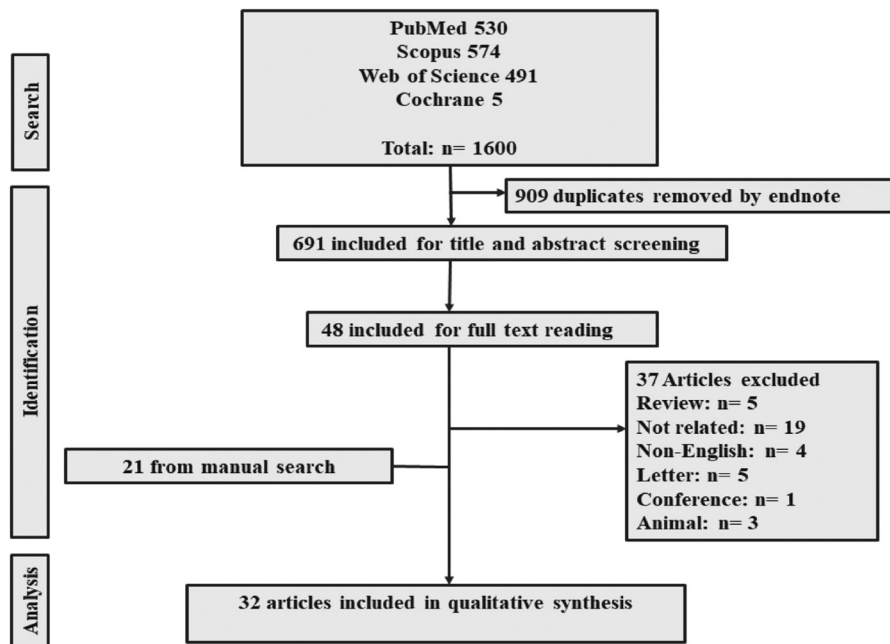


Figure 1 - Flow diagram of studies selection and screening.

female patients. The type of implant device and the basic demographic characteristics are detailed in [Table 1](#).

Incidence of meningitis and vaccination status.

Of the 27358 who performed CI, 202 (0.7%) who developed meningitis were enrolled in the study. Meningitis occurred in the period ranging from 1 day to 72 months after CI. *Streptococcus pneumoniae* was responsible for the incidence of meningitis in 42 episodes of infection. Ten patients experienced more than one episode of meningitis. A total of 55 patients had a history of immunization with the pneumococcal vaccine, while 20 patients received the *Haemophilus influenzae* type B vaccine (HIB). The remaining studies did not report any detailed data regarding the pathogenic organism or previous vaccination. Regarding potential risk factors, as per the authors' report, a large number of participants (n=47) had associated anatomical malformations. Moreover, 62 patients had otitis media before meningitis and 23 had a history of CSF leak after CI ([Table 2](#)).

Management and outcomes of meningitis cases after CI. Clinical evaluation and CSF culture were frequently reported methods of diagnosis (n=18 studies); otherwise, they were not reported. A total of 42 infection episodes were managed only through medical treatment, whereas 24 episodes required revision surgery along with medical treatment. Full recovery was the outcome reported by 19 patients in the included studies ([Table 3](#)).

Discussion. This review investigated the incidence of meningitis in CI users. The incidence has been reported in a minority of cases (0.7%) involving several risk factors. Meningitis may occur at any time point after CI, which warrants the immediate identification of suspected cases for earlier management.⁴¹ The exact mechanism underlying the incidence of meningitis after CI remains unclear. Probable mechanisms have been previously investigated in animal research that suggested that immune response, by way or another, is affected by the presence of the foreign body in the inner ear.¹² Our study showed that some patients who were vaccinated against meningitis may still have been infected. Vaccination against bacterial meningitis in developed nations has become routine practice. Moreover, it is recommended that children undergoing surgical interventions have an increased risk of post-surgery meningitis. Owing to the vast number of immunization practices, the incidence of meningitis has declined significantly, and such a reduction has been reported in European and North American countries.^{42,43} In contrast, immunization against meningitis in developing countries is still challenging and carries a financial burden as being not supported by their local healthcare systems.³² Regarding children who are CI recipients or scheduled for CI, conjugate vaccines against *S. pneumoniae* and *H. influenzae* are recommended for all patients with CI, while the meningococcal vaccine is not regularly required.^{41,44,45} Moreover, in children aged

Table 1 - Baseline demographic characters of enrolled studies.

Author/year	Country	Design	Total sample size	Meningitis patients (N)	Age mean Y (Range)	Male/Female	Device type
Cohen & Hoffman ¹⁵ /1991	USA	ROS	921	23	-	-	(23) Nucleus device
Daspir ⁶ /1991	USA	Case report	1	1	51	Male	(1) Nucleus device
Webb et al ¹⁶ /1991	Germany	ROS	712	1	-	-	(1) multichannel implant
Page & Eby ¹⁷ /1997	UK	Case report	1	1	5	Male	-
Woolley et al ¹⁸ /1998	USA	Case series	4	1	5 y 4 months	Female	(1) multichannel implant
Reefhuis. etl ¹⁹ /2003	USA	POS	4264	26	(11-6)	18/8	(4) CI24M, (11) AB-5100H, (2) AB-5100,(7) AB-5100H-11 , (2) CI24RST, (2) CI22M, (1) C40+HS
Callanan & Poje ¹⁰ /2004	USA	Case series	2	2	3-5	Male	(2) Clarion C-II device with positioner
Pettersen ²⁰ /2005	Canada	Case report	1	1	4	Male	(1) Advanced bionics with positioner
Biernath KR ²¹ /2006	USA	ROS	4265	12	(21-103) months	9/3	(8) AB-5100H-11, (1) C40HGB, (3) AB-5100H
Ahn et al ²² /2008	Korea	ROS	388	1	5.2	Male	(1) HF.1.2
Mancini et al ²³ /2008	Italy	Case series	3	3	(9-11)	2/1	(2) Clarion 1.2, 1 Clarion HiFocus CII
Torkos et al ²⁴ /2009	Hungary	Case report	1	1	2	Male	(1) Nucleus 24 R
Loundon et al ²⁵ /2010	France	ROS	434	1	6	-	-
McJunkin & Jeyakumar ²⁶ /2010	USA	ROS	165	1	5	Female	(1) Med El
Ahn et al ²⁷ /2011	Korea	ROS	11	2	-	1/1	(1) HF 1.2, (1) CI24RST
Ajallouyeen et al ⁴ /2011	Iran	Case series	262	1	1.5	-	-
Gurbuz et al ⁷ /2011	Turkey	Case report	1	1	8.5	Female	(1) Med-El© short electrode
Lalwani & Cohen ²⁸ /2011	USA	ROS	8,329	86	43 <5 y	-	(86) devices with positioner
O'Mahony et al ²⁹ /2011	USA	Case report	1	1	4	Male	(1) Implant with positioner
Mancini et al ⁸ /2013	Italy	Case series	5	5	(2.5-8)	2/3	AB devices: 2 with Clarion 1.2 HiFocus with positioner, 1 with HiFocus CII and 1 with HiResolutionTM 90K; 1 with a Med-El Combi 40+
Roman et al ³⁰ /2013	USA	Case report	1	1	2	Female	(1) Nucleus CI24 straight electrode array
Tarkan et al ³¹ /2013	Turkey	ROS	475	2	1.5-2.5	-	(1) Med-El Sonata (straight array), (1) Nucleus CI24RE (contour array)
Afsharpaiman et al ³² /2014	Iran	ROS	371	1	24 months	Female	-
Suleiman et al ³³ /2014	Nigeria	Case series	2	1	8	Female	(1) Medel Pulsar
Daneshi et al ³⁴ /2015	Iran	ROS	4346	4	-	-	-
Javia et al ³⁵ /2016	USA	ROS	478	4	(1.3-4)	3/1	(1) Clarion CII, 1 Clarion HiFocus , 2 AB HiRes 90K
Pross et al ³⁶ /2016	USA	Case report	1	1	4	Female	-
Tandon et al ³⁷ /2016	India	case report	1	1	1	Male	(1) Nucleus freedom implant CI24RE (straight electrode) with straight electrode array
Vartanyan et al ³⁸ /2018	Australia	Case report	1	1	23	Female	-
Dağkiran et al ⁵ /2020	Turkey	ROS	1452	5	-	-	-
Yan et al ³⁹ /2020	China	Case report	1	1	7	Male	-
Piromchai et al ⁴⁰ /2021	Thailand	ROS	458	9	-	-	(9) 3M device

ROS: retrospective study, POS: prospective study, USA: United States of America, Y: years, AB: advanced Bionics, HS: Houston Galveston Brazoria, HF: HiFocus electrode array

2-9 years, a conjugate pneumococcal vaccine followed by a 23-valent pneumococcal polysaccharide vaccine is recommended. Therefore, healthcare workers should check immunization records to ensure that all the required vaccines have been administered.^{20,45}

A large number of patients in the present study had otitis media before meningitis. As *S. pneumoniae* and *H. influenzae* are commonly identified organisms in the CSF, it is often proposed that these organisms take the

route of the middle ear to the cochlea and eventually to the meninges.⁴⁶ However, clinical and animal studies have reported that both implanted and non-implanted cochlea can withstand the spread of infection with similar effectiveness.⁴⁷ Nonetheless, CI doctors should discuss with infectious disease physicians regarding the optimal treatment for otitis media and its complications. Patients with CI who present with features of acute otitis media should be managed urgently to prevent

Table 2 - Reported meningitis with associated risk factors and role of vaccination.

Author/year	Meningitis cases (N)	Onset of meningitis (month)	Organism	N. of meningitis episodes	Immunization status	Anatomical deformities N of patients	Possible risk factors		
							CSF leak	Otitis media	Others
Cohen & Hoffman ¹⁵ /1991	23	-	-	-	-	-	-	-	-
Daspi ⁶ /1991	1	0.2	<i>Streptococcus pneumoniae</i>	-	-	-	1	-	Head trauma and bilateral Meniere's disease
Webb et al ¹⁶ /1991	1	-	-	-	-	-	-	-	-
Page & Eby ¹⁷ /1997	1	24	-	2	-	1	-	-	Sinusitis prior to meningitis onset
Woolley et al ¹⁸ /1998	1	7	-	1	-	1	1	1	-
Reefhuis et al ¹⁹ /2003	26	1 day-36 months	<i>Streptococcus pneumoniae</i> : 15 episodes <i>Haemophilus influenzae</i> type b: 5 episodes Acinetobacter baumannii: 2 episodes Enterococcus: 1 episode, <i>Escherichia coli</i> : 1 episode, Unknown: 5 episodes	23 patient: 1 3 patients: 2	-	9	6	13	Previous meningitis: 5 ventriculoperitoneal shunt: 1
Callanan & Poje ¹⁰ /2004	2	0.4-12	<i>Haemophilus influenzae</i> : 1 patient <i>Streptococcus pneumoniae</i> : 1 patient	1	HIB and the pneumococcal vaccines	1	1	1	-
Pettersen et al ²⁰ /2005	1	42	Group A streptococcus: 1 st episode, <i>Pseudomonas aeruginosa</i> : 2 nd episode	2	Pneumococcal and Neisseria meningitidis vaccines	-	-	2	-
Biernath et al ²¹ /2006	12	5-40	Group A Streptococcus: one patient Unknown: 2 patients	1	-	2	1	-	Previous meningitis: 3
Ahn et al ²² /2008	1	-	-	3	-	1	-	-	-
Mancini et al ²³ /2008	3	8-72	<i>Streptococcus pneumoniae</i> : 3 episodes, Negative CSF: one episode	2 patients: 1 1 patient: 2	Pneumococcal vaccine	2	-	-	-
Torkos et al ²⁴ /2009	1	6	<i>Haemophilus influenzae</i> in episode, <i>Streptococcus pneumoniae</i> in another	4	-	1	1	-	-
McJunkin & Jeyakumar ²⁶ /2010	1	12	<i>Pseudomonas aeruginosa</i>	1	-	1	-	-	-
Loundon et al ²⁵ /2010	1	48	<i>Streptococcus pneumoniae</i>	1	Complete immunization history	-	-	-	-
Ahn et al ²⁷ /2011	2	48	-	-	-	2	1	-	-
Ajallouyeen et al ⁴ /2011	1	-	-	-	-	-	-	-	-
Gurbuz et al ⁷ /2011	1	24	<i>Streptococcus pneumoniae</i>	4	HIB vaccine before first 3 episodes. Pneumococcal vaccine before the last episode	1	1	-	Head trauma

HIB: Haemophiles influenzae type B, CSF: cerebrospinal fluid

Table 2 - Reported meningitis with associated risk factors and role of vaccination (continuation).

Author/year	Meningitis cases (N)	Onset of meningitis (month)	Organism	N. of meningitis episodes	Immunization status	Possible risk factors			
						Anatomical deformities N of patients	CSF leak	Otitis media	Others
Lalwani Cohen ²⁸ /2011	86	59 within the first 24 month	-	-	Fully immunized: 31 HIB vaccine: 11 Not immunized: 22 Unknown: 22	9	-	42	Previous meningitis: 7
O'Mahony et al ²⁹ /2011	1	36	<i>Streptococcus pneumoniae</i>	1	Complete immunization history	1	-	-	-
Mancini et al ⁸ /2013	5	3 -72	<i>Streptococcus pneumoniae</i> <i>Streptococcus pneumoniae</i> in the first episode,	-	Pneumococcal vaccine	3	-	4	-
Roman et al ³⁰ /2013	1	5-10	Haemophilus influenza in the 2nd episode	2	HIB and the pneumococcal vaccines	1	1	-	-
Tarkan et al ³¹ /2013	2	1, 30	-	1 patient: 3	Pneumococcus vaccine	2	1	-	-
Afsharpaiman et al ³² /2014	1	-	-	-	Complete immunization history but not with Pneumococcal vaccine	-	-	-	-
Suleiman et al ³³ /2014	1	-	-	1	Pneumococcal vaccine	0	-	-	Previous meningitis
Daneshi et al ³⁴ /2015	4	-	-	-	Pneumococcal vaccine	-	-	-	-
Javia et al ³⁵ /2016	4	4 days-32 months	Acinetobacter baumannii: one patient, <i>Streptococcus pneumoniae</i> : one patient, Negative CSF: 2 patients	1	-	-	1	-	-
Pross et al ³⁶ /2016	1	3	<i>Streptococcus pneumoniae</i>	2	Pneumococcal vaccine	1	-	-	Previous meningitis Respiratory tract infection prior to meningitis
Tandon et al ³⁷ /2016	1	4-13	Nonhemolytic streptococci: 1 st episode, <i>Streptococcus pneumoniae</i> : 2 nd episode	3	Pneumococcal, meningococcal, and HIB vaccines	1	1	-	-
Vartanyan et al ³⁸ /2018	1	72	<i>Streptococcus pneumoniae</i>	1	HIB and Neisseria meningitidis vaccines	1	1	-	-
Dağkiran et al ⁵ /2020	5	-	-	-	-	5	4	-	-
Yan et al ³⁹ /2020	1	72	<i>Pseudomonas aeruginosa</i>	1	-	1	1	-	Head trauma
Piromchai et al ⁴⁰ /2021	9	-	-	-	-	-	-	-	-

HIB: Haemophilus influenzae type B, CSF: cerebrospinal fluid, N: number

the transmission of infection to the cochlea. Beside prophylaxis antibiotics for CI users who are known to be otitis-prone, immediate identification and aggressive management of each episode of acute otitis media should involve tympanostomy tubes and full course of antibiotic.⁴⁴ Moreover, patients undergoing CI with known cochlear malformations, such as Mondini, and enlarged vestibular aqueduct, are considered vulnerable to the risk of otogenic meningitis.⁴⁸

Our study showed that most infection episodes were managed with medical treatment alone. Parents of children with CI must be able to recognize the potential clinical features and seek medical evaluation immediately if their child's hearing is altered. The

recommended empirical antibiotics for the management of patients with CI with suspected meningitis do not differ from the routine recommendations for other cases of meningitis, as the organisms are similar. The current treatment regimen should include vancomycin combined with a broad-spectrum cephalosporin. In cases of chronic otorrhea or perforated tympanic membranes, antibiotics such as ceftazidime, cefepime, or meropenem are recommended to fight probable *P. aeruginosa*.^{46,49} Of note, antibiotics must be chosen based on the identified microorganism once CSF infection is evident; however, the course length may vary owing to a lack of consensus in the literature.²⁰

Table 3 - Management and outcomes for patients with meningitis after cochlear implant.

Author/year	N. of patients had meningitis	Diagnostic method	Management	Outcomes
Cohen & Hoffman ¹⁵ /1991	23	-	-	-
Daspi ⁶ /1991	1	Clinical+ CSF culture	Medical treatment	Full-recovery
Webb et al ¹⁶ /1991	1	-	-	Full-recovery
Page & Eby ¹⁷ /1997	1	-	1st episode medical treatment, 2nd episode Revision surgery + medical treatment	Quadriplegia
Woolley et al ¹⁸ /1998	1	Clinical+ CSF culture	Revision surgery + medical treatment	Full-recovery
Reefhuis et al ¹⁹ /2003	26	Clinical+ CSF culture	3 Revision surgery + medical treatment, 23 medical treatment	1 Died, 3 implant removed
Callanan & Poje ¹⁰ /2004	2	Clinical+ CSF culture	Medical treatment	Full-recovery
Petersen et al ²⁰ /2005	1	Clinical+ CSF culture	1st episode medical treatment, 2nd episode Revision surgery + medical treatment	Language delay with loss of his milestones
Biernath et al ²¹ /2006	12	Clinical+ CSF culture	-	2 died
Ahn et al ²² /2008	1	-	-	Re-implantation was required
Mancini et al ²³ /2008	3	Clinical+ CSF culture	3 Medical treatment, 1 medical treatment+ revision surgery in the second episode	2 Fibrosed coclea require reimplantation, 1 full recovery
Torkos et al ²⁴ /2009	1	Clinical+ CSF culture	1 Revision surgery + medical treatment	Full-recovery
McJunkin & Jeyakumar ²⁶ /2010	1	Clinical+ CSF culture	-	-
Loundon et al ²⁵ /2010	1	Clinical+ CSF culture	Medical treatment	Full-recovery
Ahn et al ²⁷ /2011	2	-	1 Revision surgery + medical treatment, 1 medical treatment	-
Ajallouyeen et al ⁴ /2011	1	-	-	-
Gurbuz et al ⁷ /2011	1	Clinical+ CSF culture	The first 3 episodes were treated with medical treatment and the last episode treated with revision surgery + medical treatment	-
O'Mahony et al ²⁹ /2011	1	Clinical+ CSF culture	medical treatment+ shunt surgery	Right hemiparesis
Lalwani & Cohen ²⁸ /2011	86	-	-	-
Mancini et al ⁸ /2013	5	-	2 Revision surgery + medical treatment	2 ipsilateral ossification, 1 ipsilateral and contralateral ossification
Roman et al ³⁰ /2013	1	Clinical+ CSF culture	1 Revision surgery + medical treatment	Full-recovery
Tarkan et al ³¹ /2013	2	-	1 Revision surgery + medical treatment, 1 medical treatment	Full-recovery
Afsharpaiman et al ³² /2014	1	-	-	-
Suleiman et al ³³ /2014	1	-	Medical treatment	Full-recovery
Daneshi et al ³⁴ /2015	4	-	-	-
Pross et al ³⁶ /2016	1	Clinical+ CSF culture	1 Medical treatment	Full-recovery
Javia et al ³⁵ /2016	4	Clinical+ CSF culture	2 Revision surgery + medical treatment, 2 medical treatment	1 Temporary hemiparesis that resolved, 3 Full-recovery
Tandon et al ³⁷ /2016	1	Clinical+ CSF culture	The first 2 episodes medical treatment, the 3rd episode revision surgery + medical treatment	Full-recovery
Vartanyan et al ³⁸ /2018	1	Clinical+ CSF culture	1 Revision surgery + medical treatment	Full-recovery
Dağkiran et al ⁵ /2020	5	-	4 Revision surgery + medical treatment, 1 medical treatment	-
Yan et al ³⁹ /2020	1	Clinical+ CSF culture	1 Revision surgery + medical treatment	Full-recovery
Piromchai et al ⁴⁰ /2021	9	-	-	-

CSF: cerebrospinal fluid

Study limitations. First is the small sample size of the included studies, with approximately half of them being case studies. However, we searched for relevant reports using four major databases, along with the manual search of relevant reference lists. Second, there is a lack of meta-analyses owing to the limited number of studies and data heterogeneity. Immunization status has not been discussed in some studies. We believe that the findings of our study should be cautiously interpreted, and future studies are recommended to further evaluate the role of vaccination and identify potential risk factors. Public health workers and

should provide proper knowledge and awareness to primary care health practitioners and family members of children with CI regarding the early symptoms of otitis media and meningitis. Thus, antibiotics can be initiated immediately once the symptoms appear.^{41,46} They should be aware of the possible bacterial invasion from the middle ear to the meninges. At all times and taking all precautions, CI candidates and their close members should be aware of the continuous risk of post-implant meningitis before implantation. Future experimental studies are recommended to evaluate the exact role of meningitis vaccination in minimizing the risk of infection or complications. Moreover, data from large multicentre studies are recommended to establish generalized measures.

In conclusion, given the current findings from the systematic literature search, CI users seem to be at possible risk of bacterial meningitis at any time after implantation. Specific factors may increase the risk of infection, such as otitis media, anatomical malformations of the cochlea, and CSF leak. Primary care physicians and family members of CI users should be aware of the presenting features of meningitis and otitis media. There are no different management guidelines for meningitis in CI users; therefore, the key element in curbing the risk is extending CI-specific vaccination campaigns to be followed and recommended. CI candidates should be educated regarding the possible risk of post-implant meningitis, particularly if they have any of the potential risk factors.

Acknowledgment. The authors gratefully acknowledge Editage (www.editage.com) for the English language editing.

References

- Clark G, Cowan RS, Dowell RC. Cochlear implantation for infants and children: Advances: Singular; 1997.
- Young N, Nguyen T, Wiet R. Cochlear implantation. *Oper Tech Otolaryngol Head Neck Surg* 2003; 14: 263-267.
- Kim CS, Chang SO, Oh SH, Lee HJ. Complications in cochlear implantation. *Int Congr Ser* 2004; 145-148.
- Ajallouyeen M, Amirsalari S, Yousefi J, Raeesi M, Radfar S, Hassanalifard M. A report of surgical complications in a series of 262 consecutive pediatric cochlear implantations in Iran. *Iran J Pediatr* 2011; 21: 455-460.
- Dagkiran M, Tarkan O, Surmelioglu O, Ozdemir S, Onan E, Tuncer U, Bayraktar S, Kiroglu M. Management of complications in 1452 pediatric and adult cochlear implantations. *Turk Arch Otorhinolaryngol* 2020; 58: 16-23.
- Daspit CP. Meningitis as a result of a cochlear implant; case report. *Otolaryngol Head Neck Surg* 1991; 105:115-116.
- Gurbuz MK, Incesulu A, Adapinar B, Erdinc M, Kecik C. Recurrent meningitis due to non-implanted ear in cochlear implant patient with bilateral inner ear abnormality: A case report. *Int Adv Otol* 2011; 7: 257-262
- Mancini P, Viccaro M, Dincer H, Covelli E, Attanasio G, Panebianco V, et al. Contralateral implantation in children affected by postimplant meningitis. *Audiol Neurootol* 2013; 18: 214-222.
- Food and Drug Administration. Public Health Web Notification: Risk of bacterial meningitis in children with cochlear implants. In.; 2003. [Updated 2003; Cited 2022 Aug 21]. Available from: <https://www.reliasmedia.com/articles/79973-fda-public-health-web-notification-cochlear-implant-recipients-may-be-at-greater-risk-for-meningitis>
- Callanan V, Poje C. Cochlear implantation and meningitis. *Int J Pediatr Otorhinolaryngol* 2004; 68: 545-550.
- Bluestone CD. Prevention of meningitis: Cochlear implants and inner ear abnormalities. *Arch Otolaryngol Head Neck Surg* 2003; 129: 279-281.
- Wei BPC, Robins-Browne RM, Shepherd RK, Clark GM, O'Leary SJ. Can we prevent cochlear implant recipients from developing pneumococcal meningitis? *Clin Infect Dis* 2008; 46: e1-e7.
- Rose M, Hey C, Kujumdshiev S, Gall V, Schubert R, Zielen S. Immunogenicity of pneumococcal vaccination of patients with cochlear implants. *J Infect Dis* 2004, 190: 551-557.
- Centers for Disease Control and Prevention. Pneumococcal vaccination for cochlear implant recipients. *MMWR* 2002; 51: 931.
- Cohen NL, Hoffman RA. Complications of cochlear implant surgery in adults and children. *Ann Otol Rhinol Laryngol* 1991; 100: 708-711.
- Webb RL, Lehnhardt E, Clark GM, Laszig R, Pyman BC, Franz BK. Surgical complications with the cochlear multiple-channel intracochlear implant: experience at Hannover and Melbourne. *Ann Otol Rhinol Laryngol* 1991; 100: 131-136.
- Page EL, Eby TL. Meningitis after cochlear implantation in Mondini malformation. *Otolaryngol Head Neck Surg* 1997; 116: 104-106.
- Woolley AL, Lusk RP, Jenison V, Bahadori RS, Stroer BS, Wippold 2nd FJ. Cochlear implantation in children with inner ear malformations. *Ann Otol Rhinol Laryngol* 1998; 107: 492-500.
- Reefhuis J, Honein MA, Whitney CG, Chamany S, Mann EA, Biernath KR, et al. Risk of bacterial meningitis in children with cochlear implants. *N Engl J Med* 2003; 349: 435-445.
- Pettersen G, Ovetchkine P, Tapiero B. Group A streptococcal meningitis in a pediatric patient following cochlear implantation: report of the first case and review of the literature. *J Clin Microbiol* 2005; 43: 5816-5818.

21. Biernath KR, Reefhuis J, Whitney CG, Mann EA, Costa P, Eichwald J, et al. Bacterial meningitis among children with cochlear implants beyond 24 months after implantation. *Pediatrics* 2006; 117: 284-289.
22. Ahn JH, Chung JW, Lee KS. Complications following cochlear implantation in patients with anomalous inner ears: experiences in Asan Medical Center. *Acta Oto-Laryngologica* 2008; 128: 38-42.
23. Mancini P, D'Elia C, Bosco E, De Seta E, Panebianco V, Vergari V, et al. Follow-up of cochlear implant use in patients who developed bacterial meningitis following cochlear implantation. *Laryngoscope* 2008; 118: 1467-1471.
24. Torkos A, Czigner J, Jarabin J, Toth F, Szamoskoezi A, Kiss JG, et al. Recurrent bacterial meningitis after cochlear implantation in a patient with a newly described labyrinthine malformation. *Int J Pediatr Otorhinolaryngol* 2009; 73: 163-171.
25. Loundon N, Blanchard M, Roger G, Denoyelle F, Garabedian EN: Medical and surgical complications in pediatric cochlear implantation. *Arch Otolaryngol Head Neck Surg* 2010; 136: 12-15.
26. McJunkin J, Jeyakumar A. Complications in pediatric cochlear implants. *Am J Otolaryngol* 2010; 31: 110-113.
27. Ahn JH, Lim HW, Lee KS. Hearing improvement after cochlear implantation in common cavity malformed cochleae: long-term follow-up results. *Acta Otolaryngol* 2011; 131: 908-913.
28. Lalwani AK, Cohen NL. Longitudinal risk of meningitis after cochlear implantation associated with the use of the positioner. *Otol Neurotol* 2011; 32: 1082-1085.
29. O'Mahony LN, Klein EJ, Walker W. A fully immunized child with a cochlear implant and Streptococcus pneumoniae meningitis 3 years after implantation. *Pediatr Emerg Care* 2011; 27: 200-202.
30. Roman BR, Coelho DH, Roland Jr JT. Implantation of the common cavity malformation may prevent meningitis. *Cochlear Implants Int* 2013; 14: 56-60.
31. Tarkan O, Tuncer U, Ozdemir S, Surmelioglu O, Cetik F, Kiroglu M, Kayikcioglu E, Kara K. Surgical and medical management for complications in 475 consecutive pediatric cochlear implantations. *Int J Pediatr Otorhinolaryngol* 2013; 77: 473-479.
32. Afsharpaiman S, Amirsalari S, Ajalloueyan M, Saburi A. Bacterial meningitis after cochlear implantation among children without polyvalent conjugate vaccine: A brief report of an Iranian cohort study on 371 cases. *Int J Prev Med* 2014; 5: 1067-1070.
33. Suleiman AO, Suleiman BM, Abdulmajid UF, Suleiman MR, Mustapha AY, Afolabi OA, Yet al. Paediatric cochlear implantation in north-western Nigeria case report and review of our challenges. *Int J Pediatr Otorhinolaryngol* 2014; 78: 363-365.
34. Daneshi A, Ajalloueyan M, Ghasemi MM, Hashemi BS, Emamjome H, Farhadi M, et al. Complications in a series of 4400 paediatric cochlear implantation. *Int J Pediatr Otorhinolaryngol* 2015; 79: 1401-1403.
35. Javia L, Brant J, Guidi J, Rameau A, Pross S, Cohn S, et al. Infectious complications and ventilation tubes in pediatric cochlear implant recipients. *Laryngoscope* 2016; 126: 1671-1676.
36. Pross SE, Cardenas RU, Ahn ES, Stewart CM. Recurrent meningitis in a child with bilateral cochlear implantation associated with a petrous apex encephalocele: a case report and literature review. *Acta Oto-Laryngol Case Rep* 2016; 1: 24-29.
37. Tandon S, Singh S, Sharma S, Lahiri AK. Use of intrathecal fluorescein in recurrent meningitis after cochlear implantation. *Iran J Otorhinolaryngol* 2016; 28: 221-226.
38. Vartanyan M, Hill F, Orimoto K, O'Leary S. Stapes footplate defect as a source of CSF leak and otogenic meningitis in a patient with a cochlear implant. *Ear Nose Throat J* 2018; 97: E31-E32.
39. Yan Z, Hao P, Wen X, Liu J, Dai J, Cao K, et al. Pseudomonas aeruginosa meningitis following head injury after cochlear implantation: A case report. *Eur J Inflamm* 2020; 18.
40. Pirochchai P, Tanamai N, Kiatthanabumrung S, Kaewsiri S, Thongyai K, Atchariyasathian V, et al. Multicentre cohort study of cochlear implantation outcomes in Thailand. *BMJ open* 2021; 11: e054041.
41. Risk of Bacterial Meningitis in Children with Cochlear Implants, CDC and FDA Recommendations. [Updated 2002, Cited 2022 Aug 21]. Available from: <https://www.cdc.gov/ncbddd/hearingloss/meningitis.html>
42. Black SB, Shinefield HR, Fireman B, Hiatt R, Polen M, Vittinghoff E. Efficacy in infancy of oligosaccharide conjugate Haemophilus influenzae type b (HbOC) vaccine in a United States population of 61,080 children. The Northern California Kaiser Permanente Vaccine Study Center Pediatrics Group. *Pediatr Infect Dis J* 1991; 10: 97-104.
43. Peltola H, Kilpi T, Anttila M: Rapid disappearance of Haemophilus influenzae type B meningitis after routine childhood immunisation with conjugate vaccines. *Lancet* 1992; 340: 592-594.
44. Bluestone CD. Cochlear implants and meningitis: Update and recommendations for prevention. *J Pediatr Infect Dis* 2003; 22: 477-478.
45. Whitney CG. Cochlear implants and meningitis in children. *J Pediatr Infect Dis* 2004; 23: 767-768.
46. Arnold W, Bredberg G, Gstottner W, Helms J, Hildmann H, Kiratzidis T, et al. Meningitis following cochlear implantation: Pathomechanisms, clinical symptoms, conservative and surgical treatments. *ORL J Otorhinolaryngol Relat Spec* 2002; 64: 382-389.
47. DeSaSouza, Sandra. A Historical Review of the Development of Cochlear Implants. *Cochlear Implants* 2022; 1-7.
48. Angeli S, Balkany T. Post-cochlear implant meningitis. *Oper Tech Otolaryngol Head Neck Surg* 2003; 14: 293-296.
49. Alper CM, Bluestone CD, Casselbrant ML, Dohar JE, Mandel EM. Advanced Therapy of Otitis Media. BC Decker Inc; Hamilton (ON): 2004. pg. 32-37