

CAS CLINIQUE / CASE REPORT

ACINETOBACTER BAUMANNII MENINGITIS

A Case Report

<http://www.lebanesemedicaljournal.org/articles/66-5/case4.pdf>

Mohamed EL HUSSEINI¹, Adnan MRAD², Taghid CHAABAN²

El Husseini M, Mrad A, Chaaban T. *Acinetobacter baumannii* meningitis: A case report. J Med Liban 2018; 66 (5): 288-291.

ABSTRACT • Background : *Acinetobacter baumannii* infection is an important cause of nosocomial infection in critically ill patients, but still rare in neurosurgical patients. The treatment of multidrug-resistant *A. baumannii* meningitis is a serious therapeutic problem due to the limited penetration of antibiotics into the cerebrospinal fluid (CSF). **Case study :** We present one case of acquired *A. baumannii* meningitis after neurosurgical operation. The 44-year-old patient presented with cranial trauma and five days from operation developed meningitis with *A. baumannii*. Despite the treatment, he died at one month from operation. **Conclusion :** *A. baumannii* has become an emerging threat for post-neurosurgical patients. This nosocomial infection has a high rate of mortality. We need to increase the awareness of *A. baumannii* meningitis among neurosurgeons.

Keywords : *Acinetobacter baumannii*; multidrug resistant; meningitis

El Husseini M, Mrad A, Chaaban T. Étude d'un cas de méningite *Acinetobacter baumannii*. J Med Liban 2018; 66 (5): 288-291.

RÉSUMÉ • Contexte : L'infection à l'*Acinetobacter baumannii* est une cause importante d'infection nosocomiale chez les patients gravement malades, mais encore rare chez les patients en neurochirurgie. Le traitement de la méningite *A. baumannii* multirésistante est un grave problème thérapeutique en raison de la pénétration limitée des antibiotiques dans le LCR. **Étude de cas :** Nous rapportons un cas de méningite *A. baumannii* acquise après une opération neurochirurgicale. Le patient âgé de 44 ans avait un traumatisme crânien et développa une méningite *A. baumannii* 5 jours après l'opération. Malgré le traitement, il est décédé un mois après l'intervention chirurgicale. **Conclusion :** L'*A. baumannii* est devenu une grave complication émergente pour les patients en neurochirurgie. Cette infection nosocomiale a un taux élevé de mortalité infantile. Sensibiliser les neurochirurgiens à la gravité de la méningite *A. baumannii* devient une priorité.

Mots-Clés : *Acinetobacter baumannii*; polypharmacorésistance; méningite

INTRODUCTION

Acinetobacter baumannii causes various serious nosocomial infections worldwide. Bacterial meningitis is a common complication after neurosurgical operation, and the percentage of *A. baumannii* meningitis is growing, especially the one resisting multiple drugs [1]. The neurosurgical patients have a high risk to suffer from bacterial meningitis caused by *A. baumannii* with potentially fatal consequences. The meningitis caused by *A. baumannii* is well recognized and has been described worldwide. Most case reports about the meningitis were associated with external ventricular drainage (EVD), cerebrospinal fluid (CSF) leaking, or head trauma [2-5]. This post-surgical meningitis is especially severe, because the selection of the antibiotic depends not only on the susceptibility of *A. baumannii*, but also on the penetration of the chosen antibiotic through the blood-brain barrier [6-7]. We report here the case of a male patient that acquired meningitis with *A. baumannii* following neurosurgical operation.

¹Department of Neurosurgery, Dar Al Amal Hospital, Lebanese University, Lebanon.

²Islamic University of Lebanon-IUL, Lebanon.

*Corresponding author: *Mohamed El Husseini, MD.*
e-mail: drhusseiny@yahoo.com

CASE PRESENTATION

We report the case of a 44-year-old male patient who presented to our department with a cranial trauma.

His physical exam revealed: coma (Glasgow score 9); afebrile, right hemiplegia, left frontal cranial-cerebral lesion.

Further inquiries included: leukocytosis: 30000/mm³ with N 90%; anemia: Hb 10 g/dl; ESR: 120 mm/h; hypoproteinemia: total protein 5.8 g/dl. CT scan indicated areas with hypodensity interesting frontal lobe and signs of left ventriculitis (Figure 1).

Five days following the neurosurgical intervention the patient presented fever, altered state, confusion and neck stiffness.

We then performed a lumbar puncture; the examination of CSF revealed a purulent liquid with PN 95%. The direct exam showed diplococci and Gram-negative coccobacilli. From CSF culture it was identified as *A. baumannii*.

The antibiogram revealed that *A. baumannii* was resistant to: amoxicilline + clavulanic acid, cefoxitine, ceftriaxone, cefuroxime, amikacin, ticarcilline, ciprofloxacine, meropenem and tienam.

The patient was given pathogenic therapy, vancomycin and colistin. He remained febrile without improving



Figure 1. CT scan showing hypodensity in frontal and left periventricular areas

his neuropsychological status. We also noticed the dehiscent cranial wound. *Repeated lumbar cultures after 3 and 7 days respectively, showed the persistence of A. baumannii*, which indicated the failure of the treatment.

Unfortunately, the patient died on day 32 from hospital admission.

In this case, some factors could have led to acquired *A. baumannii* meningitis, such as: admission to ICU, the neurosurgical intervention, the presence of venous and urinary catheters. Unfortunately, we were not able to test the sensibility of this bacterium to colistin, which proved to be effective with multidrug-resistant *A. baumannii*.

DISCUSSION

Acinetobacter baumannii is a polymorphic Gram-negative coccobacillus and tolerates long periods in both moist and dry conditions, and the easy acquisition of drug resistance makes it a great challenge in disease control.

The epidemiological profile suggests that it is of low virulence and the disease is dependent on significant host immunological impairment. It spreads through contact with contaminated medium (soil, meat and water), open wounds, skin, contact with patient, ventilators, sinks, bed rails, humidifiers.

Risk factors are represented by hospitalized patients and people with chronic lung disease, diabetes and weakened immune system. It seems plausible that this ubiquitous organism has acquired a vast array of pathogenesis islands to deal with this diversity [8]. Therefore, management of infections due to *A. baumannii* has be-

come a real public health issue in many countries.

Virulence factors that influence the pathogenesis of *A. baumannii* are its surface motility on solid/semisolid media and the ability to form biofilm on abiotic or biotic surfaces. They also include the outer membrane protein A of *Acinetobacter baumannii* (OmpA), phospholipases, membrane polysaccharide components, penicillin-binding proteins (PB 7/80) and β -lactamase PER-1, metal acquisition system and outer membrane vesicles.

Intracranial infections including ventriculitis and meningitis caused by *A. baumannii* in the neurosurgery setting have led to challenging situations. The percentage of intracranial infection caused by *A. baumannii* in postoperative infection continuously increased in recent years. Regarding the frequency of meningitis with *A. baumannii*, in a review of four studies including 281 adult patients with hospital-acquired meningitis, 3.6% had meningitis due to *Acinetobacter* spp. [8-11]. One study on nosocomial meningitis in children reported that *Acinetobacter* accounted for 11.2% (20/178) of cases [12]. In USA and Taiwan, *Acinetobacter* ranked fifth place in nosocomial meningitis [8,10], while in Turkey it was the leading cause of Gram-negative post-neurosurgical meningitis [13,14].

Risk factors for post-neurosurgical meningitis are craniotomy, cerebrospinal leakage, incision infection, prolonged duration of surgery, surgery of sinus, external and internal ventricular drain, lumbar puncture, head trauma [15]. The median time to develop *Acinetobacter* meningitis after a neurosurgical procedure is 12 days (range 1-40 days) [16].

Meningitis which develops within 3 months after neurosurgical operation is defined as "post-neurosurgical meningitis" [17].

Post-neurosurgical *A. baumannii* meningitis was diagnosed after meeting the following criteria:

- 1) CSF culture: presence of *A. baumannii*;
- 2) CSF modifications: WBC increased, protein elevating and glucose decreasing;
- 3) The patient had fever ($\geq 38^{\circ}\text{C}$), headache, vomiting, confusion, irritability or meningeal irritation;
- 4) The patient had a neurosurgical operation within 3 months [18].

Regarding treatment of *A. baumannii* meningitis, carbapenems used to be the empirical drugs for choice. However, more than 30% of *A. baumannii* strains were resistant to at least three kinds of antibiotics in many general hospitals [19]. Its resistance in the Middle East in general, and specifically in Lebanon, is increasing and many strains were found to be carbapenem resistant. In a study including hospitals from different Middle Eastern countries, it was found that *A. baumannii* isolated from a Lebanese hospital carried the oxa-58, oxa-23 and oxa-72

genes conferring the carbapenems resistance [20]. Another study, conducted in Hôtel-Dieu University Hospital in Beirut, showed an alarming increase in imipenem resistance among *A. baumannii* isolates from 7.7% in 2006 to 35.4% in 2009. In Lebanon, resistance was given by predominance of OXA-23, OXA-24, OXA-58, and OXA-143 and by spread of NDM-1 and GES-11 [21-28].

Colistin, an old antibiotic, was introduced in clinical use from 1950s, and abrogated in 1980s due to serious renal toxicity and neurovirulence. Maartens *et al.* compared colistin with carbapenems and tobramycin, and found that colistin was still effective for *A. baumannii* in resistance of other antibiotics, and no difference in renal toxicity was revealed among these antibiotics [29]. Rolain *et al.* indicated that colistin worked through modifying the negative charges of outer membrane in Gram-negative bacteria [30].

This synergistic combination was studied also in Lebanon, at Saint George Hospital, Beirut, where it showed that combining colistin with carbapenems had high rates of additive effects [31]. The reason for the increased synergy with meropenem than imipenem might be that most carbapenemases target with greater affinity imipenem as compared to meropenem. The synergistic or additive effect might be influenced by the ability of colistin to disrupt the bacterial outer membrane and increase its permeability for carbapenems [25-36] and therefore stop the cross-linking of the new synthesized polymers. Combination therapy delays the emergence of bacterial resistance and specifically the rapidly developing resistance and heteroresistance toward colistin.

CONCLUSION

We recommend an increased awareness of the significant role of the multidrug-resistant bacteria *Acinetobacter baumannii* in nosocomial infections in the field of neurosurgery. It represents a serious challenge for epidemiologists, infection control professionals, and clinicians.

REFERENCES

- Munoz-Price LS, Weinstein RA. *Acinetobacter* infection. N Engl J Med 2008; 358: 1271-81.
- Karaiskos I, Galani L, Baziaka F et al. Successful treatment of extensively drug-resistant *Acinetobacter baumannii* ventriculitis and meningitis with intraventricular colistin after application of a loading dose: a case series. Int J Antimicrob Agents 2013; 41: 480-3.
- Jimenez-Mejias ME, Pachon J, Becerril B, Palomino-Nicas J, Rodriguez-Cobacho A, Revuelta M. Treatment of multidrug-resistant *Acinetobacter baumannii* meningitis with ampicillin/sulbactam. Clin Infect Dis 1996; 24: 932-5.
- Bergogne-Berezin E, Towner KJ. *Acinetobacter* spp. as nosocomial pathogens: microbiological, clinical, and epidemiological features. Clin Microbiol Rev 1996; 9: 148-65.
- Siegmán-Igra Y, Bar Yosef S, Gorea A et al. Nosocomial *Acinetobacter* meningitis secondary to invasive procedures: report of 25 cases and review. Clin Infect Dis 1993; 17: 843-9.
- Giamarellou H, Antoniadou A, Kanellakopoulou K. *Acinetobacter baumannii*: a universal threat to public health. Int J Antimicrob Agents 2008; 32: 106-19.
- Kim HI, Kim SW, Park GY et al. The causes and treatment outcomes of 91 patients with adult nosocomial meningitis. Korean J Intern Med 2012; 27: 171-9.
- Durand ML, Calderwood SB, Weber DJ et al. Acute bacterial meningitis in adults. A review of 493 episodes. N Engl J Med 1993; 328: 21-8.
- Hussein AS, Shafran SD. Acute bacterial meningitis in adults. A 12-year review. Medicine (Baltimore) 2000; 79: 360-8.
- Lu CH, Chang WN, Chang HW. Adult bacterial meningitis in southern Taiwan: epidemiologic trend and prognostic factors. J Neurol Sci 2000; 182: 36-44.
- Weisfelt M, van de Beek D, Spanjaard L, de Gans J. Nosocomial bacterial meningitis in adults: a prospective series of 50 cases. J Hosp Infect 2007; 66: 71-8.
- Rudinsky B, Stankovic I, Kacerova A et al. Nosocomial postsurgical meningitis in children: a 12-year survey comparing data from 1993-1998 with data from 1999-2004. Infect Control Hosp Epidemiol 2006; 27: 788-90.
- Sacar S, Turgut H, Toprak S et al. A retrospective study of central nervous system shunt infections diagnosed in a university hospital during a 4-year period. BMC Infect Dis 2006; 6: 43.
- Metan G, Alp E, Aygen B, Sumerkan B. Carbapenem-resistant *Acinetobacter baumannii*: an emerging threat for patients with post-neurosurgical meningitis. Int J Antimicrob Agents 2007; 29: 112-13.
- van de Beek D, Drake JM, Tunkel AR. Nosocomial bacterial meningitis. N Engl J Med 2010; 362: 146-54.
- Siegmán-Igra Y, Bar-Yosef S, Gorea A, Avram J. Nosocomial *Acinetobacter* meningitis secondary to invasive procedures: report of 25 cases and review. Clin Infect Dis 1993; 17: 843-9.
- Lowman W, Kalk T, Menezes CN, John MA, Grobusch MP. A case of community-acquired *Acinetobacter baumannii* meningitis – has the threat moved beyond the hospital? Journal of Medical Microbiology 2008; 57: 676-8.
- Chang CJ, Ye JJ, Yang CC, Huang PY, Chiang PC, Lee MH. Influence of third-generation cephalosporin resistance on adult in-hospital mortality from post-neurosurgical bacterial meningitis. J Microbiol Immunol Infect 2010; 43: 301-9.
- Khan FY, Abukhattab M, Baager K. Nosocomial post-neurosurgical *Acinetobacter baumannii* meningitis: a retrospective study of six cases admitted to Hamad General Hospital, Qatar. J Hosp Infect 2012; 80: 176-9.
- Kim BN, Peleg AY, Lodise TP et al. Management of meningitis due to antibiotic-resistant *Acinetobacter* species. Lancet Infect Dis 2009; 9: 245-55.
- Araj GF, Avedissian AZ, Ayyash NS et al. A reflection on bacterial resistance to antimicrobial agents at a major tertiary care center in Lebanon over a decade, The Lebanese Medical Journal 2012; 60 (3): 125-35.
- Zarrilli R, Vitale D, Di Popolo A et al. A plasmid-borne

- blaOXA-58 gene confers imipenem resistance to *Acinetobacter baumannii* isolates from a Lebanese hospital. *Antimicrob Agents Chemother* 2008; 52 (11): 4115-20.
23. Rafei R, Dabboussi F, Hamze M et al. First report of blaNDM-1-producing *Acinetobacter baumannii* isolated in Lebanon from civilians wounded during the Syrian war. *Int J Infect Dis* 2014 Apr; 21: 21-3.
 24. Rafei R, Pailhoriès H, Hamze M et al. Molecular epidemiology of *Acinetobacter baumannii* in different hospitals in Tripoli, Lebanon, using bla OXA-51-like sequence based typing. *BMC Microbiology* 2015; 15: 11-17.
 25. Hammoudi D, Moubareck CA, Hakime N et al. Spread of imipenem-resistant *Acinetobacter baumannii* co-expressing OXA-23 and GES-11 carbapenemases in Lebanon. *Int J Infect Dis* 2015 Jul; 36: 56-61.
 26. Al Atrouni A, Hamze M, Jisr T et al. Wide spread of OXA-23-producing carbapenem-resistant *Acinetobacter baumannii* belonging to clonal complex II in different hospitals in Lebanon. *Int J Infect Dis* 2016 Nov; 52: 29-36.
 27. Dahdouh E, Hajjar M, Suarez M, Daoud Z. *Acinetobacter baumannii* isolated from Lebanese patients: Phenotypes and genotypes of resistance, clonality, and determinants of pathogenicity. *Front Cell Infect Microbiol* 2016; 6: 163.
 28. Soudeiha MAH, Dahdouh EA, Azar E, Sarkis DK, Daoud Z. In vitro evaluation of the colistin-carbapenem combination in clinical isolates of *A. baumannii* using the checkerboard, Etest, and Time-Kill Curve Techniques. *Front Cell Infect Microbiol* 2017; 7: 209.
 29. Gounden R, Bamford C, van Zyl-Smit R, Cohen K, Maartens G. Safety and effectiveness of colistin compared with tobramycin for multidrug resistant *Acinetobacter baumannii* infections. *BMC Infect Dis* 2009; 9: 26.
 30. Rolain JM, Roch A, Castanier M, Papazian L, Raoult D. *Acinetobacter baumannii* resistant to colistin with impaired virulence: a case report from France. *J Infect Dis* 2011; 204: 1146-7.
 31. Santella G, Pollini S, Docquier JD et al. Carbapenem resistance in *Pseudomonas aeruginosa* isolates: An example of interaction between different Mechanisms. *Rev Panam Salud Publica* 2011; 30 (6): 545-8.
 32. Oliver A, Levin BR, Juan C, Baquero F, Blazquez J. Hypermutation and the preexistence of antibiotic-resistant *Pseudomonas aeruginosa* mutants: Implications for susceptibility testing and treatment of chronic infections. *Antimicrobial Agents and Chemotherapy* 2004; 48 (11): 4226-33.
 33. Walsh TR, Toleman MA, Poirel L, Nordmann P. Metallo beta-lactamases: The clam before the storm? *Clinical Microbiology Reviews* 2005; 18 (2): 306-25.
 34. Peleg AY, Seifert H, Paterson DL. *Acinetobacter baumannii*: Emergence of a successful pathogen. *Clinical Microbiology Reviews* 2008; 21 (3): 538-82.
 35. Zhang JP, Zhu W, Tian SF, Chu YZ, Chen BY. Molecular characteristics and resistant mechanisms of imipenem-resistant *Acinetobacter baumannii* isolates in Shenyang, China. *Journal of Microbiology* 2010; 48 (5): 689-94.
 36. Falagas ME, Karageorgopoulos DE. Pandrug resistance (PDR), extensive drug resistance (XDR), and multidrug resistance (MDR) among Gram-negative bacilli: Need for international harmonization in terminology. *Clinical Infectious Diseases: An Official Publication of the Infectious Diseases Society of America*, 2008; 46 (7): 1121-2.