The prevalence and antimicrobial sensitivity of *Esbl Escherichia Coli.* in clinical isolates

Zaid I. Al-Attar .M.B.Ch.B. M.S.c. (pharmacology)*

ABSTRACT

Background: The antimicrobial resistance is one of the most serious and expanding health problems world -wide in the last decades. The esbl escherichia *coli. (extended - spectrum beta-lactamase e.coli)* represents an important aspect of it .

Objectives: To get an overview on the esbl *e.coli* prevalence profile in general. Also to assess the antibiotic sensitivity of esbl e. coli trying to specify the most effective antibiotics in combating this micro-organism.

Methods: this study tries to focus on this problem in Iraq which through a prospective study approach by taking 35 clinical samples from various sources (urine, blood, abscess, eye, vagina, stool and others), and after confirming the presence of *e.coli*, the presence of esbl *e.coli* and antibiotic sensitivity are confirmed by the use of Kirby - bauer method.

Results: results showed that esbl *e.coli* constitutes 80% of the cases, while the results of antibiotic sensitivity were as follows: ampicillin 3.3%, ampicillin/sulbactam 20%, amoxi/clav 0%pipracillin/tazobactam 89.7% meropenem 96.7%, imipenem 96.9%, cefotaxime 0%, ceftriaxone 11.8%,ceftazidime 16.1%,cefipime 14.3%, cefazolin 16.1% cefoxitin 64.7%, aztreonam 14.3%,gentamycin 50%

xtended - spectrum B-lactamase producing (esbl) clinical isolates among members of the entero bacteriaceae family, especially klebsiella pneumoniae and escherichia coli, represent one of the most important world problems of b-lactam antimicrobial resistance.¹betalactamases are enzymes produced by some bacteria and are responsible for their resistance to beta-lactam antibiotics like penicillins, cephamycins, and carbapenems (ertapenem). (cephalosporins are relatively resistant to beta-lactamase.) these antibiotics have a common element in their molecular structure: a four-atom ring known as a beta-lactam. the lactamase enzyme breaks that ring open, deactivating the molecule's antibacterial properties.² in the mid-1980s, a new group of enzymes, the extendedspectrum b-lactamase (esbls), was detected (first detected in Germany in 1983)³

The esbls are frequently plasmid encoded. plasmids are responsible for esbl production frequently carry genes encoding resistance to other drug classes (for example, aminoglycosides). therefore, antibiotic options in the treatment of esbl-producing organisms are extremely limited. carbapenems are the treatment of choice for serious infections due to esbl-producing organisms, vet been carbapenem-resistant isolates have recently reported⁴.

The correct identification of esbl-producing bacteria has important clinical-epidemiological and laboratory implications. First, patients may experience a delay in appropriate treatment if esbl-producing bacteria are not correctly detected by routine antimicrobial susceptibility tests.⁵ Second, while carbapenems are the most effective therapy for esbl bacterial infections, their routine use can select resistant strains, as the emergence of imipenem,tobramycin 64.3%, amikacin 94.3%,ciprofloxacin 58.8% ,levofloxacin,64.5%nitrofurantoin,79.2%,trimethprimesulpha methoxazole 29.6%.

Conclusion: the problem of esbl *e.coli* is expanding and there is a continuous demand for frequent monitoring of the new trends on antimicrobial resistance in different parts of the world in addition to trying to develop new antimicrobials to combat the new highly resistant strains .moreover there is a continuous need to educate the medical and the paramedical staff abot the risk of unjustified and improper prescription and use of antimicrobials.

Key words: *escherichia coli*, extended-spectrum betalactamase, kirby-bauer method, muller hinton agar

Al-Kindy College Medical Journal 2014: Vol. 10 No. 2 Pages: 96-99

*Assistant lecturer, department of pharmacology, Al-Kindy college of medicine, Baghdad University. Received 10th Nov 2014, accepted in final 20th April 2015 Corresponding to Dr Zaid Ihsan AL-Attar email:

zaidattar77@gmail.com.

resistant *acinetobacterbaumanii*, *pseudomonas aeruginosa* and *k. pneumoniae*.^{6,7}. Third, esbl genes are located on large plasmids that can harbor genes for resistance to other non-b-lactams antibiotics, and therefore, esbl-producing bacteria often exhibit multidrug-resistant phenotypes, reducing the drug arsenal even further⁸. Fourth, genes encoding esbls are typically located in conjugative plasmids or integron-like structures and can be effectively transferred to other strains and species⁹.finally, esbl-producing organisms, especially *k. pneumoniae*, but also *e.coli*, have been responsible for serious nosocomial infection outbreaks that lead to prolonged hospital stay, increased morbidity and mortality, and consequently increase healthcare associated costs¹⁰.

The objectives of this study were to get an overview on the esbl e.coli prevalence profile in general. Also to assess the antibiotic sensitivity of esbl e. coli trying to specify the most effective antibiotics in combating this micro-organism. Methods. a prospective study in which e.coli isolates are taken from various clinical samples from patients in alkhadimiya teaching hospital. These isolates are confirmed as being *e.coli* by microscopy which shows gram-negative rods, with no particular cell arrangement. Then, by macconkey agar is inoculated. on macconkey agar, deep red colonies are produced, as the organism is lactosepositive, and fermentation of this sugar will cause the medium's ph to drop, leading to darkening of the medium¹¹ if the isolates shows to be positive for e.coli then the antibiotic susceptibility and the presence of esbl e.coli is assessed using the disk diffusion susceptibility testing (kirby-bauer method) by inoculation of isolates into muller hinton media and applying the antibiotic disks: the antibiotics disks used are listed ampicillin, amoxiclav,

ampicillin/sulbactam, meropenem, imipenem, piracillin/tazobactam, aztreonam, cefazolin, cefotriaxone, ceftazidime, cefotaxime, cefoxitin, cefepime, amikacin, gentamycin, tobramycin, ciprofloxacin, levofloxacin, nitrofurantoin, and trimethoprime /sulpham ethoxazole.

After incubation at 35c for 24 h, zone of inhibition size is measured¹². Then these results (zones of inhibition) were interpreted according to the standards proposed by (performance standards for anti microbial susceptibility testing; twenty-first informational supplement 2011)¹³.

The prevalence of esbl *e.coli* and the antibiotic sensitivity are demonstrated by using percentages to show which antibiotics are better in combating the esbl *e.coli*.

Results. Various clinical samples (total number =35) were collected in AL-Khadimiya teaching hospital and after implementing the above methods the results were as follows: number of males =14, number of females =21, male to female ratio=0.66

Discussion.the prevalence of esbl *e.coli* is higher among females in this study and this can be due to that most of the isolates of the *e.coli* were obtained from urine samples (57%) and the most abundant micro-organism in utiin general is *e.coli*¹⁴, which when added to vaginal swab samples, both together represent 65% of the cases, in addition to the fact that uti is more abundant among females¹⁵.

The prevalence of esbl*e.coli* is very high in these samples (80%) which is similar to a study done in Indiain 2011 (80.64%)¹⁶. in another study done in Spainin 2011 the prevalenc of esbl*e.coli* was 70%¹⁷. this variation can be attributed to the different method used in that study which comprised the use of pcr-based replicon-typing scheme. by comparing the results of the present study with those done several years ago in different countries around the globe, we can notice the great increase in the prevalence of esbl*e.coli* . e.g. in a study done in latin America (smart) in 2003 the prevalence rate was 10%¹⁸, in 2004¹⁹ also 10% and in 2008 was 26%²⁰.

In addition to these differences with respect to time , there are differences that are related to geographic locations. In a study done in different parts of the world showed that in south America18.1% of *e.coli*were esbl positive, while only 7.5% of isolates from north America were esbl positive²¹.



Figure 1: Sources of clinical isolates.



Figure 2: Prevalence of ESB+VE E.COLI



Figure 3: The antimicrobial sensitivity of ESBL+VE.E COLI

Regarding the antimicrobial sensitivity of *e.coli* isolates in my study they were as follows:

- 1. ampicillin 3.3% , in comparison to a study done in china (2011) the sensitivity was 17.7% $^{\rm 22}$
- 2. ampicillin/sulbactam 20% ,compared to a study done in (2008) latin America the sensitivity was 2.2% 23
- 3. amoxi/clav 0% , in a study done in Świtzerland(2011) the sensitivity was 30.4%²⁴.
- pipracillin/tazobactam 89.7% ,compared to a study done in Sweden (2011) the sensitivity was 91%²⁵,compared to a study done in Italy (2010) the sensitivity was 63.2%²⁶
- 5. meropenem 96.7% ,compared to a study done in china (2011) the sensitivity was 100%22,compared to a study done in Italy (2010) the sensitivity was 100%26
- imipenem 96.9% ,compared to a study done in china (2011) the sensitivity was 100%22,also confirmed by a study done in Pakistan (2011) which showed 100% sensitivity²⁷.
- 7. cefotaxime 0% ,compared to a study done in Sweden (2011) the sensitivity was 1%25while a study done in Pakistan (2011) the sensitivity was 11%27
- 8. ceftriaxone 11.8%, compared to a study done in china (2011) the sensitivity was 3.2 %22
- 9. ceftazidime 16.1%, compared to a study done in Sweden (2011) the sensitivity was 9%25
- 10. cefipime 14.3% ,compared to a study done in Spain(2011) the sensitivity was 14.7% ²⁸ while a study done in Pakistan (2011) the sensitivity was 13%27
- cefazolin 16.1%%,compared to a study done in china (2011) the sensitivity was 1.6 % 22,compared to a study done in canada (2008) the sensitivity was 79.9%
- 12. cefoxitin 64.7%, while a study done in Pakistan (2011) the sensitivity was 60%27, compared to a study done in turkey (2008) the sensitivity was $100\%^{30}$
- aztreonam 14.3%, compared to a study done in china (2011) the sensitivity was 31.2 %22, compared to a study done in romania (2010) the sensitivity was 3.6%
- 14. gentamycin 50% ,compared to a study done in china (2011) the sensitivity was 49 %22
- tobramycin 64.3% compared to a study done in south (2008) the sensitivity was 49 %³²
- 16. amikacin 94.3% ,compared to a study done in Spain(2011) the sensitivity was 75.9%28,while compared to a study done in brazil (2011) the sensitivity was 81%³³
- 17. ciprofloxacin 58.8% ,compared to a study done in china (2011) the sensitivity was 47.1 %22,compared to a study done in Spain(2011) the sensitivity was 75.5%28.
- 18. evofloxacin64.5% ,compared to a study done in brazil (2011) the sensitivity was 50.8%34,while compared to a study done in Taiwan (2009) the sensitivity was $64\%^{34}$
- nitrofurantoin 79.2% ,compared to a study done in Sweden (2011) the sensitivity was 93%25in a study done in Switzerland (2011) the sensitivity was 85%%24
- 20. trimethprime-sulphamethoxazole 29.6%,compared to a study done in Sweden (2011) the sensitivity was 30%25,compared to a study done in Spain(2011) the sensitivity was 30.1%28 the variations mentioned above in results can be attributed to differences in geographic location and time and sometimes to different methods.

The results above show that the most effective antibiotics in treating esbl *e.coli* are imipenem , meropenem , piperacillin/tazobactam, and amikacin.

In conclusion, the problem of esbl *e.coli* is expanding and needs to be followed in terms of continuous monitoring of the new trends on antimicrobial resistance in addition to trying to develop new antimicrobials to combat the new highly resistant strains .moreover there is a continuous demand to educate the medical and the paramedical staff about the risk of unjustified and improper prescription and use of antimicrobials.

References:

- 1. *WHO*. Chronic suppurative otitis media .Geneva. Switzerland;2004:9
- 2. Roger F.Synopsis of Operative ENT Surgery. *Butterworth*, 1992:112
- Alan E Dugdale .Management of chronic suppurative otitis media.*MJA* 2004; 180 (2): 91-93
- Goycoolea MV, Hueb MM, Ruah C. Definitions and terminology. *Otolaryngol Clin North America.* 1991; 24 (4): 757-761.
- Érwin L. van der Veen, Anne G. M., et al. Predictors of Chronic Suppurative Otitis Media in Children.ArchOtolaryngol *Head Neck Surg.* 2006;132:1115-1118.
- Gibson PG, Stuart JE, WlodarczykJ,etal.Nasal inflammation and chronic ear disease in Australian Aboriginal children. *J Paediatr Child Health*.1996; 32: 143-147
- Jahn AF. Chronic otitis media: diagnosis and treatment. *Med Clin North America*. 1991;75 (6): 1277-1291.
- Wintermeyer SM, Nahata M. Chronic suppurative otitis media. *Annals Pharmacother*.1994; 28: 1089-1099.
- Adair-Bischoff CE, Sauve RS.Environmental tobacco smoke in middle ear disease in preschoolage children. *Arch Pediatr Adolesc Med.* 1998; 152 (2): 127-133.
- Fliss DM, Shoham I, Leiberman A, Dagan R. Chronic suppurative otitis media withoutcholesteatoma in children in Southern Israel: incidence and risk factors. *Pediatr Infect Dis J.*1991; 10: 895-899.
- Homoe P. Otitis media in Greenland.Studies on historical, epidemiological, microbiological and immunological aspects. *Int J Circumpolar Health.* 2001; 60 (Suppl 2): 1-54.
- Kenna MA. Treatment of chronic suppurative otitis media. *Otolaryngol Clin North Am*.1994; 27 (3): 457-472.
- Wiet RJ. Patterns of ear disease in the Southwestern American Indian. *Arch Otolaryngol.* 1979; 105: 381-385.
- Sunderman J, Dyer H. Chronic ear disease in Australian aborigines. *Med J Aust.* 1984;140: 708-711
- 15. Maynard JE, Fleshman JK, Tschopp CF. Otitis media in Alaskan Eskimo children. *JAMA*. 1972; 219: 597-599.
- Berman S. Otitis media in children. *N Eng J Med.* 1995; 332 (23): 1560-1565
- World Health Organization.Prevention of hearing impairment from chronic otitismedia. *Report of a WHO/CIBA* Foundation Workshop. Geneva. 1998.
- Wiet R, DeBlanc G et al. Natural history of otitis media in the American native. *Ann OtolRhinolLaryngol*.1980; 89: 14-19.
- Brobby GW. The discharging ear in the tropics: a guide to diagnosis and management in the district hospital. *Tropical Doctor*. 1992; 22 (1): 10-13.

- Brook I, Frazier E. Microbial dynamics of persistent purulent otitis media in children. *JPediatrics*. 1996; 128(2): 237-240.
- Mawson S, Pollack M. Special role of *Pseudomonas* aeruginosa in chronic suppurativeotitis media. *Ann OtolRhinolLaryngol Head and Neck Surg.* 1988;97 (Suppl. 130):10-13.
- Kenna M. Etiology and pathogenesis of chronic suppurative otitis media. *ArchOtolaryngol Head Neck Surg*. 1988; 97 (2) (Suppl. 137): 16-17.
- Alan E Dugdale .Management of chronic suppurative otitis media. *MJA* 2004; 180 (2): 91-93
- 24. *WHO*.Chronic suppurative otitis media .Geneva. Switzerland;2004:10
- MeyerhoffW. Pathology of chronic suppurative otitis media. Ann OtolRhinolLaryngol *Head and Neck Surg.* 1988; 97 (Suppl. 130): 21-24.
- Cuneyt M. Alper, Charles D. Advanced therapy of otitis media. PMPH-USA.2004; page 262
- Kangsanarak J, Fooanant S, Ruckphaopunt K, et al. Extracranial and intracranial complications of suppurative otitis media. Report of 102 cases. J Laryngol Otol. 1993;107(11):999-1004.
- Navacharoen N, Teotrakul S.O'Connor TE, Perry CF, Lannigan FJ.Complications of otitis media in Indigenous and non-Indigenous children. *Med J Aust.* 2009;191(9 Suppl):S60-4.

- 29. AS Adogaet al.Swab and aspiration specimen collection methods and antibiogram in chronic suppurative otitis media at Jos University Teaching Hospital: Which is superior?.Department of Surgery, Ear, Nose and Throat unit, Jos University Teaching Hospital. Ibadan. *Nigeria*.2010;9(4):230-234.
- Ching ChingWu.Disk Diffusion Susceptibility Testing (Kirby-Bauer Method)[internet].ADDL .1997 .[cited 2011AprilAvailablefrom:http://www.addl.purdue.edu/ne wsletters/1997/spring/dds.shtml
- Cholesteatoma (chronic suppurative otitis media):Treatment, symptoms, advice and help[internet][cited on 22/4/2011]. available from :http://www.privatehealth.co.uk/diseases/ear-nosethroat/cholesteatoma/
- 32. Kenneth Todar .Pseudomonas aeruginosa[internet]. Todars online textbookofbacteriology [cited on 23/4/2011].availablefrom;http://www.textbookofbacteriol ogy.net/pseudomonas.html
- 33. Pseudomonas_aeruginosa[internet].Wikepedia The Free Encyclopedia.[cited on 23/4/2011]. availablefrom:http://en.wikipedia.org/wiki/Pseudomonas _aeruginosa
- Luis M. ,De la Maza.Pseudomonas.Color atlas of medical bacteriology. *ASM* Press; 2004:133
- Cruickshank, R.J.P., Duguid, B.P. *Medical Microbiology*. Churchill Living Stone Edinburgh London and New York. 1975;: 195-200.