



# VacZine Analytics

Bringing life to vaccine strategy...

## DiseaseINFOPACK

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ESBL-producing *E.coli*

Oct 2007

## Abstract

**DiseaseINFOPACK: *ESBL-producing E.coli***

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### REPORT ABSTRACT:

Although mostly a harmless commensal bacterium, *Escherichia coli* (*E.coli*) is frequently a human pathogen. For example, certain forms of extraintestinal *E.coli*, known as ExPEC, cause urinary tract infections (UTIs) which occur in the community and/or hospital settings. In rare cases ExPEC can also cause bloodstream infections which lead to sepsis and even death. Since the 1980s, hospital *E.coli* has become multidrug resistant through the production of extended-spectrum beta-lactamase enzymes (ESBLs). The CTX-M types of the enzyme have the ability to hydrolyse many antibiotics limiting current treatment options. This **DiseaseINFOPACK** report is an expert review of the current literature providing the reader with a full easy-to-read overview of ESBL-producing *E.coli*. The report covers epidemiology, outbreak case studies along with an assessment of current treatment options and future dynamics.

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## Authors Note

This DiseaseINFOPACK focuses on ESBL-producing *Escherichia coli* (E.coli) in terms of extraintestinal diseases (ExPEC). Throughout the document there may be references to other forms of E.coli which cause intestinal diseases. The reader is referred to a different DiseaseINFOPACK for more information on this particular aspect of E.coli pathogenesis.

## Bibliography

1. Donnenberg MS. *Escherichia coli*: virulence mechanisms of a versatile pathogen (2002). San Diego, California: Academic Press
2. Herzer PJ., et al. Phylogenetic distribution of branched RNA-linked multicopy single-stranded DNA among natural isolates of *Escherichia coli*. (1990). J Bacteriol. 172:6175-81
3. Knothe H., et al. Transferable resistance to cefotaxime, cefotaxitin, cefamandole and cefuroxime in clinical isolates of *Klebsiella pneumoniae* and *Serratia marcescans* (1983). Infection. 11:315-7
4. Wong-Beringer A., et al.. Therapeutic Challenges Associated with ESBL producing *E.coli* and *Klebsiella pneumoniae* (2001). Pharmacotherapy 21:583-592
5. Pitout JDD., et al. Antimicrobial resistance with a focus on  $\beta$ -lactam resistance in gram-negative bacilli (1997). Am J Med. 103:51-9
6. Health Protection Agency Report (2005) Investigations into multi-drug resistant ESBL-producing *E.coli* strains causing infections in England. Available at: [http://www.hpa.org.uk/publications/2005/esbl\\_report\\_05/default.htm](http://www.hpa.org.uk/publications/2005/esbl_report_05/default.htm). Accessed: October 2007.
7. Paterson DL., et al.. Extended Spectrum  $\beta$ -lactamases: The European experience (2001). Curr Opin Infect Dis. 14: 697-701
8. Presentation by Graham Harvey. Extended spectrum beta-lactamase E.coli. Available at: <http://www.webbertraining.com/files/library/docs/11.pdf>. Accessed: October 2007
9. Wiener J. Multiple and antibiotic resistant *Klebsiella* and *Escherichia coli* in nursing homes (1999). JAMA. 281:563-4
10. Extended Spectrum Beta-Lactamases (ESBL) in bacteria associated with animals – Defra Position (March 2007). DEFRA. Accessed: October 2007.
11. Branger C., et al. Genetic Background of *Escherichia coli* and extended-spectrum beta-lactamase type (2005). Emerg Infect Dis. 11(1):54-61
12. Picard B., et al. The link between phylogeny and virulence in *E.coli* extraintestinal infection (1999). Infect Immun. 67:546-53
13. Johnson JR., et al. Phylogenetic background and virulence profiles of fluoroquinolone-resistant clinical isolates from the Netherlands (2002). J Infect Dis. 186:1852-6
14. Gupta K. Urinary Tract Infections (2005). Medscape ([www.medscape.com](http://www.medscape.com)). ACP medicine. WebMD Inc.
15. Stamm WE., et al. Natural History of recurrent urinary tract infection in young women (1991). Rev Infect Dis 13:77
16. Miller TE., et al. Host response in urinary tract infections (1974). Kidney International 5: 179-186
17. Weijer S., et al. Interleukin-18 facilitates the early antimicrobial host response to *Escherichia coli* peritonitis (2003). ASM 71:5488-5497
18. Cormican M., et al. ESBL production and fluoroquinolone resistance in pathogens associated with community-acquired urinary tract Infection (1998). Diagn Microbiol Infect Dis 32: 217-19
19. Harris A., et al. Risk factors for colonization with ESBL-producing bacteria and intensive care unit admission (2007). Emerg Infect Dis. 13: 1144-1149
20. Cormican M., et al. ESBL production and fluoroquinolone resistance in pathogens associated with community-acquired urinary tract Infection (1998). Diagn Microbiol Infect Dis 32: 217-19
21. Harris A., et al. Risk factors for colonization with ESBL-producing bacteria and intensive care unit admission (2007). Emerg Infect Dis. 13: 1144-1149
22. Centers for Disease Control and Prevention (CDC). Laboratory capacity to detect antimicrobial resistance, 1998. MMWR 2000;48:1167-71
23. M'Zali FH et al. Detection of ESBL in members of the family *Enterobacteriaceae*: comparison of the Mast DD test, the double disc and the Etest (2000). Journal of Antimicrob Chemother. 45, 881-5
24. Schoff WH. Pyelonephritis, Acute (2007). [www.emedicine.com](http://www.emedicine.com). WebMD. Accessed October 2007
25. Livermore D. Tigecycline. What is it, and where should it be used? (2005). J Antimicrob. Chemother. 58:1312-1314
26. Einhorn AE. Extended-Spectrum Beta-Lactamases: Frequency, Risk Factors and Outcomes (2002). Pharmacotherapy. 22:14-20

## Bibliography (cont..)

27. National Nosocomial Infections Surveillance (NNIS) System Report, data summary from January 1992 through June 2004, Issued October 2004. A report from the NNIS System, US DHHS, 2004
28. Winokur P., et al. Characterisation of strains from Europe, North and Latin America and Western Pacific that express an ESBL: Report from the SENTRY antimicrobial surveillance program (1997-1999). Presented at 40th ICAAC. Toronto, Canada, September 17-20,2000
29. Doi Y., et al. Community acquired Extended-Spectrum  $\beta$ -lactamase Producers, United States (2007). Emer Inf Dis. 13: 1121-1123
30. Jones RN., et al. Prevalence of important pathogens and antimicrobial activity of parenteral drugs at numerous medical centers in the United States (1994). Diagn Microbiol Infect Dis. 19-203-15
31. Jones RN., et al.. Antimicrobial activity and spectrum investigation of eight broad spectrum  $\beta$ -lactam drugs: a 1997 surveillance trial in 102 Medical centers in the United States (1998). Diagn Microbiol Infect Dis. 30:215-28
32. Jones RN., et al. Bacterial pathogens isolated from patients with bloodstream infection: frequencies of occurrence and antimicrobial susceptibility Patterns from the SENTRY antimicrobial surveillance program, 1997 (1998). Antimicrob Agents Chemother. 42:1762-70
33. Reynolds R., et al. Rising ESBL production and Ciprofloxacin Resistance In Invasive *Enterobacteriaceae* in the UK and Ireland (2005). Poster Presented at the 45th ICAAC Conference, Washington DC 2005

## About VacZine Analytics:

VacZine Analytics is a brand new research consultancy based in the United Kingdom. Its aim is to provide high quality-disease and commercial analysis to those working within or in collaboration with the vaccine industry.

- © With our product lines:
  - DiseaseINFOPACK
  - OpportunitySCAN
  - MarketVIEW
  - ExpertREACT
  - VaccineSTATS

Our key focus is helping clients to build the case for developing new vaccines.

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