Control of Pseudomonas aeruginosa blood stream infection outbreak in neonatal intensive care unit by quantitative antibiogram

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Abstract
Introduction: Neonates are very susceptible to Pseudomonas aeruginosa. Single case of Pseudomonas aeruginosa in preterm neonates is a sign for immediate action.
Aim: To describe control of an outbreak of Pseudomonas aeruginosa bloodstream infections (PA-BSI) by being vigilant and by immediate intervention, that occurred in the neonatal intensive care unit (NICU) of a teaching institution and hospital in Dewas, India.
Materials and Methods: The outbreak investigations included hunt for additional cases, evaluation of patient’s records, environmental and health care worker screening, immediate reporting to authority, and detection of source by minimal available aid i.e. quantitative antibiogram.
Results: Health care worker screening samples were negative on culture for Pseudomonas aeruginosa and an environmental source was detected to be a curtain near washroom.
Conclusion: This study described control of an outbreak of PA-BSI occurring over a week among neonates which was controlled following vigorous infection control measures. Quantitative antibiogram data should be routinely used for typing purposes as an essential part of hospital associated infection control procedure in source limited settings.

Keywords: Outbreak control, Pseudomonas aeruginosa, Curtain, Quantitative antibiogram.

Introduction
Pseudomonas aeruginosa is an obligate aerobe, Gram negative rod-shaped bacterium with minimal nutritional requirements. In deficiency of well-resourced surveillance systems, reports of an outbreak control can serve to high spot serious pathogens and importance of controlling them at starting itself. Such outbreaks are regularly reported in the literature, but few of these reports come from low income countries.

Pseudomonas aeruginosa is an important cause of healthcare-associated infections, particularly among infants in neonatal intensive care units (NICUs) owing to their underdeveloped immune system and the fact that such infants are often intubated, catheterized with intravascular catheters/devices in-situ, and/or receiving parenteral nutrition, which may increase risk of infection.¹² Predisposing factors for Pseudomonas aeruginosa infection are presence of indwelling devices, intensive care unit admission, prior antibiotic use, and length of hospital stay, underlying disease and reduced immunity.¹³ The higher survival of premature low birth-weight neonates has caused an increase in the incidence of hospital acquired P. aeruginosa infections.⁵ Although P. aeruginosa often only colonizes the infants, it sometimes causes infection and, despite improved treatments, P. aeruginosa bacteremia is fatal in 20% of cases.⁶ Traditional external reservoirs of P. aeruginosa include sinks, tubs, ventilation devices, incubators, and hand antiseptic solutions. What is more, this pathogen can also be isolated from walls, floors and even phototherapy equipment.⁷ There may be different transmission mechanisms, such as binding to catheters, mechanical ventilation, and the hands of medical of nurse personnel etc.⁸

The microbiology laboratory is at the centerpiece of attempts to watchdog the medical records of infections that can be hint to the arrival of a new microbial plague, an outbreak of any hospital associated infection. The clinical observation of two or more temporally related cases of nosocomial P. aeruginosa infection should raised the suspicion of an outbreak particularly in high risk pediatric patient populations (Neonatal intensive care unit, pediatric intensive care unit, oncology).⁹ We launched a outbreak investigation after getting two blood stream infection of Pseudomonas aeruginosa over a week. The objectives were to describe the outbreak, detect the source of the outbreak, and make recommendations for the prevention of future outbreaks. Our report highlights the importance of being vigilant, immediate implementation of infection control measures in stopping outbreak at its initial stage and use of qualitative antibiogram for surveillance of antimicrobial-resistant organisms in resource-poor settings.

Materials and Methods
Setting: AIMS, Dewas is a new setup in developing stage. The NICU consist of one room with 10 isolation division with a common attached bathroom and nursing station. Entry to NICU is supervised and basic infection control measures are followed.

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Data Collection: Neonates whose routine investigation for fever, yields growth of *Pseudomonas aeruginosa* on blood culture were taken into consideration. The patient folders were reviewed for the details of hospitalization and clinical history of the neonate.

Staff and Environmental Screening: A total of 76 Environmental sample were taken which includes, instrument trolley, tray, medicine trolley, weighing machine, incubator, door handle, curtain near washroom (which was wet at time of collection), sinks, tubs, swabs from taps in the patient isolation rooms, ventilation devices, water samples, water used in respiratory therapy equipment, phototherapy equipment, cleaning equipment and working station were taken. Samples from hands and apron of 18 healthcare worker posted in NICU were taken to identify the source and route of infection. 10 more neonates admitted there were also screened.

Environmental surfaces were swabbed with sterile cotton swabs, pre-moistened with sterile distilled water. Swabs were transported immediately to microbiology laboratory. Water sample (100 ml) was collected in a commercially available sterile bottle containing sodium thiosulphate powder.

Consent from health care workers was not taken as screening of staff for possible organism carriage is good clinical practice and mandatory for public health intervention in such type of outbreak investigation.

Microbiological Testing: Microbiology testing was done in AIMS microbiology laboratory. Screening specimens were plated onto nutrient agar, blood agar and MacConkey agar. *Pseudomonas aeruginosa* was identified as Gram negative bacilli, motile, oxidase positive and production of green pigment. Kirby bauer disc diffusion method was done for Antimicrobial susceptibility determination, which were then interpreted according to Clinical Laboratory Standards Institute (CLSI) guidelines. The following antibiotics were tested: amikacin, aztreonam, ceftriaxone, ceftazidime, piperacillin tazobactam, meropenem, ciprofloxacin, norfloxacin and gentamicin.

Water samples were processed by membrane filtration, 24 hrs incubation of the membrane filter in thioglycolate broth, subculture and subsequent identification.

Results
Description of Outbreak and Data Collection: In current outbreak, two consecutive bloodstream infection of *Pseudomonas aeruginosa* showing similar quantitative antibiogram in neonate over a week period 3-6-16 to 8-6-16, raised suspicion of outbreak in NICU.

Case 1: A term baby boy three days old, born to a primigravida with no adverse antenatal events was admitted to NICU with probable diagnosis of meconium aspiration syndrome. Laboratory results showed that total WBC count increased and elevated C-reactive protein. Blood culture grew *Pseudomonas aeruginosa*. Child was treated with Amikacin and Cefotaxime. There was clinical improvement and was discharged after complete antibiotic course.

Case 2: A term female baby two day old, with respiratory distress was admitted in NICU. Her WBC count was highly raised. Blood culture grew *Pseudomonas aeruginosa*. Child was treated with Cefotaxime and Gentamicin. Child improved and discharged after complete course of antibiotics.

Microbiological Testing: Both isolates produced green pigmentation and fruity smell on nutrient agar, beta hemolytic on blood agar, non lactose fermenter on MacConkey agar. They were motile, oxidase positive, urease negative and citrate positive Gram negative bacilli. Both isolates were morphologically and biochemically identical. Both isolates were having similar antibiotic sensitivity pattern i.e. *Pseudomonas aeruginosa* sensitive to Amikacin (19mm ZOI), Gentamicin (22mm ZOI), Ciprofloxacin (25mm ZOI), norfloxacin (20mm ZOI), piperacillin-tazobactam (20mm ZOI), meropenem (18mm ZOI) and were resistant to ceftazidime (11mm ZOI), ceftriaxone (10mm ZOI), aztreonam (9mm ZOI). (ZOI- zone of inhibition)

No health care worker in NICU showed isolation of *Pseudomonas aeruginosa*. *Pseudomonas aeruginosa* with similar morphology, biochemical reaction and quantitative antibiogram was isolated in a single specimen taken from the curtain near washroom. No neonate admitted showed isolation of *Pseudomonas aeruginosa*.

Discussion

Now days, outbreaks of hospital associated infections due to antibiotic resistant organisms are significant infection control problem in developed as well as resource-limited countries. Middle income countries are more likely to face these outbreaks of drug resistant pathogens, where resources allow use of broad spectrum antibiotics and not adequate infection control measures and regular surveillance. Despite recognition of this multidrug resistant pathogens as a global problem, very few outbreaks are investigated and documented in low-resourced countries.

*Pseudomonas aeruginosa* is a well known cause of invasive and non-invasive disease in the hospital setting and many studies are published linking one or more environmental samples as source. The noticeable similarity between the isolated strains in current outbreak indicates an exogenous common source. In this case, curtain near washroom was a point source of infection which was wet during collection of sample and it is well known fact that *Pseudomonas aeruginosa* is primarily an environmental organism that is adapted to survive in numerous conditions and is particularly well adapted to wet or damp conditions.

Multi-resistant organisms may transmit from one patient to another by direct contact or indirectly, by
contact of healthcare workers or fomites as vectors. In current outbreak, hands of health care worker can be considered likely to be vehicle between case and source. Though screening of staff did not reveal any carrier but it may be possibility that *Pseudomonas aeruginosa* may be a part of transient flora of health care workers hands and eliminated by the time of screening.

The control measures taken during the current outbreak were: early detection of cases, immediate reporting to higher authority and concerned health care workers, immediate microbiological assessment as well as instigating additional contact precautions for all cases (strict hand hygiene, correct use of gloves and gown) and cohorting of cases. After identification of source, the curtain was removed and health care workers were reeducated and instructed about health care practices.

The infection control procedures in place should be able to contain the infection of sporadically introduced *Pseudomonas aeruginosa* into NICU. In current outbreak, early identification and immediate decontamination of source, had lead to stoppage of transmission of infection in the first place. This indicates the significance of consistent and continuous surveillance of hospital associated infection among high risk patients. In low resource settings, surveillance of such pathogens should also incorporate molecular methods to support epidemiological data.

Passive surveillance is the strategy where problems are detected by using data obtained in the routine management of patient. Monitoring the records of laboratories allows early detection and response for mandatory changes in infection control procedures as well as implementation of additional surveillance methods to cut off new health care-associated infections at an initial stage. In light of this study, high-quality diagnostic microbiology laboratories are mandatory to support the infrastructure for help of the health care providers in management and control of health care associated infections. Modern hospital epidemiology began in the mid-1960s and from that time the clinical microbiology laboratory has played the critical roles it can serve for management and control of health care associated infectious diseases.

In general, the current outbreak studied, provide evidence that Quantitative antibiogram typing using inhibition zone diameter is a simple, rapid, readily available method. It appears to be suitable for prospective surveillance in a hospital which does not have molecular typing facilities.

**Conclusion**


This report highlights the problem of hospital associated infections and infection control challenges that they pose in a resource limited setting. It also emphasizes strict passive surveillance and use of quantitative antibiogram as a tool for microbiology laboratory to identify outbreak and its source in a low resourced and simple setup laboratory.

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