



Journal List > J Clin Microbiol > v.41(8); Aug 2003

Abstract

J Clin Microbiol. 2003 August; 41(8): 3986–3988.

■ Full Text

doi: 10.1128/JCM.41.8.3986-3988.2003.

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PDF (101K)

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Severe *Serratia liquefaciens* Sepsis following Vitamin C Infusion Treatment by a Naturopathic Practitioner

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Received October 21, 2002; Revised February 24, 2003; Accepted April 25, 2003.

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ABSTRACT

A 66-year-old female patient developed severe *Serratia liquefaciens* sepsis following infusion treatment by a naturopathic practitioner. The clinical course of the infection was characterized by several complications, and the direct costs of the hospital stay amounted to 40,000 Euro. Genotypically identical *S. liquefaciens* was isolated from the residue left in the washbasin given to the patient, as well as from the washbasin overflow and from two other locations. A careful inspection of the dispensing facilities and review of procedures used to prepare the infusion revealed several indications of poor hygiene. However, the source of contamination was not fully clarified. This case report raises questions about the local facilities and procedures required for naturopathic practitioners to conduct invasive procedures and demonstrates that poor hygiene can lead to severe morbidity and high cost.

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CASE REPORT

A 66-year-old female patient presented with symptoms of septic shock, meningitis, and decreased consciousness. Her medical history revealed a carcinoma of the cervix (stage IIIb) treated by primary radiotherapy 3 years previously. The day before, she had received an infusion containing 200 ml (i.e., 30 g) of vitamin C, 50 ml of lactopurum (homeopathic preparation of lactic acid in water for injection), and 250 ml of isotonic 0.9% sodium chloride solution. A peripherally inserted venous catheter was inserted immediately after termination of the infusion.

suffered from neck pain, vomiting, and fever. First, she was given symptomatic (antipyretic and antiemetic drugs), but during the night, her state worsened dramati-

On admission to the intensive care unit, the patient was comatose, her skin was pale and the peripheral pulses were not palpable. Clinically, we found signs of meningitis (white blood cell count was 22.8×10^9 /liter, and the level of vitamin C in serum was 55.9 mg/liter (normal range 5.0 to 15.0 mg/liter). Blood cultures revealed growth of *S. liquefaciens* that was susceptible to piperacillin plus tazobactam, cotrimoxazole, cefotaxime, ceftriaxone, and gentamicin. Empirically, ampicillin, cefazolin, cefuroxime, and nitrofurantoin. At the time of admission, ampicillin, cefazolin, cefuroxime, and nitrofurantoin were given empirically as antibiotic medication. A sample from the original infusate was obtained for culture and also revealed growth of *S. liquefaciens* by API 20E (BioMerieux). The initial antibiotic therapy was conducted until the patient's admission, and *S. liquefaciens* could not be isolated any longer after this initial period.

The further clinical course was characterized by a protracted septic shock with disseminated intravascular coagulation (D-dimers > 20 mg/liter); the clinical picture of systemic inflammatory response syndrome, with a great need for catecholamines to support circulation; a respiratory distress syndrome; reversible acute renal failure; severe anasarca, and transient hypotension. Mechanical ventilation had to be conducted for about 1 month. On the 37th day the patient was transferred to a medical unit, and on the 68th day, the patient, who was in a poor physical and psychic state (reactive depression), was dismissed from the hospital for further treatment in a rehabilitation center.

S. liquefaciens is an infrequent but increasingly recognized cause of transfusion-related sepsis associated with a high rate of mortality (1, 7). Recently, two outbreaks of *S. liquefaciens* infections in a critical care unit (4) and at a hemodialysis center (3) have been described. The environmental sources of *S. liquefaciens* remained unclear. In this paper, we report the case of severe *S. liquefaciens* sepsis following administration of a vitamin C infusion by a nurse practitioner, which was associated with a contaminated washbasin overflow and poor infection control practice.

Inspection and microbiological investigations. After notification to the local health administration, we were asked to assist in the investigation of the case. First, we conducted an inspection of the dispensing facilities. The room where the infusion was prepared was small, and there was only limited workspace for preparation of infusions. The sink was constructed with an overflow hole, a type of design that is not recommended for use in hospitals. A dispenser for hand disinfectant solution was lacking.

Second, we undertook a review of the procedure used to prepare the infusate. This included several examples of poor aseptic technique. (i) Some of the infusion components were taken from unpunctured bottles. (ii) In order to remove air bubbles, the infusion tubing was worked with the distal end of the tubing touching the washbasin overflow hole. (The washbasin is a safeguard to prevent flooding in case a tap is left running while the plug is in; the plug is located at the top of the washbasin.) (iii) Compliance with hand disinfection was poor. The puncture site of the infusion container was not disinfected.

Samples were taken from the table surface and from different sites of the washbasin (aerator, armatures, surface, and overflow hole) for microbiological analysis. Except for the overflow, none of the samples revealed growth of *S. liquefaciens*. The results from another two repeatedly punctured infusion bottles containing water for injection and a sodium chloride solution drawn from the same room also revealed growth of *S. liquefaciens*. Samples from sealed containers (identical batches) of the infusion components (i.e., lactopurum, and isotonic 0.9% sodium chloride solution) as supplied by the manufacturers showed no bacterial growth. Pulsed-field gel electrophoresis (PFGE) showed that all isolates were genotypically identical to the *S. liquefaciens* isolates from the patient's blood and the infusate given to the patient. Figure 1 shows the PFGE fingerprint patterns of clinical and environmental *S. liquefaciens* strains.

Discussion. There is no doubt that the severe *S. liquefaciens* sepsis described in this case was acquired from a contaminated naturopathic vitamin C infusion. The protracted course and complications associated with this case underline the severe morbidity associated with *S. liquefaciens* sepsis. In transfusion-related *S. liquefaciens* sepsis, the fatality rate is reported to be 100% (7). The direct cost of the hospital stay is estimated at 40,000 Euro, the range previously reported for nosocomial bloodstream infections (6).

Our findings strongly suggest that contamination of the infusion had occurred on the premises of the naturopathic practitioner, although the mode of contamination could not be fully determined. The possibility of intrinsic contamination of one of the components used to prepare the infusate was largely eliminated, since samples from other containers from the batches supplied by the manufacturers were found to be sterile. In addition, the same *S. liquefaciens* strain was found in used components from different manufacturers. Based on a careful inspection and review of the dispensing procedures, we found several indications of poor aseptic technique that likely contributed to the contamination of the infusate. Our first hypothesis was that when the infusion tubing was worked to remove air bubbles from the tubing, the distal end of the tubing could have touched the overflow hole of the washbasin, which was also found to be contaminated with genotypically identical

However, this hypothesis did not satisfactorily explain the contamination of the infusion within the bottle, which was largely protected from contamination in this chamber. Moreover, the time between preparation of the infusion and administration was too short to allow sufficient growth of *S. liquefaciens* within the bottle to produce the clinical picture: i.e., the occurrence of the febrile reaction immediately after administration of the infusion suggested the presence of considerable amounts of lipopolysaccharides. In several cases of *S. liquefaciens* contamination of blood, the implicated erythrocyte units were 1 to 3 days old, with a mean age of 28 days (7, 9).

More likely, *S. liquefaciens* was inoculated into the infusion bottle by improper handling. Grohskopf and coworkers (3) described an outbreak of *S. liquefaciens* bloodstream infections in a hemodialysis center caused by puncturing single-use vials multiple times and pooling the free residual epoetin alfa. Sharing of unlabeled heparin and insulin vials was also implicated in an outbreak of *S. liquefaciens* bloodstream infections in a critical care unit (4), although it was proven to be directly implicated in the outbreak.

To our knowledge, the identification of an overflow as an environmental reservoir for *S. liquefaciens* is unique in the scientific literature. Although its role in the contamination of the infusion is speculative, the recommendation to install washbasins without an overflow hole in the infusion room (5) is further supported by these findings. In the study of Grohskopf et al., the *S. liquefaciens* epoetin alfa probably originated from the hands of health care personnel transiently contaminated by soap or hand lotion (3). In our study, the soap was not investigated.

Our case report raises questions about the local facilities and personal qualifications of naturopathic practitioners and groups with related occupations to conduct invasive infusion treatments. Systematic data on the frequency of infectious complications in naturopathic medicine are lacking (2), and only a few case reports exist: e.g., a case of endocarditis with prosthetic heart valves caused by *Propionibacterium acnes* after a series of ‘acupuncture’ and semipermanent acupuncture needle procedures by a natural healer (8). These findings suggest that the hygienic prerequisites in naturopathic settings should be on a level comparable to those in conventional ambulatory medicine. As a consequence of the case described in this report, health authorities now offer special vocational training to natural practitioners to improve their techniques and have implemented a stronger focus on hygiene in their examinations.

In summary, our case report demonstrates that infusion therapy should be performed under hygienic conditions and that failures can lead to severe morbidity and high cost.

ACKNOWLEDGMENTS

We thank Sally Bloomfield for editorial revision and critical review of the manus for help with the clinical isolates, and W. Witte for PFGE typing.

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FIGURES AND TABLES

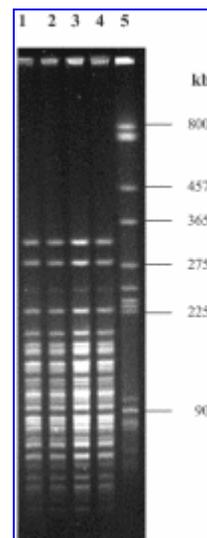


FIG. 1.

PFGE fingerprint patterns of *S. liquefaciens* strains from environmental and clinical sources. Lanes: 1 and 2, washbasin overflow hole; 3, remainder of the infusate given to patient's blood culture; 5, standard *S. typhimurium* strain LT2. (more ...)

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