

CASE REPORT



http://intl.elsevierhealth.com/journals/ijid

# Non-typhoidal Salmonella septic arthritis in an immunocompetent child with a pharyngeal streptococcal infection

Ilaria Pezone<sup>a</sup>, Maria Rosa Della Penna<sup>a</sup>, Stefano Flamini<sup>b</sup>, Giovanni Nigro<sup>a,\*</sup>

<sup>a</sup> Department of Pediatrics, University of L'Aquila, L'Aquila, Italy <sup>b</sup> Department of Orthopedics, University of L'Aquila, L'Aquila, Italy

Received 8 May 2008; accepted 2 June 2008 Corresponding Editor: William Cameron, Ottawa, Canada

### **KEYWORDS**

Salmonella enterica; Group A β-hemolytic Streptococcus; Septic arthritis **Summary** We report the case of an immunocompetent child who showed monoarticular arthritis and fever, preceded by pharyngitis and arthralgias. Because group A  $\beta$ -hemolytic Streptococcus had been detected in the pharyngeal swab, erythromycin was given on admission. However, based on ultrasound examination, therapy with ceftriaxone and joint fluid drainage were promptly performed, and a rapid and full recovery followed. Meanwhile, *Salmonella enterica* infection was revealed in blood and joint fluid. Our case suggests that septic arthritis caused by a non-typhoidal Salmonella infection may occur without gastrointestinal manifestations and concomitantly with a pharyngeal streptococcal infection.

© 2008 International Society for Infectious Diseases. Published by Elsevier Ltd. All rights reserved.

## Introduction

Non-typhoidal Salmonella serotypes (NTSS) are major causes of foodborne infections, which most often result in selflimited acute gastroenteritis.<sup>1</sup> However, approximately 5% of individuals with a gastrointestinal illness caused by NTSS develop bacteremia.<sup>2</sup> Septic arthritis is a rare consequence of Salmonella bacteremia, usually occurring in patients with underlying diseases, such as sickle cell disease and systemic lupus erythematosus (SLE).<sup>3</sup> We report on an immunocompetent child with monoarticular septic arthritis without gastrointestinal manifestations due to *Salmonella enterica*, concomitantly with group A  $\beta$ -hemolytic Streptococcus (GAS) pharyngitis.

# Case report

A 5-year-old boy was admitted because of a 7-day history of fever (38.5 °C) and arthralgias, particularly on the right elbow. There was no history of trauma or injections. On examination, the child showed signs of right elbow joint involvement including warm-skin, swelling, edema, erythema, pain, and limited range of motion. The remainder of his physical examination was unremarkable. His white blood cell count (WBC) was  $13.22 \times 10^9$ /l with 61% neutrophils, 34% lymphocytes, 4% monocytes, and 1% eosinophils. The erythrocyte sedimentation rate (ESR) was 83 mm/h and C-reactive protein (CRP) was 15.34 mg/dl. Results of serum electrolyte, blood urea nitrogen, glucose, and liver function tests were normal. Since a pharyngeal swab, which had been performed before admission, yielded a GAS infection and the anti-streptolysin O (ASO) titer was 311 IU/ml (normal <200), therapy with erythromycin

1201-9712/\$36.00 © 2008 International Society for Infectious Diseases. Published by Elsevier Ltd. All rights reserved. doi:10.1016/j.ijid.2008.06.002

<sup>\*</sup> Corresponding author. Tel. / Fax: +39 064404994; +39 0862312029. *E-mail address:* nigrogio@libero.it (G. Nigro).

was started. On the following day, a chest radiograph, electrocardiogram, and echocardiogram were unremarkable, while ultrasound examination of the right elbow showed corpuscular fluid. An arthroscopy was performed and the joint fluid drained: WBC was  $1.735 \times 10^9$ /l and Gram-negative bacilli were found in the joint fluid. S. enterica, which was found to be susceptible to third generation cephalosporins, was identified both in the joint fluid and blood cultures. The Widal-Wright test was negative. Therapy with ceftriaxone (1 g/24 hours intravenous) was given for two weeks. Meanwhile the ASO titer increased to 641 IU/ml, WBC was  $9.650 \times 10^9$ /l, ESR was 53 mm/h, and CRP became negative. Immunoglobulin levels and T- and B-cell subset lymphocytes were within normal limits. At a 1-month follow-up, the child was in good clinical condition with full recovery of the right elbow, while all laboratory tests were normal except that the ASO titer was 529 IU/ml.

#### Discussion

Acute arthritis with sepsis occurs rarely in immunocompetent children. It is generally caused by bacterial infections starting from the skin (after cutaneous lesions) or throat (i.e., Staphylococcus or Streptococcus species). In developing countries, although invasive disease due to serovar Typhi as well as NTSS is rather common in children younger than 5 years of age, septic arthritis caused by Salmonella has been reported in <1% of the cases, often resulting in serious sequelae.<sup>1,4,5</sup> In immunocompetent children, retrospective studies by Zaidi et al. and Galanakis et al. reported septic Salmonella arthritis in only two of 144 (1.38%) and one of 1087 (0.09%) patients, respectively.<sup>6,7</sup> In another retrospective review of culture-confirmed non-typhoidal Salmonella cases, Fisker et al. showed that 233 of 3328 patients (7%) had complicated diseases and 135/233 (57%) experienced an extraintestinal infection; bone and joint infections occurred in only six patients (0.2%). In a multivariate analysis, immunosuppression was a risk factor associated with the development of extraintestinal Salmonella infections (odds ratio 3.1, 95% confidence interval 1.5–6.2, p = 0.001).<sup>8</sup> In the study by Sirinavin et al., including 172 patients <15 years of age, from whom at least one nonfecal and nonurinary specimen was culture-positive for non-typhoidal Salmonella, only three cases (1.7%) of septic arthritis occurred; immunocompromising diseases were found in 19% of 74 infants and 77% of 98 children.<sup>9</sup> The absence of gastrointestinal manifestations was not reported in any case of septic arthritis occurring in immunocompetent children. Several studies have suggested the occurrence of elevated virulence and invasiveness of certain NTSS, including S. enterica and subsp enterica serovars Enteritidis and Typhimurium.<sup>8,10-12</sup> In a retrospective analysis, Hsu et al. observed that the predictors of primary bacteremia were age, presence of SLE, group B, C, or D Salmonella infection, and immunodeficiency.<sup>13</sup> In our immunocompetent child, NTSS infection, which occurred without gastrointestinal manifestations although presumably acquired by the usual oral-fecal route, spread to the bloodstream causing a septic fever with arthralgias and a subsequent monolateral arthritis. Since the boy had a pharyngeal GAS infection, we can presume that the immune defenses were decreased by the streptococcal infection and the consequent cytokine-induced inflammatory condition, making the Salmonella infection more invasive. The localization of Salmonella to the right elbow, which could have been favored by the child's activity or body position, was probably related to a local vasculitis and consequent thrombosis. A recent experimental study in mice on the dissemination of S. enterica to the brain showed the presence of thrombosed subarachnoid vessels in the areas of meningitis.<sup>14</sup> In conclusion, two interesting aspects are present in our case: (1) monoarticular arthritis by non-typhoidal Salmonella infection was associated with sepsis but not gastrointestinal manifestations; (2) an inappropriate antibiotic therapy was initially given because the pharyngeal swab yielded group A  $\beta$ -hemolytic Streptococcus, which was considered the cause of arthritis, but the yield of S. enterica from the joint fluid and blood suggested the correct therapy with ceftriaxone.

*Conflict of interest:* No conflict of interest to declare.

## References

- Graham SM. Salmonellosis in children in developing and developed countries and populations. Curr Opin Infect Dis 2002;15:507–12.
- 2. Hohmann EL. Nontyphoidal salmonellosis. *Clin Infect Dis* 2001; **32**:263–9.
- Day LJ, Qayyum QJ, Kauffman CA. Salmonella prosthetic joint septic arthritis. *Clin Microbiol Infect* 2002;8:427–30.
- Cohen JI, Bartlett JA, Corey GR. Extra-intestinal manifestations of Salmonella infections. *Medicine (Baltimore)* 1987;66: 349–88.
- Gomez HF, Cleary TG, Salmonella. In: Feigin RD, Cherry JD, editors. *Textbook of pediatric infectious diseases*. Philadelphia: WB Saunders; 1998. p. 1321–34.
- 6. Zaidi E, Bachur R, Harper M. Non-typhi Salmonella bacteremia in children. *Pediatr Infect Dis J* 1999;18:1073–7.
- Galanakis E, Bitsori M, Maraki S, Giannakopoulou C, Samonis G, Tselentis Y. Invasive non-typhoidal salmonellosis in immunocompetent infants and children. *Int J Infect Dis* 2007;11:36–9.
- Fisker N, Vinding K, Molbak K, Hornstrup MK. Clinical review of nontyphoid Salmonella infections from 1991 to 1999 in a Danish county. *Clin Infect Dis* 2003;**37**:47–52.
- Sirinavin S, Jayanetra P, Thakkinstian A. Clinical and prognostic categorization of extraintestinal nontyphoidal Salmonella infections in infants and children. *Clin Infect Dis* 1999;29: 1151–6.
- 10. Chiu CH, Ou JT. Risk factors for endovascular infection due to nontyphoid salmonellae. *Clin Infect Dis* 2003;**36**:835–6.
- Rodriguez M, de Diego I, Mendoza MC. Extraintestinal salmonellosis in a general hospital (1991 to 1996): relationships between Salmonella genomic groups and clinical presentations. J Clin Microbiol 1998;36:3291-6.
- Galofre J, Moreno A, Mensa J, Miro JM, Gatell JM, Almela M, et al. Analysis of factors influencing the outcome and development of septic metastasis or relapse in Salmonella bacteremia. *Clin Infect Dis* 1994;18:873–8.
- Hsu RB, Tsay YG, Chen RJ, Chu SH. Risk factors for primary bacteremia and endovascular infection in patients without acquired immunodeficiency syndrome who have nontyphoid salmonellosis. *Clin Infect Dis* 2003;36:829–34.
- Wickham ME, Brown NF, Provias J, Finlay BB, Coombes BK. Oral infection of mice with Salmonella enterica serovar Typhimurium causes meningitis and infection of the brain. BMC Infect Dis 2007;7:65.