In conclusion, CSF rhinorrhoea can occur more than 20 years after head trauma, and thus the patient remains at risk of developing meningitis for the rest of his life. Surgical repair of the fistula is mandatory, as spontaneous closure is unlikely.\textsuperscript{1,17} It is worth mentioning that meningitis can occur whatever the duration of CSF rhinorrhoea (3 years in this case). It is also important to remember that antibiotics prescribed before the repair is completed can select for antibiotic-resistant organisms, which may prove difficult to treat should they gain access to the CSF. The possibility of a CSF leak should be considered in any patient with a chronic clear nasal discharge. The patient should be questioned in detail about any previous head trauma, no matter when it occurred. The nasal fluid should be analysed with sensitive immunoochemical methods for the presence of CSF-specific proteins, especially the beta-2-transferrin.\textsuperscript{14} Finally, in patients with Gram-negative bacillary meningitis, in the absence of classical risk factors, the possibility of a CSF leak should be considered, as well as any associated prior antibiotic treatment.

\textbf{References}


Haemophilus aphrophilus Bacteraemia Complicated with Vertebral Osteomyelitis and Spinal Epidural Abscess in a Patient with Liver Cirrhosis

C. C. Hung\textsuperscript{1}, P. R. Hsueh\textsuperscript{2}, Y. C. Chen\textsuperscript{1}, C. T. Fang\textsuperscript{1}, S. C. Chang\textsuperscript{1}*, K. T. Luh\textsuperscript{2} and W. C. Hsieh\textsuperscript{1}

\textsuperscript{1}Section of Infectious Diseases, Department of Internal Medicine and \textsuperscript{2}Department of Laboratory Medicine, National Taiwan University Hospital, 7 Chung-Shan South Road, Taipei, Taiwan

\textbf{Haemophilus aphrophilus} is rarely implicated as an aetiology of spinal epidural abscess. A 73-year-old woman with liver cirrhosis who developed \textit{H. aphrophilus} bacteraemia complicated with vertebral osteomyelitis and spinal epidural abscess is presented. Without surgical decompression, she was successfully treated with cefotaxime for 3 weeks, followed by maintenance with ciprofloxacin for another 10 weeks. The clinical features of eight previously reported cases of vertebral osteomyelitis without epidural abscess due to \textit{H. aphrophilus} are reviewed.

\textbf{Introduction}

While \textit{Haemophilus aphrophilus} is a normal component of oral flora and can be recovered from gingival scrapings and inter-

\* Address correspondence to: Dr S. C. Chang.

Accepted for publication 10 January 1997.
Spinal epidural abscess due to *H. aphrophilus* has not been reported. Here, we present an elderly female with liver cirrhosis who developed *H. aphrophilus* bacteremia complicated with vertebral osteomyelitis and spinal epidural abscess, and was successfully treated with antibiotics without surgical intervention.

**Case Report**

A 73-year-old woman was admitted with a 2-day history of fever. She had had liver cirrhosis for 10 years but had been otherwise well. One week before admission, she had a fall from a height of three stairs, and had suffered from back pain with radiation to the right lower limb since then. Two days prior to admission high fever, numbness and weakness of the right lower extremity developed. There were no respiratory symptoms, dysuria, skin abrasion from the fall, or abdominal symptoms. She denied undergoing previous dental or urogenital procedures. On examination, the body temperature was 37.9 °C, pulse rate 110/min and BP 120/70 mmHg. Consciousness was clear. She had dental crown but there was no evidence of periodontitis. Neither tonsillar infection nor parasanal percussion tenderness were noted. Chest examination was negative. Cardiac examination revealed a grade 3/6 pansystolic murmur at the left lower sternal border. The liver was not enlarged, while spleen tip was palpable. Percussion tenderness was elicited along spinous processes of lumbar spines and right iliac area. Muscle power of the right lower leg was diminished, but deep tendon reflexes were not increased. The plantar responses were flexor bilaterally. Co-ordination, proprioceptive and sphincter functions were preserved. Hypoesthesia was noted over the saddle area. WBC count was 5.0 x 10⁹/l with 12% band forms and 81% neutrophils, haemoglobin 9.5 g/l and platelet count 33 x 10⁹/l. Serum biochemistry showed albumin 3.0 g/dl, globulin 3.6 g/dl, total bilirubin 54.7 μmol/l, aspartate aminotransferase 106 IU/l, alanine aminotransferase 75 IU/l and alkaline phosphatase 159 IU/l. ESR was 110 mm/h and C-reactive protein 6.35 mg/dl. Radiographs of thoracic and lumbar spines did not show osteolytic lesions or compression fractures. Three phase bone imaging revealed increased tracer uptake at the fourth (L4) and fifth (L5) lumbar spines. Two sets of blood culture obtained at entry both yielded Gram-negative coccobacilli after 4 days of incubation. The organism grew well by subculture onto sheep blood agar, chocolate agar, and trypticase soy agar at 35-37 °C in 5-10% CO₂ for 24 h. The organism was not V- or X-factor dependent and exhibited a positive porphyrin reaction, characteristics which were in agreement with those of *H. aphrophilus*. Minimum inhibitory concentrations (MICs) of antimicrobial agents against this isolate were determined by means of the E test (PDM Epsilometer, AB Biodisk, Solna, Sweden) on Mueller-Hinton agar supplemented with 5% sheep blood agar. *Haemophilus influenzae* ATCC 49274 was used as the control strain. Ampicillin (MIC = 0.25 mg/l), cefotaxime (MIC = 0.12 mg/l), aztreonam (MIC = 0.25 mg/l) and ciprofloxacin (MIC = 0.12 mg/l) were shown to be active against the isolate. Erythromycin (MIC = 16 mg/l) and clarithromycin (MIC = 3.2 mg/l) had poor activities against this organism. β-lactamase activity of this isolate, as checked by a cefinase disk (BBL, Microbiology System, Cockeysville, MD, U.S.A.), was negative.

Transsthoracic and transesophageal echocardiography both failed to detect vegetation or intracardiac thrombi. Because of persistent back pain, an enhanced magnetic resonance imaging (MRI) scan was performed to reveal abnormal signal intensity, with enhancement of L4 and L5 and anterior paraspinous region around L3 to L5. Prominent dural and epidural enhancement were detected (Fig. 1). The thecal sac was compressed (Figs 1 and 2). With the diagnosis of vertebral osteomyelitis, spinal dural and epidural abscess, antibiotic was started with cefotaxime 1.0 g, q 4 h. Surgical or computed tomography (CT)-guided drainage was suggested, but was not accepted by the patient. The clinical response to antibiotic
therapy was slow, though neurological symptoms did not deteriorate. After 3 weeks of cephalosporin therapy, worsening thrombocytopenia and leucopenia developed. Cefotaxime was changed to intravenous ciprofloxacin 300 mg q 12 h for the following 14 days. She was then maintained on oral ciprofloxacin 500 mg q 12 h for another 8 weeks. She was free of back pain during follow-up for 4 months. Repeated MRI scans after 4 months of therapy showed resolution of the epidural abscess and vertebral osteomyelitis.

**Discussion**

Despite the ubiquity of *H. aphrophilus* in the oral cavity, the infection is infrequent, probably due to the lower virulence of *H. aphrophilus* as compared to that of *H. influenzae*, fastidiousness of growth, and requirement of a selective medium. 1,13

Up to 1995, only eight cases of vertebral osteomyelitis due to *H. aphrophilus* has ever been reported in the English literature 5,11,12 and, to our knowledge, spinal epidural abscess caused by *H. aphrophilus* has not been reported before. Including the present case, there are five men and four women with a mean age of 54 years who have presented with vertebral osteomyelitis due to *H. aphrophilus* (Table I). Underlying diseases have been described in five patients: polyarthritis, cirrhosis, diabetes mellitus, and sciatica. The predisposing factors to vertebral osteomyelitis include laceration and penetrating injury such as epidual catheterization or myelography. The bacteria may have been introduced directly to infect the disc and vertebral bodies through the lumbar puncture. 5,7 Following laceration, 6 trivial skin breaks, or dental procedures, 5,10,12 bacteremia develops with subsequent haematogenous dissemination to the spine. In patients without evident predisposing factors, the mouth may still be a likely source of infection due to this organism. 5 Daily dental cleansing or flossing may cause bacteremia, 14 and the bacteria were not cleared efficiently by the reticuloendothelial system as our patient had liver cirrhosis with impaired phagocytosis. With comorbidity of degenerative bone and joint diseases in the elderly patients, 15 *H. aphrophilus* may be seeded to vertebral bodies and cause vertebral osteomyelitis.

While the bacteria could be isolated from the bone, disc and blood, infective endocarditis is not noted in seven patients undergoing echocardiographic evaluation. The clinical presentation of vertebral osteomyelitis due to *H. aphrophilus* can be subacute to chronic 5,7,10,13 or acute. 5,11 and is frequently confusing, as patients may have pre-existing sciatica. 5,7,9,12 Lower back pain involving lower thoracic and lumbar spines with or without fever is the most common initial manifestation. Following vertebral osteomyelitis, the clinical course may be complicated with paravertebral and psoas muscle abscesses, 5,7 or, in our patient, with spinal epidural abscess. The spinal epidural abscess impinges on the nerve root or spinal cord and therefore causes her motor or sensory deficits.

The initial correct diagnosis of vertebral osteomyelitis and spinal epidural abscess is often difficult 15,16 and delayed diagnosis and treatment may result in irreversible neurological deficits or fatalities. 15,17 In our patient, the diagnosis was prompted by the presentation of fever, persistent back pain and the associated neurological symptoms and signs. 15,16,18 With the aid of gadolinium-enhanced MRI, 17 early diagnosis, clear definition of extents and serial follow-ups are possible and potential contamination during diagnostic myelography or lumbar puncture at spinal levels of active infection can be avoided.

To treat spinal epidural abscess, immediate surgical decompression has been advocated in order to reduce the morbidity and mortality, 5,11 especially in patients with worsening neurological deficits. 5,12 Persistent severe pain, increasing body temperature, WBC count, 19 or complication of psoas abscesses. 7 In certain patients without neurological deficits or without progressing deficits, and with clinical response to antibiotics in careful daily evaluations, 7,9 or, as in our patient, with a pathogen with lower pathogenicity, antibiotic therapy can be successful.

Clinical isolates of *H. aphrophilus* have been demonstrated to be susceptible to penicillin, ampicillin, and cephalosporins by the disk diffusion susceptibility tests. 7,10 The *in vitro* activities of penicillin, ampicillin, third generation cephalosporins and ciprofloxacin against clinical isolates of *H. aphrophilus*, as determined by the microdilution susceptibility test, are excellent. 6,8,10 while those of oxacillin, erythromycin, and aminoglycosides are poor. 5,6 By using the E test, we also demonstrated similar MIC patterns of our isolate to those obtained by using the microdilution method. With or without the help of surgical intervention, with administration of antibiotics based on these *in vitro* susceptibility tests, successful treatment of *H. aphrophilus* vertebral osteomyelitis without relapse can be achieved. 7,9,10,12

Clinical failure to third generation cephalosporin and resistance of *H. aphrophilus* to ampicillin, cefuroxime, cefotaxime, and ceftazidime may occur, however, without producing β-lactamase. 12 Marked inoculum effect has been observed, and clinical improvement is not achieved until ciprofloxacin replaces the cephalosporin. 12

In summary, this unique case of *H. aphrophilus* bacteraemia complicated with spinal epidural abscess and vertebral osteomyelitis not only serves to expand the spectrum of bacteria causing spinal epidural abscess, but also highlights the possibility that medical therapy can be successful in carefully observed patients without neurological progression.

**References**


Table 1. Clinical features of nine patients with vertebral osteomyelitis due to *Haemophilus aphrophilus*.

<table>
<thead>
<tr>
<th>Case no./ (References)</th>
<th>Age/sex</th>
<th>Underlying diseases</th>
<th>Predisposing factors</th>
<th>Symptomatology</th>
<th>Endocarditis</th>
<th>Location of lesions</th>
<th>Positive cultures</th>
<th>Treatment</th>
<th>Outcome</th>
</tr>
</thead>
<tbody>
<tr>
<td>1(5) 59/F</td>
<td>Polyarthritis; sciatica</td>
<td>Epidural catheterization</td>
<td>Fever, worsening back pain</td>
<td>NM</td>
<td>L2–L3; psoas abscess</td>
<td>Abscess; blood</td>
<td>Antibiotic; surgical drainage</td>
<td>No relapse</td>
<td></td>
</tr>
<tr>
<td>2(6) 36/M</td>
<td>Nil</td>
<td>Lip laceration</td>
<td>Fever, back pain</td>
<td>Nil</td>
<td>L2–L3</td>
<td>Bone; blood</td>
<td>Antibiotic; aspiration</td>
<td>No relapse</td>
<td></td>
</tr>
<tr>
<td>3(7) 69/F</td>
<td>Sciatica</td>
<td>Myelography</td>
<td>Fever, abdominal mass, worsening back pain</td>
<td>Nil</td>
<td>L2–L4; paraspinal</td>
<td>CSF; blood; abscess</td>
<td>Antibiotic; surgical drainage</td>
<td>No relapse</td>
<td></td>
</tr>
<tr>
<td>4(8) 60/F</td>
<td>DM; sciatica</td>
<td>Nil</td>
<td>Worsening back pain</td>
<td>Nil</td>
<td>T11; paraspinal</td>
<td>Disc; bone</td>
<td>Antibiotic; surgical evacuation</td>
<td>No relapse</td>
<td></td>
</tr>
<tr>
<td>5(9) 59/M</td>
<td>Nil</td>
<td>Nil</td>
<td>Fever, back pain</td>
<td>Nil</td>
<td>L2–L3; paraspinal</td>
<td>Bone; blood</td>
<td>Antibiotic; surgical debridement</td>
<td>No relapse</td>
<td></td>
</tr>
<tr>
<td>6(10) 38/M</td>
<td>Nil</td>
<td>Trivial injury; gum bleed</td>
<td>Fever, back pain</td>
<td>NM</td>
<td>T11–T12; paraspinal</td>
<td>Blood</td>
<td>Antibiotic</td>
<td>No relapse</td>
<td></td>
</tr>
<tr>
<td>7(11) 40/M</td>
<td>Nil</td>
<td>Nil</td>
<td>Fever, back pain</td>
<td>Nil</td>
<td>T11–T12</td>
<td>Blood</td>
<td>Antibiotic</td>
<td>No relapse</td>
<td></td>
</tr>
<tr>
<td>8(12) 53/M</td>
<td>Sciatica</td>
<td>Dental procedure</td>
<td>Back pain, worsening neurological deficits</td>
<td>Nil</td>
<td>L4–L5</td>
<td>Blood; disc</td>
<td>Antibiotic; surgical decompression</td>
<td>No relapse</td>
<td></td>
</tr>
<tr>
<td>9(PR) 73/F</td>
<td>Liver cirrhosis</td>
<td>Nil</td>
<td>Fever, back pain</td>
<td>Nil</td>
<td>L1–S1; epidural paraspinal</td>
<td>Blood</td>
<td>Antibiotic</td>
<td>No relapse</td>
<td></td>
</tr>
</tbody>
</table>

CSF: cerebrospinal fluid; DM: diabetes mellitus; NM: not mentioned; L: lumbar spine; PR: present report; T: thoracic spine; S: sacrum.
Quinolone-resistant *Salmonella paratyphi* B Meningitis in a Newborn: a Case Report

Z. A. Bhutta*

Department of Pediatrics, The Aga Khan University Medical Center, Karachi, Pakistan

While there are concerns about the consequences of widespread use of quinolones, there are few reports of quinolone-resistant strains of *Salmonella typhi* or *Salmonella paratyphi* from the Indian subcontinent. We present a case report of a newborn with meningitis due to a quinolone-resistant strain of *S. paratyphi* B presenting to the Aga Khan University Hospital (AKUH).

**Introduction**

The emergence of multidrug-resistant strains of *Salmonella typhi* and *Salmonella paratyphi* in the Indian subcontinent have posed major problems in the management of these patients, especially in the paediatric age group. Although there are concerns about the use of quinolones in the paediatric age group, in view of the limited choice of antimicrobial therapy and the prohibitively expensive alternative third generation cephalosporins, quinolones, especially ciprofloxacin, have been widely recommended for use in India for the management of typhoidal salmonellosis in childhood.

**Case Report**

A 10-day-old female infant was referred to the Aga Khan University Hospital (AKUH) from another city in Northern Pakistan with a history of cyanotic spells and worsening respiratory distress. She was delivered prematurely at 36 weeks’ gestation to a gravida mother at a district headquarters hospital (DHQ) by LSCS for preterm labour and intrauterine distress. The baby weighed 2.2 kg at birth and was in a poor condition requiring resuscitation (Apgar score 4 and 2 at 1 and 5 min, respectively). She was placed in an incubator and given oxygen. She was administered ampicillin and gentamicin at the DHQ hospital and dexamethasone was administered for 48 h. Her respiratory distress improved gradually over the next 48 h, and formula feeds were given. She developed diarrhoea at 4 days of age, and her general condition deteriorated. Only a blood picture and glucose could be performed at this hospital. She was then placed on i.v. ceftriaxone, tobramycin and amoxicillin, and when she developed cyanotic spells a request for transfer to Karachi was made. She was air-transferred to the NICU at AKUH at 10 days of age.

At admission to AKUH she weighed 1.7 kg and was very sick with poor peripheral perfusion, dehydration and respiratory distress (respiratory rate 94/min). She was not icteric. The abdomen was distended and both the liver and spleen were palpable 2 cm below the costal margin. The anterior fontanelle was sunken and there was generalized hypertonia with extensor plantar responses. A blood picture and a blood smear was performed; no organisms and latex agglutination tests for *H. influenzae* were negative. Chest X-ray was unremarkable.

At 6 h after admission. Admission investigations revealed normal blood gases, and oxygen saturation (93%), haemoglobin (13.5 g/dL), haematocrit (39.8%), white blood cell count (7.5 × 10^9/L), platelets (18 × 10^11/L), sodium (129 mmol/L), potassium (5.8 mmol/L), bicarbonate (13.6 mmol/L), creatinine (0.8 mmol/L), BUN (17 mg/dL), calcium (8.1 mg/dL), glucose (189 mg/dL), PT (18 s, control 13 s), PTT (33 s, control 35 s), FDP (20 μg/mL). Cerebrospinal fluid (CSF) examination revealed turbid fluid with 480 cells/mm³ (mostly polymorphs), protein 728 mg/L and glucose of 55 mg/dL. The Gram stain did not reveal any organisms and latex agglutination tests for *Escherichia coli*, group B streptococcus, pneumococcus, *Haemophilus influenzae* and pseudomonas were negative. Chest X-ray was unremarkable. A contrast computed tomography (CT) scan revealed evidence of generalized meningoencephalitis and

---

* Address correspondence to: Dr Z. A. Bhutta, Department of Pediatrics, The Aga Khan University Hospital, Stadium Road, P.O. Box 3500, Karachi 74800, Pakistan. Accepted for publication 7 February 1997.