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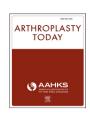
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#### Case report

## Lyme periprosthetic joint infection in total knee arthroplasty

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#### ABSTRACT

Lyme arthritis, caused by the spirochete *Borrelia burgdorferi* sensu stricto, is a common tick-borne illness in New England and the upper Midwest. Most often, the disease affects the knee and has typically been reported as a cause of native joint infection. There has been only 1 case of Lyme periprosthetic joint infection (associated with a total knee arthroplasty) reported in the literature, and to our knowledge, no other reported cases of Lyme periprosthetic joint infections exist. In this article, we report on 2 patients diagnosed with prosthetic joint infections who were subsequently found to have Lyme prosthetic joint infections, with *B burgdorferi* as the infectious organism. We discuss the medical and surgical management of these patients.

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#### Introduction

Lyme arthritis is common in New England and the upper Midwest. It typically presents with monoarticular pain and swelling, a clinical picture that is similar to a native septic joint. Before 2016, however, only 1 case of Lyme arthritis associated with a total knee periprosthetic joint infection (PJI) had been reported in the literature [1].

Prosthetic joint infections are both a personally devastating and systemically costly complication of joint arthroplasty. The management of PJI frequently necessitates staged surgical procedures and prolonged antibiotic therapy [2]. Diagnosis of the condition can be challenging and is frequently made through indirect measurements of inflammatory markers, joint fluid cell counts in addition to microbiology cultures [2,3]. The American Academy of Orthopedic Surgeons developed a clinical practice guideline [4] for the assessment of PJI, which include first screening patients who are found to have the potential for PJI with erythrocyte sedimentation rate (ESR) and C-reactive protein (CRP). In the case where one or both of these inflammatory markers are elevated, an aspiration of the knee is then recommended, which should be sent for microbiologic culture, synovial fluid and

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differential white blood cell (WBC) count. A second aspiration is recommended if there is a discrepancy between the probability of PII and the initial aspiration culture result.

The definition of PJI is also not completely straight forward and is often the source of controversy. The closest consensus definition emerged from the Musculoskeletal Infection Society workgroup in 2011 [5], which is reproduced in Table 1.

Once the diagnosis of knee PJI is made, timely surgical intervention can be crucial. The algorithm used for treatment of PJI differs based on institutional preferences, but a distinction is usually made between acute and chronic infections. In the case of chronic infections, a two-stage revision is nearly always indicated. This involves removal of all hardware, insertion of an antibiotic-loaded dynamic or static spacer, a 6-week course of intravenous (IV) antibiotics, and reimplantation of the joint replacement once the infection is proven eradicated [6]. If the clinical assessment shows an acute infection, however, with symptoms less than 3 weeks duration, a one-stage revision or irrigation and debridement with retention of hardware is often attempted [7].

We report 2 cases of patients initially diagnosed with a PJI, who were subsequently found to have Lyme arthritis PJI.

#### Case histories

Case 1

An 89-year-old woman with a past surgical history of a right total knee arthroplasty for degenerative joint disease 16 years

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#### Table 1

Musculoskeletal Infection Society consensus definition for periprosthetic joint infection.

- 1. There is a sinus tract communicating with the prosthesis; or
- A pathogen is isolated by culture from at least 2 separate tissue or fluid samples obtained from the affected prosthetic joint; or
- 3. Four of the following 6 criteria exist:
  - a. Elevated serum ESR and CRP
  - b. Elevated synovial leukocyte count
  - c. Elevated synovial neutrophil percentage
  - d. Presence of purulence in the affected joint
  - e. Isolation of a microorganism in a culture of periprosthetic tissue or fluid
  - f. Greater than 5 neutrophils per high-power field in 5 high-power field observed from histologic analysis of periprosthetic tissue at  $\times 400$  magnification

before presentation, presented to the Emergency Department with 24-hour history of right knee pain, swelling, and stiffness. She denied fevers, chills, sweats, or other constitutional symptoms. She denied any recent trauma but did report a recent dental cleaning 1 month prior, which was performed without antibiotic prophylaxis. She had a past medical history significant for monoclonal gammopathy of undetermined significance, chronic kidney disease, duodenal ulcer, hypertension, a right bundle branch block, and glaucoma. She was an active independent woman who lived alone and ambulated with a walker with no prior problem with her right total knee arthroplasty.

On examination, she had a right knee effusion with range of motion limited by pain to a 10-40 degree flexion arc. The skin overlying the knee was moderately warm and intact without drainage. All vital signs were within normal limits. Laboratory data included a serum white blood count of  $13.4\times1000/\mu L$ , 74% neutrophils, CRP of 101.0 mg/L (normal 0.1-3), and ESR of 19 mm/h (normal 0-20). A urinalysis was remarkable for pyuria and bacteriuria. An arthrocentesis of the right knee demonstrated 66,100 cells (WBC)/ $\mu L$ , 93% granulocytes. The aspirate was also sent for culture and sensitivities. No organisms were seen on gram stain. A moderate number of calcium pyrophosphate crystals were also noted.

The patient was admitted to the hospital and started on vancomycin, ceftriaxone, and colchicine for treatment of presumed prosthetic joint infection, urinary tract infection, and concomitant pseudogout. The following day, she was taken to the operating room for a right knee irrigation and debridement and liner exchange for presumed PJI. During the surgery, a small amount of clear joint fluid was noted, without frank purulence. All components were noted to be well fixed. Synovial fluid and tissue cultures were obtained. She was continued on IV antibiotics postoperatively. Due to clinical suspicion, serum samples were taken during the operative day for Lyme antibodies. On postoperative day 1, serum Lyme returned positive at 1.71 (normal range <0.9). At this point, vancomycin and colchicine were discontinued, and ceftriaxone was continued for presumed Lyme PJI. On postoperative day 4, Western blot confirmed the positive Lyme enzyme-linked immunosorbent assay (ELISA) with a positive IgM and negative IgG. Admission arthrocentesis fluid was reprocessed for Lyme polymerase chain reaction (PCR), which was found to be positive. Ceftriaxone was discontinued, and the patient was started on oral doxycycline to complete a 28-day course. The patient's mobility, range of motion, and pain improved significantly, and she was discharged to a short-term rehabilitation facility to complete rehabilitation and the course of oral antibiotics. Both admission arthrocentesis cultures and operative cultures were finalized as negative for acid fast, fungal and sonicated bacterial cultures at 8 weeks.

At 6-week follow-up, 2 weeks after completion of her antibiotic treatment, the patient's knee pain and swelling had completely

resolved. The patient missed further follow-up appointments. She expired 5 months after this hospitalization from heart disease. No autopsy was performed.

Case 2

An 80-year-old woman with hypertension and atrial fibrillation underwent an uncomplicated right knee arthroplasty for osteoarthritis 4 months before presenting to her surgeon's outpatient orthopaedic office with anterior proximal tibia erythema and tenderness. The patient denied fevers, chills, sweats, or constitutional symptoms. She denied trauma to the leg, and she stated that she had a pedicure 6 weeks before presentation. In the office, she was prescribed oral cephalexin. Over the following 4 days, the erythema resolved; however, she experienced an increase in right knee pain and swelling. She underwent an arthrocentesis of the right knee, which revealed 87,830 WBC/ $\mu$ L, 93% polys with a negative gram stain. She was admitted to the hospital for a presumed acute prosthetic joint infection.

On arrival to the hospital, the patient had normal vital signs, a serum white blood count of 8.8  $\times$  1000/ $\mu$ L, ESR of 108 mm/h, and CRP of 161 mg/L. IV cefazolin was started, and she was brought to the operating room on hospital day 3 for presumed PJI. Exploration of the prosthetic knee revealed well-fixed components with a small amount of intra-articular purulence and fibrinous exudate with minimal synovial hypertrophy. The polyethylene liner was exchanged, additional fluid and tissue samples were sent for analysis, and the wound was irrigated with bacitracin solution. Lyme PCR was performed on the synovial fluid obtained during the surgery, and this was found to be positive. Subsequent Lyme ELISA performed on serum was also strongly positive at 4.65 units (normal <0.9). Western blot IgM and IgG were negative. The patient was diagnosed with Lyme disease and switched to ceftriaxone. All preoperative and intraoperative cultures were negative for growth at end of a 7-day incubation period.

The patient was discharged on postoperative day 3. Pain and swelling rapidly resolved. She was maintained on ceftriaxone for the following 4 weeks and was then switched to oral doxycycline, which she took for an additional 4 weeks. Three months after the procedure, ESR and CRP serum levels had normalized, and the patient's pain had resolved.

She underwent an uncomplicated contralateral, left knee replacement 18 months later. At 24-month follow-up, the patient remained asymptomatic without further problems with her right knee.

#### Discussion

Lyme disease is endemic to the United States, concentrated around the Northeast and upper Midwest, with as many as 30,000 cases reported annually [8]. The disease is caused by the spirochete *Borrelia burgdorferi* sensu stricto, delivered through the bite of an Ixodes tick and is concentrated in regions where human populations coexist with white-tailed deer populations [9]. The classic initial presentation involves a target lesion (erythema migrans) centered on the tick bite. The bacteria then disseminates hematogenously. Patients may present with multisystem involvement, resulting in fatigue, headaches, myalgias, and arthralgias [10,11]. Arthritis is often a late sign of Lyme disease and manifests in a majority of untreated cases [12].

Although common, the pathophysiology of Lyme arthritis is not fully understood but thought to be a product of hematogenous dissemination to the synovial tissue. Once in the synovium, the organism likely stimulates an immune response that includes recruitment of inflammatory cells, immune complexes, cytokines,

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