

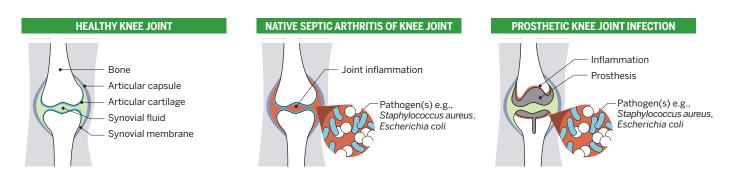


SEPTIC ARTHRITIS

Septic arthritis, also known as infectious arthritis, is a severe type of orthopedic infection. It occurs when bacteria, mycobacteria, fungi, or viruses infect an articulating joint, causing inflammation of the synovial membrane and accumulation of purulent fluid within the joint capsule.

SEPTIC ARTHRITIS

- Septic arthritis is a medical emergency which, if not treated rapidly, can lead to irreversible damage to the joint resulting in significant disability and an increased risk of death.
- Septic arthritis most commonly affects young children, the elderly, anyone with an artificial joint or existing joint disease, and immunocompromised people.
- **Prosthetic joint infection (PJI)** is a subcategory of septic arthritis where the infection involves an artificial joint that has been surgically implanted. It is useful to think of PJI and **native joint septic arthritis (NSA)** separately because the evaluation, differential diagnosis, and treatment strategies are somewhat different.¹



THE BURDEN OF SEPTIC ARTHRITIS

Joint infections represent a **heavy health and economic burden** for patients and society, and prosthetic joint infections are particularly costly to treat.

NATIVE SEPTIC ARTHRITIS¹⁻³

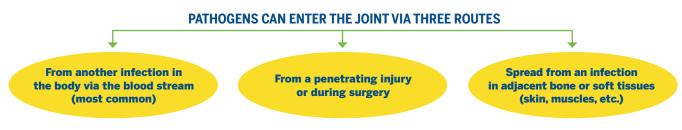
- 6-10 new cases/year/100,000 population
- 84% admitted to hospital
- ~2.5% mortality rate during hospitalization
- 55% discharged to skilled care facility or
- require home health care
- Cost = \$759 million

PROSTHETIC JOINT INFECTION 1,4-6

- Number of arthroplasties rising in developed countries
- Examples of infection rates:
 - 2.2% US
 - 0.85% Germany
 - 1.41% Finland
 - 0.76% Taiwan
- Estimated combined costs >\$1.85 billion by 2030 in US

CAUSES AND CLINICAL PRESENTATION OF SEPTIC ARTHRITIS 1,2,4

Septic arthritis is caused by bacteria, mycobacteria, fungi and, less commonly, viruses.





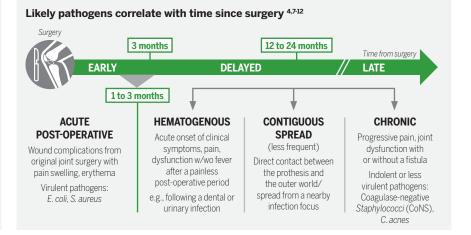
NATIVE SEPTIC ARTHRITIS

Many gram-positive and gram-negative bacteria can cause NSA:²

- S. aureus is the most common for all ages
- K. kingae is significant in children <5 years
 N. gonorrhoeae is an important cause in
- sexually active patients

Signs and symptoms:

- joint pain/tenderness
- erythema
- warmth
- edema/joint effusion
- fever



PROSTHETIC JOINT INFECTION

DIAGNOSTIC APPROACH 1,2,4,7

- **Diagnosis of joint infections is complicated** as they are often associated with hard-to-grow organisms. In PJI, biofilm-forming organisms and polymicrobial infections may also be present. Traditional testing methods may therefore require multiple tests and it can take up to two weeks to identify the pathogen(s). Also, it is not uncommon for the diagnosis to be made without ever identifying the pathogen.
- Molecular diagnostics such as polymerase chain reaction (PCR) tests are emerging as new and powerful diagnostic tools. The molecular syndromic testing approach enables rapid accurate identification of multiple target pathogens and antimicrobial resistance markers. It can also detect fastidious organisms, hard-to-grow anaerobes and polymicrobial infections. This may be possible even when the patient has been pre-treated with antibiotics.
- Syndromic tests may help to more rapidly and easily identify the cause of non-specific symptoms, such as joint pain, red or swollen joints in a **clinically actionable timeframe**. More rapid results may potentially **reduce the time to targeted therapy** and **avoid unnecessary antibiotic treatments**.

NATIVE SEPTIC ARTHRITIS

A positive synovial fluid culture is the only definitive diagnosis^{2,13}

Blood culture bottles are frequently used to increase the sensitivity of synovial fluid culture.

- Arthrocentesis/surgery to collect synovial fluid for gram stain, culture and chemical/cellular analysis
- · Radiologic assessment of the joint
- Blood tests for markers of inflammation
- Differential diagnosis includes rheumatoid arthritis, reactive arthritis, gout/pseudogout, trauma, degenerative joint disease.

PROSTHETIC JOINT INFECTION

Diagnosis is criteria-based and various organizations have published slightly different criteria 7,14-18

- Presence of a sinus tract is a common, definitive criteria.
- In addition to synovial fluid analysis/culture, blood tests, and radiologic evaluation:
 - Multiple peri-prosthetic tissue biopsies are obtained for histopathology and culture
 - Sonicate fluid culture (useful for detecting biofilm-producing organisms)
- Two or more positive cultures with the same organism are recommended to differentiate a true infection from a contamination.



TREATMENT STRATEGIES

NATIVE SEPTIC ARTHRITIS^{2,13}

- · Surgery to drain the infection and wash out the joint
- Antibiotics are usually given for 3 6 weeks, based on susceptibility testing when available or local resistance patterns and trends.

PROSTHETIC JOINT INFECTION 1,4,11-13,16

Along with antibiotics and drainage, a decision must be made in regard to **retaining vs removing/replacing the prosthesis**. While this treatment decision is not standardized, the table below lists different options and their considerations.

SURGICAL OPTIONS	CONSIDERATION	TREATMENT / PROCEDURE
DAIR*	Presentation <30 days	ATBX** (e.g., Rifampin + Fluoroquinolone) x 2-6 weeks
One-Stage Replacement Arthroplasty (More common in hip)	No sinus tract Healthy patient and soft tissue Prolonged ATBX use No bone graft Low-virulence organism with good ATBX sensitivity	Remove and replace prosthesis ATBX Impregnated cement IV*** ATBX x 4-6 weeks
Two-Stage Replacement Arthroplasty	Gold standard for infected joint >4 weeks Must be medically fit for multiple surgeries Requires adequate bone stock	Remove prosthesis \rightarrow ATBX spacer \rightarrow IV ATBX x 4-6 weeks \rightarrow new prosthesis implanted
Resection Arthroplasty	Poor bone and soft tissue quality Recurrent infection with MDR**** organism	Remove infected tissue and hardware without reimplantation Joint fused
Unfit for Surgery	Refuse surgery	Suppressive ATBX

* DAIR: Debridement, antibiotics and implant retention; **ATBX: Antibiotics; ***IV: Intravenous; ****MDR: Multi-drug resistant.

References:

- Roerdink RL, Huijbregts HJ, van Lieshout AW, et al. The difference between native septic arthritis and prosthetic joint infections: A review of literature. J. Orthop. Surg. 2019,27(2):230949901986046.
- Krogstad PA. Bacterial arthritis: Epidemiology, pathogenesis, and microbiology in infants and children - UpToDate. Oct 2019.
- Singh JA, Yu S. The burden of septic arthritis on the U.S. inpatient care: A national study. PLoS One. 2017;12(8):e0182577.
- 4. Tande AJ & Patel R. Prosthetic Joint Infection. Clin Microbiol Rev. 2014:27(2):302-345.
- Premkumar A, Kolin DA, Farley KX, et al. Projected Economic Burden of Periprosthetic Joint Infection of the Hip and Knee in the United States. J Arthroplasty. 2021;36(5):1484-1489.e3.
- Lenguerrand E, Whitehouse MR, Beswick AD, et al. Description of the rates, trends and surgical burden associated with revision for prosthetic joint infection following primary and revision knee replacements in England and Wales: an analysis of the National Joint Registry for England, Wales, Northern Ireland and the Isle of Man. BMJ Open. 2017;7(7):e014056.
- Signore A, Sconfienza LM, Borens O, et al. Consensus document for the diagnosis of prosthetic joint infections: a joint paper by the EANM, EBJIS, and ESR (with ESCMID endorsement). Eur J Nucl Med Mol Imaging 2019;46(4):971-988.
- Zeller V, Kerroumi Y, Meyssonnier V, et al. Analysis of postoperative and hematogenous prosthetic jointinfection microbiological patterns in a large cohort. J Infect. 2018;76(4):328-334.
- Parvizi J, Tan TL, Goswami K, et al. The 2018 Definition of Periprosthetic Hip and Knee Infection: An Evidence-Based and Validated Criteria. J Arthroplasty 2018;33(5):1309-1314.e2.

- Rakow A, Perka C, Trampuz A, Renz N. Origin and characteristics of haematogenous periprosthetic joint infection. *Clin Microbiol Infect*. 2019;25(7):845-850.
- Izakovicova P, Borens O, Trampuz A. Periprosthetic joint infection: current concepts and outlook. EFORT Open Rev. 2019;4(7):482-494.
- Wouthuyzen-Bakker M.et al; ESCMID Study Group for Implant-Associated Infections (ESGIAI). Lower Success Rate of Débridement and Implant Retention in Late Acute versus Early Acute Periprosthetic Joint Infection Caused by Staphylococcus spp. Results from a Matched Cohort Study. Clin Orthop Relat Res. 2020;478(6):1348-1355.
- 13. Zimmerli W. Bone and Joint Infections: From Microbiology to Diagnostics and Treatment. 2015.
- McNally M, Sousa R, Wouthuyzen-Bakker M, et al. The EBJIS definition of periprosthetic joint infection. Bone Joint J. 2021;103-B(1):18-25.
- Parvizi J, Tan TL, Goswami K, et al. The 2018 Definition of Periprosthetic Hip and Knee Infection: An Evidence-Based and Validated Criteria. J Arthroplasty. 2018;33(5):1309-1314.
- Osmon DR, Berbari EF, Berendt AR, et al. Diagnosis and management of prosthetic joint infection: clinical practice guidelines by the Infectious Diseases Society of America. *Clin Infect Dis.* 2013;56: e1–e25.
- Parvizi J, Zmistowski B, Berbari EF, et al. New definition for periprosthetic joint infection: from the Workgroup of the Musculoskeletal Infection Society. Clin Orthop Relat Res. 2011;469(11):2992-2994.
- Parvizi J, Gehrke T, Chen AF. Proceedings of the International Consensus on Periprosthetic Joint Infection. Bone Joint J. 2013;95-B(11):1450-2.

10-22 / This document is not legally binding, bioMérieux reserves the right to modify specifications without notice / BIOMÉRIEUX and the BIOMÉRIEUX logo are used, pending and/or registered trademarks belonging to bioMérieux, or one of its subsidiaries, or one of its companies / bioMérieux S.A. RCS Lyon 673 620 399

bioMérieux, S.A. • 69280 Marcy l'Étoile • France • Tel: +33 (0)4 78 87 20 00 • www.biomerieux.com