

studies should aim to further investigate these potential differences in the organism and resistance profiles in hips and knees diagnosed with SSI and PJI.

Groff *et al.* recently examined 1,214 PJI cases (501 hips and 713 knees) over a 17-year timeframe and found significant differences in pathogens causing PJI in the hip and the knee. A higher incidence of *Streptococcal* species (odds ratio (OR) 1.82, 95% confidence interval (CI), 1.23-2.67) and culture-negative PJI (OR 1.53, 95% CI, 1.12-2.09) were identified in TKA compared to THA. In contrast, *Pseudomonas* (OR 2.123, 95% CI, 1.04-4.34), *Enterococcus* (OR 1.72, 95% CI, 1.03-2.86), resistant species (OR 1.64, 95% CI, 1.19-2.25), *Staphylococcus aureus* (OR 1.40, 95% CI, 1.11-1.77) and gram-positive (OR 1.37, 95% CI, 1.05-1.78) organisms were more prevalent in hips. The authors suggested that the higher rates of urogenital-associated pathogens causing PJI in hips may have been related to the close proximity of the incision to the flexural creases and the groin region.

Although most studies have not demonstrated a definitive difference in organism profile between hips and knees, some have identified differences in virulence patterns, culture-negative rates, urogenital and fecal bacteria, as well as the overall rates of PJI in bilateral compared to unilateral TKA [12-14,17]. It is important to further delineate the differences in organism profile at these anatomic sites in order to establish adequate protocols and select antimicrobials accordingly, that may account for potential differences in the pathogenic flora and mitigate the risk of SSI/PJI.

## REFERENCES

- [1] Bori G, Navarro G, Morata L, Fernández-Valencia JA, Soriano A, Gallart X. Preliminary results after changing from two-stage to one-stage revision arthroplasty protocol using cementless arthroplasty for chronic infected hip replacements. *J Arthroplasty*. 2018;33:527-532. doi:10.1016/j.arth.2017.08.033.
- [2] George DA, Logoluso N, Castellini G, Gianola S, Scarponi S, Haddad FS, et al. Does cemented or cementless single-stage exchange arthroplasty of chronic periprosthetic hip infections provide similar infection rates to a two-stage? A systematic review. *BMC Infect Dis*. 2016;16:553. doi:10.1186/s12879-016-1869-4.
- [3] Mahmud T, Lyons MC, Naudie DD, Macdonald SJ, McCalden RW. Assessing the gold standard: a review of 253 two-stage revisions for infected TKA. *Clin Orthop Relat Res*. 2012;470:2730-2736. doi:10.1007/s11999-012-2358-8.
- [4] Sheehan E, McKenna J, Mulhall KJ, Marks P, McCormack D. Adhesion of *Staphylococcus* to orthopaedic metals, an in vivo study. *J Orthop Res*. 2004;22:39-43. doi:10.1016/s0736-0266(03)00152-9.
- [5] Moran E, Masters S, Berendt AR, McLardy-Smith P, Byren I, Atkins BL. Guiding empirical antibiotic therapy in orthopaedics: The microbiology of prosthetic joint infection managed by debridement, irrigation and prosthesis retention. *J Infect*. 2007;55:1-7. doi:10.1016/j.jinf.2007.01.007.
- [6] Holleyman RJ, Baker P, Charlett A, Gould K, Deehan DJ. Microorganisms responsible for periprosthetic knee infections in England and Wales. *Knee Surg Sports Traumatol Arthrosc*. 2016;24:3080-3087. doi:10.1007/s00167-015-3539-2.
- [7] Phillips JE, Crane TP, Noy M, Elliott TSJ, Grimer RJ. The incidence of deep prosthetic infections in a specialist orthopaedic hospital: a 15-year prospective survey. *J Bone Joint Surg Br*. 2006;88:943-948. doi:10.1302/0301-620X.88B7.17150.
- [8] Arciola CR, Campoccia D, Ehrlich GD, Montanaro L. Biofilm-based implant infections in orthopaedics. *Adv Exp Med Biol*. 2015;830:29-46. doi:10.1007/978-3-319-11038-7\_2.
- [9] Tetrycz D, Ferry T, Lew D, Stern R, Assal M, Hoffmeyer P, et al. Outcome of orthopedic implant infections due to different staphylococci. *Int J Infect Dis*. 2010;14:e913-e918. doi:10.1016/j.ijid.2010.05.014.
- [10] Ravi S, Zhu M, Luey C, Young SW. Antibiotic resistance in early periprosthetic joint infection. *ANZ J Surg*. 2016;86:1014-1018. doi:10.1111/ans.13720.
- [11] Joshy S, Gogi N, Thomas B, Mahale A, Singh BK. Delayed onset of deep infection after total knee arthroplasty: comparison based on the infecting organism. *J Orthop Surg Hong Kong*. 2007;15:154-158. doi:10.1177/230949900701500205.
- [12] Aggarwal VK, Bakhshi H, Ecker NU, Parvizi J, Gehrke T, Kendoff D. Organism profile in periprosthetic joint infection: pathogens differ at two arthroplasty infection referral centers in Europe and in the United States. *J Knee Surg*. 2014;27:399-406. doi:10.1055/s-0033-1364102.
- [13] Pulido L, Ghanem E, Joshi A, Purtill JJ, Parvizi J. Periprosthetic joint infection: the incidence, timing, and predisposing factors. *Clin Orthop Relat Res*. 2008;466:1710-1715. doi:10.1007/s11999-008-0209-4.
- [14] Bjerke-Kroll BT, Christ AB, McLawhorn AS, Sculco PK, Jules-Elysée KM, Sculco TP. Periprosthetic joint infections treated with two-stage revision over 14 years: an evolving microbiology profile. *J Arthroplasty*. 2014;29:877-882. doi:10.1016/j.arth.2013.09.053.
- [15] Nickinson RSJ, Board TN, Gambhir AK, Porter ML, Kay PR. The microbiology of the infected knee arthroplasty. *Int Orthop*. 2010;34:505-510. doi:10.1007/s00264-009-0797-y.
- [16] Uçkay I, Harbarth S, Ferry T, Lübbecke A, Emonet S, Hoffmeyer P, et al. Methicillin resistance in orthopaedic coagulase-negative staphylococcal infections. *J Hosp Infect*. 2011;79:248-253. doi:10.1016/j.jhin.2011.06.014.
- [17] Drago L, De Vecchi E, Bortolin M, Zagra L, Romanò CL, Cappelletti L. Epidemiology and antibiotic resistance of late prosthetic knee and hip infections. *J Arthroplasty*. 2017;32:2496-2500. doi:10.1016/j.arth.2017.03.005.



**Authors:** Paul M. Courtney, Nemandra A. Sandiford, Daniel Kendoff

## QUESTION 4: Is there a difference in the organism profile that causes periprosthetic joint infections (PJIs) in different countries?

**RECOMMENDATION:** Yes, there is a difference in the organism profile causing PJIs in different countries and regions of this world. There seems to be a higher incidence of PJI caused by methicillin-resistant *Staphylococcus aureus* (MRSA) in the United States and Australia compared to Europe.

**LEVEL OF EVIDENCE:** Moderate

**DELEGATE VOTE:** Agree: 97%, Disagree: 1%, Abstain: 2% (Unanimous, Strongest Consensus)

## RATIONALE

General strategies to prevent occurrence of PJIs have become more relevant over the last few years. As one recommendation of the International Consensus Meeting on Periprosthetic Joint Infection in 2013, surgical antibiotic prophylaxis with either single or 24-hour dose of cephalosporin should be performed. However, antibiotics (prophylactic and therapeutic) should be selected to cover the most frequently encountered pathogens, which might vary regionally, nationally and internationally (and could be affected as well by other factors) and not simply be administered empirically.

To date, several authors have described the bacterial incidence in isolated series of PJI with either single- or multicenter studies. However, the comparison of organism profiles causing PJI between countries or world regions has been evaluated by relatively few studies.

A study comparing organism profiles between PJI referral centers in the United States (US) (Rothman Institute) and Europe (HELIOS ENDO-Klinik) found that the percentage of MRSA pathogens was significantly higher in the US than in Europe [1]. In addition,

tion, a higher incidence of more virulent organisms was found in the US patient cohort in this study. Stefansdottir et al. and Phillips et al. in their study also found a higher incidence of coagulase-negative *Staphylococcus* (CoNS) and *Streptococcus* pathogens compared with *Staphylococcus aureus* (*S. aureus*) within various European registries (United Kingdom (UK) and Sweden) [2,3].

Peel et al. [4] showed that causative pathogens in PJI differ significantly in Australia compared to other reported studies and geographic regions such as the US, Sweden and the UK. In particular, the rates of polymicrobial infections showed high differences (36 vs. 14%), as did the isolation of MRSA (over 40% of all cases), as compared to previous European and US reports.

Pakroo et al. [5] reported similar geographic variation in organisms causing spinal infections in patients presenting to a tertiary referral center in the UK. The epidemiology and antibiotic susceptibility of bacteria causing skin and soft tissue infections do show geographic variation (e.g., between US, Germany, Italy and Spain) differentiating between MRSA, methicillin-sensitive *Staphylococcus aureus* (MSSA) and CoNS pathogens [6]. Although these data which relate predominantly to general skin infections cannot be easily transferred to PJI, it has been well accepted that such local infections (at the time of surgery or after) subsequently might lead to PJI.

Furthermore, it has been shown that community-acquired soft tissue MRSA infections have a much higher incidence in the US compared to Europe [7]. While a large percentage of soft-tissue infections are caused by community-acquired MRSA in the US, the community-acquired MRSA cutaneous infection rate in Europe only accounts for between 1 and 3% of presenting wound infections [8].

Along with this geographic variability, Anthony et al. [9] found a seasonal variability of surgical site infection (SSI) in total knee arthroplasty (TKA) and total hip arthroplasty (THA), with seasonal increase of SSI between 30 and 19% in patients with TKA or THA procedures respectively in the summer months, suggesting the possibility that geographic temperature conditions might influence the inci-

dence and etiology of PJI. This data was extracted from a US National Database.

Data from several multicenter, retrospective studies has demonstrated that the organisms causing PJI vary by country or region of the world. An increasing number of PJIs are being caused by more virulent and resistant organisms such as MRSA in the US and Australia. With the literature lacking large prospective studies, we assign a moderate recommendation.

## REFERENCES

- [1] Aggarwal VK, Bakhshi H, Ecker NU, Parvizi J, Gehrke T, Kendoff D. Organism profile in periprosthetic joint infection: pathogens differ at two arthroplasty infection referral centers in Europe and in the United States. *J Knee Surg.* 2014;27:399–406. doi:10.1055/s-0033-1364102.
- [2] Stefánsdóttir A, Johansson D, Knutson K, Lidgren L, Robertsson O. Microbiology of the infected knee arthroplasty: report from the Swedish Knee Arthroplasty Register on 426 surgically revised cases. *Scand J Infect Dis.* 2009;41:831–840. doi:10.3109/00365540903186207.
- [3] Phillips JE, Crane TP, Noy M, Elliott TSJ, Grimer RJ. The incidence of deep prosthetic infections in a specialist orthopaedic hospital: a 15-year prospective survey. *J Bone Joint Surg Br.* 2006;88:943–948. doi:10.1302/0301-620X.88B7.17150.
- [4] Peel TN, Cheng AC, Buising KL, Choong PFM. Microbiological aetiology, epidemiology, and clinical profile of prosthetic joint infections: are current antibiotic prophylaxis guidelines effective? *Antimicrob Agents Chemother.* 2012;56:2386–2391. doi:10.1128/AAC.06246-11.
- [5] Pakroo N, Mahendra M, Hemsley C, Back D, Lucas J, Sandiford N. Microbiology of spinal infections in a national tertiary referral, London: 2017.
- [6] Jones ME, Karlowsky JA, Draghi DC, Thornsberry C, Sahm DF, Nathwani D. Epidemiology and antibiotic susceptibility of bacteria causing skin and soft tissue infections in the USA and Europe: a guide to appropriate antimicrobial therapy. *Int J Antimicrob Agents.* 2003;22:406–419.
- [7] Ferry T, Etienne J. Community acquired MRSA in Europe. *BMJ.* 2007;335:947–948. doi:10.1136/bmj.39373.465903.BE.
- [8] Bode LGM, Wertheim HFL, Kluytmans J a. JW, Bogaers-Hofman D, Vandembroucke-Grauls CMJE, Roosendaal R, et al. Sustained low prevalence of methicillin-resistant *Staphylococcus aureus* upon admission to hospital in The Netherlands. *J Hosp Infect.* 2011;79:198–201. doi:10.1016/j.jhin.2011.05.009.
- [9] Anthony T, Murray BW, Sum-Ping JT, Lenkovsky F, Vornik VD, Parker BJ, et al. Evaluating an evidence-based bundle for preventing surgical site infection: a randomized trial. *Arch Surg.* 2011 Mar;146:263–269.

