

REFERENCES

- [1] Duwelius PJ, Burkhart B, Carnahan C, Branam G, Ko LM, Wu Y, et al. Modular versus nonmodular neck femoral implants in primary total hip arthroplasty: which is better? *Clin Orthop Relat Res.* 2014;472:1240-1245. doi:10.1007/s11999-013-3361-4.
- [2] Krishnan H, Krishnan SP, Blunn G, Skinner JA, Hart AJ. Modular neck femoral stems. *Bone Joint J.* 2013;95-B:1011-1021. doi:10.1302/0301-620X.95B8.31525.
- [3] U.S. Food and Drug Administration. Class 2 Device Recall Rejuvenate Modular Stems. <https://www.accessdata.fda.gov/scripts/cdrh/cfdocs/cfres/res.cfm?id=110699>. Accessed August 4, 2018.
- [4] Colas S, Allalou A, Poichotte A, Piriou P, Dray-Spira R, Zureik M. Exchangeable femoral neck (dual-modular) THA prostheses have poorer survivorship than other designs: a nationwide cohort of 324,108 patients. *Clin Orthop Relat Res.* 2017;475:2046-2059. doi:10.1007/s11999-017-5260-6.
- [5] Australian Orthopaedic Association. Annual Report 2017. <https://aoanjr.sahmri.com/en/annual-reports-2017>. Accessed August 4, 2018.
- [6] Walsh CP, Hubbard JC, Nessler JP, Markel DC. Revision of recalled modular neck rejuvenate and ABG femoral implants. *J Arthroplasty.* 2015;30:82-86. doi:10.1016/j.arth.2014.12.002.
- [7] Bernstein DT, Meflah M, Paraniilam J, Incavo SJ. Eighty-six percent failure rate of a modular-neck femoral stem design at 3 to 5 years: lessons learned. *J Bone Joint Surg Am.* 2016;98:e49. doi:10.2106/JBJS.15.01082.
- [8] Graves SE, de Steiger R, Davidson D, Donnelly W, Rainbird S, Lorimer MF, et al. The use of femoral stems with exchangeable necks in primary total hip arthroplasty increases the rate of revision. *Bone Joint J.* 2017;99-B:766-773. doi:10.1302/0301-620X.99B6.38020.
- [9] Dangles CJ, Altstetter CJ. Failure of the modular neck in a total hip arthroplasty. *J Arthroplasty.* 2010;25:1169.e5-7. doi:10.1016/j.arth.2009.07.015.
- [10] Skendzel JG, Blaha JD, Urquhart AG. Total hip arthroplasty modular neck failure. *J Arthroplasty.* 2011;26:338.e1-4. doi:10.1016/j.arth.2010.03.011.
- [11] Wilson DAJ, Dunbar MJ, Amiralet JD, Farhat Z. Early failure of a modular femoral neck total hip arthroplasty component: a case report. *J Bone Joint Surg Am.* 2010;92:1514-1517. doi:10.2106/JBJS.I.01017.
- [12] Wright G, Sporer S, Urban R, Jacobs J. Fracture of a modular femoral neck after total hip arthroplasty. *J Bone Joint Surg Am.* 2010;92:1518-1521. doi:10.2106/JBJS.I.01033.
- [13] Pelayo-de-Tomás JM, Rodrigo-Pérez JL, Novoa-Parra CD, Lizaur-Utrilla A, Morales-Suárez-Varela M, Blas-Dobón JA. Cementless modular neck stems: are they a safe option in primary total hip arthroplasty? *Eur J Orthop Surg Traumatol.* 2018;28:463-469. doi:10.1007/s00590-017-2071-0.
- [14] Cooper HJ, Urban RM, Wixson RL, Meneghini RM, Jacobs JJ. Adverse local tissue reaction arising from corrosion at the femoral neck-body junction in a dual-taper stem with a cobalt-chromium modular neck. *J Bone Joint Surg Am.* 2013;95:865-872. doi:10.2106/JBJS.L.01042.
- [15] De Martino I, Assini JB, Elpers ME, Wright TM, Westrich GH. Corrosion and fretting of a modular hip system: a retrieval analysis of 60 rejuvenate stems. *J Arthroplasty.* 2015;30:1470-1475. doi:10.1016/j.arth.2015.03.010.
- [16] Gill IPS, Webb J, Sloan K, Beaver RJ. Corrosion at the neck-stem junction as a cause of metal ion release and pseudotumour formation. *J Bone Joint Surg Br.* 2012;94:895-900. doi:10.1302/0301-620X.94B7.29122.
- [17] Grupp TM, Weik T, Bloemer W, Knaebel H-P. Modular titanium alloy neck adapter failures in hip replacement - failure mode analysis and influence of implant material. *BMC Musculoskelet Disord.* 2010;11:3. doi:10.1186/1471-2474-11-3.
- [18] Kop AM, Swarts E. Corrosion of a hip stem with a modular neck taper junction: a retrieval study of 16 cases. *J Arthroplasty.* 2009;24:1019-1023. doi:10.1016/j.arth.2008.09.009.
- [19] Restrepo C, Ross D, Restrepo S, Heller S, Goyal N, Moore R, et al. Adverse clinical outcomes in a primary modular neck/stem system. *J Arthroplasty.* 2014;29:173-178. doi:10.1016/j.arth.2014.01.040.
- [20] Su SL, Koch CN, Nguyen TM, Burket JC, Wright TM, Westrich GH. Retrieval analysis of neck-stem coupling in modular hip prostheses. *J Arthroplasty.* 2017;32:2301-2306. doi:10.1016/j.arth.2017.02.016.
- [21] Werner SD, Bono JV, Nandi S, Ward DM, Talmo CT. Adverse tissue reactions in modular exchangeable neck implants: a report of two cases. *J Arthroplasty.* 2013;28:543.e13-e15. doi:10.1016/j.arth.2012.07.026.
- [22] Nawabi DH, Do HT, Ruel A, Lurie B, Elpers ME, Wright T, et al. Comprehensive analysis of a recalled modular total hip system and recommendations for management. *J Bone Joint Surg Am.* 2016;98:40-47. doi:10.2106/JBJS.N.01121.
- [23] Panagiotidou A, Meswania J, Hua J, Muirhead-Allwood S, Hart A, Blunn G. Enhanced wear and corrosion in modular tapers in total hip replacement is associated with the contact area and surface topography. *J Orthop Res.* 2013;31:2032-2039. doi:10.1002/jor.22461.
- [24] Haddad FS, Thakrar RR, Hart AJ, Skinner JA, Nargol AVF, Nolan JF, et al. Metal-on-metal bearings: the evidence so far. *J Bone Joint Surg Br.* 2011;93:572-579. doi:10.1302/0301-620X.93B4.26429.
- [25] Bozic KJ, Lau EC, Ong KL, Vail TP, Rubash HE, Berry DJ. Comparative effectiveness of metal-on-metal and metal-on-polyethylene bearings in Medicare total hip arthroplasty patients. *J Arthroplasty.* 2012;27:37-40. doi:10.1016/j.arth.2012.03.031.
- [26] Bozic KJ, Ong K, Lau E, Kurtz SM, Vail TP, Rubash HE, et al. Risk of complication and revision total hip arthroplasty among medicare patients with different bearing surfaces. *Clin Orthop Relat Res.* 2010;468:2357-2362. doi:10.1007/s11999-010-1262-3.
- [27] Browne JA, Bechtold CD, Berry DJ, Hanssen AD, Lewallen DG. Failed metal-on-metal hip arthroplasties: a spectrum of clinical presentations and operative findings. *Clin Orthop Relat Res.* 2010;468:2313-2320. doi:10.1007/s11999-010-1419-0.
- [28] Judd KT, Noiseux N. Concomitant infection and local metal reaction in patients undergoing revision of metal on metal total hip arthroplasty. *Iowa Orthop J.* 2011;31:59-63.
- [29] Prieto HA, Berbari EF, Sierra RJ. Acute delayed infection: increased risk in failed metal on metal total hip arthroplasty. *J Arthroplasty.* 2014;29:1808-1812. doi:10.1016/j.arth.2014.04.008.
- [30] Hosman AH, van der Mei HC, Bulstra SK, Busscher HJ, Neut D. Effects of metal-on-metal wear on the host immune system and infection in hip arthroplasty. *Acta Orthop.* 2010;81:526-534. doi:10.3109/17453674.2010.519169.
- [31] de Steiger RN, Hang JR, Miller LN, Graves SE, Davidson DC. Five-year results of the ASR XL Acetabular System and the ASR Hip Resurfacing System: an analysis from the Australian Orthopaedic Association National Joint Replacement Registry. *J Bone Joint Surg Am.* 2011;93:2287-2293. doi:10.2106/JBJS.I.01727.
- [32] Ogunwale B, Schmidt-Ott A, Meek RMD, Brewer JM. Investigating the immunologic effects of CoCr nanoparticles. *Clin Orthop Relat Res.* 2009;467:3010-3016. doi:10.1007/s11999-009-0949-9.
- [33] Engh CA, Ho H, Engh CA. Metal-on-metal hip arthroplasty: does early clinical outcome justify the chance of an adverse local tissue reaction? *Clin Orthop Relat Res.* 2010;468:406-412. doi:10.1007/s11999-009-1063-8.
- [34] Mikhael MM, Hanssen AD, Sierra RJ. Failure of metal-on-metal total hip arthroplasty mimicking hip infection. A report of two cases. *J Bone Joint Surg Am.* 2009;91:443-446. doi:10.2106/JBJS.H.00603.
- [35] Bonanzinga T, Zahar A, Dütsch M, Lausmann C, Kendoff D, Gehrke T. How reliable is the alpha-defensin immunoassay test for diagnosing periprosthetic joint infection? A prospective study. *Clin Orthop Relat Res.* 2017;475:408-415. doi:10.1007/s11999-016-4906-0.
- [36] Okroj KT, Calkins TE, Kayupov E, Kheir MM, Bingham JS, Beauchamp CP, et al. The alpha-defensin test for diagnosing periprosthetic joint infection in the setting of an adverse local tissue reaction secondary to a failed metal-on-metal bearing or corrosion at the head-neck junction. *J Arthroplasty.* 2018;33:1896-1898.
- [37] Aljaniipour P, Adeli B, Hansen EN, Chen AF, Parvizi J. Intraoperative purulence is not reliable for diagnosing periprosthetic joint infection. *J Arthroplasty.* 2015;30:1403-1406. doi:10.1016/j.arth.2015.03.005.
- [38] Parvizi J, Tan TL, Goswami K, Higuera C, Della Valle C, Chen AF, et al. The 2018 definition of periprosthetic hip and knee infection: an evidence-based and validated criteria. *J Arthroplasty.* 2018;33:1309-1314.e2. doi:10.1016/j.arth.2018.02.078.

● ● ● ● ●

Authors: Kevin Perry, Alisina Shahi

QUESTION 6: Can implant factors (i.e., type of bearing) influence the thresholds for serum and synovial markers in acute and chronic periprosthetic joint infections (PJIs)?

RECOMMENDATION: Yes. Different bearing surfaces such as metal-on-metal (MoM), metal-on-polyethylene and dual taper modular stems in the setting of taper corrosion can influence the serum and synovial markers. Metal debris may interfere with automated cell counts. Manual cell counts are preferred when evaluating patients for PJIs who have elevated synovial fluid metal levels. Optimal thresholds for serum and synovial markers for diagnosing PJIs in these settings still need to be established.

LEVEL OF EVIDENCE: Moderate

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

RATIONALE

Implant factors such as bearing surfaces can influence serum and synovial markers when evaluating for PJIs. This has been mostly studied in MoM bearings and dual taper modular stems [1–3]. It can be difficult to discern adverse local tissue reactions (ALTRs) with associated metal ion release from inflammatory response to infection [4,5]. However, it is important to determine the presence of infection as it will alter treatment [6,7]. Serum erythrocyte sedimentation rate (ESR), C-reactive protein (CRP) and white blood cell (WBC) count with differential are important tests in helping determine presence of PJI [8].

There have been various recommendations regarding the parameters for serum and synovial markers for diagnosing PJI in the presence of MoM corrosion, but most studies have demonstrated that the type of bearing surface and other implant factors can affect the thresholds for serum and synovial markers in PJI. Still, no literature has clearly delineated the specific parameters that should be utilized for differing bearing surfaces to diagnose PJI [9,10].

Automated synovial cell counts and differentials in the setting of a failed MoM THA have been reported to be inaccurate [2,3,11]. It has been theorized that the automated cell counting machine may be incorrectly identifying particulate debris and counting it as cellular [2]. As such, many surgeons propose utilizing a manual cell count and differential when analyzing the synovial WBC and differential [1].

Wyles et al. [2] found that the sensitivity of the synovial WBC count could be maintained at 100% while improving specificity to 71% if the cutoff to diagnose infection was moved from >3,000 to >15,000 cells/microliter. Additionally, the authors found the sensitivity of neutrophil percentage could be maintained at 100% and improved specificity to 100% by elevating the cutoff percentage from 82 to 92% neutrophils. Regarding CRP, the authors found that the sensitivity of CRP could be maintained at 75% while improving the specificity of CRP to 97% if the cutoff value of CRP was raised from >8 to >54 mg/L. The authors demonstrated that changing the cutoff value for the ESR did not change specificity as significantly.

In contrast, Yi et al. [3] studied PJI in patients with failed MoM bearing surfaces and after excluding what they deemed to be inaccuracies, recommended a synovial WBC cutoff of 4,350 WBC/microliter with 100% sensitivity and 95% specificity. The authors, however, reported low positive predictive values of 43% and 39% for ESR and CRP, respectively, in the setting of MoM bearings.

Kwon et al. reported that ESR and CRP have a limited value in the diagnosis of PJI in dual taper modular implants with evidence of corrosion, but acknowledged the utility of ESR and CRP in excluding PJI [1]. The authors demonstrated, however, that synovial WBC and differential were useful markers for diagnosing infection. Specifically, the authors demonstrated a sensitivity and specificity of 86% and 80%, respectively, when utilizing a synovial WBC cutoff of 730 cells/microliter. A synovial polymorphonuclear (PMN) % cutoff of 65% yielded a 100% sensitivity and a 70% sensitivity.

Okroj et al. in a multicenter study evaluated the alpha-defensin test to diagnose PJI in the setting of ALTRs. Twenty-six patients were reviewed with one of 26 (3.8%) meeting the MusculoSkeletal Infection Society (MSIS) criteria for PJI. The one patient with PJI had a metal-on-polyethylene bearing surface with head-neck taper corrosion. Of note, there were 8 falsely positive alpha-defensin tests. The authors concluded that in the setting on ALTRs, alpha-defensin testing can lead to a high rate of false positives [12].

Though the exact parameters to diagnose PJI in the setting of different implant factors need further elucidation, given the existing literature, we conclude that various implant factors can influence both synovial and serum markers in the setting of PJI. We strongly urge the orthopaedic community to be cognizant of the influence of bearing surfaces, especially in the setting of MoM implants or potential metal corrosion, and to consider using a combination of diagnostic tests along with manual cell counts as part of their PJI diagnostic workup.

REFERENCES

- [1] Kwon YM, Antoci V, Leone WA, Tsai TY, Dimitriou D, Liow MHL. Utility of serum inflammatory and synovial fluid counts in the diagnosis of infection in taper corrosion of dual taper modular stems. *J Arthroplasty*. 2016;31:1997–2003. doi:10.1016/j.arth.2016.02.020.
- [2] Wyles CC, Larson DR, Houdek MT, Sierra RJ, Trousdale RT. Utility of synovial fluid aspirations in failed metal-on-metal total hip arthroplasty. *J Arthroplasty*. 2013;28:818–823. doi:10.1016/j.arth.2012.11.006.
- [3] Yi PH, Cross MB, Moric M, Levine BR, Sporer SM, Paprosky WG, et al. Do serologic and synovial tests help diagnose infection in revision hip arthroplasty with metal-on-metal bearings or corrosion? *Clin Orthop Relat Res*. 2015;473:498–505. doi:10.1007/s11999-014-3902-5.
- [4] Judd KT, Noiseux N. Concomitant infection and local metal reaction in patients undergoing revision of metal on metal total hip arthroplasty. *Iowa Orthop J*. 2011;31:59–63.
- [5] Watters TS, Eward WC, Hallows RK, Dodd LG, Wellman SS, Bolognesi MP. Pseudotumor with superimposed periprosthetic infection following metal-on-metal total hip arthroplasty: a case report. *J Bone Joint Surg Am*. 2010;92:1666–1669. doi:10.2106/JBJS.I.01208.
- [6] Mabilletau G, Kwon Y-M, Pandit H, Murray DW, Sabokbar A. Metal-on-metal hip resurfacing arthroplasty: a review of periprosthetic biological reactions. *Acta Orthop*. 2008;79:734–747. doi:10.1080/17453670810016795.
- [7] Darouiche RO. Treatment of infections associated with surgical implants. *N Engl J Med*. 2004;350:1422–1429. doi:10.1056/NEJMra035415.
- [8] Parvizi J, Zmistowski B, Berbari EF, Bauer TW, Springer BD, Della Valle CJ, et al. New definition for periprosthetic joint infection: from the Workgroup of the Musculoskeletal Infection Society. *Clin Orthop Relat Res*. 2011;469:2992–2994. doi:10.1007/s11999-011-2102-9.
- [9] Schinsky MF, Della Valle CJ, Sporer SM, Paprosky WG. Perioperative testing for joint infection in patients undergoing revision total hip arthroplasty. *JBJS*. 2008;90:1869. doi:10.2106/JBJS.G.01255.
- [10] Parvizi J, Adeli B, Zmistowski B, Restrepo C, Greenwald AS. Management of periprosthetic joint infection: the current knowledge: AAOS exhibit selection. *J Bone Joint Surg Am*. 2012;94:e104. doi:10.2106/JBJS.K.01417.
- [11] Alijanipour P, Adeli B, Hansen EN, Chen AF, Parvizi J. Intraoperative purulence is not reliable for diagnosing periprosthetic joint infection. *J Arthroplasty*. 2015;30:1403–1406. doi:10.1016/j.arth.2015.03.005.
- [12] Okroj KT, Calkins TE, Kayupov E, Kheir MM, Bingham JS, Beauchamp CP, et al. The alpha-defensin test for diagnosing periprosthetic joint infection in the setting of an adverse local tissue reaction secondary to a failed metal-on-metal bearing or corrosion at the head-neck junction. *J Arthroplasty*. 2018;33:1896–1898.

