G29 – Are former COVID-19 patients undergoing major orthopedic procedures at risk for subsequent periprosthetic joint infection (PJI) and surgical site infection (SSI)?

# **Response/Recommendation:**

Former COVID-19 patients undergoing major orthopedic procedures have a significantly increased risk of periprosthetic joint infection (PJI) and surgical site infection (SSI), particularly when COVID-19 infection occurs preoperatively or intraoperatively. Postoperative COVID-19 infection appears to have no statistically significant association with PJI risk but remains associated with a higher probability of SSI.

**Strength of recommendation:** Strong

### **Delegate Vote:**

### **Rationale:**

The COVID-19 pandemic is one of the greatest health challenges of the 21st century, imposing a significant social and economic burden, particularly on vulnerable populations such as the elderly, with chronic diseases, and those requiring urgent medical interventions, including trauma care (1-4). Despite several articles have highlighted the negative impact of the pandemic on the outcomes of orthopedic surgeries including a high morality rate (5-9), the effect of COVID-19 on the incidence of PJI and SSI is still unclear and no comprehensive review has investigated the effect of COVID-19 on PJI and SSI after orthopedic surgeries (7, 10-17). We conducted a systematic review and meta-analysis to investigate the effect of COVID-19 based on the time of infection (Pre-op, during and post-op) on the outcomes of PJI and SSI after orthopedic surgeries. The inclusion criteria consisted of observational studies examining the effect of patients presenting with COVID-19 on PJI and SSI complications following major orthopedic surgeries, including all fractures and joint replacements with at least one month of follow-up. Preoperative, intraoperative, and postoperative COVID-19 were defined based on a positive test within 90 days before surgery, on the day of surgery, and within 90 days after surgery, respectively. Seventeen studies met the inclusion criteria for analysis. 19157(2.1%) of the patients had a positive perioperative COVID-19 test. The pooled prevalence of PJI and SSI in COVID-19-positive patients who underwent orthopedic surgery was 1.8% (95% CI: 1.2%-2.5%) and 2% (95% CI: 1.5%–2.5%), regardless of the timing of COVID-19 positivity. Pooled estimation of 13 studies showed that, regardless of the time of infection, COVID-19 positivity was associated with an increased probability of PJI (OR: 2.25, 95% CI: 2.68, 2.83, I<sup>2</sup>:93.7%), and SSI (OR: 2.01, 95% CI: 1.74, 2.28, I<sup>2</sup>:73.6%), in major orthopedic surgeries. The probability of PJI occurrence was significantly higher in patients who tested positive for COVID-19 preoperatively and intraoperatively compared to those without COVID-19. However, the probability of PJI was not significantly associated with post-operative COVID-19 positivity (OR: 1.5, 95% CI: 0.67–2.33, I<sup>2</sup>: 76.3%). The probability of SSI in patients with COVID-19 preoperatively, intraoperatively, and postoperatively was significantly higher compared to those without COVID-19.

Several studies provide insights into the potential risks and mechanisms by which a history of COVID-19 infection impacts SSI and PJI in orthopedic patients. COVID-19 leads to prolonged immune system dysregulation, including hyperinflammation, T-cell dysfunction, and increased prothrombotic states. These factors collectively impair wound healing and increase infection susceptibility(18, 19). A recent study showed significantly elevated risks for venous thromboembolism (VTE), sepsis, and 30-day mortality in patients undergoing spinal surgery within 2 weeks of a COVID-19 diagnosis. However, patients operated on beyond 6 weeks showed no significant differences in complications, highlighting the importance of delaying surgery(19). COVID-19-associated systemic inflammation and vascular damage may predispose patients to SSI and PJI. For example, one study reported a 4.65-fold increase in PJI risk after total joint arthroplasty in patients with a postoperative COVID-19 diagnosis(18). An analysis of Japan's national surgical database found that heightened hygiene protocols during the pandemic had no significant effect on overall SSI rates in orthopedic surgeries, suggesting the need for targeted infection prevention strategies (20). Studies recommend postponing elective orthopedic procedures for at least 6-12 weeks post-COVID-19 diagnosis to allow for immune recovery and stabilization. Another study evaluating trauma patients undergoing surgery demonstrated that the presence of COVID-19 significantly increases the likelihood of perioperative complications, including infections, due to systemic inflammation(21). Finally, COVID-19 protocols implemented during primary and revision total hip arthroplasty highlighted the importance of institutional infection control measures and their role in mitigating risk(21, 22). These findings emphasize the need for a cautious approach in managing former COVID-19 patients undergoing major orthopedic procedures, balancing surgical urgency with the risks of postoperative complications.

Meta-analyses show that COVID-19 significantly increases postoperative mortality and complications in orthopedic trauma patients. KC Wang et al. (7) reported a 32.6% early mortality rate in hip fracture patients, 5.66 times higher than non-COVID-19 patients, while another study found a 7.72-fold increase in mortality (8). SK Tripathy et al. (9) and VK Jain et al. (23) confirmed higher risks of mortality, pneumonia, severe complications, and prolonged hospitalization in COVID-19-positive patients. These outcomes are linked to advanced age, immunosuppression, inactivity, and limited rehabilitation, increasing risks of PE, DVT, PJI, and SSI (24-31).

## **Conclusion:**

This meta-analysis revealed a higher prevalence of PJI and SSI in COVID-19-infected patients undergoing orthopedic surgery compared to those without COVID-19. Infection during the preoperative and perioperative periods increased the risk of PJI in major orthopedic procedures by over twofold, while postoperative infection showed no significant association. Additionally, COVID-19 increased the likelihood of SSI across all periods. Future research should explore the optimal timing for elective surgeries after COVID-19 and develop preventive strategies to reduce PJI and SSI risks.

#### **References**:

- 1. Mohammadpour M, Yazdi H, Bagherifard A, Jabalameli M, Moghtadaei M, Torkaman A, et al. Evaluation of early complications, outcome, and mortality in Coronavirus Disease 2019 (COVID-19) infection in patients who underwent orthopedic surgery. BMC Musculoskeletal Disorders. 2022;23(1):64.
- 2. O'Connor CM, Anoushiravani AA, DiCaprio MR, Healy WL, Iorio R. Economic recovery after the COVID-19 pandemic: resuming elective orthopedic surgery and total joint arthroplasty. The Journal of arthroplasty. 2020;35(7):S32-S6.
- 3. Park C, Sugand K, Nathwani D, Bhattacharya R, Sarraf KM. Impact of the COVID-19 pandemic on orthopedic trauma workload in a London level 1 trauma center: the "golden month" The COVid Emergency Related Trauma and orthopaedics (COVERT) Collaborative. Acta orthopaedica. 2020;91(5):556-61.
- 4. Global guidance for surgical care during the COVID-19 pandemic. Journal of British Surgery. 2020;107(9):1097-103.
- 5. North T, Bullock MW, Danoff JR, Saxena A, Fischer SJ, Stronach BM, et al. Arthroplasty during the COVID-19 pandemic. Arthroplasty Today. 2020;6(3):427-30.
- 6. Humphrey T, Daniell H, Chen AF, Hollenbeck B, Talmo C, Fang CJ, et al. Effect of the COVID-19 Pandemic on Rates of Ninety-Day Peri-Prosthetic Joint and Surgical Site Infections after Primary Total Joint Arthroplasty: A Multicenter, Retrospective Study. Surgical Infections. 2022;23(5):458-64.
- 7. Wang KC, Xiao R, Cheung ZB, Barbera JP, Forsh DA. Early mortality after hip fracture surgery in COVID-19 patients: A systematic review and meta-analysis. Journal of Orthopaedics. 2020;22:584-91.
- 8. Putera HD, Halim V, Panghiyangani R. The impact of COVID-19 on mortality in trauma patients undergoing orthopedic surgery: a systematic review and meta-analysis. Clin Exp Emerg Med. 2023;10(3):315-26.
- 9. Tripathy SK, Varghese P, Panigrahi S, Panda BB, Srinivasan A, Sen RK. Perioperative mortality and morbidity of hip fractures among COVID-19 infected and non-infected patients: A systematic review and meta-analysis. Chinese Journal of Traumatology. 2023;26(03):162-73.
- 10. Forlenza EM, Higgins JDD, Burnett RA, Serino J, Della Valle CJ. COVID-19 Infection After Total Joint Arthroplasty Is Associated With Increased Complications. The Journal of Arthroplasty. 2022;37(7, Supplement):S457-S64.
- 11. Chokshi SN, Somerson JS. Preoperative COVID-19 Infection Increases Risk for 60-Day Complications Following Total Shoulder Arthroplasty: A Propensity-Matched Analysis. Journal of Shoulder and Elbow Surgery. 2024.
- 12. Wenzel AN, Marrache M, Schmerler J, Kinney J, Khanuja HS, Hegde V. Impact of Postoperative COVID-19 Infection Status on Outcomes in Elective Primary Total Joint Arthroplasty. The Journal of Arthroplasty. 2024;39(4):871-7.
- 13. Jao W-Y, Lai C-C, Chien C-S, Li Y-J. The Impact of Coronavirus Disease 2019 on the Postoperative Complication of Pediatric Orthopedic Surgery: A Multicenter Retrospective Cohort Study. Formosan Journal of Musculoskeletal Disorders. 2024;15(1):28-35.
- 14. Rosas S, Pollock DC, Roche MW, Najafi F, Hollingsworth N, Buller LT, et al. Patients With Previous COVID-19 Infection Can Safely Undergo Primary Total Joint Arthroplasty. The Journal of Arthroplasty. 2023;38(4):649-54.
- 15. Mercier MR, Koucheki R, Lex JR, Khoshbin A, Park SS, Daniels TR, et al. The association between preoperative COVID-19-positivity and acute postoperative complication

- risk among patients undergoing orthopedic surgery: a matched cohort analysis. Bone & Joint Open. 2023;4(9):704-12.
- 16. Bains SS, Hameed D, Dubin JA, Chen Z, Rizzo Jr SA, Van Nielen DL, et al. The temporal effects of asymptomatic COVID-19 infection on peri-operative complications in patients receiving total joint arthroplasty. Archives of Orthopaedic and Trauma Surgery. 2023;143(11):6799-804.
- 17. Baldwin AJ, Jackowski A, Jamal A, Vaz J, Rodrigues JN, Tyler M, et al. Risk of surgical site infection in hand trauma, and the impact of the SARS-CoV-2 pandemic: A cohort study. Journal of Plastic, Reconstructive & Aesthetic Surgery. 2021;74(11):3080-6.
- 18. Heckmann ND, Wang JC, Piple AS, Bouz GJ, Chung BC, Oakes DA, et al. Positive COVID-19 diagnosis following primary elective total joint arthroplasty: increased complication and mortality rates. The Journal of Arthroplasty. 2023;38(9):1682-92. e2.
- 19. Chan JP, Hoang H, Hashmi SZ, Lee Y-P, Bhatia NN, Consortium NC. A temporal analysis of perioperative complications following COVID-19 infection in patients undergoing lumbar spinal fusion: When is it safe to proceed? North American Spine Society Journal (NASSJ). 2023;16:100262.
- 20. Mimura T, Matsumoto G, Natori T, Ikegami S, Uehara M, Oba H, et al. Impact of the COVID-19 pandemic on the incidence of surgical site infection after orthopaedic surgery: an interrupted time series analysis of the nationwide surveillance database in Japan. Journal of Hospital Infection. 2024;146:160-5.
- 21. Sybert M, Oakley CT, Christensen T, Bosco J, Schwarzkopf R, Slover J. Impact of COVID-19 protocols on primary and revision total hip arthroplasty. The Journal of Arthroplasty. 2022;37(11):2193-8.
- 22. Rincón-Hoyos JA, Vallejo-Yepes P, Restrepo-Giraldo JN, Torres-Valencia H, Buitrago-Vanegas M, Valencia-Rivas M, et al. Morbidity and mortality in hip surgery patients due to fracture during the COVID-19 pandemic. Injury. 2023;54:110731.
- 23. Jain VK, Lal H, Patralekh MK, Vaishya R. Fracture management during COVID-19 pandemic: a systematic review. Journal of clinical orthopaedics and trauma. 2020;11:S431-S41.
- 24. Goodman SB, Konttinen YT, Takagi M. Joint replacement surgery and the innate immune system. Journal of long-term effects of medical implants. 2014;24(4).
- 25. Cusack B, Buggy D. Anaesthesia, analgesia, and the surgical stress response. BJA education. 2020;20(9):321-8.
- 26. Marik PE, Flemmer M. The immune response to surgery and trauma: Implications for treatment. Journal of Trauma and Acute Care Surgery. 2012;73(4):801-8.
- 27. Desborough J. The stress response to trauma and surgery. British journal of anaesthesia. 2000;85(1):109-17.
- 28. Finnerty CC, Mabvuure NT, Ali A, Kozar RA, Herndon DN. The surgically induced stress response. Journal of parenteral and enteral nutrition. 2013;37:21S-9S.
- 29. Stone J, Hangge P, Albadawi H, Wallace A, Shamoun F, Knuttien MG, et al. Deep vein thrombosis: pathogenesis, diagnosis, and medical management. Cardiovascular diagnosis and therapy. 2017;7(Suppl 3):S276.
- 30. McLendon K, Goyal A, Attia M. Deep venous thrombosis risk factors. 2017.

31. Groff D, Sun A, Ssentongo AE, Ba DM, Parsons N, Poudel GR, et al. Short-term and long-term rates of postacute sequelae of SARS-CoV-2 infection: a systematic review. JAMA network open. 2021;4(10):e2128568-e.