HK68: Is there a role for abbreviated 2 stage exchange arthroplasty?

James Cashman, Scott Matthews, Bryan Springer, Thomas Hilton, Christian Hipfl, Tobias Winkler, Vasily Artyukh

RESPONSE/RECOMMENDATION – A number of studies demonstrate that early reimplantation provides similar outcomes to the traditional two stage exchange with a reimplantation window of 4-12 weeks. The role of high-dose local antibiotics to improve clinical outcomes and criteria for selecting suitable candidates have yet to be defined.

LEVEL OF EVIDENCE: Limited

DELEGATE VOTE:

RATIONALE

The growing popularity for single stage exchange arthroplasty, raises some questions regarding two-stage exchange arthroplasty and the protocols associated with the latter surgical procedure. One issue related to two-stage exchange arthroplasty relates to the optimal timing of reimplantation. No definite timing window for reimplantation has been identified in the literature to date. With most studies quoting between 4-12 weeks as the optimal window 2, a common classification by Trampuz and Zimmerli defines intervals of 2–4 weeks (a short interval) and 6–8 weeks (a long interval) until reimplantation. Some studies suggest an interval period of 4–6 weeks or 9 weeks between the stages. Clinical studies range from a few days to several hundred days or even years, however, an interval of around 60–100 days is mostly reported. 6-13 This heterogeneity in clinical practice indicates that the optimal duration of the interval between the stages has yet to be defined. 2

There is a paucity of evidence to support earlier than standard long interval reimplantation at 2nd stage. Even in literature, where this early reimplantation has been investigated, there is variable timing to 2nd stage with no consistency across the board. In a cohort of 32 periprosthetic hip infections, Lieberman et al. report no difference between reimplantation at 6 weeks versus 1 year. ¹⁴ Haddad et al showed no compromise of success with early reimplantation at 3 weeks in 32 total hip replacements. ¹⁵ Burnett et al also reports similar success with a shortened interval of 3-4 weeks compared to 4-6 weeks. ¹⁶

Winkler et al. prospectively compared reimplantation at less than 4 weeks (mean 19.3 days, range,7-27 days) versus longer than 4 weeks (mean 63 days, range, 28-204 days) for 38 patients undergoing second stage PJI. Their group used non biofilm active antibiotics between stages and highly active biofilm- antibiotic treatment following reimplantation. Half the patients were included in the short interval group with no reinfections at a mean follow up of 40 months. One reinfection occurred in the longer interval group.⁸

However, some studies contradict the above and suggest that early reimplantation may compromise outcome. Vielgut et al. retrospectively analysed 72 hip PJI and 4 native hip infection patients operated between 2005 and 2010. All cases were infected but 4 were native joint infections. They found that 90% of a cohort of 40 patients undergoing reimplantation between 4 to 11 weeks remained infection free. The reinfection was higher in patients undergoing early reimplantation (< 4 weeks) or later than 11 weeks. Five patients were in the less than 4 weeks group all of which got reinfected and 23 in the greater than 11 weeks group

Commented [WT1]: Here we should suggest more refs as there is also only one ref given but the text says "some"

Commented [WT2]: This has to be cross-checked as we have seen a lot of intervals of 6-8 weeks, which would be 42-56 days and not 80-100, which seems too long for this statement. I also would suggest to give the references for the statement "Clinical studies range from a few days to several hundred days or even several years " and "an interval of around 80–100 days is mostly reported" separately.

with a 47.8% reinfection rate. Since the study was retrospective, no clear inclusion and exclusion criteria could be applied making the results difficult to interpret. ¹⁷

In recent years the role of abbreviated two-stage exchange arthroplasty has been investigated in a number of prospective studies. A study has evaluated implantation of an antibiotic drug eluting device, instead of a spacer, following resection arthroplasty ¹⁸. In these cases intra-articular antibiotics, as well as systemic antibiotics were administered. After one week the device was then removed and reimplantation was carried out. The outcome of this protocol has been very encouraging. Numerous presentations and pending publications demonstrate that the local delivery of antibiotics allows for early reimplantation without compromising the outcome of two-stage exchange.

Based on the current available literature, the optimal timing for reimplantation remains unknown. One of the until now used standard protocols of administering six weeks of antibiotics, followed by a two-week drug holiday, and subsequent reimplantation needs to be examined since shorter intervals might have an advantage. The two-week drug holiday is based on little evidence and has also been shown in some studies to be outdated. ^{19,20} Accurate and cost-effective tests for optimal reimplantation timing are still needed. Beyond that, however, there is a need to work on selection criteria for choosing patients, in which an abbreviated two-stage protocol yield similar or improved clinical outcomes compared to traditional long interval reimplantation.

REFERENCES

- 1. Zhao, Y., Fan, S., Wang, Z., Yan, X. and Luo, H. (2024). Systematic review and meta-analysis of single-stage vs two-stage revision for periprosthetic joint infection: a call for a prospective randomized trial. *BMC musculoskeletal disorders*, 25(1). doi:https://doi.org/10.1186/s12891-024-07229-z.
- Puetzler, J., Hofschneider, M., Georg Gosheger, Theil, C., Schulze, M., Schwarze, J., Koch, R. and Burkhard Moellenbeck (2024). Evaluation of time to reimplantation as a risk factor in two-stage revision with static spacers for periprosthetic knee joint infection. *Journal of Orthopaedics and Traumatology*, 25(1). doi:https://doi.org/10.1186/s10195-024-00745-7.
- Trampuz, A. and Zimmerli, W. (2008). Diagnosis and treatment of implant-associated septic arthritis and osteomyelitis. *Current Infectious Disease Reports*, 10(5), pp.394– 403. doi:https://doi.org/10.1007/s11908-008-0064-1.
- Warth, L.C., Hadley, C.J. and Grossman, E.L. (2020). Two-Stage Treatment for Total Knee Arthroplasty Infection Utilizing an Articulating Prefabricated Antibiotic Spacer. *The Journal of Arthroplasty*, 35(3), pp.S57–S62. doi:https://doi.org/10.1016/j.arth.2019.10.049.
- Cooper, H.J. and Della Valle, C.J. (2013). The two-stage standard in revision total hip replacement. *The Bone & Joint Journal*, 95-B(11_Supple_A), pp.84–87. doi:https://doi.org/10.1302/0301-620x.95b11.32906.
- Dieckmann, R., Schmidt-Braekling, T., Georg Gosheger, Theil, C., Jendrik Hardes and Burkhard Moellenbeck (2019). Two stage revision with a proximal femur

Commented [WT3]: the 4 cases with native hip joint infections are not really biasing heavily since they are only 4 out of 76 which is why i suggest to leave the last sentence out. If these were among the less than 4 weeks group then it would be relevant and makes this group uninterpretable.

Commented [WT4]: the 4 cases with native hip joint infections are not really biasing heavily since they are only 4 out of 76 which is why i suggest to leave the last sentence out. If these were among the less than 4 weeks group then it would be relevant and makes this group uninterpretable.

- replacement. *BMC Musculoskeletal Disorders*, 20(1). doi:https://doi.org/10.1186/s12891-019-2442-2.
- Vasarhelyi, E., Sidhu, S.P., Somerville, L., Lanting, B., Naudie, D. and Howard, J. (2022). Static vs Articulating Spacers for Two-Stage Revision Total Knee Arthroplasty: Minimum Five-Year Review. *Arthroplasty Today*, 13, pp.171–175. doi:https://doi.org/10.1016/j.artd.2021.10.010.
- Winkler T et al (2019) Outcome of short versus long interval in two-stage exchange for periprosthetic joint infection: a prospective cohort study. Arch Orthop Trauma Surg 139(3):295–303
- 9. Aali Rezaie A et al (2018) Time to reimplantation: waiting longer confers no added benefit. J Arthroplasty 33(6):1850–1854
- 10. Sabry FY et al (2014) Preoperative prediction of failure following two-stage revision for knee prosthetic joint infections. J Arthroplasty 29(1):115–121
- Hart WJ, Jones RS (2006) Two-stage revision of infected total knee replacements using articulating cement spacers and short-term antibiotic therapy. J Bone Joint Surg Br 88(8):1011–1015
- 12. Babis GC et al (2015) Two-stage revision protocol in multidrug resistant periprosthetic infection following total hip arthroplasty using a long interval between stages. J Arthroplasty 30(9):1602–1606
- 13. Choi HR et al (2011) Can implant retention be recommended for treat ment of infected TKA? Clin Orthop Relat Res 469(4):961–969
- 14. Lieberman, J.R., Callaway, G.H., Salvati, E.A., Pellicci, P.M. and Brause, B.D. (1994). Treatment of the infected total hip arthroplasty with a two-stage reimplantation protocol. *PubMed*, (301), pp.205–12.
- Haddad, F.S., Muirhead-Allwood, S., Manktelow, A. and Bacarese-Hamilton, I. (2000). Two-stage uncemented revision hip arthroplasty for infection. *The journal of bone and joint surgery*, 82(5), pp.689–694. doi:https://doi.org/10.1302/0301-620x.82b5.9668.
- Burnett, R., Kelly, M.A., Hanssen, A.D. and Barrack, R.L. (2007). Technique and Timing of Two-stage Exchange for Infection in TKA. 464, pp.164–178. doi:https://doi.org/10.1097/blo.0b013e318157eb1e.
- 17. Vielgut, I., Sadoghi, P., Wolf, M., Holzer, L., Leithner, A., Schwantzer, G., Poolman, R., Frankl, B. and Glehr, M. (2015). Two-stage revision of prosthetic hip joint infections using antibiotic-loaded cement spacers: When is the best time to perform the second stage? *International Orthopaedics*, 39(9), pp.1731–1736. doi:https://doi.org/10.1007/s00264-015-2751-5.
- 18. Springer BD, Higuera-Rueda CA, de Beaubien BC, Warner KD, Glassman AH, Parvataneni HK, Piuzzi NS. Safety Profile of Seven-Day Intra-articular Antibiotic Irrigation for the Treatment of Chronic Periprosthetic Joint Infection: A Prospective Randomized Phase II Comparative Study. J Arthroplasty. 2024 Sep;39(9S1):S229-S235.e1. doi: 10.1016/j.arth.2024.03.069. Epub 2024 Apr 9. PMID: 38604274.
- 19. Janz V, Bartek B, Wassilew GI, Stuhlert M, Perka CF, Winkler T. Validation of Synovial Aspiration in Girdlestone Hips for Detection of Infection Persistence in

- Patients Undergoing 2-Stage Revision Total Hip Arthroplasty. J Arthroplasty. 2016 Mar;31(3):684-7. doi: 10.1016/j.arth.2015.09.053. Epub 2015 Oct 9. PMID: 26521130.
- 20. Preininger B, Janz V, von Roth P, Trampuz A, Perka CF, Pfitzner T. Inadequacy of Joint Aspiration for Detection of Persistent Periprosthetic Infection During Two-Stage Septic Revision Knee Surgery. Orthopedics. 2017 Jul 1;40(4):231-234. doi: 10.3928/01477447-20170411-04. Epub 2017 Apr 18. PMID: 28418574.