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Systematic review of single stage revision for prosthetic joint infection

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Author contributions: Lum ZC designed and performed the research; Lum ZC and Holland CT analyzed the data; Lum ZC, Meehan JP, and Holland CT wrote and were responsible for revising the paper; Lum ZC and Meehan PJ supervised the manuscript. All authors read and approved the manuscript.

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Abstract

While advanced technology, increased medical knowledge and improved surgical technique has improved patient outcomes in total joint arthroplasty, prosthetic joint infection still remains one of the leading causes of increased healthcare costs, medical resources and societal burdens in orthopaedic care. Two stage arthroplasty revision remains the gold standard for treatment of prosthetic joint infection. Proponents of single stage revision arthroplasty for infection argue that it results in lower healthcare costs while improving patient reported functional outcomes and with equivalent success rates. Here we review the history of single stage revision arthroplasty, discuss the key principles, highlight the indications and contraindications, and review the reported outcomes with a focus on future developments of single stage revision arthroplasty for hip and knee periprosthetic joint infections.

Key Words: Prosthetic joint infection; Single stage; Revision total joint arthroplasty; Total knee arthroplasty; Infection

Core Tip: Single stage revision for periprosthetic joint infection can be a successful operation with careful selection of the patient, infecting organism and precise surgical technique. The 3 key principles of bacterial sensitivities, thorough radial debridement and delivery of local and systemic antibiotics can result in similar infection free survivorship to two stage exchange. Future developments into this technique include its practice in culture negative infections or use of cementless implants. Randomized
INTRODUCTION

In the early 1970s Dr. Hans Wilhelm Buchholz published extensive research into the properties of antibiotics and polymethylmethacrylate with respect to hip and knee replacement surgery. There, he was able to report consistently lower infection rates with addition of gentamycin antibiotics to the bone cement [1]. He was one of the first to report successful prophylactic treatment of endoprostheses with antibiotic loaded bone cement as well as surgical treatment by single exchange arthroplasty for infected prostheses [2]. Buchholz’s experience was unique as treatment of prosthetic joint infections (PJI) were evolving during this time and two-stage exchange arthroplasty was proving to have improved infection control in other parts of the world.

In the modern treatment paradigm, two-stage exchange arthroplasty for PJI is the gold standard treatment. However, more centers around the world are challenging that concept by reporting their successful results of single stage exchange arthroplasty, with techniques and principles similar to the original ones described by Buchholz’s center at the ENDO Klinik in Hamburg, Germany [3-6]. With a globalization of education and technology, improvements to this process has allowed for a better understanding of single stage exchange arthroplasty. We report the current literature regarding key principles, indications, contraindications, outcomes and future direction of single stage revision arthroplasty for prosthetic hip and knee infections.

REASONING FOR SINGLE STAGE

Periprosthetic joint infection is one of the most common reasons for early, less than 2 years, and all-cause revisions in total knee arthroplasty and total hip arthroplasty [7-9]. With a rising total number of both primary joint procedures and periprosthetic joint infections, understanding surgical management options are important, especially in a value-based healthcare system [10]. Single stage revision (SSR) for PJI (SSR PJI) has been reported to have equivalent infection-free success compared to two stage revision, with lower mortality and morbidity, with less hospitalizations, shorter antibiotic treatment duration, and lower overall healthcare costs [11-15] (Table 1). The ability to perform one less operation appeals to patients on many levels and naturally will lend itself to lower the risks of an additional operation.

Two stage revision is not without its complications. Gomez et al [16] reviewed 504 patients undergoing two stage revision for PJI at 56 mo. Only 82% of patients underwent final implantation. In patients with gram negative infection, the re-implantation rate was even lower at 64%. 11% of patients required a spacer exchange from reinfection, and 10.3% of patients died at 2 years. The option for the surgeon to perform single stage revision may help reduce the risk of some devastating outcomes from a second operation.

KEY PRINCIPLES OF SINGLE STAGE

There are 3 key principles of SSR PJI [17]. The first is the organism must be known with sensitivities and bacterial minimally inhibitory concentrations. Single stage revision should not be performed without this information as antibiotic treatment cannot be tailored to combat the infection. Aspiration of the joint is performed with the patient off antibiotics for at least 14 d, using an “as sterile as possible technique” and with a culture incubation period of 14 d. Studies have suggested anesthetics such as controlled trials may help further our understanding of single stage revision compared with two stage.

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Table 1 Single stage revision outcomes for total knee arthroplasty and total hip arthroplasty

<table>
<thead>
<tr>
<th>Ref.</th>
<th>Year</th>
<th>PJI</th>
<th>Cases</th>
<th>Reinfection</th>
<th>Follow-up (in year)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Freeman et al[1]</td>
<td>1985</td>
<td>Knee</td>
<td>8</td>
<td>0</td>
<td>2</td>
</tr>
<tr>
<td>Scott et al[3]</td>
<td>1993</td>
<td>Knee</td>
<td>10</td>
<td>3 (30%)</td>
<td></td>
</tr>
<tr>
<td>Buechel et al[5]</td>
<td>2004</td>
<td>Knee</td>
<td>22</td>
<td>2 (9%)</td>
<td>10</td>
</tr>
<tr>
<td>Whiteside et al[6]</td>
<td>2011</td>
<td>Knee</td>
<td>18</td>
<td>1 (6%)</td>
<td>5.2</td>
</tr>
<tr>
<td>Parkinson et al[7]</td>
<td>2011</td>
<td>Knee</td>
<td>22</td>
<td>0</td>
<td>2</td>
</tr>
<tr>
<td>Singer et al[8]</td>
<td>2012</td>
<td>Knee</td>
<td>63</td>
<td>3 (5%)</td>
<td>3</td>
</tr>
<tr>
<td>Klatte et al[9]</td>
<td>2014</td>
<td>Knee</td>
<td>4</td>
<td>1 (25%)</td>
<td>2.1</td>
</tr>
<tr>
<td>Tibrewal et al[10]</td>
<td>2014</td>
<td>Knee</td>
<td>50</td>
<td>1 (2%)</td>
<td>10.5</td>
</tr>
<tr>
<td>Labruyère et al[12]</td>
<td>2015</td>
<td>Knee</td>
<td>9</td>
<td>0</td>
<td>5</td>
</tr>
<tr>
<td>Holland et al[13]</td>
<td>2019</td>
<td>Knee</td>
<td>26</td>
<td>1 (4%)</td>
<td>5.3</td>
</tr>
<tr>
<td>Siddiqi et al[14]</td>
<td>2019</td>
<td>TKA</td>
<td>57</td>
<td>8 (14%)</td>
<td>4.4</td>
</tr>
<tr>
<td>Abdelaziz et al[15]</td>
<td>2020</td>
<td>TKA</td>
<td>72</td>
<td>8 (11%)</td>
<td>4.2</td>
</tr>
<tr>
<td>Ji et al[16]</td>
<td>2020</td>
<td>Both</td>
<td>51</td>
<td>5 (10%)</td>
<td>4.4</td>
</tr>
<tr>
<td>Carlsson et al[17]</td>
<td>1978</td>
<td>Hip</td>
<td>77</td>
<td>17 (22%)</td>
<td></td>
</tr>
<tr>
<td>Hughes et al[18]</td>
<td>1979</td>
<td>Hip</td>
<td>13</td>
<td>4 (31%)</td>
<td></td>
</tr>
<tr>
<td>Buchholz et al[19]</td>
<td>1981</td>
<td>Hip</td>
<td>640</td>
<td>130 (20%)</td>
<td>4.3</td>
</tr>
<tr>
<td>Miley et al[20]</td>
<td>1982</td>
<td>Hip</td>
<td>47</td>
<td>6 (13%)</td>
<td>4</td>
</tr>
<tr>
<td>Weber et al[21]</td>
<td>1986</td>
<td>Hip</td>
<td>8</td>
<td>2 (25%)</td>
<td>6</td>
</tr>
<tr>
<td>Wroblewski et al[22]</td>
<td>1986</td>
<td>Hip</td>
<td>102</td>
<td>9 (9%)</td>
<td>3.2</td>
</tr>
<tr>
<td>Sanzén et al[23]</td>
<td>1988</td>
<td>Hip</td>
<td>102</td>
<td>25 (25%)</td>
<td>7</td>
</tr>
<tr>
<td>Hope et al[24]</td>
<td>1989</td>
<td>Hip</td>
<td>72</td>
<td>9 (13%)</td>
<td>3.75</td>
</tr>
<tr>
<td>Raut et al[25]</td>
<td>1994</td>
<td>Hip</td>
<td>57</td>
<td>8 (14%)</td>
<td>7.3</td>
</tr>
<tr>
<td>Mulkady et al[26]</td>
<td>1996</td>
<td>Hip</td>
<td>15</td>
<td>0</td>
<td>4</td>
</tr>
<tr>
<td>Ure et al[27]</td>
<td>1998</td>
<td>Hip</td>
<td>22</td>
<td>0</td>
<td>10.5</td>
</tr>
<tr>
<td>Callaghan et al[28]</td>
<td>1999</td>
<td>Hip</td>
<td>24</td>
<td>2 (8%)</td>
<td>10</td>
</tr>
<tr>
<td>Rudelli et al[29]</td>
<td>2008</td>
<td>Hip</td>
<td>32</td>
<td>2 (6%)</td>
<td>5</td>
</tr>
<tr>
<td>Winkler et al[30]</td>
<td>2008</td>
<td>Hip</td>
<td>37</td>
<td>3 (8%)</td>
<td>4.4</td>
</tr>
<tr>
<td>Yoo et al[31]</td>
<td>2008</td>
<td>Hip</td>
<td>12</td>
<td>1 (8%)</td>
<td>7.2</td>
</tr>
<tr>
<td>Oussedik et al[32]</td>
<td>2010</td>
<td>Hip</td>
<td>11</td>
<td>0</td>
<td>6.8</td>
</tr>
<tr>
<td>De Man et al[33]</td>
<td>2011</td>
<td>Hip</td>
<td>22</td>
<td>0</td>
<td>3.8</td>
</tr>
<tr>
<td>Klooche et al[34]</td>
<td>2012</td>
<td>Hip</td>
<td>38</td>
<td>0</td>
<td>2</td>
</tr>
<tr>
<td>Choi et al[35]</td>
<td>2013</td>
<td>Hip</td>
<td>17</td>
<td>3 (18%)</td>
<td>5.1</td>
</tr>
<tr>
<td>Hansen et al[36]</td>
<td>2013</td>
<td>Hip</td>
<td>27</td>
<td>8 (30%)</td>
<td>2.25</td>
</tr>
<tr>
<td>Bori et al[37]</td>
<td>2014</td>
<td>Hip</td>
<td>24</td>
<td>1 (4%)</td>
<td>3.6</td>
</tr>
<tr>
<td>Jenny et al[38]</td>
<td>2014</td>
<td>Hip</td>
<td>65</td>
<td>17 (26%)</td>
<td>5</td>
</tr>
<tr>
<td>Klatte et al[39]</td>
<td>2014</td>
<td>Hip</td>
<td>6</td>
<td>0</td>
<td>2.1</td>
</tr>
<tr>
<td>Wolf et al[40]</td>
<td>2014</td>
<td>Hip</td>
<td>37</td>
<td>24 (65%)</td>
<td>2</td>
</tr>
<tr>
<td>Zeller et al[41]</td>
<td>2014</td>
<td>Hip</td>
<td>157</td>
<td>8 (5%)</td>
<td>3.4</td>
</tr>
</tbody>
</table>
Table 2

| Study author listed, year of publication, joint investigated, number of cases, reinfection rate, and follow up reported. PJI: Prosthetic joint infections. |
|-------------------------|-----------------|____________|-----------------|-----------------|-----------------|-----------------|-----------------|
| Ebied et al[25]         | 2016            | Hip         | 33              | 1 (3%)          | 6               |
| Ichman et al[26]        | 2016            | Hip         | 39              | 0               | 6.6             |
| Born et al[27]          | 2016            | Hip         | 28              | 0               | 7               |
| Lange et al[28]         | 2017            | Hip         | 56              | 5 (91%)         | 4               |
| Whiteside et al[29]     | 2017            | Hip         | 21              | 1 (95%)         | 5.2             |
| Zahar et al[30]         | 2018            | Hip         | 85              | 5 (6%)          | 10.5            |
| Bori et al[31]          | 2018            | Hip         | 19              | 1 (5%)          |                 |
| Wolff et al[32]         | 2019            | Hip         | 26              | 1 (96%)         | 15              |
| Ji et al[33]            | 2019            | Hip         | 111             | 12 (11%)        | 4.8             |

lindocaine and ethyl chloride spray can affect bacterial growth[34,35]. Because of this, our institution does not use local anesthetic while aspirating the joint.

The second principle is the debridement. This may be the most important principle of SSR. Mortality after PJI has been worse than the 5 most common cancers, with mortalities reaching up to 5% per year postoperatively[20,21]. Treating the debridement similarly to cancer necessitates a radial resection of all infected tissue in order to obtain clean margins and healthy tissue. Aggressive debridement includes anterior and posterior capsule, and even the abductors when addressing PJI in the hip. In knees, the entire synovial membrane is removed en bloc so as to not contaminate the joint and to minimize risk of contaminating the entire surgical field. The incision excises the previous scar if able, and the debridement removes all peri-incisional tissue towards the joint. This is culminated in removal of the entire block of infected tissue. This style of debridement is in contrast to two-stage revision, where intra-lesional debridement occurs and may leave infected tissue throughout the surgical wound. In certain instances, if there is a sinus tract that can be incorporated in the planned incision then it will be excised as well. Unresectable sinus tracts, and involvement of neurovascular bundles are contraindications for single stage revision. Lastly, while some studies report use of methylene blue injected into the joint prior to resection to ensure adequate resection works for two stage, it has not been studied in single stage as the resection margins need to be greater and the goal of the debridement is not intra-lesional[22].

The third principle is both local and systemic antibiotic delivery tailored to the known pathologic organism. Local antibiotic delivery can be in the form of cement, powder or intra-articular infusion[23,24]. The delivery of antibiotics locally is known to achieve much higher tissue concentrations at the site of infection. Naturally, the use of bactericidal antibiotics such as aminoglycosides, cephalosporins, fluoroquinolones, metronidazole, penicillin and vancomycin will provide better treatment than bacteriostatic antibiotics (Table 2)[25]. Some authors advocate for Clindamycin as an acceptable bacteriostatic antibiotic[26]. Up to 10% of the dry crystalline weight of antibiotics can be added to the cement without significant mechanical loss[27]. Addition of slightly more (approximately 1/8th bottle per batch) liquid monomer may allow better viscosity characteristics for mixing. While local antibiotics generate high intra-articular concentrations with lower systemic risks, there are case reports of systemic complications such as renal or hepatic failure, and allergic reaction[28]. Surgeons must be mindful of dosage in patients with these risk factors. However, local antibiotic use is a key principle. Failure to add local antibiotics has resulted in lower infection free survival rates[29].

Combination of these three principles in single stage revision for periprosthetic joint infection has resulted in midterm infection free survival rates in the 80%-95% range; similar to two-stage revision arthroplasty[30].

### INDICATIONS

Single stage revision for PJI is a viable treatment option that has gradually increased in acceptance and popularity at certain specialized centers around the world. There has been an increasing body of published literature on this topic with favorable results[31].
Table 2 Bactericidal antibiotics vs bacteriostatic antibiotics

<table>
<thead>
<tr>
<th>Bactericidal antibiotics</th>
<th>Bacteriostatic</th>
</tr>
</thead>
<tbody>
<tr>
<td>Aminoglycosides, cephalosporins, fluoroquinolones, metronidazole, penicillin, vancomycin</td>
<td>Chloramphenicol, clindamycin, erythromycin, sulfamethoxazole, tetracycline, trimethoprim</td>
</tr>
</tbody>
</table>

It is important to use local bactericidal antibiotics to combat periprosthetic joint infection in single stage revision. The exception is clindamycin where it has been reported to have success in single stage revision for infection.

The first goal is to confirm the presence of PJI in the patient. The International Consensus Meeting (ICM) in Philadelphia 2018 gathered world leaders in orthopaedic surgery in their subspecialized fields and created consensus guidelines based upon available current published literature. The musculoskeletal infection society, part of the ICM, moderately agreed on criteria for diagnosis of PJI based upon patient laboratory tests, cultures and synovial aspiration. Patients are considered positive for PJI if they fulfill one of two major criteria: Two positive cultures of the same organism or a sinus tract with evidence of communication to the joint or prosthesis. Patients are also considered positive if they have 6 or more points, inconclusive if 3-5 points, and negative if less than 3 points on the minor criteria system. Elevated C-reactive protein or D-Dimer (2 points), elevated ESR (1 point), elevated synovial white blood cell or ++ leukocyte esterase test stripe or positive alpha defensin (3 points), elevated synovial polymorphonuclear percentage (2 points), single positive culture (2 points), positive histology such as frozen section (3 points), intraoperative purulence (3 points). A definition and threshold of acute and chronic infection is defined in Table 3. Difference from the 2013 criteria is the addition of leukocyte esterase, alpha defensin, and D-Dimer which may improve sensitivity compared to the previous criteria.

Although some authors report chronic PJI to be a strict timeframe of more than 4 wk, the key to diagnosis of chronic infection is the establishment of biofilm and the inability to eradicate the bacterial burden without removal of foreign material including the metallic implants, polymethylmethacrylate, and polyethylene bearings. Biofilm is created when groups of the infectious micro-organism(s) aggregate together on a surface embedded in a glycosylated extra-cellular matrix. This matrix prevents penetration of antibiotics, and allows enhanced cell to cell communication for microbe protection and nutrition. Although true biofilm establishment by microbes vary based on organism and conditions, early in vitro studies have suggested formation may occur as early as 8 h. However, 4 wk is usually the acceptable limit for the ability to debride, lower bacterial burden, and remove sufficient biofilm material, while retaining implants.

Theoretically patients with established biofilm and PJI can be candidates for single stage revision (Figure 1). Although some authors consider virulent organisms a relative contraindication, we further discuss that it may not be a strict contraindication. Patients in the acutely infected period may be better off treated with debridement, antibiotics, irrigation and prosthesis retention. This method assumes biofilm has not been established and bacterial burden can be lowered enough for the host’s own immune system to clear infection. Surgical options for chronic infection includes single stage revision, two stage revision, chronic suppression, or amputation/disarticulation.

A host’s comorbidities play an important role in outcomes in primary and revision total joint arthroplasty. The increase in count and severity of comorbidities can result in higher rates of medical and surgical complications, including infection, reoperation and death. McPherson et al. classified periprosthetic joint infections in hips into infection type, host class, and local extremity grade. Host class correlated with the ability to clear infection, undergo successful reimplantation, and avoid amputation or permanent resection arthroplasty. Host class A has no comorbidities, host class B has 1-2 comorbidities and host class C has 3 or more comorbidities or were essentially immunocompromised with absolute neutrophil count < 1000, CD4 count below 100, intravenous drug abuse, chronic active infection in another site, or dysplasia or neoplasm of immune system. Comorbidities are listed in Table 4. Specific to single stage revision for PJI, one study suggested a host class A had better success of infection clearance compared to host class B or C. Although this study did not follow one of the principles of single stage, which is local antibiotic delivery, they were still able to achieve 100% eradication of infection in that subset of patients. European centers more selective in their single stage revision criteria may exclude immunocompromised individuals. Although there is some variance in patient selection criteria, reports
Table 3 International Consensus Meeting 2018 Guidelines for prosthetic joint infection for hip and knee arthroplasty

<table>
<thead>
<tr>
<th>Major criteria</th>
<th>Minor criteria</th>
</tr>
</thead>
<tbody>
<tr>
<td>Two positive cultures of the same organism / Sinus tract with communication to the joint or prosthesis</td>
<td>Criteria</td>
</tr>
<tr>
<td>CRP (mg/L) or D Dimer (ug/L)</td>
<td>Acute threshold</td>
</tr>
<tr>
<td>100; unknown</td>
<td>10; 860</td>
</tr>
<tr>
<td>ESR (mm/h)</td>
<td>No role</td>
</tr>
<tr>
<td>Synovial WBC or leukocyte esterase or alpha defensin</td>
<td>10000; ++; 1.0</td>
</tr>
<tr>
<td>Synovial PMN %</td>
<td>90</td>
</tr>
<tr>
<td>Single positive culture</td>
<td></td>
</tr>
<tr>
<td>Positive histology</td>
<td></td>
</tr>
<tr>
<td>Positive intraoperative purulence</td>
<td></td>
</tr>
</tbody>
</table>

A positive prosthetic joint infection is one of two major criteria, or 6 or more points for minor criteria. 3-5 points for minor criteria is inconclusive and less than 3 points is negative for prosthetic joint infection. WBC: White blood cell; PJI: Prosthetic joint infection; CRP: C-reactive protein; ESR: Erythrocyte sedimentation rate; PMN: Polymorphonuclear leukocyte.

Table 4 McPherson host class comorbidities

<table>
<thead>
<tr>
<th>Systemic host compromising factors</th>
</tr>
</thead>
<tbody>
<tr>
<td>Age ≥ 80 yr</td>
</tr>
<tr>
<td>Chronic indwelling catheter</td>
</tr>
<tr>
<td>Diabetes</td>
</tr>
<tr>
<td>Malignancy (history or active)</td>
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<tr>
<td>Systemic inflammatory disease</td>
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McPherson hip infection classification comprises of infection type, host type and local factors. Host factors are graded A, B or C. A has none of the listed factors, B has 2 or fewer factors, C has 3 or more factors or severe immunocompromise defined by: absolute neutrophil count < 1000/mm$^3$, CD4 T cell count < 100/mm$^3$, intravenous drug abuse, or dysplasia or neoplasm of the immune system. HIV: Human immunodeficiency virus; AIDS: Acquired immune deficiency syndrome.

that followed the 3 listed principles had higher infection free success that those that did not$^{30,31}$. Further determination of patient selection will be discussed later in this article.

Knowledge of the infecting organism and appropriate antibiotic sensitivities is of critical importance. It is worth repeating, a culture negative infection is a contraindication to single stage revision for PJI. Highly virulent organisms such as methicillin resistant staphylococcus aureus or methicillin resistant staphylococcus epidermidis (MRSA)/(MRSE) or enterococcus, multi-drug resistance, and rare flora may make infection eradication more difficult. Additionally, inability to add local antibiotics to combat the infection due to high drug resistance or inactivation in cement may be a contraindication to single stage revision$^{31}$. Other factors that may make single stage revision difficult would be repeat surgeries, repeat infections and enterococcus infection. ENDO Klinik retrospectively reviewed their subsequent failures of single stage revision in hip and knee arthroplasty. For knees, failures correlated to previous failed revision for infection, enterococcus or streptococcus infection. For hips, wound drainage > 1 wk, previous operation and enterococcus infection were risk factors for reoperation or re-infection$^{16,19}$. Although these were not contraindications for single stage, surgeons considering performing single stage revision arthroplasty should acknowledge the difficulty eradicating infection in these situations.
ICM 2018 indications

In July 2018, the International Consensus Meeting on Periprosthetic Joint Infection was held in Philadelphia, PA with over 850 international delegates specializing in the field of orthopaedic surgery. There they developed consensus guidelines on management of various episodes of periprosthetic joint infection based upon current literature. Single stage revision for infection was discussed and 4 consensus agreements were published[31].

The potential advantages of a 1 stage exchange arthroplasty are multiple, including lower surgical morbidity and mortality, earlier functional return, lower healthcare and global economic costs, and increase in health related quality adjusted life years. This statement had an 89% consensus agreement with a moderate level of evidence (LOE).

The indications and contraindications listed below had 93% consensus agreement with a moderate LOE (Table 5): (1) Indications: Non-immunocompromised host, absence of systemic sepsis, minimal bone/soft tissue loss allowing primary wound closure, known pathologic organism with sensitivities preoperatively; and (2) Contraindications: Severe soft tissue defect or unresectable sinus tract, culture negative PJI, unable to perform a radical debridement, unable to deliver local antibiotic treatment, lack of bone stock for fixation of new implant.

There is a role for single stage exchange arthroplasty in the setting of acute cementless total hip arthroplasty (THA) PJI. This statement had an 89% consensus agreement with a moderate LOE.

Putting aside the effect on successful treatment of PJs, it is logical that a single surgical procedure puts patients at lower risk of both mortality and morbidity.
Table 5 International consensus meeting indications and contra-indications for single stage revision for prosthetic joint infection

<table>
<thead>
<tr>
<th>Indications</th>
<th>Contra-indications</th>
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<tbody>
<tr>
<td>Non-immunocompromised host</td>
<td>Severe damage to soft tissues were direct closure not possible or inexcisable complex sinus tract with old scar</td>
</tr>
<tr>
<td>Absence of systemic sepsis</td>
<td>Culture negative PJI</td>
</tr>
<tr>
<td>Minimal bone/soft tissue loss allowing primary closure</td>
<td>Inability to perform radical debridement</td>
</tr>
<tr>
<td>Isolation of pathologic organism preoperatively</td>
<td>Unable to give local antibiotic treatment</td>
</tr>
<tr>
<td>Known sensitivities to bactericidal treatment</td>
<td>No proper bone stock available for fixation</td>
</tr>
</tbody>
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PJI: Prosthetic joint infection.

compared to a 2-stage exchange arthroplasty that involves 2 separate operations. 83% of the ICM agreed using moderate LOE.

These 4 agreements suggested single stage revision for PJI may result in similar outcomes with one less operation in the appropriately selected patient with guidance to whom may be good candidates.

CONTRAINDICATIONS

Although certain centers have attested to performing SSR in a majority of their infection cases, they have a few specific criteria that are absolute contraindications for single stage revision. It is worth mentioning, a culture negative infection where bacterial sensitivities are unknown does not allow for properly targeted antibiotic treatment. Inability to obtain adequate primary soft tissue coverage, infection involving the neurovascular bundles, and other unresectable infected areas are also contraindications for single stage revision. Other authors successfully performing single stage revision for PJI have their own indications and contraindications. Haddad et al. reported on 102 patients undergoing treatment for total knee arthroplasty (TKA) PJI. 28 underwent single stage revision while 74 underwent two stage revision. At 6.5 years, no patients in the single stage developed reinfection compared to 5 (9%) of the two stage. In addition to the ICM contraindication criteria, they excluded patients with peripheral vascular disease, polymicrobial infection, and multi-resistant organisms such as MRSA/MRSE. Their retrospective results should not be interpreted as single stage revision is better than two stage, as there is inherent patient selection bias, but that in properly selected individuals it can work as successfully.

OUTCOMES OF SINGLE STAGE PROSTHETIC JOINT INFECTIONS

Infection free success for properly indicated single stage revision varies from 77%-100% by a variety of studies. Failures of single stage for aseptic loosening on long term follow-up can range from 5%-18%, although this is not dramatically different than aseptic loosening in two stage revisions. Evaluating reoperations from single stage revisions, Citak et al. reported on their reasons for re-revision in single stage TKAs in 91 patients out of 697 patients at the ENDO Klinik. There were 47/697 (6.7%) reinfections and 37/697 (5.3%) revisions for aseptic loosening, three patients underwent revision for patellar mal-tracking, three underwent revision for periprosthetic fracture and one for knee dislocation. Abdelaziz et al. also reported on their reasons for re-revision in single stage THA PJI. Out of 121 reoperations, 53 (40%) were for instability, 40 (33%) reinfection, 16 (13%) aseptic loosening, seven (6%) periprosthetic fracture, three (2%) implant failure, and one each for wound healing disorder and heterotopic ossification. These reasons for revision arthroplasty were similar to those who underwent two stage revisions.

Patient reported outcomes after single stage revision have been suggested to be comparable to two stage. Haddad et al. reported their single stage TKA revisions had knee society scores of 88 compared to 76 in their two stage (P < 0.02), and better visual analogy scores satisfaction scores of 7.82 compared to 6.18 in the two stage cohort (P < 0.01). Comparative studies investigating outcomes after single stage
THA revision had equivalent Harris hip scores (HHS) in three studies and better in one\(^{39-42}\). Oussedik\(^{39}\) et al. reported on 50 patients undergoing revision for THA PJI with 11 patients undergoing single stage compared with 39 with two stage revision. HHS in the single stage was 87.8 compared with 75.5 in the two stage (\(P < 0.001\)) as well as change in HHS favoring single stage (\(P = 0.027\)). Authors recognized the retrospective selection bias of single stage, but suggested for appropriately indicated patients, single stage may provide improved functional outcomes. In additional, Markov model decision tree analyses and quality of life adjusted values after single stage revision has also been calculated to have favorability over two stage revision\(^{15,43}\). Although healthcare cost savings, outcomes and improved morbidity from one less operation may by appealing for single stage revision, the larger effort of work and lower compensatory reward may be de-incentivizing for the surgeon. Fehring\(^{44}\) et al. found single stage revision required more time and effort and less compensation compared to primary arthroplasty. They encouraged payors to improve reimbursement to incentivize and align the goals of the healthcare system.

Lastly, failure of a previous single revision is not a contraindication for a repeat single stage revision. Knowledge of the three principles, organism type, antimicrobial sensitivities, and thorough debridement is crucial. However, previous risk factors for single stage revision failure such as poor host class factors, enterococcus infection, virulent organism infection, polymicrobial infection or multiple previous operations may guide the treating surgeon towards two stage revision gold standard.

**TECHNICAL PEARLS**

After obtaining a diagnosis of PJI, with cultures and antibiotic sensitivities, a meticulous debridement occurs. Preoperative intravenous antibiotics are given, tailored to the organism. The operative theatre where the single stage revision is performed is not followed by another primary joint procedure, and a deep extended cleaning of the room is performed after. We typically will use a sterile tourniquet for a total knee unless the planned incision and reconstruction does not give enough operative space. The tourniquet is deflated after removal of implants and re-inflated at the start of the second re-implantation procedure. Hemostasis should be achieved at the deflation. The previous skin incision is utilized with incorporation of a sinus tract if present. Excision of the scar, sinus tract, surgical tract, and all fibrous infected tissue down to the capsule is performed in a single piece leaving normal tissue to remain. Exposure of the capsule and all unhealthy tissue is removed en bloc. Removal of the capsule going down to bone is performed in an attempt to minimize surgical field contamination. Meticulous hemostasis is achieved. Removal of implants is performed routinely with marrow reaming to remove any infectious material. Pulsatile lavage with at least 6 liters of normal saline solution is then performed, an optional povidone-iodine solution diluted to 0.3% with irrigation and then the wound is packed with chlorohexidine soaked sponges. Povidone iodine solution is not routinely used at our center as the infection rates are lower with simple saline irrigation. The wound is loosely closed at the skin, all drapes, gowns, gloves, table with all previous instruments are removed. The patient is then re-prepped, re-draped, all new instruments, new table, new gowns and gloves, and new scrubs for the surgical staff is performed for the second stage of the procedure. The second stage includes another irrigation, a change of gloves and suction tips, and proceeding to re-implantation. At this point, the procedure is similar to the second stage of a two-stage revision. Local antibiotics are mixed in cement if cementation is occurring, or topical antibiotics are added down the canals, in the acetabulum and in the joint prior to deep closure. Cementless implants in total hip arthroplasty have been described with success, with all authors adding topical and intravenous antibiotics to their treatment regimen. Antibiotics in calcium sulfate beads are an optional way to deliver the intra-articular concentrations of antibiotics. Dosages of more than 20 cc of beads may cause increased drainage, or other complications such as heterotopic ossification, or hyper-calcemia\(^{45}\). Postoperatively, specific targeted intravenous antibiotics are given for 6 wk with assistance in management by an infectious disease specialist with sub-specialization in periprosthetic joint infections.
FUTURE DEVELOPMENTS

Single stage revision arthroplasty for infection has been in development since the 1970s, initially pioneered by Buchholz et al., it has now been furthered by surgeons as technology and surgical techniques continue to improve. One area of progress is in the use of cementless THA implants in single stage revision. The ICM 2018 suggested potential viability of this technique in response to several articles reporting successful infection free survivorship. Bori et al. reported on single stage THA revision in 24 patients with cementless femoral components, but antibiotic cemented acetabular components in nine patients. Only one patient (95.8%) developed an infection at 44 mo follow-up. Ji et al. reported on 111 patients with various infecting organisms using both cementless femur and acetabular components with 89.2% infection free survivorship at 58 mo. They applied intra-articular antibiotic infusions in MRSE/MRSA, fungal or culture negative in addition to powdered local antibiotics. An important note is that these patients still received local antibiotic delivery by other methods such as intra-articular infusion or topical powder application, a key principle for this technique.

To date, there is no algorithm for predicting what specific patient factors, organism factors, or surgical factors lead to infection free success in single stage revision for PJI. Although host factors such as McPherson Host Class C, virulent organisms such as fungal, MRSA/MRSE or culture negative, or surgical factors such as inadequate radical debridement may lead to lower infection free survival, it is not known how many factors will lead to failure or success of single stage revision. These are potential future research goals for improvement.

Naturally, single stage revision for PJI has an inherent retrospective selection bias due to less virulent organisms and healthier hosts when comparing to two stage revision. Undergoing a single operation will typically be less costly and have lower morbidity and mortality for the patient. Ultimately, a randomized control trial may be necessary to fully realize who may benefit from this procedure and by how much.

CONCLUSION

Single stage revision for periprosthetic joint infection can be a successful operation with careful selection of the patient, infecting organism and precise surgical technique. The 3 key principles of bacterial sensitivities, thorough radial debridement and delivery of local and systemic antibiotics can result in similar infection free survivorship to two stage exchange. Future developments into this technique include its practice in culture negative infections or use of cementless implants. Randomized controlled trials may help further our understanding of single stage revision compared with two stage.

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